

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（8/21）

| System Organ Class / Adverse Event |                 | Min - 29.9kg/m <sup>2</sup> |         |                     |         |         |                     |          |         |                     |          |         |                       |          |         |          |
|------------------------------------|-----------------|-----------------------------|---------|---------------------|---------|---------|---------------------|----------|---------|---------------------|----------|---------|-----------------------|----------|---------|----------|
|                                    |                 | Placebo (n=82)              |         | ATL-962 15mg (n=46) |         |         | ATL-962 30mg (n=53) |          |         | ATL-962 60mg (n=58) |          |         | ATL-962 120mg (n=402) |          |         |          |
|                                    |                 | 関連あり                        | 関連なし    | 合計                  | 関連あり    | 関連なし    | 合計                  | 関連あり     | 関連なし    | 合計                  | 関連あり     | 関連なし    | 合計                    | 関連あり     | 関連なし    | 合計       |
| 腎および尿路障害（続き）                       | 水腎症             |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 2 (0.5)  | 2 (0.5) |          |
|                                    | 血尿              |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 2 (0.5)  | 2 (0.5) |          |
|                                    | 着色尿             | 1 (1.2)                     |         | 1 (1.2)             |         |         |                     |          |         |                     |          |         |                       |          |         |          |
| 生殖系および乳房障害                         | (SOC sub-total) |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 2 (0.5)  | 2 (0.5) |          |
|                                    | 良性前立腺肥大症        |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 1 (0.2)  | 1 (0.2) |          |
|                                    | 閉経期症状           |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 1 (0.2)  | 1 (0.2) |          |
| 一般・全身障害および投与部位の状態                  | (SOC sub-total) | 1 (1.2)                     | 3 (3.7) | 4 (4.9)             | 1 (2.2) | 1 (2.2) | 1 (1.9)             | 2 (3.8)  | 3 (5.7) | 3 (5.2)             | 3 (5.2)  | 2 (0.5) | 22 (5.5)              | 24 (6.0) |         |          |
|                                    | 末梢性浮腫           | 1 (1.2)                     |         | 1 (1.2)             |         |         |                     | 1 (1.9)  |         | 1 (1.9)             | 1 (1.7)  | 1 (0.2) | 5 (1.2)               | 6 (1.5)  |         |          |
|                                    | 倦怠感             |                             |         |                     |         |         |                     | 1 (1.9)  | 1 (1.9) | 2 (3.4)             |          | 2 (3.4) | 3 (0.7)               | 3 (0.7)  |         |          |
|                                    | 発熱              |                             |         |                     |         |         |                     |          |         |                     |          |         | 5 (1.2)               | 5 (1.2)  |         |          |
|                                    | 胸痛              |                             |         |                     | 1 (2.2) | 1 (2.2) |                     | 1 (1.9)  | 1 (1.9) |                     |          |         | 2 (0.5)               | 2 (0.5)  |         |          |
|                                    | 医療機器破損          |                             |         |                     |         |         |                     |          |         |                     |          |         | 1 (0.2)               | 1 (0.2)  |         |          |
|                                    | 異常感             |                             | 1 (1.2) | 1 (1.2)             |         |         |                     |          |         |                     |          |         | 1 (0.2)               | 1 (0.2)  |         |          |
|                                    | 悪寒              |                             |         |                     |         |         |                     |          |         |                     |          |         | 1 (0.2)               | 1 (0.2)  |         |          |
|                                    | 疼痛              |                             |         |                     |         |         |                     |          |         |                     |          |         | 1 (0.2)               | 1 (0.2)  |         |          |
|                                    | 胸部不快感           |                             |         |                     |         |         |                     |          |         |                     |          |         | 1 (0.2)               | 1 (0.2)  |         |          |
|                                    | 冷感              |                             | 1 (1.2) | 1 (1.2)             |         |         |                     |          |         |                     |          |         |                       |          |         |          |
|                                    | 全身性浮腫           |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 1 (0.2)  | 1 (0.2) |          |
|                                    | 空腹              |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 1 (0.2)  | 1 (0.2) |          |
|                                    | 易刺激性            |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 1 (0.2)  | 1 (0.2) |          |
|                                    | 臨床検査            | 口渇                          |         | 1 (1.2)             | 1 (1.2) |         |                     |          |         |                     |          |         |                       |          |         |          |
|                                    |                 | (SOC sub-total)             | 2 (2.4) | 2 (2.4)             | 3 (3.7) | 2 (4.3) | 3 (6.5)             | 5 (10.9) | 3 (5.7) | 4 (7.5)             | 7 (13.2) | 2 (3.4) | 3 (5.2)               | 5 (8.6)  | 7 (1.7) | 38 (9.5) |
| 血中クレアチンホスホキナーゼ増加                   |                 |                             |         |                     |         | 1 (2.2) | 1 (2.2)             |          |         |                     |          | 2 (3.4) | 2 (3.4)               | 1 (0.2)  | 9 (2.2) | 10 (2.5) |
| アラニンアミノトランスフェラーゼ増加                 |                 | 2 (2.4)                     | 1 (1.2) | 2 (2.4)             |         | 1 (2.2) | 1 (2.2)             |          | 2 (3.8) | 2 (3.8)             | 2 (3.4)  |         | 2 (3.4)               | 3 (0.7)  | 3 (0.7) | 6 (1.5)  |
| アスパラギン酸アミノトランスフェラーゼ増加              |                 | 1 (1.2)                     | 1 (1.2) | 2 (2.4)             |         | 1 (2.2) | 1 (2.2)             |          |         |                     |          | 1 (1.7) | 1 (1.7)               | 3 (0.7)  | 3 (0.7) | 6 (1.5)  |
| γ-グルタミルトランスフェラーゼ増加                 |                 | 1 (1.2)                     |         | 1 (1.2)             | 1 (2.2) |         | 1 (2.2)             |          | 1 (1.9) | 1 (1.9)             |          |         |                       | 3 (0.7)  | 3 (0.7) | 6 (1.5)  |
| 血中尿酸増加                             |                 |                             |         | 1 (1.2)             | 1 (1.2) |         |                     |          |         |                     |          |         |                       |          | 7 (1.7) | 7 (1.7)  |
| サイトケラチン18増加                        |                 | 1 (1.2)                     |         | 1 (1.2)             |         | 1 (2.2) | 1 (2.2)             |          |         |                     |          |         |                       | 1 (0.2)  |         | 1 (0.2)  |
| 肝機能検査異常                            |                 |                             |         |                     |         |         |                     |          |         |                     |          |         |                       |          | 4 (1.0) | 4 (1.0)  |
| 尿中血陽性                              |                 |                             |         |                     |         |         |                     |          |         |                     |          |         |                       |          | 5 (1.2) | 5 (1.2)  |
| m-A S T 増加                         |                 | 1 (1.2)                     | 1 (1.2) | 2 (2.4)             |         | 1 (2.2) | 1 (2.2)             |          |         |                     |          |         |                       | 1 (0.2)  |         | 1 (0.2)  |
| 血中乳酸脱水素酵素増加                        |                 |                             |         |                     |         | 1 (2.2) | 1 (2.2)             |          |         |                     |          | 1 (1.7) | 1 (1.7)               | 1 (0.2)  |         | 1 (0.2)  |
| 白血球数増加                             |                 |                             |         |                     |         |         |                     | 1 (1.9)  |         | 1 (1.9)             |          |         |                       |          | 4 (1.0) | 4 (1.0)  |
| 好酸球数増加                             |                 |                             |         |                     |         |         |                     |          | 1 (1.9) | 1 (1.9)             |          |         |                       |          |         |          |
| 血中アルカリホスファターゼ増加                    |                 |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 1 (0.2)  |         | 1 (0.2)  |
| 血中ビリルビン増加                          |                 |                             |         |                     |         |         |                     | 1 (1.9)  |         | 1 (1.9)             |          |         |                       | 1 (0.2)  |         | 1 (0.2)  |
| 血圧上昇                               |                 |                             |         |                     |         |         |                     |          | 1 (1.9) | 1 (1.9)             |          |         |                       |          | 1 (0.2) | 1 (0.2)  |
| 尿中ブドウ糖陽性                           |                 |                             |         |                     |         |         |                     |          |         |                     |          |         |                       |          | 2 (0.5) | 2 (0.5)  |
| 血中カリウム増加                           |                 |                             |         |                     |         |         |                     |          |         |                     |          |         |                       |          | 1 (0.2) | 1 (0.2)  |
| 血中トリグリセリド増加                        |                 |                             |         |                     |         | 1 (2.2) | 1 (2.2)             |          |         |                     |          |         |                       |          |         |          |
| カロチン減少                             |                 |                             |         |                     |         |         |                     |          |         |                     |          |         |                       | 1 (0.2)  |         | 1 (0.2)  |
| 心電図T波逆転                            |                 |                             |         |                     |         |         |                     |          |         |                     |          |         |                       |          | 1 (0.2) | 1 (0.2)  |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（9/21）

| System Organ Class / Adverse Event | Min - 29.9kg/m <sup>2</sup> |         |                        |          |                        |         |                        |         |                          |           |           |         |
|------------------------------------|-----------------------------|---------|------------------------|----------|------------------------|---------|------------------------|---------|--------------------------|-----------|-----------|---------|
|                                    | Placebo<br>(n=82)           |         | ATL-962 15mg<br>(n=46) |          | ATL-962 30mg<br>(n=53) |         | ATL-962 60mg<br>(n=58) |         | ATL-962 120mg<br>(n=402) |           |           |         |
|                                    | 関連あり                        | 関連なし    | 合計                     | 関連あり     | 関連なし                   | 合計      | 関連あり                   | 関連なし    | 合計                       | 関連あり      | 関連なし      | 合計      |
| 臨床検査（続き）                           |                             |         |                        |          |                        |         |                        |         |                          |           |           |         |
| リンパ球数減少                            |                             |         |                        |          |                        |         |                        |         |                          |           | 1 (0.2)   | 1 (0.2) |
| 好中球数減少                             |                             |         |                        |          |                        | 1 (1.9) | 1 (1.9)                |         |                          |           |           |         |
| 好中球数増加                             |                             |         |                        |          |                        |         |                        |         |                          |           | 1 (0.2)   | 1 (0.2) |
| 血小板数減少                             |                             |         |                        |          |                        |         |                        |         |                          |           | 1 (0.2)   | 1 (0.2) |
| プロトロンビン時間延長                        |                             |         |                        |          |                        |         |                        | 1 (1.7) | 1 (1.7)                  |           |           |         |
| 赤血球数減少                             |                             |         |                        |          |                        |         |                        |         |                          |           | 1 (0.2)   | 1 (0.2) |
| 白血球数減少                             |                             |         |                        | 1 (2.2)  | 1 (2.2)                |         |                        |         |                          |           |           |         |
| 便潜血陽性                              |                             |         |                        |          |                        |         |                        |         |                          |           | 1 (0.2)   | 1 (0.2) |
| 傷害、中毒および処置合併症 (SOC sub-total)      | 7 (8.5)                     | 7 (8.5) | 6 (13.0)               | 6 (13.0) | 4 (7.5)                | 4 (7.5) | 5 (8.6)                | 5 (8.6) | 1 (0.2)                  | 41 (10.2) | 41 (10.2) |         |
| 転倒                                 | 1 (1.2)                     | 1 (1.2) | 4 (8.7)                | 4 (8.7)  |                        |         | 3 (5.2)                | 3 (5.2) | 1 (0.2)                  | 17 (4.2)  | 17 (4.2)  |         |
| 挫傷                                 | 3 (3.7)                     | 3 (3.7) |                        |          |                        |         |                        |         |                          | 7 (1.7)   | 7 (1.7)   |         |
| 靭帯捻挫                               | 1 (1.2)                     | 1 (1.2) |                        |          |                        |         |                        |         |                          | 3 (0.7)   | 3 (0.7)   |         |
| 事故                                 |                             |         | 1 (2.2)                | 1 (2.2)  | 2 (3.8)                | 2 (3.8) | 2 (3.4)                | 2 (3.4) |                          | 2 (0.5)   | 2 (0.5)   |         |
| 交通事故                               |                             |         |                        |          |                        |         |                        |         |                          | 3 (0.7)   | 3 (0.7)   |         |
| 節足動物刺傷                             | 1 (1.2)                     | 1 (1.2) |                        |          | 1 (1.9)                | 1 (1.9) |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 足骨折                                |                             |         |                        |          |                        |         |                        |         |                          | 2 (0.5)   | 2 (0.5)   |         |
| 擦過傷                                |                             |         |                        |          |                        |         |                        |         |                          | 4 (1.0)   | 4 (1.0)   |         |
| 熱傷                                 |                             |         |                        |          |                        |         |                        |         |                          | 2 (0.5)   | 2 (0.5)   |         |
| 家庭内事故                              |                             |         |                        |          | 1 (1.9)                | 1 (1.9) |                        |         |                          |           |           |         |
| 靭帯損傷                               |                             |         |                        |          |                        |         |                        |         |                          | 2 (0.5)   | 2 (0.5)   |         |
| 歯牙破折                               |                             |         |                        |          |                        |         |                        |         |                          | 3 (0.7)   | 3 (0.7)   |         |
| 熱中症                                | 1 (1.2)                     | 1 (1.2) |                        |          |                        |         |                        |         |                          |           |           |         |
| 上腕骨骨折                              |                             |         |                        |          |                        |         |                        |         |                          | 2 (0.5)   | 2 (0.5)   |         |
| 関節脱臼                               |                             |         |                        |          |                        |         |                        |         |                          | 2 (0.5)   | 2 (0.5)   |         |
| 脊椎圧迫骨折                             |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 創傷                                 |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 節足動物咬傷                             |                             |         |                        |          | 1 (2.2)                | 1 (2.2) |                        |         |                          |           |           |         |
| 肉離れ                                | 1 (1.2)                     | 1 (1.2) |                        |          |                        |         |                        |         |                          |           |           |         |
| 第2度熱傷                              |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 凍瘡                                 |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 損傷                                 |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 刺創                                 |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 腱損傷                                |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 歯牙損傷                               |                             |         |                        |          |                        |         |                        | 1 (1.7) | 1 (1.7)                  |           |           |         |
| 手首関節骨折                             |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 半月板障害                              |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 口唇損傷                               |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 外科および内科処置 (SOC sub-total)          |                             |         |                        |          | 1 (1.9)                | 1 (1.9) |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |
| 抜歯                                 |                             |         |                        |          | 1 (1.9)                | 1 (1.9) |                        |         |                          |           |           |         |
| 歯科補てつ物装着                           |                             |         |                        |          |                        |         |                        |         |                          | 1 (0.2)   | 1 (0.2)   |         |

表中の数字：発現例数（発現率 (%)）

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（10/21）

| System Organ Class / Adverse Event | 30.0 - 34.9kg/m <sup>2</sup> |          |          |                        |         |          |                        |          |          |                        |          |          |                          |           |           |           |
|------------------------------------|------------------------------|----------|----------|------------------------|---------|----------|------------------------|----------|----------|------------------------|----------|----------|--------------------------|-----------|-----------|-----------|
|                                    | Placebo<br>(n=57)            |          |          | ATL-962 15mg<br>(n=25) |         |          | ATL-962 30mg<br>(n=29) |          |          | ATL-962 60mg<br>(n=24) |          |          | ATL-962 120mg<br>(n=319) |           |           |           |
|                                    | 関連あり                         | 関連なし     | 合計       | 関連あり                   | 関連なし    | 合計       | 関連あり                   | 関連なし     | 合計       | 関連あり                   | 関連なし     | 合計       | 関連あり                     | 関連なし      | 合計        |           |
| Over All                           | (Number of Patients)         | 16(28.1) | 45(78.9) | 50(87.7)               | 4(16.0) | 16(64.0) | 17(68.0)               | 10(34.5) | 22(75.9) | 25(86.2)               | 12(50.0) | 17(70.8) | 21(87.5)                 | 190(59.6) | 237(74.3) | 280(87.8) |
|                                    | (Number of AEs)              | 21       | 121      | 142                    | 7       | 30       | 37                     | 13       | 49       | 62                     | 24       | 48       | 72                       | 266       | 742       | 1008      |
| 感染症および寄生虫症                         | (SOC sub-total)              | 23(40.4) | 23(40.4) |                        | 6(24.0) | 6(24.0)  |                        | 13(44.8) | 13(44.8) |                        | 8(33.3)  | 8(33.3)  |                          | 143(44.8) | 143(44.8) |           |
|                                    | 鼻咽頭炎                         | 14(24.6) | 14(24.6) |                        | 6(24.0) | 6(24.0)  |                        | 7(24.1)  | 7(24.1)  |                        | 6(25.0)  | 6(25.0)  |                          | 104(32.6) | 104(32.6) |           |
|                                    | 咽頭炎                          | 2(3.5)   | 2(3.5)   |                        |         |          |                        | 1(3.4)   | 1(3.4)   |                        |          |          |                          | 8(2.5)    | 8(2.5)    |           |
|                                    | 膀胱炎                          | 2(3.5)   | 2(3.5)   |                        |         |          |                        | 2(6.9)   | 2(6.9)   |                        |          |          |                          | 7(2.2)    | 7(2.2)    |           |
|                                    | 気管支炎                         | 1(1.8)   | 1(1.8)   |                        |         |          |                        | 2(6.9)   | 2(6.9)   |                        | 1(4.2)   | 1(4.2)   |                          | 8(2.5)    | 8(2.5)    |           |
|                                    | インフルエンザ                      |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 8(2.5)    | 8(2.5)    |           |
|                                    | 胃腸炎                          | 2(3.5)   | 2(3.5)   |                        |         |          |                        |          |          |                        |          |          |                          | 4(1.3)    | 4(1.3)    |           |
|                                    | 足部白癬                         |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 2(0.6)    | 2(0.6)    |           |
|                                    | 麦粒腫                          | 2(3.5)   | 2(3.5)   |                        |         |          |                        |          |          |                        |          |          |                          | 3(0.9)    | 3(0.9)    |           |
|                                    | 口腔ヘルペス                       |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 3(0.9)    | 3(0.9)    |           |
|                                    | 帯状疱疹                         | 1(1.8)   | 1(1.8)   |                        |         |          |                        |          |          |                        |          |          |                          | 2(0.6)    | 2(0.6)    |           |
|                                    | 蜂巣炎                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 爪囲炎                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 3(0.9)    | 3(0.9)    |           |
|                                    | 鼻炎                           | 2(3.5)   | 2(3.5)   |                        |         |          |                        |          |          |                        |          |          |                          |           |           |           |
|                                    | 中耳炎                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 2(0.6)    | 2(0.6)    |           |
|                                    | 扁桃炎                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 3(0.9)    | 3(0.9)    |           |
|                                    | 毛包炎                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 2(0.6)    | 2(0.6)    |           |
|                                    | ウイルス性胃腸炎                     |          |          |                        |         |          |                        | 1(3.4)   | 1(3.4)   |                        |          |          |                          |           |           |           |
|                                    | 爪真菌症                         |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 2(0.6)    | 2(0.6)    |           |
|                                    | 外耳炎                          |          |          |                        |         |          |                        |          |          |                        | 1(4.2)   | 1(4.2)   |                          |           |           |           |
|                                    | 白癬感染                         | 1(1.8)   | 1(1.8)   |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 感染性皮膚膿腫                      |          |          |                        |         |          |                        |          |          |                        | 1(4.2)   | 1(4.2)   |                          |           |           |           |
|                                    | 肺炎                           |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 2(0.6)    | 2(0.6)    |           |
|                                    | 菌髄炎                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 3(0.9)    | 3(0.9)    |           |
|                                    | 副鼻腔炎                         |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 2(0.6)    | 2(0.6)    |           |
|                                    | 尿路感染                         | 1(1.8)   | 1(1.8)   |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | せつ                           |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 2(0.6)    | 2(0.6)    |           |
|                                    | ヘリコクター感染                     |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 急性副鼻腔炎                       |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 虫垂炎                          | 1(1.8)   | 1(1.8)   |                        |         |          |                        |          |          |                        |          |          |                          |           |           |           |
|                                    | 肛門膿瘍                         |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 憩室炎                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 感染性湿疹                        |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 真菌感染                         |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | リンパ管炎                        | 1(1.8)   | 1(1.8)   |                        |         |          |                        |          |          |                        |          |          |                          |           |           |           |
|                                    | 乳腺炎                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 尿道炎                          |          |          |                        |         |          |                        | 1(3.4)   | 1(3.4)   |                        |          |          |                          |           |           |           |
|                                    | 膣感染                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 四肢膿瘍                         |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | ウイルス性気管支炎                    |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 頸膿瘍                          |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 菌肉感染                         |          |          |                        |         |          |                        | 1(3.4)   | 1(3.4)   |                        |          |          |                          |           |           |           |
|                                    | レンサ球菌感染                      |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | ウイルス性腸炎                      |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |
|                                    | 耳帯状疱疹                        |          |          |                        |         |          |                        |          |          |                        |          |          |                          | 1(0.3)    | 1(0.3)    |           |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（11/21）

| System Organ Class / Adverse Event     | 30.0 - 34.9kg/m <sup>2</sup> |      |                        |      |                        |      |                        |    |                          |      |          |  |
|--|------------------------------|------|------------------------|------|------------------------|------|------------------------|----|--------------------------|------|----------|--|
|  | Placebo<br>(n=57)            |      | ATL-962 15mg<br>(n=25) |      | ATL-962 30mg<br>(n=29) |      | ATL-962 60mg<br>(n=24) |    | ATL-962 120mg<br>(n=319) |      |          |  |
|  | 関連あり                         | 関連なし | 関連あり                   | 関連なし | 合計                     | 関連あり | 関連なし                   | 合計 | 関連あり                     | 関連なし |          |  |
| 良性、悪性および詳細不明の<br>新生物(嚢胞およびポリープ<br>を含む) | (SOC sub-total)              |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    | 6 (1.9)                  |      | 6 (1.9)  |  |
|  | 脂肪腫                          |      |                        |      |                        |      |                        |    | 2 (0.6)                  |      | 2 (0.6)  |  |
|  | 膀胱新生物                        |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    |                          |      |          |  |
|  | 子宮癌                          |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
|  | 転移性結腸癌                       |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
|  | 肺新生物                         |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
|  | 甲状腺癌                         |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
| 血液およびリンパ系障害                            | (SOC sub-total)              |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    |                          |      |          |  |
|  | 脾腫                           |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    |                          |      |          |  |
| 免疫系障害                                  | (SOC sub-total)              |      |                        |      |                        |      |                        |    | 8 (2.5)                  |      | 8 (2.5)  |  |
|  | 季節性アレルギー                     |      |                        |      |                        |      |                        |    | 8 (2.5)                  |      | 8 (2.5)  |  |
| 代謝および栄養障害                              | (SOC sub-total)              |      | 1 (1.8)                |      | 6 (10.5)               |      | 7 (12.3)               |    | 1 (3.4)                  |      | 1 (3.4)  |  |
|  | 糖尿病                          |      | 1 (1.8)                |      | 5 (8.8)                |      | 6 (10.5)               |    | 1 (4.2)                  |      | 1 (4.2)  |  |
|  | 低血糖症                         |      |                        |      |                        |      |                        |    | 5 (1.6)                  |      | 24 (7.5) |  |
|  | 高尿酸血症                        |      |                        |      |                        |      |                        |    | 2 (0.6)                  |      | 7 (2.2)  |  |
|  | 2型糖尿病                        |      |                        |      |                        |      |                        |    | 2 (0.6)                  |      | 6 (1.9)  |  |
|  | コントロール不良の糖尿病                 |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 2 (0.6)  |  |
|  | 食欲減退                         |      |                        |      |                        |      |                        |    | 4 (1.3)                  |      | 4 (1.3)  |  |
|  | 脱水                           |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
|  | 痛風                           |      |                        |      |                        |      |                        |    | 2 (0.6)                  |      | 2 (0.6)  |  |
|  | 高血糖                          |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
|  | 高トリグリセリド血症                   |      |                        |      |                        |      | 1 (3.4)                |    | 1 (3.4)                  |      |          |  |
|  | 脂質異常症                        |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
|  | 高脂血症                         |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
| 精神障害                                   | (SOC sub-total)              |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    | 1 (4.2)                  |      | 1 (4.2)  |  |
|  | 不眠症                          |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 3 (0.9)  |  |
|  | 不快感                          |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    | 1 (4.2)                  |      | 1 (4.2)  |  |
|  | 幻覚                           |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 3 (0.9)  |  |
| 神経系障害                                  | (SOC sub-total)              |      | 1 (1.8)                |      | 2 (3.5)                |      | 3 (5.3)                |    | 3 (12.0)                 |      | 3 (12.0) |  |
|  | 頭痛                           |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    | 1 (3.4)                  |      | 2 (6.9)  |  |
|  | 浮動性めまい                       |      |                        |      |                        |      | 1 (4.0)                |    | 1 (4.0)                  |      | 3 (10.3) |  |
|  | 感覚鈍麻                         |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    | 2 (8.3)                  |      | 1 (4.2)  |  |
|  | 頸腕症候群                        |      |                        |      |                        |      |                        |    | 3 (12.5)                 |      | 1 (0.3)  |  |
|  | 体位性めまい                       |      |                        |      |                        |      |                        |    | 29 (9.1)                 |      | 30 (9.4) |  |
|  | 片頭痛                          |      |                        |      |                        |      | 1 (4.0)                |    | 1 (4.0)                  |      | 10 (3.1) |  |
|  | 手根管症候群                       |      |                        |      |                        |      | 1 (3.4)                |    | 1 (3.4)                  |      | 10 (3.1) |  |
|  | 糖尿病性ニューロパチー                  |      |                        |      |                        |      |                        |    | 2 (8.3)                  |      | 1 (4.2)  |  |
|  | 緊張性頭痛                        |      |                        |      |                        |      | 1 (4.0)                |    | 1 (4.0)                  |      | 3 (12.5) |  |
|  | 振戦                           |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 4 (1.3)  |  |
|  | 第7脳神経麻痺                      |      |                        |      |                        |      |                        |    | 4 (1.3)                  |      | 5 (1.6)  |  |
|  | ラクナ梗塞                        |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
|  | 脳幹梗塞                         |      |                        |      |                        |      |                        |    | 2 (0.6)                  |      | 2 (0.6)  |  |
|  | 味覚異常                         |      | 1 (1.8)                |      | 1 (1.8)                |      |                        |    | 2 (0.6)                  |      | 2 (0.6)  |  |
|  | 末梢性ニューロパチー                   |      |                        |      |                        |      | 1 (4.0)                |    | 1 (4.0)                  |      | 1 (0.3)  |  |
|  | 三叉神経痛                        |      |                        |      |                        |      |                        |    | 2 (0.6)                  |      | 2 (0.6)  |  |
|  | 自律神経ニューロパチー                  |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
|  | 顔面痙攣                         |      |                        |      |                        |      |                        |    | 1 (0.3)                  |      | 1 (0.3)  |  |
|  |                              |      |                        |      |                        |      | 1 (3.4)                |    | 1 (3.4)                  |      |          |  |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（12/21）

| System Organ Class / Adverse Event | 30.0 - 34.9kg/m <sup>2</sup> |         |                        |         |                        |         |                        |         |                          |          |          |         |
|------------------------------------|------------------------------|---------|------------------------|---------|------------------------|---------|------------------------|---------|--------------------------|----------|----------|---------|
|                                    | Placebo<br>(n=57)            |         | ATL-962 15mg<br>(n=25) |         | ATL-962 30mg<br>(n=29) |         | ATL-962 60mg<br>(n=24) |         | ATL-962 120mg<br>(n=319) |          |          |         |
|                                    | 関連あり<br>関連なし                 | 合計      | 関連あり<br>関連なし           | 合計      | 関連あり<br>関連なし           | 合計      | 関連あり<br>関連なし           | 合計      | 関連あり<br>関連なし             | 合計       |          |         |
| 眼障害                                |                              |         |                        |         |                        |         |                        |         |                          |          |          |         |
| (SOC sub-total)                    | 1 (1.8)                      | 2 (3.5) | 3 (5.3)                | 2 (8.0) | 2 (8.0)                | 1 (3.4) | 1 (3.4)                | 5(20.8) | 5(20.8)                  | 24 (7.5) | 24 (7.5) |         |
| 白内障                                |                              |         |                        | 1 (4.0) | 1 (4.0)                |         |                        | 1 (4.2) | 1 (4.2)                  | 5 (1.6)  | 5 (1.6)  |         |
| 糖尿病網膜症                             |                              |         |                        |         |                        |         |                        | 1 (4.2) | 1 (4.2)                  | 6 (1.9)  | 6 (1.9)  |         |
| 結膜炎                                |                              | 2 (3.5) | 2 (3.5)                |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| アレルギー性結膜炎                          |                              |         |                        | 1 (4.0) | 1 (4.0)                |         |                        |         |                          | 2 (0.6)  | 2 (0.6)  |         |
| 網膜出血                               |                              |         |                        | 1 (4.0) | 1 (4.0)                |         |                        | 2 (8.3) | 2 (8.3)                  | 1 (0.3)  | 1 (0.3)  |         |
| 眼精疲労                               |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 結膜出血                               | 1 (1.8)                      |         | 1 (1.8)                |         |                        |         |                        |         |                          | 2 (0.6)  | 2 (0.6)  |         |
| 眼乾燥                                |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 糖尿病性網膜浮腫                           |                              |         |                        |         |                        |         |                        |         |                          | 2 (0.6)  | 2 (0.6)  |         |
| 視力低下                               |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 眼瞼炎                                |                              | 1 (1.8) | 1 (1.8)                |         |                        |         |                        |         |                          |          |          |         |
| 眼脂                                 |                              |         |                        |         |                        |         | 1 (3.4)                | 1 (3.4) |                          |          |          |         |
| 角膜炎                                |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 眼充血                                |                              |         |                        |         |                        |         |                        | 1 (4.2) | 1 (4.2)                  |          |          |         |
| 点状角膜炎                              |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 網膜動脈閉塞                             |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 網膜変性                               |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 耳および迷路障害                           |                              |         |                        |         |                        |         |                        | 3(12.5) | 3(12.5)                  | 8 (2.5)  | 8 (2.5)  |         |
| (SOC sub-total)                    |                              |         |                        |         |                        |         |                        | 2 (8.3) | 2 (8.3)                  | 2 (0.6)  | 2 (0.6)  |         |
| 回転性めまい                             |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 耳鳴                                 |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 感音性難聴                              |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 鼓膜穿孔                               |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 迷路性回転性めまい                          |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 耳不快感                               |                              |         |                        |         |                        |         |                        | 1 (4.2) | 1 (4.2)                  |          |          |         |
| 突発難聴                               |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 耳管狭窄                               |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 心臓障害                               |                              |         |                        |         |                        |         |                        |         |                          | 2 (0.6)  | 7 (2.2)  |         |
| (SOC sub-total)                    | 2 (3.5)                      | 2 (3.5) | 1 (4.0)                | 1 (4.0) | 1 (3.4)                | 1 (3.4) |                        |         |                          |          | 1 (0.3)  | 1 (0.3) |
| 狭心症                                |                              |         |                        |         |                        |         |                        |         |                          |          |          |         |
| 心房細動                               | 1 (1.8)                      | 1 (1.8) |                        |         |                        |         |                        |         |                          |          | 1 (0.3)  | 1 (0.3) |
| 上室性期外収縮                            |                              |         |                        | 1 (4.0) | 1 (4.0)                |         |                        |         |                          | 2 (0.6)  | 2 (0.6)  |         |
| 心室性期外収縮                            |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 頻脈                                 | 1 (1.8)                      | 1 (1.8) |                        |         |                        |         |                        |         |                          |          |          |         |
| 急性心筋梗塞                             |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 不安定狭心症                             |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 右脚ブロック                             |                              |         |                        |         |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3)  |         |
| 期外収縮                               |                              |         |                        |         |                        | 1 (3.4) | 1 (3.4)                |         |                          |          |          |         |
| 血管障害                               |                              |         |                        |         |                        |         |                        |         |                          | 4 (1.3)  | 4 (1.3)  |         |
| (SOC sub-total)                    | 1 (1.8)                      | 1 (1.8) |                        |         |                        |         |                        |         |                          |          | 3 (0.9)  | 3 (0.9) |
| 高血圧                                | 1 (1.8)                      | 1 (1.8) |                        |         |                        |         |                        |         |                          |          | 1 (0.3)  | 1 (0.3) |
| 深部静脈血栓症                            |                              |         |                        |         |                        |         |                        |         |                          |          |          |         |
| 呼吸器、胸郭および縦隔障害                      | 7(12.3)                      | 7(12.3) | 2 (8.0)                | 2 (8.0) | 6(20.7)                | 6(20.7) | 4(16.7)                | 4(16.7) | 32(10.0)                 | 32(10.0) |          |         |
| (SOC sub-total)                    | 4 (7.0)                      | 4 (7.0) | 1 (4.0)                | 1 (4.0) | 4(13.8)                | 4(13.8) | 2 (8.3)                | 2 (8.3) | 19 (6.0)                 | 19 (6.0) | 2 (0.6)  | 2 (0.6) |
| 上気道の炎症                             |                              |         |                        |         |                        | 1 (3.4) | 1 (3.4)                | 1 (4.2) | 1 (4.2)                  | 4 (1.3)  | 4 (1.3)  |         |
| 咳嗽                                 |                              |         |                        |         |                        |         |                        |         |                          | 2 (0.6)  | 2 (0.6)  |         |
| アレルギー性鼻炎                           |                              |         |                        |         |                        |         |                        |         |                          | 4 (1.3)  | 4 (1.3)  |         |
| 口腔咽喉痛                              | 1 (1.8)                      | 1 (1.8) |                        |         |                        |         |                        |         |                          |          | 2 (0.6)  | 2 (0.6) |

表中の数字：発現例数（発現率 (%)）

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（13/21）

| System Organ Class / Adverse Event | 30.0 - 34.9kg/m <sup>2</sup> |          |          |                        |         |                        |         |                        |          |                          |         |          |           |          |           |
|------------------------------------|------------------------------|----------|----------|------------------------|---------|------------------------|---------|------------------------|----------|--------------------------|---------|----------|-----------|----------|-----------|
|                                    | Placebo<br>(n=57)            |          |          | ATL-962 15mg<br>(n=25) |         | ATL-962 30mg<br>(n=29) |         | ATL-962 60mg<br>(n=24) |          | ATL-962 120mg<br>(n=319) |         |          |           |          |           |
|                                    | 関連あり                         | 関連なし     | 合計       | 関連あり                   | 関連なし    | 合計                     | 関連あり    | 関連なし                   | 合計       | 関連あり                     | 関連なし    | 合計       |           |          |           |
| 呼吸器、胸郭および縦隔障害<br>(続き)              |                              |          |          |                        |         |                        |         |                        |          |                          |         |          |           |          |           |
| 喘息                                 |                              |          |          | 1 (4.0)                | 1 (4.0) |                        |         |                        |          |                          | 2 (0.6) | 2 (0.6)  |           |          |           |
| 鼻出血                                | 1 (1.8)                      |          | 1 (1.8)  |                        |         |                        |         |                        |          |                          | 2 (0.6) | 2 (0.6)  |           |          |           |
| 鼻漏                                 | 1 (1.8)                      |          | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          |           |          |           |
| 睡眠時無呼吸症候群                          |                              |          |          |                        |         |                        |         |                        |          |                          | 1 (0.3) | 1 (0.3)  |           |          |           |
| 湿性咳嗽                               |                              |          |          |                        |         |                        |         |                        |          |                          | 2 (0.6) | 2 (0.6)  |           |          |           |
| 間質性肺疾患                             |                              |          |          |                        |         | 1 (3.4)                | 1 (3.4) |                        |          |                          |         |          |           |          |           |
| 肺塞栓症                               |                              |          |          |                        |         |                        |         |                        |          |                          | 1 (0.3) | 1 (0.3)  |           |          |           |
| 痰貯留                                |                              |          |          |                        |         |                        |         |                        |          |                          | 1 (0.3) | 1 (0.3)  |           |          |           |
| 副鼻腔囊胞                              |                              |          |          |                        |         |                        |         | 1 (4.2)                | 1 (4.2)  |                          |         |          |           |          |           |
| 胃腸障害                               |                              |          |          |                        |         |                        |         |                        |          |                          |         |          |           |          |           |
| (SOC sub-total)                    | 11(19.3)                     | 14(24.6) | 23(40.4) | 4(16.0)                | 3(12.0) | 7(28.0)                | 7(24.1) | 5(17.2)                | 11(37.9) | 11(45.8)                 | 5(20.8) | 14(58.3) | 183(57.4) | 63(19.7) | 201(63.0) |
| 脂肪便                                | 1 (1.8)                      |          | 1 (1.8)  | 1 (4.0)                |         | 1 (4.0)                | 3(10.3) |                        | 3(10.3)  | 2 (8.3)                  |         | 2 (8.3)  | 120(37.6) |          | 120(37.6) |
| 下痢                                 | 5 (8.8)                      | 4 (7.0)  | 8(14.0)  | 3(12.0)                | 1 (4.0) | 4(16.0)                | 4(13.8) |                        | 4(13.8)  | 10(41.7)                 | 2 (8.3) | 11(45.8) | 73(22.9)  | 10 (3.1) | 83(26.0)  |
| 便秘                                 | 2 (3.5)                      | 2 (3.5)  | 4 (7.0)  |                        |         |                        |         | 1 (3.4)                | 1 (3.4)  | 1 (4.2)                  | 2 (8.3) | 3(12.5)  | 7 (2.2)   | 6 (1.9)  | 13 (4.1)  |
| 真菌                                 |                              | 2 (3.5)  | 2 (3.5)  |                        |         |                        |         | 1 (3.4)                | 1 (3.4)  |                          |         |          | 7 (2.2)   |          | 7 (2.2)   |
| 腹部膨満                               |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 6 (1.9)   | 1 (0.3)  | 7 (2.2)   |
| 腹痛                                 |                              | 1 (1.8)  | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 3 (0.9)  | 4 (1.3)   |
| 悪心                                 |                              |          |          |                        | 1 (4.0) | 1 (4.0)                | 1 (3.4) |                        | 1 (3.4)  | 1 (4.2)                  |         | 1 (4.2)  | 2 (0.6)   | 3 (0.9)  | 5 (1.6)   |
| 腹部不快感                              | 1 (1.8)                      |          | 1 (1.8)  |                        |         |                        |         |                        |          |                          | 1 (4.2) | 1 (4.2)  | 2 (0.6)   | 6 (1.9)  | 7 (2.2)   |
| 上腹部痛                               |                              | 1 (1.8)  | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 4 (1.3)  | 5 (1.6)   |
| 菌周病                                |                              | 1 (1.8)  | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 2 (0.6)   | 2 (0.6)  |           |
| 嘔吐                                 | 1 (1.8)                      |          | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 4 (1.3)  | 5 (1.6)   |
| 菌周炎                                |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 10 (3.1)  | 10 (3.1) |           |
| 痔核                                 |                              |          |          |                        |         |                        | 1 (3.4) | 1 (3.4)                |          | 1 (4.2)                  | 1 (4.2) |          | 3 (0.9)   | 3 (0.9)  |           |
| 口内炎                                |                              | 1 (1.8)  | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 4 (1.3)  | 5 (1.6)   |
| 菌肉炎                                |                              |          |          |                        |         |                        | 1 (3.4) | 1 (3.4)                |          |                          |         |          | 1 (0.3)   | 1 (0.3)  |           |
| 消化不良                               |                              |          |          |                        |         |                        | 1 (3.4) | 1 (3.4)                | 1 (4.2)  |                          | 1 (4.2) |          | 1 (0.3)   | 1 (0.3)  |           |
| 胃炎                                 |                              | 1 (1.8)  | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 4 (1.3)  | 5 (1.6)   |
| 鼓腸                                 |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 2 (0.6)   | 2 (0.6)  |           |
| 菌痛                                 |                              | 1 (1.8)  | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 1 (0.3)  |           |
| 結腸ポリープ                             |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 2 (0.6)   | 2 (0.6)  |           |
| 排便回数増加                             | 1 (1.8)                      |          | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 3 (0.9)   | 1 (0.3)  | 4 (1.3)   |
| びらん性胃炎                             |                              |          |          |                        | 1 (4.0) | 1 (4.0)                |         |                        |          |                          |         |          | 1 (0.3)   | 1 (0.3)  |           |
| 胃食道逆流性疾患                           |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 1 (0.3)  |           |
| 下腹部痛                               |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 3 (0.9)   | 2 (0.6)  | 5 (1.6)   |
| 腸炎                                 |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 1 (0.3)  |           |
| 口唇炎                                |                              | 1 (1.8)  | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 1 (0.3)  | 2 (0.6)   |
| 十二指腸潰瘍                             |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 1 (0.3)  |           |
| 胃ポリープ                              |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 2 (0.6)   | 2 (0.6)  |           |
| 裂孔ヘルニア                             |                              | 1 (1.8)  | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 1 (0.3)  |           |
| 変色便                                |                              |          |          |                        |         |                        |         | 1 (3.4)                | 1 (3.4)  |                          |         |          |           |          |           |
| 胃腸音異常                              |                              |          |          |                        |         |                        |         | 1 (3.4)                | 1 (3.4)  |                          |         |          |           |          |           |
| 便失禁                                |                              |          |          |                        |         |                        |         |                        |          |                          |         |          | 1 (0.3)   | 1 (0.3)  |           |
| 白色便                                | 1 (1.8)                      |          | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          |           |          |           |
| 萎縮性胃炎                              |                              | 1 (1.8)  | 1 (1.8)  |                        |         |                        |         |                        |          |                          |         |          |           |          |           |
| 血便排泄                               |                              |          |          |                        | 1 (4.0) | 1 (4.0)                |         |                        |          |                          |         |          | 1 (0.3)   | 1 (0.3)  |           |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（14/21）

| System Organ Class / Adverse Event | 30.0 - 34.9kg/m <sup>2</sup> |                 |                        |           |                        |         |                        |         |                          |          |         |          |           |           |
|------------------------------------|------------------------------|-----------------|------------------------|-----------|------------------------|---------|------------------------|---------|--------------------------|----------|---------|----------|-----------|-----------|
|                                    | Placebo<br>(n=57)            |                 | ATL-962 15mg<br>(n=25) |           | ATL-962 30mg<br>(n=29) |         | ATL-962 60mg<br>(n=24) |         | ATL-962 120mg<br>(n=319) |          |         |          |           |           |
|                                    | 関連あり                         | 関連なし            | 合計                     | 関連あり      | 関連なし                   | 合計      | 関連あり                   | 関連なし    | 合計                       | 関連あり     | 関連なし    | 合計       |           |           |
| 胃腸障害（続き）                           | 過敏性腸症候群                      |                 |                        |           |                        |         |                        |         |                          | 1 (0.3)  |         | 1 (0.3)  |           |           |
|                                    | 裂肛                           |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 腸憩室                          |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 出血性腸憩室                       |                 |                        |           |                        |         |                        |         |                          | 1 (0.3)  |         | 1 (0.3)  |           |           |
|                                    | 歯肉腫脹                         |                 |                        |           |                        |         |                        | 1 (4.2) | 1 (4.2)                  |          |         |          |           |           |
|                                    | 肛門周囲炎                        |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 排便障害                         |                 |                        |           |                        |         |                        |         |                          | 1 (0.3)  |         | 1 (0.3)  |           |           |
| 肝胆道系障害                             | (SOC sub-total)              | 1 (1.8)         | 3 (5.3)                | 4 (7.0)   |                        |         |                        |         |                          | 1 (0.3)  | 6 (1.9) | 7 (2.2)  |           |           |
|                                    | 肝機能異常                        | 1 (1.8)         | 1 (1.8)                | 2 (3.5)   |                        |         |                        |         |                          | 1 (0.3)  | 1 (0.3) | 2 (0.6)  |           |           |
|                                    | 脂肪肝                          |                 | 1 (1.8)                | 1 (1.8)   |                        |         |                        |         |                          |          | 2 (0.6) | 2 (0.6)  |           |           |
|                                    | 胆石症                          |                 | 1 (1.8)                | 1 (1.8)   |                        |         |                        |         |                          |          | 2 (0.6) | 2 (0.6)  |           |           |
|                                    | 肝障害                          |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | (SOC sub-total)              | 1 (1.8)         | 1 (1.8)                | 2 (3.5)   | 2 (8.0)                | 2 (8.0) | 2 (6.9)                | 2 (6.9) | 3 (12.5)                 | 3 (12.5) | 6 (1.9) | 28 (8.8) | 34 (10.7) |           |
| 皮膚および皮下組織障害                        | 湿疹                           |                 |                        |           | 1 (4.0)                | 1 (4.0) | 1 (3.4)                | 1 (3.4) | 1 (4.2)                  | 1 (4.2)  | 2 (0.6) | 10 (3.1) | 12 (3.8)  |           |
|                                    | 接触性皮膚炎                       |                 | 1 (1.8)                | 1 (1.8)   |                        |         |                        |         |                          |          | 1 (0.3) | 2 (0.6)  | 3 (0.9)   |           |
|                                    | そう痒症                         | 1 (1.8)         |                        | 1 (1.8)   | 1 (4.0)                | 1 (4.0) |                        |         |                          |          |         | 4 (1.3)  | 4 (1.3)   |           |
|                                    | 蕁麻疹                          |                 |                        |           |                        |         | 1 (3.4)                | 1 (3.4) | 1 (4.2)                  | 1 (4.2)  | 1 (0.3) | 3 (0.9)  | 4 (1.3)   |           |
|                                    | 発疹                           |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 乾皮症                          |                 |                        |           |                        |         |                        |         |                          |          | 2 (0.6) | 2 (0.6)  |           |           |
|                                    | 紅色汗疹                         |                 |                        |           |                        |         |                        |         | 1 (4.2)                  | 1 (4.2)  | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | ざ瘡                           |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 皮膚囊腫                         |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 嵌入爪                          |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 脂漏性皮膚炎                       |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 皮膚乾燥                         |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 光線過敏性反応                      |                 |                        |           |                        |         |                        |         |                          |          | 2 (0.6) | 2 (0.6)  |           |           |
|                                    | 全身性そう痒症                      |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 冷汗                           |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 中毒性皮疹                        |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  |           |           |
|                                    | 筋骨格系および結合組織障害                | (SOC sub-total) | 13 (22.8)              | 13 (22.8) |                        | 1 (4.0) | 1 (4.0)                | 2 (6.9) | 2 (6.9)                  | 1 (4.2)  | 1 (4.2) | 3 (0.9)  | 63 (19.7) | 65 (20.4) |
|                                    |                              | 背部痛             | 4 (7.0)                | 4 (7.0)   |                        |         |                        |         |                          |          |         |          | 12 (3.8)  | 12 (3.8)  |
|                                    |                              | 関節痛             | 4 (7.0)                | 4 (7.0)   |                        |         |                        |         |                          |          |         |          | 12 (3.8)  | 12 (3.8)  |
|                                    |                              | 変形性関節症          |                        |           |                        |         |                        |         |                          |          |         |          | 7 (2.2)   | 7 (2.2)   |
| 筋肉痛                                |                              | 1 (1.8)         | 1 (1.8)                |           |                        |         |                        |         |                          |          |         | 8 (2.5)  | 8 (2.5)   |           |
| 筋骨格硬直                              |                              | 2 (3.5)         | 2 (3.5)                |           | 1 (4.0)                | 1 (4.0) |                        |         |                          |          |         | 4 (1.3)  | 4 (1.3)   |           |
| 腱鞘炎                                |                              | 2 (3.5)         | 2 (3.5)                |           |                        |         |                        |         |                          |          |         | 2 (0.6)  | 2 (0.6)   |           |
| 四肢痛                                |                              | 1 (1.8)         | 1 (1.8)                |           |                        |         | 1 (3.4)                | 1 (3.4) |                          |          |         | 3 (0.9)  | 3 (0.9)   |           |
| 関節周囲炎                              |                              | 1 (1.8)         | 1 (1.8)                |           |                        |         |                        |         |                          |          |         | 5 (1.6)  | 5 (1.6)   |           |
| 変形性脊椎症                             |                              |                 |                        |           |                        |         |                        |         |                          |          |         | 1 (0.3)  | 1 (0.3)   |           |
| 椎間板突出                              |                              |                 |                        |           |                        |         | 1 (3.4)                | 1 (3.4) |                          |          |         | 5 (1.6)  | 5 (1.6)   |           |
| 関節炎                                |                              |                 |                        |           |                        |         |                        |         |                          |          |         | 1 (0.3)  | 1 (0.3)   |           |
| 筋痙攣                                |                              |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 2 (0.6)  | 3 (0.9)   |           |
| 筋骨格痛                               |                              |                 |                        |           |                        |         |                        |         |                          |          |         | 3 (0.9)  | 3 (0.9)   |           |
| 弾発指                                |                              |                 |                        |           |                        |         |                        |         |                          |          |         | 1 (0.3)  | 1 (0.3)   |           |
| 滑液嚢腫                               |                              |                 |                        |           |                        |         |                        |         |                          |          | 1 (0.3) | 1 (0.3)  | 2 (0.6)   |           |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（15/21）

| System Organ Class / Adverse Event | 30.0 - 34.9kg/m <sup>2</sup> |         |                     |          |                     |         |                     |         |                       |         |          |          |          |          |
|------------------------------------|------------------------------|---------|---------------------|----------|---------------------|---------|---------------------|---------|-----------------------|---------|----------|----------|----------|----------|
|                                    | Placebo (n=57)               |         | ATL-962 15mg (n=25) |          | ATL-962 30mg (n=29) |         | ATL-962 60mg (n=24) |         | ATL-962 120mg (n=319) |         |          |          |          |          |
|                                    | 関連あり                         | 関連なし    | 合計                  | 関連あり     | 関連なし                | 合計      | 関連あり                | 関連なし    | 合計                    | 関連あり    | 関連なし     | 合計       |          |          |
| 筋骨格系および結合組織障害（続き）                  | 側腹部痛                         |         |                     |          |                     |         |                     |         |                       | 1 (0.3) | 1 (0.3)  |          |          |          |
|                                    | 脊椎靭帯骨化症                      |         |                     |          |                     |         |                     |         |                       | 1 (0.3) | 1 (0.3)  |          |          |          |
|                                    | 筋膜炎                          |         |                     |          |                     |         |                     |         |                       | 1 (0.3) | 1 (0.3)  |          |          |          |
|                                    | 足底筋膜炎                        |         |                     |          |                     |         |                     |         |                       | 1 (0.3) | 1 (0.3)  |          |          |          |
|                                    | 脊椎すべり症                       |         |                     |          |                     |         |                     | 1 (4.2) | 1 (4.2)               |         |          |          |          |          |
|                                    | 腰部脊柱管狭窄症                     |         |                     |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | リウマチ性多発筋痛                    |         |                     |          |                     |         |                     |         |                       | 1 (0.3) | 1 (0.3)  |          |          |          |
|                                    | (SOC sub-total)              | 1 (1.8) | 1 (1.8)             |          |                     |         | 2 (6.9)             | 2 (6.9) | 2 (8.3)               | 2 (8.3) | 1 (0.3)  | 6 (1.9)  | 7 (2.2)  |          |
| 腎および尿路障害                           | 腎結石症                         |         |                     |          |                     |         |                     | 1 (4.2) | 1 (4.2)               | 1 (0.3) | 3 (0.9)  | 4 (1.3)  |          |          |
|                                    | 尿路結石                         |         |                     |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | 排尿困難                         | 1 (1.8) | 1 (1.8)             |          |                     |         |                     |         | 1 (4.2)               | 1 (4.2) |          |          |          |          |
|                                    | 水腎症                          |         |                     |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | 腎嚢胞                          |         |                     |          |                     |         | 1 (3.4)             | 1 (3.4) | 1 (4.2)               | 1 (4.2) |          |          |          |          |
|                                    | 神経因性膀胱                       |         |                     |          |                     |         | 1 (3.4)             | 1 (3.4) |                       |         |          |          |          |          |
|                                    | 非感染性膀胱炎                      |         |                     |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | (SOC sub-total)              | 1 (1.8) | 1 (1.8)             |          | 1 (4.0)             | 1 (4.0) |                     |         |                       |         | 1 (0.3)  | 3 (0.9)  | 4 (1.3)  |          |
|                                    | 良性前立腺肥大症                     |         |                     |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | 乳房痛                          |         |                     |          | 1 (4.0)             | 1 (4.0) |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
| 生殖系および乳房障害                         | 月経困難症                        |         |                     |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | 陰部そう痒症                       |         |                     |          |                     |         |                     |         |                       | 1 (0.3) | 1 (0.3)  |          |          |          |
|                                    | 乳房障害                         | 1 (1.8) | 1 (1.8)             |          |                     |         |                     |         |                       |         |          |          |          |          |
|                                    | (SOC sub-total)              | 1 (1.8) | 2 (3.5)             | 3 (5.3)  | 1 (4.0)             | 1 (4.0) | 2 (6.9)             | 2 (6.9) | 4 (13.8)              | 1 (4.2) | 1 (4.2)  | 14 (4.4) | 14 (4.4) |          |
|                                    | 末梢性浮腫                        | 1 (1.8) | 1 (1.8)             |          | 1 (4.0)             | 1 (4.0) |                     |         |                       | 1 (4.2) | 1 (4.2)  | 3 (0.9)  | 3 (0.9)  |          |
|                                    | 倦怠感                          |         |                     |          |                     |         | 1 (3.4)             |         | 1 (3.4)               |         |          | 1 (0.3)  | 1 (0.3)  |          |
|                                    | 発熱                           |         |                     |          |                     |         |                     | 1 (3.4) | 1 (3.4)               |         |          | 2 (0.6)  | 2 (0.6)  |          |
|                                    | 胸痛                           | 1 (1.8) | 1 (1.8)             |          |                     |         |                     |         |                       |         |          | 2 (0.6)  | 2 (0.6)  |          |
|                                    | 医療機器破損                       |         |                     |          |                     |         |                     |         |                       |         |          | 2 (0.6)  | 2 (0.6)  |          |
|                                    | 異常感                          |         |                     |          |                     |         | 1 (3.4)             |         | 1 (3.4)               |         |          |          |          |          |
| 一般・全身障害および投与部位の状態                  | 浮腫                           | 1 (1.8) | 1 (1.8)             |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | 痒痛                           |         |                     |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | 医療機器不具合                      |         |                     |          |                     |         |                     |         |                       |         | 2 (0.6)  | 2 (0.6)  |          |          |
|                                    | 無力症                          |         |                     |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | 石灰沈着症                        |         |                     |          |                     |         |                     |         |                       |         | 1 (0.3)  | 1 (0.3)  |          |          |
|                                    | 炎症                           |         |                     |          |                     |         |                     |         | 1 (3.4)               | 1 (3.4) |          |          |          |          |
|                                    | (SOC sub-total)              | 1 (1.8) | 6 (10.5)            | 7 (12.3) | 1 (4.0)             | 1 (4.0) | 1 (3.4)             | 1 (3.4) | 2 (8.3)               | 2 (8.3) | 3 (12.5) | 8 (2.5)  | 23 (7.2) | 29 (9.1) |
|                                    | 血中クレアチンホスホキナーゼ増加             |         | 4 (7.0)             | 4 (7.0)  |                     |         |                     |         |                       | 1 (4.2) | 1 (4.2)  | 1 (0.3)  | 9 (2.8)  | 10 (3.1) |
|                                    | アラニンアミノトランスフェラーゼ増加           | 1 (1.8) | 2 (3.5)             | 3 (5.3)  | 1 (4.0)             | 1 (4.0) |                     |         |                       | 1 (4.2) | 1 (4.2)  | 1 (0.3)  | 2 (0.6)  | 3 (0.9)  |
|                                    | アスパラギン酸アミノトランスフェラーゼ増加        | 1 (1.8) | 2 (3.5)             | 3 (5.3)  | 1 (4.0)             | 1 (4.0) |                     |         |                       | 1 (4.2) | 1 (4.2)  |          | 3 (0.9)  | 3 (0.9)  |
| γ-グルタミルトランスフェラーゼ増加                 |                              | 1 (1.8) | 1 (1.8)             |          |                     |         |                     |         |                       |         | 1 (0.3)  | 2 (0.6)  | 3 (0.9)  |          |
| 臨床検査                               | 血中尿酸増加                       |         | 1 (1.8)             | 1 (1.8)  |                     |         |                     |         |                       |         | 1 (0.3)  | 2 (0.6)  | 3 (0.9)  |          |
|                                    | サイトケラチン 18 増加                | 1 (1.8) | 1 (1.8)             | 2 (3.5)  |                     |         |                     |         |                       |         | 1 (0.3)  | 4 (1.3)  | 5 (1.6)  |          |
|                                    | 肝機能検査異常                      |         |                     |          |                     |         |                     |         |                       |         |          | 2 (0.6)  | 2 (0.6)  |          |

表中の数字：発現例数（発現率（%））



表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（16/21）

| System Organ Class / Adverse Event | 30.0 - 34.9kg/m <sup>2</sup> |         |                        |         |                        |          |                        |         |                          |         |           |           |
|------------------------------------|------------------------------|---------|------------------------|---------|------------------------|----------|------------------------|---------|--------------------------|---------|-----------|-----------|
|                                    | Placebo<br>(n=57)            |         | ATL-962 15mg<br>(n=25) |         | ATL-962 30mg<br>(n=29) |          | ATL-962 60mg<br>(n=24) |         | ATL-962 120mg<br>(n=319) |         |           |           |
|                                    | 関連あり                         | 関連なし    | 合計                     | 関連あり    | 関連なし                   | 合計       | 関連あり                   | 関連なし    | 合計                       | 関連あり    | 合計        |           |
| 臨床検査（続き）                           |                              |         |                        |         |                        |          |                        |         |                          |         |           |           |
| 尿中血陽性                              |                              |         |                        |         |                        |          |                        |         |                          | 2 (0.6) | 2 (0.6)   |           |
| m-A S T 増加                         |                              |         |                        |         |                        |          | 1 (4.2)                | 1 (4.2) |                          | 1 (0.3) | 1 (0.3)   |           |
| 血中乳酸脱水素酵素増加                        | 1 (1.8)                      |         | 1 (1.8)                |         |                        |          |                        |         |                          | 1 (0.3) | 1 (0.3)   |           |
| 好酸球数増加                             |                              |         |                        |         |                        |          |                        |         | 1 (0.3)                  |         | 1 (0.3)   |           |
| 血中アルカリホスファターゼ増加                    |                              |         |                        |         |                        |          | 1 (4.2)                | 1 (4.2) | 1 (0.3)                  |         | 1 (0.3)   |           |
| 血中クレアチニン増加                         |                              |         |                        |         |                        |          |                        | 1 (4.2) | 1 (4.2)                  | 1 (0.3) | 1 (0.3)   |           |
| 血圧低下                               |                              |         |                        |         |                        |          |                        |         |                          | 1 (0.3) | 1 (0.3)   |           |
| グリコヘモグロビン増加                        |                              |         |                        |         |                        | 1 (3.4)  | 1 (3.4)                |         |                          | 1 (0.3) | 1 (0.3)   |           |
| 尿中ケトン体陽性                           |                              |         |                        |         |                        |          |                        |         |                          | 1 (0.3) | 1 (0.3)   |           |
| 血中尿素増加                             | 1 (1.8)                      |         | 1 (1.8)                |         |                        |          |                        |         |                          |         |           |           |
| 単球数減少                              | 1 (1.8)                      |         | 1 (1.8)                |         |                        |          |                        |         |                          |         |           |           |
| 腎機能検査異常                            |                              |         |                        |         |                        |          |                        |         |                          | 1 (0.3) | 1 (0.3)   |           |
| 傷害、中毒および処置合併症 (SOC sub-total)      | 3 (5.3)                      | 3 (5.3) | 3 (5.3)                | 1 (4.0) | 1 (4.0)                | 4 (13.8) | 4 (13.8)               | 2 (8.3) | 2 (8.3)                  | 1 (0.3) | 39 (12.2) | 39 (12.2) |
| 転倒                                 | 3 (5.3)                      | 3 (5.3) | 3 (5.3)                | 1 (4.0) | 1 (4.0)                | 1 (3.4)  | 1 (3.4)                | 2 (8.3) | 2 (8.3)                  | 1 (0.3) | 17 (5.3)  | 17 (5.3)  |
| 挫傷                                 |                              |         |                        |         |                        |          |                        |         |                          |         | 13 (4.1)  | 13 (4.1)  |
| 靭帯捻挫                               |                              |         |                        |         |                        | 1 (3.4)  | 1 (3.4)                |         |                          |         | 5 (1.6)   | 5 (1.6)   |
| 事故                                 |                              |         |                        |         |                        | 1 (3.4)  | 1 (3.4)                |         |                          |         |           |           |
| 交通事故                               |                              |         |                        |         |                        |          |                        |         |                          |         | 5 (1.6)   | 5 (1.6)   |
| 節足動物刺傷                             |                              |         |                        |         |                        |          |                        |         |                          |         | 2 (0.6)   | 2 (0.6)   |
| 足骨折                                |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 熱傷                                 |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 家庭内事故                              |                              |         |                        |         |                        | 1 (3.4)  | 1 (3.4)                | 1 (4.2) | 1 (4.2)                  |         |           |           |
| 靭帯損傷                               |                              |         |                        |         |                        |          |                        |         |                          |         | 2 (0.6)   | 2 (0.6)   |
| 歯牙破折                               |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 熱中症                                |                              |         |                        |         |                        |          |                        |         |                          |         | 2 (0.6)   | 2 (0.6)   |
| 関節脱臼                               |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 脊椎圧迫骨折                             |                              |         |                        |         |                        |          |                        |         |                          |         | 2 (0.6)   | 2 (0.6)   |
| 創傷                                 |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 節足動物咬傷                             |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 鎖骨骨折                               |                              |         |                        |         |                        |          |                        |         |                          |         | 2 (0.6)   | 2 (0.6)   |
| 上顎炎                                |                              |         |                        |         |                        |          |                        |         |                          |         | 2 (0.6)   | 2 (0.6)   |
| 手骨折                                |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 大腿骨骨折                              |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 筋損傷                                |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 橈骨骨折                               |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 処置による疼痛                            |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 外傷後頸部症候群                           |                              |         |                        |         |                        |          |                        |         |                          |         | 1 (0.3)   | 1 (0.3)   |
| 外科および内科処置 (SOC sub-total)          | 1 (1.8)                      | 1 (1.8) | 1 (1.8)                |         |                        |          |                        |         |                          |         |           |           |
| 抜歯                                 | 1 (1.8)                      |         | 1 (1.8)                |         |                        |          |                        |         |                          |         |           |           |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（17/21）

| System Organ Class / Adverse Event     |                      | 35.0kg/m <sup>2</sup> - Max |          |                        |         |                        |          |                        |          |                          |         |          |          |          |           |           |
|--|----------------------|-----------------------------|----------|------------------------|---------|------------------------|----------|------------------------|----------|--------------------------|---------|----------|----------|----------|-----------|-----------|
|  |                      | Placebo<br>(n=24)           |          | ATL-962 15mg<br>(n=18) |         | ATL-962 30mg<br>(n=11) |          | ATL-962 60mg<br>(n=12) |          | ATL-962 120mg<br>(n=140) |         |          |          |          |           |           |
|  |                      | 関連あり<br>関連なし                | 合計       | 関連あり<br>関連なし           | 合計      | 関連あり<br>関連なし           | 合計       | 関連あり<br>関連なし           | 合計       | 関連あり<br>関連なし             | 合計      |          |          |          |           |           |
| Over All                               | (Number of Patients) | 8(33.3)                     | 20(83.3) | 20(83.3)               | 6(33.3) | 12(66.7)               | 12(66.7) | 5(45.5)                | 7(63.6)  | 9(81.8)                  | 7(58.3) | 11(91.7) | 11(91.7) | 90(64.3) | 110(78.6) | 128(91.4) |
|  | (Number of AEs)      | 11                          | 68       | 79                     | 8       | 32                     | 40       | 8                      | 16       | 24                       | 14      | 36       | 50       | 115      | 355       | 470       |
| 感染症および寄生虫症                             | (SOC sub-total)      | 1 (4.2)                     | 15(62.5) | 15(62.5)               | 9(50.0) | 9(50.0)                | 5(45.5)  | 5(45.5)                | 10(83.3) | 10(83.3)                 | 1 (0.7) | 70(50.0) | 71(50.7) | 46(32.9) | 46(32.9)  |           |
|  | 鼻咽頭炎                 |                             | 10(41.7) | 10(41.7)               | 7(38.9) | 7(38.9)                | 2(18.2)  | 2(18.2)                | 10(83.3) | 10(83.3)                 |         | 46(32.9) | 46(32.9) |          |           |           |
|  | 咽頭炎                  |                             | 3(12.5)  | 3(12.5)                | 2(11.1) | 2(11.1)                |          |                        |          |                          | 1 (0.7) | 5 (3.6)  | 6 (4.3)  |          |           |           |
|  | 膀胱炎                  |                             | 1 (4.2)  | 1 (4.2)                |         |                        |          |                        |          |                          |         | 4 (2.9)  | 4 (2.9)  |          |           |           |
|  | 気管支炎                 |                             | 2 (8.3)  | 2 (8.3)                | 1 (5.6) | 1 (5.6)                | 1 (9.1)  | 1 (9.1)                |          |                          |         | 1 (0.7)  | 1 (0.7)  |          |           |           |
|  | インフルエンザ              |                             |          |                        |         |                        |          | 1 (9.1)                | 1 (9.1)  |                          |         | 4 (2.9)  | 4 (2.9)  |          |           |           |
|  | 胃腸炎                  |                             |          |                        |         |                        |          |                        |          |                          |         | 3 (2.1)  | 3 (2.1)  |          |           |           |
|  | 足部白癬                 |                             | 1 (4.2)  | 1 (4.2)                |         |                        |          |                        |          |                          |         | 2 (1.4)  | 2 (1.4)  |          |           |           |
|  | 麦粒腫                  |                             | 1 (4.2)  | 1 (4.2)                |         |                        |          |                        |          |                          |         | 5 (3.6)  | 5 (3.6)  |          |           |           |
|  | 蜂巣炎                  |                             | 1 (4.2)  | 1 (4.2)                |         |                        |          |                        |          |                          |         | 2 (1.4)  | 2 (1.4)  |          |           |           |
|  | 爪囲炎                  |                             | 1 (4.2)  | 1 (4.2)                |         |                        |          |                        |          |                          |         |          |          |          |           |           |
|  | 鼻炎                   |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 2 (1.4)  | 2 (1.4)   |           |
|  | 中耳炎                  |                             |          |                        |         |                        |          | 1 (9.1)                | 1 (9.1)  |                          |         |          |          |          |           |           |
|  | 扁桃炎                  |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 3 (2.1)  | 3 (2.1)   |           |
|  | 急性扁桃炎                |                             |          |                        |         |                        |          | 1 (9.1)                | 1 (9.1)  |                          |         |          |          | 2 (1.4)  | 2 (1.4)   |           |
|  | ウイルス性胃腸炎             |                             |          |                        | 1 (5.6) | 1 (5.6)                |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 爪真菌症                 |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 外耳炎                  |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 白癬感染                 |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 感染性皮膚膿腫              |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 肺炎                   |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 歯髄炎                  |                             |          | 1 (4.2)                | 1 (4.2) |                        |          |                        |          |                          |         |          |          |          |           |           |
|  | 副鼻腔炎                 |                             | 1 (4.2)  | 1 (4.2)                |         |                        |          |                        |          |                          |         |          |          |          |           |           |
|  | せつ                   |                             |          | 1 (4.2)                | 1 (4.2) |                        |          |                        |          |                          |         |          |          |          |           |           |
|  | 感染                   |                             |          |                        |         |                        |          |                        |          |                          | 1 (8.3) | 1 (8.3)  |          |          |           |           |
|  | 急性腎盂腎炎               |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 肛門膿瘍                 |                             |          |                        |         | 1 (5.6)                | 1 (5.6)  |                        |          |                          |         |          |          |          |           |           |
|  | 感染性腸炎                |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 癰                    |                             |          | 1 (4.2)                | 1 (4.2) |                        |          |                        |          |                          |         |          |          |          |           |           |
|  | 慢性副鼻腔炎               |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 慢性扁桃炎                |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 皮膚真菌感染               |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 歯膿瘍                  |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 細菌性咽頭炎               |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
| 良性、悪性および詳細不明の<br>新生物(嚢胞およびポリープ<br>を含む) | (SOC sub-total)      |                             |          |                        |         |                        |          | 1 (9.1)                | 1 (9.1)  |                          |         |          |          | 1 (0.7)  | 3 (2.1)   | 4 (2.9)   |
|  | 子宮平滑筋腫               |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   | 2 (1.4)   |
|  | 皮膚乳頭腫                |                             |          |                        |         |                        |          | 1 (9.1)                | 1 (9.1)  |                          |         |          |          |          |           |           |
|  | 乳癌                   |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 縦隔新生物                |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
| 血液およびリンパ系障害                            | (SOC sub-total)      |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
|  | 鉄欠乏性貧血               |                             |          |                        |         |                        |          |                        |          |                          |         |          |          | 1 (0.7)  | 1 (0.7)   |           |
| 免疫系障害                                  | (SOC sub-total)      |                             |          |                        |         |                        |          |                        |          |                          | 1 (8.3) | 1 (8.3)  |          | 2 (1.4)  | 2 (1.4)   |           |
|  | 季節性アレルギー             |                             |          |                        |         |                        |          |                        |          |                          | 1 (8.3) | 1 (8.3)  |          | 2 (1.4)  | 2 (1.4)   |           |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（18/21）

| System Organ Class / Adverse Event | 35.0kg/m <sup>2</sup> - Max |          |    |                        |         |    |                        |         |          |                        |         |                          |          |
|------------------------------------|-----------------------------|----------|----|------------------------|---------|----|------------------------|---------|----------|------------------------|---------|--------------------------|----------|
|                                    | Placebo<br>(n=24)           |          |    | ATL-962 15mg<br>(n=18) |         |    | ATL-962 30mg<br>(n=11) |         |          | ATL-962 60mg<br>(n=12) |         | ATL-962 120mg<br>(n=140) |          |
|                                    | 関連あり                        | 関連なし     | 合計 | 関連あり                   | 関連なし    | 合計 | 関連あり                   | 関連なし    | 合計       | 関連あり                   | 関連なし    | 合計                       |          |
| 内分泌障害                              | (SOC sub-total)             |          |    |                        |         |    |                        |         |          |                        |         |                          |          |
| 甲状腺腫                               |                             |          |    |                        |         |    |                        |         |          | 1 (0.7)                | 1 (0.7) |                          |          |
| 代謝および栄養障害                          | (SOC sub-total)             |          |    |                        |         |    |                        |         |          |                        |         |                          |          |
| 糖尿病                                |                             |          |    | 1 (5.6)                | 1 (5.6) |    |                        |         | 1 (8.3)  | 1 (8.3)                | 3 (2.1) | 11 (7.9)                 | 13 (9.3) |
| 低血糖症                               |                             |          |    |                        |         |    |                        |         |          |                        | 2 (1.4) | 3 (2.1)                  | 5 (3.6)  |
| 高尿酸血症                              |                             |          |    |                        |         |    |                        |         | 1 (8.3)  | 1 (8.3)                |         | 2 (1.4)                  | 2 (1.4)  |
| 2型糖尿病                              |                             |          |    |                        |         |    |                        |         |          |                        |         | 3 (2.1)                  | 3 (2.1)  |
| コントロール不良の糖尿病                       |                             |          |    |                        |         |    |                        |         |          |                        | 1 (0.7) | 2 (1.4)                  | 3 (2.1)  |
| 過食                                 |                             |          |    | 1 (5.6)                | 1 (5.6) |    |                        |         |          |                        |         |                          |          |
| 精神障害                               | (SOC sub-total)             |          |    |                        |         |    |                        |         |          |                        |         |                          |          |
| 不眠症                                |                             |          |    |                        |         |    |                        |         |          |                        |         | 4 (2.9)                  | 4 (2.9)  |
| うつ病                                |                             |          |    |                        |         |    |                        |         |          |                        |         | 3 (2.1)                  | 3 (2.1)  |
| 睡眠障害                               |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 神経系障害                              | (SOC sub-total)             |          |    |                        |         |    |                        |         |          |                        |         |                          |          |
| 頭痛                                 | 4 (16.7)                    | 4 (16.7) |    | 1 (5.6)                | 1 (5.6) |    |                        |         | 2 (16.7) | 2 (16.7)               |         | 13 (9.3)                 | 13 (9.3) |
| 感覚鈍麻                               | 3 (12.5)                    | 3 (12.5) |    | 1 (5.6)                | 1 (5.6) |    |                        |         | 2 (16.7) | 2 (16.7)               |         | 7 (5.0)                  | 7 (5.0)  |
| 坐骨神経痛                              | 1 (4.2)                     | 1 (4.2)  |    |                        |         |    |                        |         | 1 (8.3)  | 1 (8.3)                |         |                          |          |
| 脳梗塞                                |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 自律神経失調                             |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 頸髄症                                | 1 (4.2)                     | 1 (4.2)  |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 振戦                                 |                             |          |    |                        |         |    |                        |         | 1 (8.3)  | 1 (8.3)                |         |                          |          |
| 第7脳神経麻痺                            |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 大脳動脈狭窄                             |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 眼障害                                | (SOC sub-total)             |          |    |                        |         |    |                        |         |          |                        |         |                          |          |
| 白内障                                | 2 (8.3)                     | 2 (8.3)  |    | 1 (5.6)                | 1 (5.6) |    |                        |         | 1 (8.3)  | 1 (8.3)                |         | 11 (7.9)                 | 11 (7.9) |
| 糖尿病網膜症                             |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 結膜炎                                |                             |          |    | 1 (5.6)                | 1 (5.6) |    |                        |         | 1 (8.3)  | 1 (8.3)                |         | 3 (2.1)                  | 3 (2.1)  |
| アレルギー性結膜炎                          | 1 (4.2)                     | 1 (4.2)  |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 網膜出血                               |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 眼精疲労                               |                             |          |    |                        |         |    |                        |         |          |                        |         | 2 (1.4)                  | 2 (1.4)  |
| 眼乾燥                                |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 硝子体浮遊物                             | 1 (4.2)                     | 1 (4.2)  |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 霰粒腫                                |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 緑内障                                |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 耳および迷路障害                           | (SOC sub-total)             |          |    |                        |         |    |                        |         |          |                        |         |                          |          |
| 回転性めまい                             |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 2 (1.4)  |
| 頭位性回転性めまい                          |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 心臓障害                               | (SOC sub-total)             |          |    |                        |         |    |                        |         |          |                        |         |                          |          |
| 動悸                                 |                             |          |    | 1 (5.6)                | 1 (5.6) |    |                        | 1 (9.1) | 1 (9.1)  |                        |         | 4 (2.9)                  | 4 (2.9)  |
| 心室性期外収縮                            |                             |          |    | 1 (5.6)                | 1 (5.6) |    |                        | 1 (9.1) | 1 (9.1)  |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 左脚ブロック                             |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 心不全                                |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |
| 血管障害                               | (SOC sub-total)             |          |    |                        |         |    |                        |         |          |                        |         |                          |          |
| 高血圧                                |                             |          |    |                        |         |    |                        |         |          |                        |         | 1 (0.7)                  | 1 (0.7)  |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（19/21）

| System Organ Class / Adverse Event | 35.0kg/m <sup>2</sup> - Max |         |                        |         |                        |         |                        |         |                          |         |          |          |          |          |          |        |
|------------------------------------|-----------------------------|---------|------------------------|---------|------------------------|---------|------------------------|---------|--------------------------|---------|----------|----------|----------|----------|----------|--------|
|                                    | Placebo<br>(n=24)           |         | ATL-962 15mg<br>(n=18) |         | ATL-962 30mg<br>(n=11) |         | ATL-962 60mg<br>(n=12) |         | ATL-962 120mg<br>(n=140) |         |          |          |          |          |          |        |
|                                    | 関連あり                        | 関連なし    | 合計                     | 関連あり    | 関連なし                   | 合計      | 関連あり                   | 関連なし    | 合計                       | 関連あり    | 合計       |          |          |          |          |        |
| 呼吸器、胸郭および縦隔障害 (SOC sub-total)      | 3(12.5)                     | 3(12.5) | 3(12.5)                | 1(5.6)  | 1(5.6)                 | 1(5.6)  | 1(8.3)                 | 1(8.3)  | 1(8.3)                   | 1(8.3)  | 22(15.7) | 22(15.7) |          |          |          |        |
| 上気道の炎症                             | 3(12.5)                     | 3(12.5) | 3(12.5)                | 1(5.6)  | 1(5.6)                 | 1(5.6)  |                        |         |                          |         | 12(8.6)  | 12(8.6)  |          |          |          |        |
| 咳嗽                                 |                             |         |                        |         |                        |         |                        |         |                          |         | 5(3.6)   | 5(3.6)   |          |          |          |        |
| アレルギー性鼻炎                           | 1(4.2)                      | 1(4.2)  | 1(4.2)                 |         |                        |         |                        |         |                          |         | 1(0.7)   | 1(0.7)   |          |          |          |        |
| 喘息                                 |                             |         |                        |         |                        |         |                        |         |                          |         | 1(0.7)   | 1(0.7)   |          |          |          |        |
| 鼻出血                                |                             |         |                        |         |                        |         | 1(8.3)                 | 1(8.3)  |                          |         |          |          |          |          |          |        |
| 睡眠時無呼吸症候群                          |                             |         |                        |         |                        |         |                        |         |                          |         | 1(0.7)   | 1(0.7)   |          |          |          |        |
| 湿性咳嗽                               |                             |         |                        |         |                        |         |                        |         |                          |         | 1(0.7)   | 1(0.7)   |          |          |          |        |
| 肺水腫                                |                             |         |                        |         |                        |         |                        |         |                          |         | 1(0.7)   | 1(0.7)   |          |          |          |        |
| 気管軟化症                              |                             |         |                        |         |                        |         |                        |         |                          |         | 1(0.7)   | 1(0.7)   |          |          |          |        |
| びまん性汎細気管支炎                         |                             |         |                        |         |                        |         |                        |         |                          |         | 1(0.7)   | 1(0.7)   |          |          |          |        |
| 胃腸障害 (SOC sub-total)               | 8(33.3)                     | 3(12.5) | 11(45.8)               | 5(27.8) | 3(16.7)                | 7(38.9) | 5(45.5)                | 2(18.2) | 7(63.6)                  | 7(58.3) | 2(16.7)  | 8(66.7)  | 83(59.3) | 21(15.0) | 88(62.9) |        |
| 脂肪便                                | 2(8.3)                      |         | 2(8.3)                 | 1(5.6)  |                        | 1(5.6)  | 3(27.3)                |         | 3(27.3)                  | 3(25.0) |          | 3(25.0)  | 51(36.4) |          | 51(36.4) |        |
| 下痢                                 | 3(12.5)                     | 1(4.2)  | 4(16.7)                | 3(16.7) |                        | 3(16.7) | 2(18.2)                |         | 2(18.2)                  | 3(25.0) | 1(8.3)   | 4(33.3)  | 33(23.6) | 1(0.7)   | 34(24.3) |        |
| 便秘                                 | 2(8.3)                      | 1(4.2)  | 3(12.5)                |         |                        |         |                        |         |                          |         |          |          | 2(1.4)   | 2(1.4)   | 4(2.9)   |        |
| 膈疝                                 |                             |         |                        |         |                        | 1(5.6)  | 1(5.6)                 |         |                          |         |          |          | 2(1.4)   | 2(1.4)   | 2(1.4)   |        |
| 腹部膨満                               | 1(4.2)                      | 1(4.2)  | 1(4.2)                 | 1(5.6)  | 1(5.6)                 | 1(5.6)  | 1(9.1)                 |         | 1(9.1)                   | 1(8.3)  |          | 1(8.3)   | 1(0.7)   |          | 1(0.7)   |        |
| 腹痛                                 | 1(4.2)                      | 1(4.2)  | 1(4.2)                 | 1(5.6)  |                        | 1(5.6)  |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 悪心                                 |                             |         |                        |         |                        |         |                        |         | 1(9.1)                   | 1(9.1)  | 1(8.3)   |          | 1(8.3)   | 1(0.7)   | 2(1.4)   | 3(2.1) |
| 腹部不快感                              | 1(4.2)                      | 1(4.2)  | 2(8.3)                 |         |                        |         |                        | 1(9.1)  | 1(9.1)                   | 1(8.3)  |          | 1(8.3)   | 1(0.7)   |          | 1(0.7)   |        |
| 上腹部痛                               |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 歯周病                                |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 3(2.1)   | 4(2.9) |
| 嘔吐                                 |                             |         |                        |         |                        |         |                        |         |                          | 1(8.3)  | 1(8.3)   |          |          |          |          |        |
| 歯周炎                                |                             |         |                        |         |                        |         |                        | 1(9.1)  | 1(9.1)                   |         |          |          |          |          |          |        |
| 痔核                                 |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 2(1.4)   | 2(1.4)   | 2(1.4)   |        |
| 口内炎                                |                             |         |                        |         |                        | 1(5.6)  | 1(5.6)                 |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 歯肉炎                                |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 消化不良                               |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 胃炎                                 |                             |         |                        |         |                        | 1(5.6)  | 1(5.6)                 |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 歯痛                                 |                             |         |                        |         |                        |         |                        |         |                          | 1(8.3)  | 1(8.3)   |          | 1(0.7)   |          | 1(0.7)   |        |
| 結腸ポリープ                             | 1(4.2)                      |         | 1(4.2)                 |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   | 2(1.4)   | 2(1.4)   |        |
| 排便回数増加                             |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 2(1.4)   |          | 2(1.4)   |        |
| びらん性胃炎                             |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 胃食道逆流性疾患                           |                             |         |                        |         |                        | 1(5.6)  | 1(5.6)                 |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 腸炎                                 |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 口唇炎                                |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 十二指腸潰瘍                             |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 直腸分泌物                              |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 食中毒                                |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| メレナ                                |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 便秘不規則                              | 1(4.2)                      |         | 1(4.2)                 |         |                        |         |                        |         |                          |         |          |          |          |          |          |        |
| 肝胆道系障害 (SOC sub-total)             |                             | 1(4.2)  | 1(4.2)                 |         |                        |         |                        | 1(9.1)  | 1(9.1)                   |         |          |          | 2(1.4)   | 4(2.9)   | 6(4.3)   |        |
| 肝機能異常                              |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   | 2(1.4)   | 3(2.1)   |        |
| 脂肪肝                                |                             |         |                        |         |                        |         |                        | 1(9.1)  | 1(9.1)                   |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 胆石症                                |                             | 1(4.2)  | 1(4.2)                 |         |                        |         |                        |         |                          |         |          |          |          |          |          |        |
| 肝障害                                |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |
| 肝嚢胞                                |                             |         |                        |         |                        |         |                        |         |                          |         |          |          | 1(0.7)   |          | 1(0.7)   |        |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（20/21）

| System Organ Class / Adverse Event | 35.0kg/m <sup>2</sup> - Max |          |                        |          |                        |          |                        |          |                          |           |           |
|------------------------------------|-----------------------------|----------|------------------------|----------|------------------------|----------|------------------------|----------|--------------------------|-----------|-----------|
|                                    | Placebo<br>(n=24)           |          | ATL-962 15mg<br>(n=18) |          | ATL-962 30mg<br>(n=11) |          | ATL-962 60mg<br>(n=12) |          | ATL-962 120mg<br>(n=140) |           |           |
|                                    | 関連あり                        | 関連なし     | 合計                     | 関連あり     | 関連なし                   | 合計       | 関連あり                   | 関連なし     | 合計                       | 関連あり      | 関連なし      |
| 皮膚および皮下組織障害 (SOC sub-total)        | 1 (4.2)                     | 1 (4.2)  | 2 (4.2)                | 1 (5.6)  | 1 (5.6)                | 2 (18.2) | 2 (16.7)               | 2 (16.7) | 4 (2.9)                  | 21 (15.0) | 24 (17.1) |
| 湿疹                                 | 1 (4.2)                     | 1 (4.2)  | 2 (4.2)                |          |                        |          | 2 (16.7)               | 2 (16.7) |                          | 5 (3.6)   | 5 (3.6)   |
| 接触性皮膚炎                             |                             |          |                        |          |                        |          |                        |          | 1 (0.7)                  | 4 (2.9)   | 5 (3.6)   |
| そう痒症                               |                             |          |                        |          |                        |          |                        |          | 1 (0.7)                  | 1 (0.7)   | 1 (0.7)   |
| 蕁麻疹                                |                             |          |                        | 1 (5.6)  | 1 (5.6)                | 2 (18.2) |                        |          | 1 (0.7)                  | 1 (0.7)   | 2 (1.4)   |
| 発疹                                 |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 過角化                                |                             |          |                        |          |                        |          |                        |          | 1 (0.7)                  | 2 (1.4)   | 3 (2.1)   |
| 皮脂欠乏性湿疹                            |                             |          |                        |          |                        |          |                        |          | 1 (0.7)                  | 2 (1.4)   | 3 (2.1)   |
| 皮脂欠乏症                              |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 水疱                                 |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 皮膚剥脱                               |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| うっ滞性皮膚炎                            |                             |          |                        |          |                        |          |                        |          |                          | 2 (1.4)   | 2 (1.4)   |
| 全身性そう痒症                            |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 褥瘡性潰瘍                              |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 老人性そう痒症                            |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 筋骨格系および結合組織障害 (SOC sub-total)      | 8 (33.3)                    | 8 (33.3) | 16 (33.3)              | 3 (16.7) | 3 (16.7)               | 6 (54.5) | 3 (27.3)               | 3 (27.3) | 1 (8.3)                  | 1 (8.3)   | 22 (15.7) |
| 背部痛                                | 1 (4.2)                     | 1 (4.2)  | 2 (4.2)                |          |                        |          |                        |          |                          | 6 (4.3)   | 6 (4.3)   |
| 関節痛                                | 4 (16.7)                    | 4 (16.7) | 8 (16.7)               | 1 (5.6)  | 1 (5.6)                | 2 (18.2) | 1 (9.1)                | 1 (9.1)  | 3 (2.1)                  | 3 (2.1)   | 3 (2.1)   |
| 変形性関節症                             |                             |          |                        |          |                        |          | 1 (9.1)                | 1 (9.1)  |                          | 2 (1.4)   | 2 (1.4)   |
| 筋肉痛                                |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 筋骨格硬直                              | 2 (8.3)                     | 2 (8.3)  | 4 (8.3)                |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 関節周囲炎                              |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 変形性脊椎症                             |                             |          |                        | 1 (5.6)  | 1 (5.6)                | 2 (18.2) |                        |          |                          | 2 (1.4)   | 2 (1.4)   |
| 椎間板突出                              |                             |          |                        |          |                        |          |                        |          |                          | 2 (1.4)   | 2 (1.4)   |
| 関節炎                                |                             |          |                        |          |                        |          |                        |          | 1 (8.3)                  | 1 (8.3)   | 2 (1.4)   |
| 筋痙攣                                |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 筋骨格痛                               |                             |          |                        |          |                        |          | 1 (9.1)                | 1 (9.1)  |                          | 1 (0.7)   | 1 (0.7)   |
| 滑液嚢腫                               |                             |          |                        | 1 (5.6)  | 1 (5.6)                | 2 (18.2) |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 脊椎靭帯骨化症                            | 1 (4.2)                     | 1 (4.2)  | 2 (4.2)                |          |                        |          |                        |          |                          |           |           |
| 筋膜炎                                |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 足底筋膜炎                              |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 脊柱管狭窄症                             |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 椎間板障害                              |                             |          |                        | 1 (5.6)  | 1 (5.6)                | 2 (18.2) |                        |          |                          |           |           |
| 横紋筋融解症                             |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 腎および尿路障害 (SOC sub-total)           |                             |          |                        |          |                        |          |                        |          |                          | 2 (1.4)   | 2 (1.4)   |
| 頻尿                                 |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 腎萎縮                                |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 腎腫大                                |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 生殖系および乳房障害 (SOC sub-total)         |                             |          |                        |          |                        |          |                        |          |                          | 2 (1.4)   | 2 (1.4)   |
| 乳房石灰化                              |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 乳房不快感                              |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 先天性、家族性および遺伝性障害 (SOC sub-total)    |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 朝顔症候群                              |                             |          |                        |          |                        |          |                        |          |                          | 1 (0.7)   | 1 (0.7)   |

表中の数字：発現例数（発現率（%））

表 2.7.4.7-付録-7 BMI 別・治験薬との因果関係別、有害事象発現頻度（国内臨床試験）（21/21）

| System Organ Class / Adverse Event | 35.0kg/m <sup>2</sup> - Max |          |                        |         |                        |         |                        |          |                          |           |           |
|------------------------------------|-----------------------------|----------|------------------------|---------|------------------------|---------|------------------------|----------|--------------------------|-----------|-----------|
|                                    | Placebo<br>(n=24)           |          | ATL-962 15mg<br>(n=18) |         | ATL-962 30mg<br>(n=11) |         | ATL-962 60mg<br>(n=12) |          | ATL-962 120mg<br>(n=140) |           |           |
|                                    | 関連あり<br>関連なし                | 合計       | 関連あり<br>関連なし           | 合計      | 関連あり<br>関連なし           | 合計      | 関連あり<br>関連なし           | 合計       | 関連あり<br>関連なし             | 合計        |           |
| 一般・全身障害および投与部位の状態 (SOC sub-total)  | 1 (4.2)                     | 1 (4.2)  | 1 (5.6)                | 1 (5.6) | 1 (8.3)                | 1 (8.3) | 1 (8.3)                | 1 (8.3)  | 1 (0.7)                  | 5 (3.6)   | 6 (4.3)   |
| 末梢性浮腫                              |                             |          | 1 (5.6)                | 1 (5.6) |                        |         | 1 (8.3)                | 1 (8.3)  |                          | 3 (2.1)   | 3 (2.1)   |
| 倦怠感                                |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 発熱                                 |                             |          |                        |         |                        |         |                        |          | 1 (0.7)                  |           | 1 (0.7)   |
| 悪寒                                 | 1 (4.2)                     | 1 (4.2)  |                        |         |                        |         |                        |          |                          |           |           |
| 医療機器関連損傷                           |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 臨床検査 (SOC sub-total)               | 3 (12.5)                    | 3 (12.5) | 1 (5.6)                | 1 (5.6) | 1 (9.1)                | 1 (9.1) | 1 (8.3)                | 1 (8.3)  | 2 (16.7)                 | 5 (3.6)   | 12 (8.6)  |
| 血中クレアチンホスホキナーゼ増加                   | 2 (8.3)                     | 2 (8.3)  |                        |         |                        |         |                        |          |                          |           | 3 (2.1)   |
| アラニンアミノトランスフェラーゼ増加                 |                             |          |                        |         |                        |         | 1 (8.3)                | 1 (8.3)  | 1 (0.7)                  | 3 (2.1)   | 4 (2.9)   |
| アスパラギン酸アミノトランスフェラーゼ増加              |                             |          |                        |         |                        |         |                        |          |                          | 3 (2.1)   | 3 (2.1)   |
| γ-グルタミルトランスフェラーゼ増加                 |                             |          |                        |         |                        |         |                        |          |                          | 2 (1.4)   | 2 (1.4)   |
| 血中尿酸増加                             |                             |          |                        |         |                        |         |                        |          | 1 (0.7)                  | 1 (0.7)   | 2 (1.4)   |
| サイトケラチン 18 増加                      |                             |          |                        |         | 1 (9.1)                | 1 (9.1) |                        |          |                          |           |           |
| 肝機能検査異常                            |                             |          |                        |         |                        |         |                        |          | 1 (0.7)                  | 3 (2.1)   | 4 (2.9)   |
| 尿中血陽性                              |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| m-A S T 増加                         | 1 (4.2)                     | 1 (4.2)  | 1 (5.6)                | 1 (5.6) |                        |         |                        |          |                          |           |           |
| 好酸球数増加                             |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 尿中ケトン体陽性                           |                             |          |                        |         |                        |         |                        |          | 1 (0.7)                  |           | 1 (0.7)   |
| 血中コリンエステラーゼ増加                      |                             |          |                        |         |                        |         |                        |          | 1 (0.7)                  |           | 1 (0.7)   |
| ヘマトクリット減少                          |                             |          |                        |         |                        |         | 1 (8.3)                | 1 (8.3)  |                          |           |           |
| ヘモグロビン減少                           |                             |          |                        |         |                        |         | 1 (8.3)                | 1 (8.3)  |                          |           |           |
| 傷害、中毒および処置合併症 (SOC sub-total)      | 3 (12.5)                    | 3 (12.5) |                        |         |                        |         | 3 (25.0)               | 3 (25.0) |                          | 15 (10.7) | 15 (10.7) |
| 転倒                                 | 2 (8.3)                     | 2 (8.3)  |                        |         |                        |         | 1 (8.3)                | 1 (8.3)  |                          | 6 (4.3)   | 6 (4.3)   |
| 挫傷                                 |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 靭帯捻挫                               |                             |          |                        |         |                        |         |                        |          |                          | 3 (2.1)   | 3 (2.1)   |
| 事故                                 | 2 (8.3)                     | 2 (8.3)  |                        |         |                        |         |                        |          |                          |           |           |
| 交通事故                               |                             |          |                        |         |                        |         |                        |          |                          | 2 (1.4)   | 2 (1.4)   |
| 節足動物刺傷                             | 1 (4.2)                     | 1 (4.2)  |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 足骨折                                |                             |          |                        |         |                        |         |                        |          |                          | 2 (1.4)   | 2 (1.4)   |
| 擦過傷                                |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 熱傷                                 |                             |          |                        |         |                        |         | 1 (8.3)                | 1 (8.3)  |                          | 1 (0.7)   | 1 (0.7)   |
| 家庭内事故                              |                             |          |                        |         |                        |         | 1 (8.3)                | 1 (8.3)  |                          |           |           |
| 熱中症                                |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 上腕骨骨折                              |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 創傷                                 |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 手骨折                                |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 肉離れ                                |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 皮下血腫                               |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 術後貧血                               |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 骨盤骨折                               |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |
| 四肢損傷                               | 1 (4.2)                     | 1 (4.2)  |                        |         |                        |         |                        |          |                          |           |           |
| 骨亀裂                                |                             |          |                        |         |                        |         |                        |          |                          | 1 (0.7)   | 1 (0.7)   |

表中の数字：発現例数（発現率 (%)）

表 2.7.4.7-付録-8 治験薬投与中止（治験中止）に至った有害事象の一覧（CCT-001 試験）（1/3）

| 投与群        | 被験者識別コード   | 人種  | 性別 | 年齢     | 有害事象名<br>基本語 <sup>1)</sup> | 発現日<br>(投与後日数) | 因果関係       | 程度  | 転帰 |
|------------|------------|-----|----|--------|----------------------------|----------------|------------|-----|----|
| プラセボ群      | [REDACTED] | 日本人 | 女性 | 56     | 胃不快感*                      | 3              | 関連あるかもしれない | 軽度  | 回復 |
|            |            | 日本人 | 女性 | 65     | 下痢*                        | 3              | 関連あるかもしれない | 中等度 | 回復 |
|            |            |     |    |        | 嘔吐*                        | 14             | 関連あるかもしれない | 軽度  | 回復 |
|            |            |     |    |        | AST 増加                     | 15             | 関連あるかもしれない | 軽度  | 回復 |
|            |            |     |    |        | ALT 増加                     | 15             | 関連あるかもしれない | 軽度  | 回復 |
|            |            |     |    | 臨床検査異常 | 15                         | 関連あるかもしれない     | 軽度         | 回復  |    |
| 15 mg 群    | [REDACTED] | 日本人 | 男性 | 76     | 膀胱新生物*                     | 29             | 関連なし       | 中等度 | 回復 |
|            |            |     |    |        | 排尿困難                       | 51             | 関連なし       | 軽度  | 回復 |
|            | [REDACTED] | 日本人 | 男性 | 54     | 下痢                         | 1              | 多分関連あり     | 軽度  | 回復 |
|            |            |     |    |        | 脂肪便*                       | 12             | 多分関連あり     | 中等度 | 回復 |
|            | [REDACTED] | 日本人 | 女性 | 60     | 鼻咽頭炎                       | 12             | 関連なし       | 軽度  | 回復 |
|            |            |     |    |        | 脳梗塞*                       | 96             | 関連なし       | 高度  | 死亡 |
| 30 mg 群    | [REDACTED] | 日本人 | 男性 | 47     | 下痢*                        | 5              | 多分関連あり     | 中等度 | 回復 |
|            |            |     |    |        | 鼻咽頭炎                       | 16             | 関連なし       | 軽度  | 回復 |
|            | [REDACTED] | 日本人 | 女性 | 51     | 下痢                         | 36             | 多分関連あり     | 軽度  | 回復 |
|            |            |     |    |        | 急性腎盂腎炎*                    | 40             | 関連なし       | 中等度 | 回復 |
|            |            |     |    |        | 尿管結石*                      | 40             | 関連なし       | 中等度 | 回復 |
|            |            |     |    |        | 腎結石症*                      | 40             | 関連なし       | 中等度 | 回復 |
| [REDACTED] | 日本人        | 女性  | 76 | 鼻咽頭炎   | 16                         | 関連なし           | 軽度         | 回復  |    |
|            |            |     |    | 転倒     | 26                         | 関連なし           | 軽度         | 回復  |    |
|            |            |     |    | 異常感*   | 61                         | 関連あるかもしれない     | 軽度         | 回復  |    |
|            |            |     |    | 腎嚢胞    | 76                         | 関連なし           | 軽度         | 未回復 |    |
|            |            |     |    | 下痢*    | 3                          | 多分関連あり         | 軽度         | 回復  |    |
| 60 mg 群    | [REDACTED] | 日本人 | 女性 | 75     | 下痢*                        | 3              | 多分関連あり     | 軽度  | 回復 |
|            |            | 日本人 | 女性 | 64     | 悪心*                        | 1              | 多分関連あり     | 軽度  | 回復 |
|            | [REDACTED] | 日本人 | 男性 | 49     | 鼻咽頭炎                       | 10             | 関連なし       | 軽度  | 回復 |
|            |            |     |    |        | 下痢*                        | 2              | 多分関連あり     | 軽度  | 回復 |
|            |            |     |    | 粘液便*   | 30                         | 多分関連あり         | 軽度         | 回復  |    |
| 60 mg 群    | [REDACTED] | 日本人 | 女性 | 58     | ALT 増加                     | 30             | 関連あるかもしれない | 軽度  | 回復 |
|            |            |     |    |        | 血中アルカリホスファターゼ増加            | 30             | 関連あるかもしれない | 軽度  | 回復 |
|            | [REDACTED] | 日本人 | 女性 | 44     | 血中クレアチニン増加*                | 30             | 関連なし       | 軽度  | 回復 |
|            |            |     |    | 下痢*    | 16                         | 明らかに関連あり       | 中等度        | 回復  |    |

事象名：MedDRA/J Ver.10.1、総括報告書（5.3.5.1-1）表 12.3.2、症例一覧表（5.3.7.1-1）から引用

有害事象発現日は、投与開始日を1日として起算

\*治験薬投与中止に至った有害事象

プラセボ群及び120 mg 群の各1例（被験者識別コード：[REDACTED] 及び [REDACTED]）の有害事象名「臨床検査異常」は、MedDRA/J Ver. 15.0で「サイトケラチン18増加」へ基本語が変更された。

15 mg 群の1例（被験者識別コード：[REDACTED]）は治験薬投与完了後に有害事象が発現した。

1) アラニン・アミノトランスフェラーゼを ALT、アスパラギン酸アミノトランスフェラーゼを AST、γ-グルタミルトランスフェラーゼを γ-GTP と表記した。

表 2.7.4.7-付録-8 治験薬投与中止（治験中止）に至った有害事象の一覧（CCT-001 試験）（2/3）

| 投与群     | 被験者識別コード | 人種  | 性別 | 年齢      | 有害事象名<br>基本語 <sup>1)</sup> | 発現日<br>(投与後日数) | 因果関係       | 程度  | 転帰  |
|---------|----------|-----|----|---------|----------------------------|----------------|------------|-----|-----|
| 60 mg 群 | ■        | 日本人 | 女性 | 73      | 胃腸潰瘍*                      | 2              | 関連あるかもしれない | 高度  | 回復  |
|         |          |     |    |         | 腹膜炎                        | 23             | 関連あるかもしれない | 中等度 | 回復  |
|         |          |     |    |         | 虫垂炎                        | 23             | 関連あるかもしれない | 中等度 | 回復  |
|         |          |     |    |         | 出血性腸炎                      | 23             | 関連あるかもしれない | 中等度 | 回復  |
|         |          |     |    |         | イレウス                       | 34             | 関連あるかもしれない | 中等度 | 回復  |
|         |          |     |    |         | カテーテル関連感染                  | 49             | 関連なし       | 軽度  | 回復  |
|         | ■        | 日本人 | 女性 | 57      | 下痢                         | 1              | 関連あるかもしれない | 軽度  | 回復  |
|         |          |     |    |         | 鼻咽頭炎*                      | 40             | 関連なし       | 軽度  | 回復  |
|         |          |     |    |         | 悪心*                        | 90             | 関連あるかもしれない | 軽度  | 回復  |
|         |          |     |    |         | 頭痛*                        | 90             | 関連あるかもしれない | 軽度  | 回復  |
|         |          |     |    |         | 転倒                         | 128            | 関連なし       | 高度  | 回復  |
|         |          |     |    |         | AST 増加                     | 12             | 多分関連あり     | 軽度  | 回復  |
|         |          |     |    |         | m-AST 増加                   | 12             | 多分関連あり     | 軽度  | 回復  |
|         | ■        | 日本人 | 女性 | 65      | 鼻咽頭炎                       | 82             | 関連なし       | 軽度  | 回復  |
|         |          |     |    |         | 糖尿病性腎症*                    | 85             | 関連なし       | 中等度 | 未回復 |
|         | ■        | 日本人 | 男性 | 58      | 鼻咽頭炎                       | 2              | 関連なし       | 軽度  | 回復  |
|         |          |     |    |         | 高血圧                        | 15             | 関連なし       | 軽度  | 回復  |
|         |          |     |    |         | 脳梗塞*                       | 15             | 関連なし       | 中等度 | 軽快  |
|         |          |     |    |         | アルツハイマー病型痴呆                | 15             | 関連なし       | 軽度  | 軽快  |
| 下痢      |          |     |    |         | 28                         | 関連なし           | 軽度         | 回復  |     |
| ■       | 日本人      | 女性  | 55 | 網膜出血*   | 15                         | 関連なし           | 中等度        | 回復  |     |
| ■       | 日本人      | 女性  | 33 | ALT 増加* | 8                          | 関連あるかもしれない     | 軽度         | 未回復 |     |
|         |          |     |    | 上腹部痛    | 25                         | 関連なし           | 軽度         | 回復  |     |
|         |          |     |    | 湿疹      | 26                         | 関連なし           | 軽度         | 回復  |     |
| ■       | 日本人      | 男性  | 55 | 下痢      | 3                          | 関連あるかもしれない     | 軽度         | 回復  |     |
|         |          |     |    | 下痢*     | 50                         | 関連あるかもしれない     | 中等度        | 回復  |     |
|         |          |     |    | 食欲不振*   | 50                         | 関連あるかもしれない     | 軽度         | 回復  |     |

事象名：MedDRA/J Ver.10.1、総括報告書（5.3.5.1-1）表 12.3.2、症例一覧表（5.3.7.1-1）から引用

有害事象発現日は、投与開始日を1日として起算

\*治験薬投与中止に至った有害事象

プラセボ群及び120 mg 群の各1例（被験者識別コード：■及び■）の有害事象名「臨床検査異常」は、MedDRA/J Ver. 15.0で「サイトケラチン18増加」へ基本語が変更された。

15 mg 群の1例（被験者識別コード：■）は治験薬投与完了後に有害事象が発現した。

1) アラニン・アミノトランスフェラーゼをALT、アスパラギン酸アミノトランスフェラーゼをAST、 $\gamma$ -グルタミルトランスフェラーゼを $\gamma$ -GTPと表記した。



表 2.7.4.7-付録-8 治験薬投与中止（治験中止）に至った有害事象の一覧（CCT-001 試験）（3/3）

| 投与群          | 被験者識別コード   | 人種  | 性別 | 年齢     | 有害事象名<br>基本語 <sup>1)</sup> | 発現日<br>(投与後日数) | 因果関係       | 程度  | 転帰  |
|--------------|------------|-----|----|--------|----------------------------|----------------|------------|-----|-----|
| 120 mg 群     | [REDACTED] | 日本人 | 女性 | 69     | 鼻咽頭炎                       | 39             | 関連なし       | 軽度  | 回復  |
|              |            |     |    |        | 腎結石症                       | 73             | 関連なし       | 軽度  | 未回復 |
|              |            |     |    |        | 帯状疱疹*                      | 70             | 関連なし       | 中等度 | 回復  |
|              | [REDACTED] | 日本人 | 女性 | 64     | 排便回数増加                     | 1              | 多分関連あり     | 軽度  | 回復  |
|              |            |     |    |        | 全身性そう痒症*                   | 25             | 多分関連あり     | 軽度  | 回復  |
|              |            |     |    |        | 網膜変性                       | 52             | 関連なし       | 軽度  | 未回復 |
|              | [REDACTED] | 日本人 | 男性 | 49     | 下痢*                        | 1              | 多分関連あり     | 中等度 | 回復  |
|              |            |     |    |        | 歯痛                         | 22             | 関連なし       | 軽度  | 回復  |
|              |            |     |    |        | 血圧低下                       | 29             | 関連あるかもしれない | 軽度  | 回復  |
|              |            |     |    |        | 蕁麻疹*                       | 52             | 関連あるかもしれない | 軽度  | 回復  |
|              | [REDACTED] | 日本人 | 女性 | 60     | 下痢*                        | 34             | 多分関連あり     | 高度  | 回復  |
|              | [REDACTED] | 日本人 | 女性 | 69     | 転倒*                        | 93             | 関連なし       | 中等度 | 未回復 |
|              |            |     |    |        | 膀胱炎                        | 114            | 関連なし       | 軽度  | 回復  |
|              |            |     |    |        | 不眠症                        | 94             | 関連なし       | 軽度  | 回復  |
|              | [REDACTED] | 日本人 | 男性 | 42     | 下痢                         | 3              | 多分関連あり     | 軽度  | 回復  |
|              |            |     |    |        | 脂肪便                        | 3              | 明らかに関連あり   | 軽度  | 回復  |
| 末梢性浮腫*       |            |     |    |        | 60                         | 関連あるかもしれない     | 軽度         | 回復  |     |
| 下痢*          |            |     |    |        | 60                         | 明らかに関連あり       | 中等度        | 回復  |     |
| 悪心*          |            |     |    |        | 73                         | 多分関連あり         | 軽度         | 回復  |     |
| AST 増加*      |            |     |    |        | 13                         | 多分関連あり         | 軽度         | 回復  |     |
| ALT 増加*      |            |     |    |        | 13                         | 多分関連あり         | 軽度         | 回復  |     |
| γ-GTP 増加*    |            |     |    |        | 13                         | 多分関連あり         | 軽度         | 回復  |     |
| 血中乳酸脱水素酵素増加* |            |     |    |        | 13                         | 多分関連あり         | 軽度         | 回復  |     |
| 臨床検査異常*      |            |     |    |        | 13                         | 多分関連あり         | 軽度         | 回復  |     |
| [REDACTED]   | 日本人        | 女性  | 49 | 鼻咽頭炎   | 85                         | 関連なし           | 軽度         | 回復  |     |
|              |            |     |    | 下痢     | 2                          | 多分関連あり         | 軽度         | 回復  |     |
|              |            |     |    | 頭痛     | 97                         | 関連なし           | 軽度         | 回復  |     |
|              |            |     |    | 肝機能異常* | 57                         | 明らかに関連あり       | 軽度         | 回復  |     |
| [REDACTED]   | 日本人        | 男性  | 54 | 下痢*    | 2                          | 多分関連あり         | 軽度         | 回復  |     |
| [REDACTED]   | 日本人        | 男性  | 69 | 腹痛*    | 27                         | 関連あるかもしれない     | 軽度         | 回復  |     |
| [REDACTED]   | 日本人        | 男性  | 51 | 脂肪便*   | 2                          | 多分関連あり         | 中等度        | 回復  |     |

事象名：MedDRA/J Ver.10.1、総括報告書（5.3.5.1-1）表 12.3.2、症例一覧表（5.3.7.1-1）から引用

有害事象発現日は、投与開始日を1日として起算

\*治験薬投与中止に至った有害事象

プラセボ群及び120 mg 群の各1例（被験者識別コード：[REDACTED]及び[REDACTED]）の有害事象名「臨床検査異常」は、MedDRA/J Ver. 15.0で「サイトケラチン18増加」へ基本語が変更された。

15 mg 群の1例（被験者識別コード：[REDACTED]）は治験薬投与完了後に有害事象が発現した。

1) アラニン・アミノトランスフェラーゼをALT、アスパラギン酸アミノトランスフェラーゼをAST、γ-グルタミルトランスフェラーゼをγ-GTPと表記した。

表 2.7.4.7-付録-9 治験薬投与中止（治験中止）に至った有害事象の一覧（CCT-002 試験）（1/3）

| 投与群        | 被験者識別コード   | 人種  | 性別   | 年齢      | 有害事象名<br>基本語 <sup>1)</sup> | 発現日<br>(投与後日数) | 因果関係       | 程度  | 転帰  |
|------------|------------|-----|------|---------|----------------------------|----------------|------------|-----|-----|
| プラセボ群      | [REDACTED] | 日本人 | 男性   | 64      | 鼻咽頭炎                       | 11             | 関連なし       | 軽度  | 回復  |
|            |            |     |      |         | うっ血性心不全                    | 27             | 関連なし       | 中等度 | 回復  |
|            |            |     |      |         | 肺炎*                        | 61             | 関連なし       | 高度  | 回復  |
|            |            |     |      |         | 関節リウマチ                     | 85             | 関連なし       | 軽度  | 未回復 |
|            |            |     |      |         | 乳管内乳頭腫*                    | 13             | 関連なし       | 軽度  | 未回復 |
| 120 mg 群   | [REDACTED] | 日本人 | 男性   | 67      | 口腔咽頭痛                      | 32             | 関連なし       | 軽度  | 回復  |
|            |            |     |      |         | 心房細動*                      | 195            | 関連なし       | 中等度 | 回復  |
|            | [REDACTED] | 日本人 | 男性   | 63      | 糖尿病*                       | 1              | —          | 中等度 | 軽快  |
|            |            |     |      |         | そう痒症                       | 43             | 関連なし       | 軽度  | 回復  |
|            | [REDACTED] | 日本人 | 女性   | 58      | 脂肪便*                       | 2              | 関連あるかもしれない | 中等度 | 回復  |
|            |            |     |      |         | [REDACTED]                 | 日本人            | 女性         | 42  | 下痢  |
|            | 子宮平滑筋腫*    | 169 | 関連なし | 中等度     |                            |                |            |     | 未回復 |
|            | [REDACTED] | 日本人 | 女性   | 63      | 下痢                         | 2              | 多分関連あり     | 軽度  | 回復  |
|            |            |     |      |         | 口腔の良性新生物                   | 37             | 関連なし       | 中等度 | 回復  |
|            |            |     |      |         | 上気道の炎症                     | 89             | 関連なし       | 軽度  | 回復  |
|            |            |     |      |         | 上気道の炎症                     | 246            | 関連なし       | 軽度  | 回復  |
|            |            |     |      |         | 心不全*                       | 281            | 関連なし       | 中等度 | 未回復 |
|            | [REDACTED] | 日本人 | 女性   | 57      | 糖尿病*                       | -56            | —          | 中等度 | 軽快  |
| 浮動性めまい     |            |     |      |         | -24                        | —              | 軽度         | 回復  |     |
| 不安定狭心症     |            |     |      |         | 5                          | 関連なし           | 中等度        | 回復  |     |
| 鼻咽頭炎       |            |     |      |         | 12                         | 関連なし           | 軽度         | 回復  |     |
| [REDACTED] | 日本人        | 男性  | 58   | 下痢*     | 2                          | 関連あるかもしれない     | 軽度         | 回復  |     |
|            |            |     |      | 湿疹      | 94                         | 関連なし           | 軽度         | 回復  |     |
| [REDACTED] | 日本人        | 女性  | 65   | 水腎症*    | 17                         | 関連なし           | 中等度        | 回復  |     |
| [REDACTED] | 日本人        | 女性  | 68   | 脂肪便     | 2                          | 多分関連あり         | 軽度         | 回復  |     |
|            |            |     |      | 弾発指     | 219                        | 関連なし           | 軽度         | 軽快  |     |
|            |            |     |      | 感染*     | 229                        | 多分関連あり         | 中等度        | 回復  |     |
|            |            |     |      | 舌炎      | 248                        | 関連なし           | 軽度         | 回復  |     |
|            |            |     |      | 糖尿病性網膜症 | 253                        | 関連なし           | 軽度         | 未回復 |     |
|            |            |     |      | 白内障     | 253                        | 関連なし           | 軽度         | 未回復 |     |

事象名：MedDRA/J Ver.13.0、総括報告書（5.3.5.1-2）表 12.3.2、症例一覧表（5.3.7.1-2）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象、—：Pretreatment event のため該当せず

プラセボ群及び120 mg 群の各1例（被験者識別コード：[REDACTED] 及び [REDACTED]）は、Pretreatment event の悪化により治験薬の投与を中止（治験を中止）した。

1) クレアチンホスホキナーゼをCPK、アラニン・アミノトランスフェラーゼをALT、アスパラギン酸アミノトランスフェラーゼをASTと表記

表 2.7.4.7-付録-9 治験薬投与中止（治験中止）に至った有害事象の一覧（CCT-002 試験）（2/3）

| 投与群      | 被験者識別コード | 人種  | 性別 | 年齢    | 有害事象名<br>基本語 <sup>1)</sup> | 発現日<br>(投与後日数) | 因果関係       | 程度  | 転帰  |
|----------|----------|-----|----|-------|----------------------------|----------------|------------|-----|-----|
| 120 mg 群 | ■        | 日本人 | 男性 | 54    | 下痢*                        | 2              | 多分関連あり     | 中等度 | 回復  |
|          |          |     |    |       | 血中 CPK 増加                  | 141            | 関連なし       | 軽度  | 回復  |
|          | ■        | 日本人 | 男性 | 42    | 脂肪便                        | 3              | 関連あるかもしれない | 軽度  | 回復  |
|          |          |     |    |       | 口内炎                        | 159            | 関連なし       | 軽度  | 回復  |
|          |          |     |    |       | ALT 増加*                    | 169            | 関連なし       | 軽度  | 回復  |
|          |          |     |    |       | AST 増加*                    | 169            | 関連なし       | 軽度  | 回復  |
|          | ■        | 日本人 | 男性 | 51    | 鼻咽頭炎                       | 28             | 関連なし       | 軽度  | 回復  |
|          |          |     |    |       | 痔核                         | 132            | 関連なし       | 軽度  | 回復  |
|          |          |     |    |       | 胸部不快感                      | 281            | 関連なし       | 軽度  | 回復  |
|          |          |     |    |       | 糖尿病性ニューロパチー                | 281            | 関連なし       | 軽度  | 未回復 |
|          |          |     |    |       | 甲状腺腫                       | 309            | 関連なし       | 軽度  | 未回復 |
|          |          |     |    |       | 血中尿酸増加                     | 337            | 関連なし       | 軽度  | 回復  |
|          |          |     |    |       | 尿路結石*                      | 352            | 関連なし       | 中等度 | 回復  |
|          |          |     |    |       | 水腎症*                       | 352            | 関連なし       | 中等度 | 回復  |
|          |          |     |    |       | 左室肥大                       | 362            | 関連なし       | 軽度  | 未回復 |
|          | ■        | 日本人 | 男性 | 38    | 鼻咽頭炎                       | 113            | 関連なし       | 軽度  | 回復  |
|          |          |     |    |       | 上腹部痛                       | 253            | 関連なし       | 軽度  | 回復  |
|          |          |     |    |       | 肝機能異常*                     | 260            | 関連なし       | 中等度 | 軽快  |
|          |          |     |    |       | 胆石症                        | 260            | 関連なし       | 軽度  | 未回復 |
|          | ■        | 日本人 | 男性 | 41    | 下痢                         | 4              | 関連あるかもしれない | 中等度 | 回復  |
|          |          |     |    |       | 鼻咽頭炎                       | 14             | 関連なし       | 軽度  | 回復  |
| 頭痛       |          |     |    |       | 26                         | 関連なし           | 軽度         | 回復  |     |
| メレナ*     |          |     |    |       | 33                         | 関連あるかもしれない     | 中等度        | 回復  |     |
| 痔核       |          |     |    |       | 35                         | 関連なし           | 軽度         | 未回復 |     |
| 結腸ポリープ   |          |     |    |       | 35                         | 関連なし           | 軽度         | 回復  |     |
| ■        | 日本人      | 男性  | 60 | 脂肪便   | 6                          | 多分関連あり         | 軽度         | 回復  |     |
|          |          |     |    | 血尿    | 14                         | 関連あるかもしれない     | 軽度         | 回復  |     |
|          |          |     |    | 左室肥大  | 15                         | 関連なし           | 軽度         | 未回復 |     |
|          |          |     |    | 心房頻脈* | 139                        | 関連あるかもしれない     | 中等度        | 回復  |     |

事象名：MedDRA/J Ver.13.0、総括報告書（5.3.5.1-2）表 12.3.2、症例一覧表（5.3.7.1-2）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象、－：Pretreatment event のため該当せず

プラセボ群及び120 mg 群の各1例（被験者識別コード：■及び■）は、Pretreatment event の悪化により治験薬の投与を中止（治験を中止）した。

1) クレアチンホスホキナーゼを CPK、アラニン・アミノトランスフェラーゼを ALT、アスパラギン酸アミノトランスフェラーゼを AST と表記

表 2.7.4.7-付録-9 治験薬投与中止（治験中止）に至った有害事象の一覧（CCT-002 試験）（3/3）

| 投与群      | 被験者識別コード | 人種  | 性別 | 年齢      | 有害事象名<br>基本語 <sup>1)</sup> | 発現日<br>(投与後日数) | 因果関係       | 程度  | 転帰  |
|----------|----------|-----|----|---------|----------------------------|----------------|------------|-----|-----|
| 120 mg 群 | ■■■■■    | 日本人 | 男性 | 67      | 脂肪便*                       | 16             | 多分関連あり     | 中等度 | 回復  |
|          |          |     |    |         | 鼻咽頭炎                       | 46             | 関連なし       | 軽度  | 回復  |
|          |          |     |    |         | 腎結石症                       | 50             | 関連なし       | 軽度  | 未回復 |
|          |          |     |    |         | 鼻咽頭炎                       | 102            | 関連なし       | 軽度  | 回復  |
|          |          |     |    |         | 良性前立腺肥大症                   | 169            | 関連なし       | 軽度  | 未回復 |
|          | ■■■■■    | 日本人 | 女性 | 58      | 脂肪便                        | 2              | 多分関連あり     | 軽度  | 回復  |
|          |          |     |    |         | アレルギー性結膜炎                  | 54             | 関連なし       | 軽度  | 回復  |
|          |          |     |    |         | 視力低下                       | 144            | 関連なし       | 軽度  | 回復  |
|          |          |     |    |         | 浮動性めまい                     | 202            | 関連あるかもしれない | 軽度  | 回復  |
|          |          |     |    |         | 甲状腺癌*                      | 203            | 関連なし       | 中等度 | 軽快  |
|          | ■■■■■    | 日本人 | 男性 | 49      | 下痢*                        | 2              | 多分関連あり     | 軽度  | 回復  |
|          |          |     |    |         | 脂肪便*                       | 2              | 多分関連あり     | 軽度  | 回復  |
|          | ■■■■■    | 日本人 | 女性 | 62      | 中毒性皮疹*                     | 12             | 多分関連あり     | 軽度  | 回復  |
|          | ■■■■■    | 日本人 | 男性 | 52      | 下痢                         | 1              | 多分関連あり     | 軽度  | 回復  |
| 肝機能異常*   |          |     |    |         | 15                         | 多分関連あり         | 中等度        | 回復  |     |
| 自律神経失調   |          |     |    |         | 126                        | 関連なし           | 軽度         | 回復  |     |
| ■■■■■    | 日本人      | 男性  | 60 | 脂肪便     | 1                          | 多分関連あり         | 軽度         | 回復  |     |
| ■■■■■    | 日本人      | 男性  | 62 | 急性心筋梗塞* | 110                        | 関連なし           | 高度         | 回復  |     |
|          |          |     |    | 下痢*     | 22                         | 多分関連あり         | 軽度         | 回復  |     |
|          |          |     |    | 脂肪便*    | 80                         | 多分関連あり         | 軽度         | 回復  |     |

事象名：MedDRA/J Ver.13.0、総括報告書（5.3.5.1-2）表 12.3.2、症例一覧表（5.3.7.1-2）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象、－：Pretreatment eventのため該当せず

プラセボ群及び120 mg 群の各1例（被験者識別コード：■■■■■及び■■■■■）は、Pretreatment eventの悪化により治験薬の投与を中止（治験を中止）した。

1) クレアチンホスホキナーゼをCPK、アラニン・アミノトランスフェラーゼをALT、アスパラギン酸アミノトランスフェラーゼをASTと表記

表 2.7.4.7-付録-10 治験薬投与中止（治験中止）に至った有害事象の一覧（OCT-001 試験）

| 投与群      | 被験者識別<br>コード | 人種  | 性別 | 年齢 | 有害事象名<br>基本語 | 発現日<br>(投与後日数) | 因果関係       | 程度  | 転帰 |
|----------|--------------|-----|----|----|--------------|----------------|------------|-----|----|
| 120 mg 群 | ■            | 日本人 | 男性 | 44 | 下痢           | 2              | 多分関連あり     | 軽度  | 回復 |
|          |              |     |    |    | 腹痛           | 84             | 関連なし       | 軽度  | 回復 |
|          |              |     |    |    | 糖尿病*         | 115            | 関連あるかもしれない | 中等度 | 軽快 |
|          | ■            | 日本人 | 女性 | 74 | 下痢*          | 11             | 多分関連あり     | 中等度 | 回復 |
|          | ■            | 日本人 | 男性 | 54 | 下痢           | 6              | 関連あるかもしれない | 軽度  | 回復 |
|          |              |     |    |    | 脂肪便          | 6              | 関連あるかもしれない | 軽度  | 回復 |
|          |              |     |    |    | 神経鞘腫*        | 105            | 関連なし       | 軽度  | 回復 |
|          | ■            | 日本人 | 女性 | 47 | 胆石症*         | 29             | 関連なし       | 軽度  | 軽快 |
|          |              |     |    |    | 鼻咽頭炎         | 39             | 関連なし       | 軽度  | 回復 |
|          | ■            | 日本人 | 女性 | 58 | ラクナ梗塞*       | 48             | 関連なし       | 中等度 | 回復 |

事象名：MedDRA/J Ver.13.0、総括報告書（5.3.5.2-1）表 12.3.2、症例一覧表（5.3.7.1-3）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象

表 2.7.4.7-付録-11 治験薬投与中止（治験中止）に至った有害事象の一覧（OCT-002 試験）（1/8）

| 投与群      | 被験者識別コード | 人種  | 性別 | 年齢     | 有害事象名<br>基本語 | 発現日<br>(投与後日数) | 因果関係 | 程度  | 転帰  |
|----------|----------|-----|----|--------|--------------|----------------|------|-----|-----|
| 120 mg 群 | ■        | 日本人 | 女性 | 52     | 下痢*          | 68             | 関連あり | 軽度  | 回復  |
|          | ■        | 日本人 | 女性 | 61     | 消化不良         | 43             | 関連なし | 軽度  | 回復  |
|          |          |     |    |        | 嘔吐           | 43             | 関連なし | 軽度  | 回復  |
|          |          |     |    |        | 感音性難聴        | 53             | 関連なし | 軽度  | 軽快  |
|          |          |     |    |        | 転移性結腸癌*      | 93             | 関連なし | 中等度 | 回復  |
|          | ■        | 日本人 | 男性 | 72     | 上気道の炎症       | 33             | 関連なし | 軽度  | 回復  |
|          |          |     |    |        | 第二度房室ブロック*   | 59             | 関連なし | 中等度 | 回復  |
|          |          |     |    |        | 結腸ポリープ       | 73             | 関連なし | 軽度  | 回復  |
|          |          |     |    |        | 乾皮症          | 104            | 関連なし | 軽度  | 回復  |
|          |          |     |    |        | 鼻咽頭炎         | 141            | 関連なし | 軽度  | 回復  |
|          | ■        | 日本人 | 男性 | 67     | 転移性肺癌*       | 331            | 関連なし | 高度  | 未回復 |
|          | ■        | 日本人 | 男性 | 53     | 蕁麻疹*         | 1              | 関連なし | 軽度  | 回復  |
|          | ■        | 日本人 | 男性 | 75     | 便秘*          | 1              | 関連あり | 軽度  | 回復  |
|          |          |     |    |        | 下痢*          | 3              | 関連あり | 軽度  | 回復  |
|          | ■        | 日本人 | 男性 | 43     | 脂肪便*         | 6              | 関連あり | 軽度  | 回復  |
|          |          |     |    |        | 便秘*          | 27             | 関連あり | 軽度  | 回復  |
| ■        | 日本人      | 男性  | 45 | 蕁麻疹*   | 43           | 関連あり           | 軽度   | 回復  |     |
| ■        | 日本人      | 男性  | 63 | 発疹*    | 10           | 関連あり           | 軽度   | 回復  |     |
| ■        | 日本人      | 女性  | 75 | 末梢性浮腫* | 9            | 関連なし           | 軽度   | 回復  |     |
| ■        | 日本人      | 男性  | 79 | 転倒     | 6            | 関連なし           | 軽度   | 回復  |     |
|          |          |     |    | 挫傷     | 6            | 関連なし           | 軽度   | 回復  |     |
|          |          |     |    | 擦過傷    | 6            | 関連なし           | 軽度   | 回復  |     |
|          |          |     |    | 接触性皮膚炎 | 26           | 関連なし           | 軽度   | 未回復 |     |
|          |          |     |    | 心血管障害* | 172          | 関連なし           | 高度   | 死亡  |     |

事象名：MedDRA/J Ver.15.0、総括報告書（5.3.5.2-2）表 12.3.3、症例一覧表（5.3.7.1-4）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象

1例（被験者識別コード：■）は、Pretreatment eventにより治験薬の投与を中止（治験を中止）した。

表 2.7.4.7-付録-11 治験薬投与中止（治験中止）に至った有害事象の一覧（OCT-002 試験）（2/8）

| 投与群                     | 被験者識別コード | 人種  | 性別   | 年齢 | 有害事象名<br>基本語 | 発現日<br>(投与後日数) | 因果関係 | 程度  | 転帰                |
|-------------------------|----------|-----|------|----|--------------|----------------|------|-----|-------------------|
| 120 mg 群                | ■        | 日本人 | 男性   | 81 | 脂肪便          | 2              | 関連あり | 軽度  | 回復                |
|                         |          |     |      |    | ざ瘡           | 11             | 関連なし | 軽度  | 回復                |
|                         |          |     |      |    | 変形性脊椎症       | 42             | 関連なし | 軽度  | 未回復               |
|                         |          |     |      |    | 歯周炎          | 94             | 関連なし | 軽度  | 回復                |
|                         |          |     |      |    | 急性心不全*       | 277            | 関連なし | 高度  | 回復                |
|                         |          |     |      |    | うつ血性心不全*     | 277            | 関連なし | 高度  | 軽快                |
|                         |          |     |      |    | 大動脈弁狭窄*      | 277            | 関連なし | 高度  | 回復                |
|                         | ■        | 日本人 | 男性   | 62 | 下痢*          | 2              | 関連あり | 中等度 | 回復                |
|                         | ■        | 日本人 | 男性   | 54 | 脂肪便          | 2              | 関連あり | 軽度  | 回復                |
|                         |          |     |      |    | 不眠症          | 43             | 関連なし | 軽度  | 回復                |
|                         |          |     |      |    | 末梢性浮腫        | 78             | 関連なし | 軽度  | 回復                |
|                         |          |     |      |    | 2型糖尿病*       | 253            | 関連なし | 軽度  | 回復                |
|                         |          |     |      |    | 脳梗塞*         | 262            | 関連なし | 中等度 | 後遺症 <sup>1)</sup> |
|                         | 糖尿病網膜症   | 295 | 関連なし | 軽度 | 軽快           |                |      |     |                   |
|                         | ■        | 日本人 | 男性   | 63 | 低血糖症         | 5              | 関連なし | 軽度  | 回復                |
|                         |          |     |      |    | 脂肪便          | 14             | 関連あり | 軽度  | 回復                |
|                         |          |     |      |    | 齲歯           | 83             | 関連なし | 軽度  | 回復                |
|                         |          |     |      |    | 出血性腸憩室*      | 111            | 関連あり | 軽度  | 回復                |
|                         | ■        | 日本人 | 男性   | 37 | 脂肪便          | 3              | 関連あり | 軽度  | 回復                |
|                         |          |     |      |    | 頭痛           | 66             | 関連なし | 軽度  | 回復                |
| ALT 増加* <sup>2)</sup>   |          |     |      |    | 99           | 関連なし           | 軽度   | 未回復 |                   |
| AST 増加* <sup>2)</sup>   |          |     |      |    | 99           | 関連なし           | 軽度   | 未回復 |                   |
| γ-GTP 増加* <sup>2)</sup> |          |     |      |    | 99           | 関連なし           | 軽度   | 未回復 |                   |
| 鼻咽頭炎                    |          |     |      |    | 106          | 関連なし           | 軽度   | 回復  |                   |
| 鼻咽頭炎                    |          |     |      |    | 195          | 関連なし           | 軽度   | 回復  |                   |
| 頭痛                      |          |     |      |    | 218          | 関連なし           | 軽度   | 未回復 |                   |
| うつ病                     |          |     |      |    | 246          | 関連なし           | 軽度   | 未回復 |                   |
| 鼻咽頭炎                    |          |     |      |    | 252          | 関連なし           | 軽度   | 回復  |                   |

事象名：MedDRA/J Ver.15.0、総括報告書（5.3.5.2-2）表 12.3.3、症例一覧表（5.3.7.1-4）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象

1例（被験者識別コード：■）は、Pretreatment eventにより治験薬の投与を中止（治験を中止）した。

1) 回復したが後遺症あり

2) アラニン・アミノトランスフェラーゼを ALT、アスパラギン酸アミノトランスフェラーゼを AST、γ-グルタミルトランスフェラーゼを γ-GTP と表記した。

表 2.7.4.7-付録-11 治験薬投与中止（治験中止）に至った有害事象の一覧（OCT-002 試験）（3/8）

| 投与群      | 被験者識別コード | 人種  | 性別 | 年齢      | 有害事象名<br>基本語 | 発現日<br>(投与後日数) | 因果関係 | 程度  | 転帰  |
|----------|----------|-----|----|---------|--------------|----------------|------|-----|-----|
| 120 mg 群 | ■        | 日本人 | 男性 | 65      | 下痢           | 8              | 関連あり | 軽度  | 回復  |
|          |          |     |    |         | 下痢*          | 10             | 関連あり | 軽度  | 回復  |
|          | ■        | 日本人 | 女性 | 69      | 脂肪便*         | 15             | 関連あり | 軽度  | 回復  |
|          |          |     |    |         | 低血糖症         | 47             | 関連なし | 軽度  | 回復  |
|          |          |     |    |         | 転倒           | 72             | 関連なし | 軽度  | 回復  |
|          |          |     |    |         | 脊椎圧迫骨折*      | 72             | 関連なし | 中等度 | 回復  |
|          |          |     |    |         | 上腕骨骨折*       | 72             | 関連なし | 中等度 | 回復  |
|          |          |     |    |         | 不眠症          | 77             | 関連なし | 軽度  | 回復  |
|          | ■        | 日本人 | 男性 | 72      | 脂肪便          | 8              | 関連あり | 軽度  | 回復  |
|          |          |     |    |         | 歯周病          | 214            | 関連なし | 軽度  | 回復  |
|          |          |     |    |         | 直腸癌*         | 249            | 関連なし | 中等度 | 回復  |
|          | ■        | 日本人 | 女性 | 71      | 湿疹*          | 5              | 関連なし | 軽度  | 回復  |
|          |          |     |    |         | 脂肪便          | 8              | 関連あり | 軽度  | 回復  |
|          | ■        | 日本人 | 男性 | 52      | 脂肪便          | 4              | 関連あり | 軽度  | 回復  |
|          |          |     |    |         | 腎結石症*        | 105            | 関連なし | 中等度 | 未回復 |
|          |          |     |    |         | ニコチン依存       | 193            | 関連なし | 軽度  | 未回復 |
|          | ■        | 日本人 | 女性 | 34      | 脂肪便          | 3              | 関連あり | 軽度  | 回復  |
|          |          |     |    |         | 歯髄炎          | 9              | 関連なし | 中等度 | 回復  |
|          |          |     |    |         | 第7脳神経麻痺      | 170            | 関連なし | 中等度 | 回復  |
|          |          |     |    |         | 糖尿病          | 173            | 関連なし | 軽度  | 軽快  |
| 腹部不快感    |          |     |    |         | 194          | 関連なし           | 中等度  | 回復  |     |
| 肝障害*     |          |     |    |         | 197          | 関連なし           | 軽度   | 回復  |     |
| ■        | 日本人      | 女性  | 64 | 肺炎      | 25           | 関連なし           | 軽度   | 回復  |     |
|          |          |     |    | 膀胱炎     | 36           | 関連なし           | 軽度   | 未回復 |     |
|          |          |     |    | 鼻咽頭炎    | 138          | 関連なし           | 軽度   | 回復  |     |
|          |          |     |    | 下痢      | 141          | 関連あり           | 軽度   | 未回復 |     |
|          |          |     |    | 交通事故    | 175          | 関連なし           | 高度   | 回復  |     |
|          |          |     |    | 鎖骨骨折*   | 175          | 関連なし           | 高度   | 軽快  |     |
|          |          |     |    | 脊椎圧迫骨折* | 175          | 関連なし           | 高度   | 軽快  |     |

事象名：MedDRA/J Ver.15.0、総括報告書（5.3.5.2-2）表 12.3.3、症例一覧表（5.3.7.1-4）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象

1例（被験者識別コード：■）は、Pretreatment eventにより治験薬の投与を中止（治験を中止）した。



表 2.7.4.7-付録-11 治験薬投与中止（治験中止）に至った有害事象の一覧（OCT-002 試験）（4/8）

| 投与群      | 被験者識別コード   | 人種   | 性別 | 年齢 | 有害事象名<br>基本語 | 発現日<br>(投与後日数) | 因果関係 | 程度  | 転帰  |
|----------|------------|------|----|----|--------------|----------------|------|-----|-----|
| 120 mg 群 | [REDACTED] | 日本人  | 女性 | 64 | 脂肪便          | 5              | 関連あり | 軽度  | 回復  |
|          |            |      |    |    | 十二指腸潰瘍       | 29             | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 椎間板突出        | 49             | 関連なし | 中等度 | 回復  |
|          |            |      |    |    | 上気道の炎症       | 69             | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 上腹部痛         | 71             | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 便秘           | 79             | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 下痢           | 119            | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 筋痙縮          | 161            | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 甲状腺腫         | 260            | 関連なし | 軽度  | 未回復 |
|          |            |      |    |    | 縦隔新生物        | 260            | 関連なし | 軽度  | 未回復 |
|          |            |      |    |    | 肝嚢胞          | 260            | 関連なし | 軽度  | 未回復 |
|          |            |      |    |    | 脊柱管狭窄症       | 260            | 関連なし | 軽度  | 未回復 |
|          |            |      |    |    | 骨盤骨折*        | 272            | 関連なし | 高度  | 回復  |
|          |            |      |    |    | 擦過傷          | 272            | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 交通事故         | 272            | 関連なし | 高度  | 回復  |
|          |            |      |    |    | 褥瘡性潰瘍        | 275            | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 皮膚剥脱         | 279            | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 回転性めまい       | 281            | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 下痢           | 282            | 関連なし | 軽度  | 回復  |
|          |            |      |    |    | 腹部不快感        | 283            | 関連なし | 軽度  | 回復  |
| 上腹部痛     | 290        | 関連なし | 軽度 | 回復 |              |                |      |     |     |
| 120 mg 群 | [REDACTED] | 日本人  | 男性 | 37 | 鼓腸           | 2              | 関連あり | 軽度  | 回復  |
|          |            |      |    |    | 脂肪便*         | 2              | 関連あり | 軽度  | 回復  |
|          |            |      |    |    | 血中尿酸増加       | 29             | 関連なし | 軽度  | 軽快  |

事象名：MedDRA/J Ver.15.0、総括報告書（5.3.5.2-2）表 12.3.3、症例一覧表（5.3.7.1-4）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象

1例（被験者識別コード：[REDACTED]）は、Pretreatment eventにより治験薬の投与を中止（治験を中止）した。

表 2.7.4.7-付録-11 治験薬投与中止（治験中止）に至った有害事象の一覧（OCT-002 試験）（5/8）

| 投与群                    | 被験者識別コード   | 人種  | 性別 | 年齢 | 有害事象名<br>基本語 | 発現日<br>(投与後日数) | 因果関係 | 程度  | 転帰  |
|------------------------|------------|-----|----|----|--------------|----------------|------|-----|-----|
| 120 mg 群               | [REDACTED] | 日本人 | 女性 | 65 | 四肢痛          | 2              | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 体位性めまい       | 2              | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 下痢           | 6              | 関連あり | 軽度  | 回復  |
|                        |            |     |    |    | 下痢           | 9              | 関連あり | 軽度  | 回復  |
|                        |            |     |    |    | 脂肪便          | 21             | 関連あり | 軽度  | 回復  |
|                        |            |     |    |    | 四肢痛*         | 113            | 関連なし | 軽度  | 軽快  |
|                        |            |     |    |    | 腹部不快感        | 166            | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 眼乾燥          | 205            | 関連なし | 軽度  | 回復  |
|                        | [REDACTED] | 日本人 | 男性 | 70 | 悪心           | 1              | 関連あり | 軽度  | 回復  |
|                        |            |     |    |    | 下痢           | 1              | 関連あり | 軽度  | 回復  |
|                        |            |     |    |    | 嘔吐           | 4              | 関連あり | 軽度  | 回復  |
|                        |            |     |    |    | 嘔吐*          | 8              | 関連あり | 軽度  | 回復  |
|                        | [REDACTED] | 日本人 | 男性 | 41 | 下痢*          | 1              | 関連あり | 中等度 | 回復  |
|                        |            | 日本人 | 女性 | 64 | 爪真菌症         | 29             | 関連なし | 軽度  | 未回復 |
|                        | [REDACTED] | 日本人 | 女性 | 64 | 鼻咽頭炎         | 58             | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 発熱*          | 124            | 関連なし | 高度  | 回復  |
|                        |            |     |    |    | 不眠症          | 128            | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 背部痛          | 128            | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 幻覚           | 130            | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 眼精疲労         | 42             | 関連なし | 軽度  | 回復  |
|                        | [REDACTED] | 日本人 | 女性 | 63 | 上気道の炎症       | 54             | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 上気道の炎症       | 107            | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 子宮癌*         | 180            | 関連なし | 高度  | 回復  |
|                        |            |     |    |    | 下痢*          | 2              | 関連あり | 中等度 | 回復  |
|                        | [REDACTED] | 日本人 | 男性 | 60 | 鼻咽頭炎         | 13             | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 関節周囲炎        | 20             | 関連なし | 軽度  | 回復  |
|                        |            |     |    |    | 脂肪便          | 8              | 関連あり | 軽度  | 回復  |
|                        | [REDACTED] | 日本人 | 男性 | 43 | 毛巣嚢胞         | 32             | 関連なし | 中等度 | 回復  |
| 肺結核*                   |            |     |    |    | 256          | 関連なし           | 中等度  | 軽快  |     |
| γ-GTP 増加 <sup>1)</sup> |            |     |    |    | 319          | 関連なし           | 軽度   | 回復  |     |
| 血中尿酸増加                 |            |     |    |    | 319          | 関連なし           | 軽度   | 回復  |     |

事象名：MedDRA/J Ver.15.0、総括報告書（5.3.5.2-2）表 12.3.3、症例一覧表（5.3.7.1-4）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象

1例（被験者識別コード：[REDACTED]）は、Pretreatment eventにより治験薬の投与を中止（治験を中止）した。

1) γ-グルタミルトランスフェラーゼをγ-GTPと表記した。

表 2.7.4.7-付録-11 治験薬投与中止（治験中止）に至った有害事象の一覧（OCT-002 試験）（6/8）

| 投与群      | 被験者識別コード | 人種  | 性別   | 年齢 | 有害事象名<br>基本語 | 発現日<br>(投与後日数) | 因果関係 | 程度  | 転帰  |
|----------|----------|-----|------|----|--------------|----------------|------|-----|-----|
| 120 mg 群 | ██████   | 日本人 | 男性   | 69 | 気管軟化症*       | 80             | 関連なし | 高度  | 未回復 |
|          |          |     |      |    | 肺炎*          | 80             | 関連なし | 高度  | 回復  |
|          |          |     |      |    | 肺水腫*         | 86             | 関連なし | 高度  | 軽快  |
|          |          |     |      |    | 心不全*         | 86             | 関連なし | 高度  | 軽快  |
|          | ██████   | 日本人 | 女性   | 75 | 口内炎          | 7              | 関連なし | 軽度  | 回復  |
|          |          |     |      |    | 血中カリウム増加     | 71             | 関連なし | 軽度  | 回復  |
|          |          |     |      |    | 神経痛          | 84             | 関連なし | 軽度  | 回復  |
|          |          |     |      |    | 不眠症          | 221            | 関連なし | 軽度  | 回復  |
|          |          |     |      |    | 鼻咽頭炎         | 221            | 関連なし | 軽度  | 回復  |
|          |          |     |      |    | 高血圧性脳症       | 222            | 関連なし | 中等度 | 回復  |
|          |          |     |      |    | 出血性胃炎        | 222            | 関連なし | 軽度  | 回復  |
|          |          |     |      |    | 高血圧          | 222            | 関連なし | 中等度 | 回復  |
|          |          |     |      |    | 睡眠時無呼吸症候群    | 231            | 関連なし | 中等度 | 未回復 |
|          |          |     |      |    | 赤血球数減少       | 239            | 関連なし | 軽度  | 未回復 |
|          |          |     |      |    | 不眠症          | 251            | 関連なし | 中等度 | 回復  |
|          |          |     |      |    | 糖尿病網膜症       | 251            | 関連なし | 軽度  | 未回復 |
|          |          |     |      |    | 嘔吐           | 253            | 関連なし | 軽度  | 回復  |
|          |          |     |      |    | 全身性浮腫*       | 254            | 関連なし | 軽度  | 軽快  |
|          |          |     |      |    | 下痢           | 271            | 関連あり | 軽度  | 回復  |
|          |          |     |      |    | 高血圧          | 291            | 関連なし | 中等度 | 回復  |
|          |          |     |      |    | 脊椎すべり症       | 294            | 関連なし | 軽度  | 未回復 |
|          |          |     |      |    | 血中カリウム増加     | 299            | 関連なし | 軽度  | 軽快  |
|          |          |     |      |    | 擦過傷          | 299            | 関連なし | 軽度  | 回復  |
|          |          |     |      |    | 弾発指          | 299            | 関連なし | 軽度  | 未回復 |
|          | 挫傷       | 299 | 関連なし | 軽度 | 回復           |                |      |     |     |
|          | 転倒       | 299 | 関連なし | 軽度 | 回復           |                |      |     |     |
|          | ██████   | 日本人 | 女性   | 68 | 上気道の炎症       | 29             | 関連なし | 軽度  | 回復  |
|          |          |     |      |    | 下痢*          | 31             | 関連あり | 中等度 | 回復  |
|          |          |     |      |    | 血圧上昇         | 86             | 関連なし | 軽度  | 回復  |

事象名：MedDRA/J Ver.15.0、総括報告書（5.3.5.2-2）表 12.3.3、症例一覧表（5.3.7.1-4）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象

1例（被験者識別コード：██████）は、Pretreatment eventにより治験薬の投与を中止（治験を中止）した。

表 2.7.4.7-付録-11 治験薬投与中止（治験中止）に至った有害事象の一覧（OCT-002 試験）（7/8）

| 投与群      | 被験者識別コード | 人種  | 性別 | 年齢           | 有害事象名<br>基本語 | 発現日<br>(投与後日数) | 因果関係 | 程度  | 転帰  |
|----------|----------|-----|----|--------------|--------------|----------------|------|-----|-----|
| 120 mg 群 | ██████   | 日本人 | 女性 | 73           | 肛門そう痒症       | 2              | 関連あり | 軽度  | 回復  |
|          |          |     |    |              | 胃腸音異常        | 2              | 関連あり | 軽度  | 回復  |
|          |          |     |    |              | 腹部膨満*        | 37             | 関連あり | 軽度  | 回復  |
|          | ██████   | 日本人 | 女性 | 62           | 腹痛*          | 4              | 関連あり | 中等度 | 回復  |
|          |          |     |    |              | 下痢*          | 4              | 関連あり | 軽度  | 回復  |
|          | ██████   | 日本人 | 男性 | 49           | 脂肪便*         | 1              | 関連あり | 軽度  | 回復  |
|          | ██████   | 日本人 | 女性 | 75           | 便秘*          | 27             | 関連なし | 軽度  | 回復  |
|          |          |     |    |              | 腸憩室          | 78             | 関連なし | 軽度  | 未回復 |
|          | ██████   | 日本人 | 女性 | 60           | 便秘*          | 20             | 関連あり | 軽度  | 回復  |
|          | ██████   | 日本人 | 男性 | 35           | 肝機能検査異常*     | 36             | 関連なし | 軽度  | 回復  |
|          | ██████   | 日本人 | 女性 | 60           | 湿性咳嗽         | 15             | 関連なし | 軽度  | 回復  |
|          |          |     |    |              | 歯肉炎          | 81             | 関連なし | 軽度  | 回復  |
|          |          |     |    |              | 口腔ヘルペス       | 95             | 関連なし | 軽度  | 回復  |
|          |          |     |    |              | 耳帯状疱疹        | 112            | 関連なし | 軽度  | 回復  |
|          |          |     |    |              | 脂肪便*         | 131            | 関連あり | 軽度  | 回復  |
|          | ██████   | 日本人 | 女性 | 65           | 背部痛          | 77             | 関連なし | 軽度  | 回復  |
|          |          |     |    |              | 転倒           | 109            | 関連なし | 中等度 | 回復  |
|          |          |     |    |              | 大腿骨骨折*       | 109            | 関連なし | 中等度 | 回復  |
|          |          |     |    |              | 肺塞栓症         | 133            | 関連なし | 中等度 | 軽快  |
| ██████   | 日本人      | 男性  | 37 | 深部静脈血栓症      | 133          | 関連なし           | 中等度  | 軽快  |     |
|          |          |     |    | コントロール不良の糖尿病 | 57           | 関連あり           | 軽度   | 回復  |     |
| ██████   | 日本人      | 男性  | 57 | 発熱*          | 224          | 関連あり           | 中等度  | 回復  |     |
|          |          |     |    | 便失禁*         | 131          | 関連あり           | 軽度   | 回復  |     |
| ██████   | 日本人      | 男性  | 71 | 脂肪便*         | 11           | 関連あり           | 軽度   | 回復  |     |
|          |          |     |    | 低血糖症         | 27           | 関連なし           | 軽度   | 回復  |     |
|          |          |     |    | 末梢冷感         | 56           | 関連なし           | 軽度   | 回復  |     |
|          |          |     |    | 筋痙縮          | 91           | 関連なし           | 軽度   | 回復  |     |
|          |          |     |    | 低血糖症         | 139          | 関連なし           | 軽度   | 回復  |     |

事象名：MedDRA/J Ver.15.0、総括報告書（5.3.5.2-2）表 12.3.3、症例一覧表（5.3.7.1-4）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象

1例（被験者識別コード：██████）は、Pretreatment eventにより治験薬の投与を中止（治験を中止）した。

表 2.7.4.7-付録-11 治験薬投与中止（治験中止）に至った有害事象の一覧（OCT-002 試験）（8/8）

| 投与群      | 被験者識別コード | 人種  | 性別 | 年齢 | 有害事象名<br>基本語 | 発現日<br>(投与後日数) | 因果関係 | 程度  | 転帰  |
|----------|----------|-----|----|----|--------------|----------------|------|-----|-----|
| 120 mg 群 | ■        | 日本人 | 女性 | 64 | 下痢*          | 7              | 関連あり | 中等度 | 回復  |
|          | ■        | 日本人 | 女性 | 65 | 便秘           | 2              | 関連あり | 軽度  | 回復  |
|          |          |     |    |    | 鼻咽頭炎         | 35             | 関連なし | 軽度  | 回復  |
|          |          |     |    |    | 子宮平滑筋腫*      | 54             | 関連なし | 中等度 | 回復  |
|          | ■        | 日本人 | 女性 | 56 | 脂肪便          | 2              | 関連あり | 軽度  | 回復  |
|          |          |     |    |    | 下痢           | 2              | 関連あり | 軽度  | 回復  |
|          |          |     |    |    | そう痒症*        | 263            | 関連なし | 軽度  | 未回復 |
|          | ■        | 日本人 | 女性 | 54 | 脂肪便          | 1              | 関連あり | 軽度  | 回復  |
|          |          |     |    |    | 鼻咽頭炎         | 18             | 関連なし | 軽度  | 回復  |
|          |          |     |    |    | 鼻咽頭炎         | 33             | 関連なし | 軽度  | 回復  |
| 乳癌*      |          |     |    |    | 140          | 関連なし           | 高度   | 回復  |     |

事象名：MedDRA/J Ver.15.0、総括報告書（5.3.5.2-2）表 12.3.3、症例一覧表（5.3.7.1-4）から引用

有害事象発現日は、投与開始日を1日として起算

\*：投与中止に至った有害事象

1例（被験者識別コード：■）は、Pretreatment eventにより治験薬の投与を中止（治験を中止）した。

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (1/14)

| Item              | Treatment | Summary Statistics |       |       |       |                      |       |        |       |       |         | One-sample t-test |         |
|-------------------|-----------|--------------------|-------|-------|-------|----------------------|-------|--------|-------|-------|---------|-------------------|---------|
|                   |           | Baseline Mean      | N     | Mean  | SD    | Change from Baseline |       |        |       |       | t-value | p-value           |         |
|                   |           |                    |       |       |       | Min                  | Q1    | Median | Q3    | Max   |         |                   |         |
| Hemoglobin (g/dL) | Week 1    | Placebo            | 14.35 | 39    | -0.23 | 0.489                | -1.6  | -0.50  | -0.20 | 0.10  | 0.7     | -2.8836           | 0.0064  |
|                   |           | 15 mg              | 14.12 | 37    | -0.17 | 0.475                | -1.1  | -0.50  | -0.20 | 0.10  | 0.8     | -2.1785           | 0.0360  |
|                   |           | 30 mg              | 14.25 | 39    | -0.05 | 0.619                | -1.7  | -0.40  | -0.10 | 0.30  | 1.2     | -0.5174           | 0.6079  |
|                   |           | 60 mg              | 14.65 | 37    | -0.23 | 0.580                | -1.6  | -0.50  | -0.10 | 0.20  | 0.8     | -2.3807           | 0.0227  |
|                   |           | 120 mg             | 14.51 | 47    | -0.18 | 0.387                | -1.1  | -0.40  | -0.20 | 0.20  | 0.5     | -3.1295           | 0.0030  |
|                   | Week 2    | Placebo            | 14.89 | 97    | -0.29 | 0.570                | -1.5  | -0.70  | -0.30 | 0.10  | 0.9     | -4.9519           | <0.0001 |
|                   |           | 15 mg              | 14.45 | 89    | -0.13 | 0.496                | -1.3  | -0.50  | -0.10 | 0.20  | 1.0     | -2.5449           | 0.0127  |
|                   |           | 30 mg              | 14.57 | 91    | -0.17 | 0.630                | -3.0  | -0.50  | -0.10 | 0.30  | 1.7     | -2.5460           | 0.0126  |
|                   |           | 60 mg              | 14.74 | 93    | -0.17 | 0.616                | -3.0  | -0.40  | -0.10 | 0.20  | 1.4     | -2.6093           | 0.0106  |
|                   |           | 120 mg             | 14.61 | 88    | -0.30 | 0.530                | -1.4  | -0.70  | -0.30 | 0.10  | 1.3     | -5.2316           | <0.0001 |
|                   | Week 3    | Placebo            | 14.43 | 37    | -0.20 | 0.504                | -1.1  | -0.60  | -0.20 | 0.10  | 1.3     | -2.3805           | 0.0227  |
|                   |           | 15 mg              | 14.17 | 36    | -0.12 | 0.452                | -1.1  | -0.45  | -0.10 | 0.20  | 0.7     | -1.5849           | 0.1220  |
|                   |           | 30 mg              | 14.26 | 38    | -0.22 | 0.747                | -2.2  | -0.70  | -0.10 | 0.20  | 1.8     | -1.8234           | 0.0763  |
|                   |           | 60 mg              | 14.64 | 32    | -0.36 | 0.522                | -1.5  | -0.65  | -0.30 | -0.15 | 1.2     | -3.8922           | 0.0005  |
|                   |           | 120 mg             | 14.50 | 42    | -0.35 | 0.506                | -1.7  | -0.60  | -0.35 | 0.00  | 0.7     | -4.5117           | <0.0001 |
|                   | Week 4    | Placebo            | 14.94 | 96    | -0.18 | 0.541                | -2.1  | -0.50  | -0.20 | 0.10  | 1.2     | -3.3006           | 0.0014  |
|                   |           | 15 mg              | 14.45 | 89    | -0.17 | 0.525                | -1.3  | -0.60  | -0.20 | 0.20  | 0.9     | -2.9676           | 0.0039  |
|                   |           | 30 mg              | 14.60 | 89    | -0.23 | 0.672                | -2.6  | -0.70  | -0.10 | 0.10  | 1.6     | -3.2974           | 0.0014  |
|                   |           | 60 mg              | 14.79 | 89    | -0.21 | 0.569                | -2.5  | -0.50  | -0.20 | 0.20  | 1.1     | -3.5036           | 0.0007  |
|                   |           | 120 mg             | 14.58 | 86    | -0.37 | 0.639                | -2.4  | -0.70  | -0.40 | 0.00  | 1.1     | -5.3699           | <0.0001 |
|                   | Week 6    | Placebo            | 14.57 | 40    | -0.18 | 0.488                | -1.4  | -0.40  | -0.20 | 0.10  | 0.8     | -2.3026           | 0.0267  |
|                   |           | 15 mg              | 14.27 | 40    | -0.25 | 0.612                | -2.4  | -0.50  | -0.10 | 0.10  | 1.1     | -2.5326           | 0.0155  |
|                   |           | 30 mg              | 14.33 | 38    | -0.17 | 0.836                | -2.3  | -0.70  | -0.20 | 0.40  | 1.4     | -1.2219           | 0.2295  |
|                   |           | 60 mg              | 14.81 | 31    | -0.20 | 0.609                | -1.6  | -0.70  | -0.10 | 0.20  | 1.2     | -1.8290           | 0.0774  |
|                   |           | 120 mg             | 14.53 | 47    | -0.32 | 0.645                | -3.3  | -0.60  | -0.30 | 0.00  | 1.5     | -3.4368           | 0.0013  |
|                   | Week 8    | Placebo            | 14.92 | 94    | -0.22 | 0.598                | -1.8  | -0.60  | -0.25 | 0.10  | 1.3     | -3.5199           | 0.0007  |
|                   |           | 15 mg              | 14.47 | 86    | -0.12 | 0.643                | -1.7  | -0.50  | -0.10 | 0.30  | 1.7     | -1.7789           | 0.0788  |
|                   |           | 30 mg              | 14.60 | 89    | -0.17 | 0.646                | -2.2  | -0.60  | -0.20 | 0.20  | 1.4     | -2.5431           | 0.0127  |
|                   |           | 60 mg              | 14.75 | 85    | -0.11 | 0.643                | -1.6  | -0.50  | -0.10 | 0.30  | 1.2     | -1.5868           | 0.1163  |
|                   |           | 120 mg             | 14.61 | 83    | -0.35 | 0.731                | -3.3  | -0.80  | -0.30 | 0.10  | 1.2     | -4.4011           | <0.0001 |
|                   | Week 12   | Placebo            | 14.94 | 95    | -0.27 | 0.654                | -1.7  | -0.60  | -0.30 | 0.10  | 1.2     | -3.9667           | 0.0001  |
|                   |           | 15 mg              | 14.47 | 86    | -0.26 | 0.634                | -1.9  | -0.70  | -0.20 | 0.20  | 1.3     | -3.8259           | 0.0002  |
|                   |           | 30 mg              | 14.64 | 86    | -0.21 | 0.597                | -2.0  | -0.50  | -0.20 | 0.20  | 1.1     | -3.3043           | 0.0014  |
|                   |           | 60 mg              | 14.74 | 82    | -0.13 | 0.648                | -1.8  | -0.60  | -0.10 | 0.30  | 1.2     | -1.7732           | 0.0800  |
|                   |           | 120 mg             | 14.56 | 82    | -0.30 | 0.755                | -3.7  | -0.70  | -0.30 | 0.20  | 1.2     | -3.6437           | 0.0005  |
|                   | Week 16   | Placebo            | 14.94 | 95    | -0.27 | 0.746                | -3.5  | -0.60  | -0.20 | 0.20  | 1.5     | -3.5781           | 0.0005  |
|                   |           | 15 mg              | 14.44 | 84    | -0.17 | 0.612                | -1.8  | -0.40  | -0.20 | 0.20  | 1.4     | -2.5859           | 0.0115  |
|                   |           | 30 mg              | 14.65 | 85    | -0.22 | 0.794                | -3.4  | -0.60  | -0.20 | 0.30  | 1.7     | -2.5135           | 0.0139  |
|                   |           | 60 mg              | 14.86 | 80    | -0.14 | 0.651                | -2.1  | -0.60  | -0.20 | 0.25  | 1.8     | -1.9403           | 0.0559  |
|                   |           | 120 mg             | 14.58 | 81    | -0.29 | 0.744                | -3.4  | -0.70  | -0.20 | 0.10  | 1.1     | -3.4947           | 0.0008  |
|                   | Week 20   | Placebo            | 14.94 | 95    | -0.17 | 0.768                | -3.3  | -0.70  | -0.10 | 0.30  | 1.6     | -2.1113           | 0.0374  |
|                   |           | 15 mg              | 14.44 | 84    | -0.15 | 0.736                | -2.2  | -0.55  | -0.15 | 0.30  | 1.6     | -1.8237           | 0.0718  |
|                   |           | 30 mg              | 14.65 | 85    | -0.29 | 0.879                | -4.8  | -0.70  | -0.30 | 0.30  | 1.8     | -3.0495           | 0.0031  |
|                   |           | 60 mg              | 14.81 | 79    | -0.20 | 0.704                | -2.2  | -0.70  | -0.30 | 0.30  | 1.4     | -2.4755           | 0.0155  |
|                   |           | 120 mg             | 14.60 | 79    | -0.35 | 0.770                | -2.8  | -0.90  | -0.30 | 0.20  | 1.5     | -4.0031           | 0.0001  |
|                   | Week 24   | Placebo            | 14.96 | 93    | -0.38 | 0.825                | -4.2  | -0.70  | -0.30 | -0.10 | 1.8     | -4.4016           | <0.0001 |
|                   |           | 15 mg              | 14.42 | 83    | -0.18 | 0.665                | -2.0  | -0.60  | -0.20 | 0.30  | 1.6     | -2.4743           | 0.0154  |
|                   |           | 30 mg              | 14.67 | 84    | -0.30 | 0.833                | -4.0  | -0.75  | -0.20 | 0.20  | 1.4     | -3.2872           | 0.0015  |
| 60 mg             |           | 14.83              | 78    | -0.27 | 0.747 | -2.8                 | -0.80 | -0.30  | 0.20  | 1.3   | -3.1999 | 0.0020            |         |
| 120 mg            |           | 14.61              | 78    | -0.38 | 0.813 | -3.1                 | -0.70 | -0.30  | 0.00  | 1.9   | -4.0803 | 0.0001            |         |
| Week 28           | Placebo   | 14.96              | 93    | -0.34 | 0.764 | -3.0                 | -0.70 | -0.30  | 0.10  | 1.7   | -4.3031 | <0.0001           |         |
|                   | 15 mg     | 14.41              | 83    | -0.17 | 0.694 | -1.8                 | -0.60 | -0.20  | 0.30  | 1.7   | -2.2136 | 0.0296            |         |
|                   | 30 mg     | 14.67              | 84    | -0.25 | 0.736 | -2.8                 | -0.60 | -0.20  | 0.20  | 1.3   | -3.1411 | 0.0023            |         |
|                   | 60 mg     | 14.81              | 77    | -0.34 | 0.710 | -3.4                 | -0.70 | -0.30  | 0.10  | 0.8   | -4.1431 | <0.0001           |         |
|                   | 120 mg    | 14.59              | 77    | -0.28 | 0.893 | -4.0                 | -0.80 | -0.20  | 0.20  | 1.9   | -2.7060 | 0.0084            |         |
| Follow-up         | Placebo   | 14.89              | 97    | -0.34 | 0.753 | -3.0                 | -0.70 | -0.30  | 0.10  | 1.7   | -4.3796 | <0.0001           |         |
|                   | 15 mg     | 14.39              | 86    | -0.20 | 0.720 | -2.1                 | -0.70 | -0.20  | 0.30  | 1.7   | -2.5168 | 0.0137            |         |
|                   | 30 mg     | 14.59              | 92    | -0.25 | 0.708 | -2.8                 | -0.60 | -0.20  | 0.20  | 1.3   | -3.3415 | 0.0012            |         |
|                   | 60 mg     | 14.74              | 87    | -0.35 | 0.692 | -3.4                 | -0.70 | -0.30  | 0.10  | 0.8   | -4.6649 | <0.0001           |         |
|                   | 120 mg    | 14.59              | 87    | -0.29 | 0.851 | -4.0                 | -0.80 | -0.30  | 0.20  | 1.9   | -3.1738 | 0.0021            |         |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (2/14)

| Item           | Treatment | Summary Statistics |       |       |       |                      |       |        |       |      |         | One-sample t-test |         |
|----------------|-----------|--------------------|-------|-------|-------|----------------------|-------|--------|-------|------|---------|-------------------|---------|
|                |           | Baseline Mean      | N     | Mean  | SD    | Change from Baseline |       |        |       |      | t-value | p-value           |         |
|                |           |                    |       |       |       | Min                  | Q1    | Median | Q3    | Max  |         |                   |         |
| Hematocrit (%) | Week 1    | Placebo            | 43.27 | 39    | -0.17 | 1.830                | -4.7  | -1.10  | -0.20 | 0.80 | 4.2     | -0.5950           | 0.5554  |
|                |           | 15 mg              | 43.27 | 37    | -0.78 | 1.816                | -5.0  | -1.50  | -0.80 | 0.10 | 3.3     | -2.6072           | 0.0132  |
|                |           | 30 mg              | 43.17 | 39    | -0.24 | 2.227                | -6.1  | -1.70  | 0.10  | 0.80 | 4.1     | -0.6688           | 0.5077  |
|                |           | 60 mg              | 44.60 | 37    | -0.72 | 1.702                | -4.0  | -2.10  | -0.60 | 0.20 | 2.9     | -2.5689           | 0.0145  |
|                |           | 120 mg             | 43.72 | 47    | -0.33 | 1.352                | -3.7  | -1.10  | -0.20 | 0.40 | 2.9     | -1.6719           | 0.1013  |
|                | Week 2    | Placebo            | 44.89 | 97    | -0.81 | 1.741                | -5.0  | -2.10  | -0.90 | 0.30 | 3.5     | -4.5666           | <0.0001 |
|                |           | 15 mg              | 43.85 | 89    | -0.39 | 1.801                | -8.0  | -1.40  | -0.30 | 0.80 | 4.2     | -2.0487           | 0.0435  |
|                |           | 30 mg              | 44.11 | 91    | -0.49 | 1.962                | -10.2 | -1.70  | -0.30 | 0.60 | 4.4     | -2.3828           | 0.0193  |
|                |           | 60 mg              | 44.37 | 93    | -0.50 | 1.794                | -7.5  | -1.60  | -0.40 | 0.80 | 3.3     | -2.7056           | 0.0081  |
|                |           | 120 mg             | 43.94 | 88    | -0.67 | 1.699                | -4.7  | -1.75  | -0.50 | 0.30 | 5.9     | -3.6946           | 0.0004  |
|                | Week 3    | Placebo            | 43.55 | 37    | -0.58 | 1.608                | -4.0  | -1.70  | -0.80 | 0.50 | 3.5     | -2.1776           | 0.0361  |
|                |           | 15 mg              | 43.33 | 36    | -0.48 | 1.676                | -5.4  | -1.60  | -0.35 | 0.35 | 3.3     | -1.7307           | 0.0923  |
|                |           | 30 mg              | 43.30 | 38    | -0.60 | 2.281                | -6.7  | -2.00  | -0.35 | 0.80 | 4.1     | -1.6213           | 0.1134  |
|                |           | 60 mg              | 44.46 | 32    | -0.92 | 1.862                | -4.5  | -1.90  | -1.10 | 0.20 | 3.8     | -2.7812           | 0.0091  |
|                |           | 120 mg             | 43.69 | 42    | -0.75 | 1.585                | -4.7  | -1.70  | -0.60 | 0.20 | 3.2     | -3.0559           | 0.0039  |
|                | Week 4    | Placebo            | 44.98 | 96    | -0.52 | 1.788                | -6.9  | -1.70  | -0.60 | 0.45 | 4.4     | -2.8606           | 0.0052  |
|                |           | 15 mg              | 43.85 | 89    | -0.61 | 1.663                | -6.5  | -1.50  | -0.60 | 0.30 | 3.3     | -3.4609           | 0.0008  |
|                |           | 30 mg              | 44.17 | 89    | -0.70 | 2.058                | -8.1  | -2.00  | -0.80 | 0.70 | 4.1     | -3.1878           | 0.0020  |
|                |           | 60 mg              | 44.47 | 89    | -0.57 | 1.918                | -7.8  | -1.60  | -0.50 | 0.60 | 4.0     | -2.7856           | 0.0065  |
|                |           | 120 mg             | 43.88 | 86    | -0.92 | 2.037                | -6.9  | -2.10  | -0.90 | 0.50 | 3.8     | -4.1988           | <0.0001 |
| Week 6         | Placebo   | 43.78              | 40    | -0.32 | 1.688 | -4.7                 | -1.50 | -0.30  | 0.65  | 2.6  | -1.2080 | 0.2343            |         |
|                | 15 mg     | 43.51              | 40    | -0.75 | 2.108 | -6.6                 | -1.55 | -0.25  | 0.45  | 3.5  | -2.2498 | 0.0302            |         |
|                | 30 mg     | 43.28              | 38    | -0.52 | 2.087 | -7.5                 | -1.50 | -0.50  | 0.70  | 3.4  | -1.5236 | 0.1361            |         |
|                | 60 mg     | 44.95              | 31    | -0.64 | 2.072 | -5.0                 | -2.50 | -0.40  | 0.60  | 4.3  | -1.7250 | 0.0948            |         |
|                | 120 mg    | 43.88              | 47    | -0.85 | 2.147 | -8.7                 | -1.90 | -0.80  | 0.20  | 6.9  | -2.7104 | 0.0094            |         |
| Week 8         | Placebo   | 44.93              | 94    | -0.75 | 1.975 | -6.9                 | -2.10 | -0.70  | 0.40  | 3.3  | -3.6605 | 0.0004            |         |
|                | 15 mg     | 43.86              | 86    | -0.39 | 1.933 | -5.3                 | -1.80 | -0.40  | 1.00  | 4.1  | -1.8636 | 0.0658            |         |
|                | 30 mg     | 44.17              | 89    | -0.58 | 2.098 | -7.5                 | -1.90 | -0.60  | 0.70  | 4.0  | -2.6173 | 0.0104            |         |
|                | 60 mg     | 44.36              | 85    | -0.29 | 1.906 | -4.6                 | -1.40 | -0.10  | 1.10  | 3.1  | -1.3999 | 0.1652            |         |
|                | 120 mg    | 43.93              | 83    | -0.79 | 2.134 | -9.3                 | -2.00 | -0.70  | 0.60  | 4.2  | -3.3898 | 0.0011            |         |
| Week 12        | Placebo   | 44.97              | 95    | -0.79 | 2.234 | -6.1                 | -2.30 | -0.80  | 0.80  | 4.1  | -3.4447 | 0.0009            |         |
|                | 15 mg     | 43.86              | 86    | -0.78 | 1.941 | -5.0                 | -2.10 | -0.70  | 0.40  | 3.8  | -3.7112 | 0.0004            |         |
|                | 30 mg     | 44.26              | 86    | -0.80 | 1.952 | -7.5                 | -1.80 | -0.60  | 0.50  | 3.0  | -3.7897 | 0.0003            |         |
|                | 60 mg     | 44.35              | 82    | -0.38 | 1.848 | -5.6                 | -1.40 | -0.45  | 1.10  | 3.4  | -1.8580 | 0.0668            |         |
|                | 120 mg    | 43.84              | 82    | -0.73 | 2.250 | -9.1                 | -2.00 | -0.45  | 0.60  | 3.6  | -2.9400 | 0.0043            |         |
| Week 16        | Placebo   | 44.97              | 95    | -0.86 | 2.472 | -10.1                | -2.30 | -0.50  | 0.90  | 4.7  | -3.3948 | 0.0010            |         |
|                | 15 mg     | 43.78              | 84    | -0.65 | 1.956 | -5.8                 | -1.70 | -0.60  | 0.50  | 4.0  | -3.0291 | 0.0033            |         |
|                | 30 mg     | 44.30              | 85    | -0.94 | 2.466 | -10.6                | -2.00 | -1.10  | 0.30  | 5.0  | -3.5225 | 0.0007            |         |
|                | 60 mg     | 44.65              | 80    | -0.44 | 1.886 | -5.0                 | -1.70 | -0.40  | 0.70  | 4.7  | -2.0930 | 0.0396            |         |
|                | 120 mg    | 43.89              | 81    | -0.76 | 2.164 | -7.2                 | -2.10 | -0.80  | 0.50  | 3.5  | -3.1684 | 0.0022            |         |
| Week 20        | Placebo   | 44.97              | 95    | -0.70 | 2.463 | -8.8                 | -2.20 | -0.40  | 0.80  | 4.2  | -2.7741 | 0.0067            |         |
|                | 15 mg     | 43.78              | 84    | -0.60 | 2.231 | -6.4                 | -2.10 | -0.50  | 1.00  | 4.4  | -2.4599 | 0.0160            |         |
|                | 30 mg     | 44.30              | 85    | -1.20 | 2.669 | -12.2                | -2.40 | -1.00  | 0.20  | 4.6  | -4.1409 | <0.0001           |         |
|                | 60 mg     | 44.56              | 79    | -0.66 | 2.076 | -6.7                 | -2.30 | -0.70  | 0.60  | 4.4  | -2.8126 | 0.0062            |         |
|                | 120 mg    | 43.92              | 79    | -0.84 | 2.235 | -6.3                 | -2.30 | -1.10  | 0.70  | 5.0  | -3.3424 | 0.0013            |         |
| Week 24        | Placebo   | 45.03              | 93    | -1.27 | 2.742 | -14.8                | -2.40 | -1.00  | 0.40  | 3.7  | -4.4662 | <0.0001           |         |
|                | 15 mg     | 43.75              | 83    | -0.92 | 2.159 | -6.3                 | -2.30 | -1.00  | 0.40  | 4.2  | -3.8800 | 0.0002            |         |
|                | 30 mg     | 44.36              | 84    | -1.20 | 2.637 | -11.5                | -2.80 | -0.80  | 0.60  | 3.2  | -4.1838 | <0.0001           |         |
|                | 60 mg     | 44.62              | 78    | -1.07 | 1.990 | -7.4                 | -2.40 | -1.25  | 0.20  | 2.6  | -4.7634 | <0.0001           |         |
|                | 120 mg    | 43.98              | 78    | -1.20 | 2.422 | -8.4                 | -2.80 | -1.15  | 0.20  | 5.2  | -4.3656 | <0.0001           |         |
| Week 28        | Placebo   | 45.01              | 93    | -1.20 | 2.294 | -6.9                 | -2.40 | -0.90  | 0.30  | 3.9  | -5.0398 | <0.0001           |         |
|                | 15 mg     | 43.73              | 83    | -0.73 | 2.069 | -5.4                 | -2.10 | -0.40  | 0.60  | 4.7  | -3.2310 | 0.0018            |         |
|                | 30 mg     | 44.36              | 84    | -1.16 | 2.309 | -9.0                 | -2.35 | -1.10  | -0.05 | 4.2  | -4.5974 | <0.0001           |         |
|                | 60 mg     | 44.57              | 77    | -1.07 | 2.041 | -8.1                 | -2.20 | -1.10  | 0.30  | 4.0  | -4.5958 | <0.0001           |         |
|                | 120 mg    | 43.95              | 77    | -0.77 | 2.553 | -10.1                | -2.40 | -0.70  | 0.70  | 5.3  | -2.6609 | 0.0095            |         |
| Follow-up      | Placebo   | 44.87              | 97    | -1.19 | 2.276 | -6.9                 | -2.40 | -1.00  | 0.30  | 3.9  | -5.1654 | <0.0001           |         |
|                | 15 mg     | 43.69              | 86    | -0.80 | 2.121 | -5.7                 | -2.20 | -0.50  | 0.60  | 4.7  | -3.4973 | 0.0007            |         |
|                | 30 mg     | 44.16              | 92    | -1.12 | 2.223 | -9.0                 | -2.30 | -1.10  | -0.05 | 4.2  | -4.8122 | <0.0001           |         |
|                | 60 mg     | 44.38              | 87    | -1.11 | 1.970 | -8.1                 | -2.40 | -1.10  | 0.30  | 4.0  | -5.2356 | <0.0001           |         |
|                | 120 mg    | 43.92              | 87    | -0.81 | 2.449 | -10.1                | -2.40 | -0.80  | 0.70  | 5.3  | -3.0683 | 0.0029            |         |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (3/14)

| Item                                 | Treatment | Summary Statistics |       |       |       |                      |       |        |       |      |         | One-sample t-test |         |
|--------------------------------------|-----------|--------------------|-------|-------|-------|----------------------|-------|--------|-------|------|---------|-------------------|---------|
|                                      |           | Baseline Mean      | N     | Mean  | SD    | Change from Baseline |       |        |       |      | t-value | p-value           |         |
|                                      |           |                    |       |       |       | Min                  | Q1    | Median | Q3    | Max  |         |                   |         |
| RBC<br>( $\times 10^3/\mu\text{L}$ ) | Week 1    | Placebo            | 470.9 | 39    | -3.2  | 17.77                | -50   | -16.0  | -2.0  | 8.0  | 37      | -1.1264           | 0.2670  |
|                                      |           | 15 mg              | 476.6 | 37    | -8.1  | 16.09                | -39   | -20.0  | -8.0  | 3.0  | 19      | -3.0757           | 0.0040  |
|                                      |           | 30 mg              | 470.5 | 39    | -2.7  | 20.32                | -63   | -15.0  | -2.0  | 9.0  | 39      | -0.8198           | 0.4175  |
|                                      |           | 60 mg              | 480.8 | 37    | -6.8  | 18.69                | -43   | -20.0  | -8.0  | 6.0  | 29      | -2.1987           | 0.0344  |
|                                      |           | 120 mg             | 471.9 | 47    | -3.8  | 12.43                | -31   | -8.0   | -4.0  | 6.0  | 24      | -2.1127           | 0.0401  |
|                                      | Week 2    | Placebo            | 486.8 | 97    | -8.1  | 16.19                | -40   | -21.0  | -8.0  | 3.0  | 31      | -4.9551           | <0.0001 |
|                                      |           | 15 mg              | 480.5 | 89    | -4.6  | 16.67                | -55   | -15.0  | -4.0  | 8.0  | 41      | -2.5752           | 0.0117  |
|                                      |           | 30 mg              | 480.0 | 91    | -5.6  | 20.53                | -113  | -17.0  | -3.0  | 8.0  | 43      | -2.5833           | 0.0114  |
|                                      |           | 60 mg              | 481.8 | 93    | -5.2  | 18.19                | -76   | -18.0  | -3.0  | 8.0  | 42      | -2.7530           | 0.0071  |
|                                      |           | 120 mg             | 476.3 | 88    | -9.7  | 15.51                | -48   | -18.0  | -7.5  | -2.0 | 28      | -5.8782           | <0.0001 |
|                                      | Week 3    | Placebo            | 472.7 | 37    | -5.5  | 16.05                | -33   | -17.0  | -7.0  | 7.0  | 43      | -2.0691           | 0.0458  |
|                                      |           | 15 mg              | 478.3 | 36    | -7.8  | 13.87                | -38   | -16.0  | -5.5  | 1.5  | 13      | -3.3522           | 0.0019  |
|                                      |           | 30 mg              | 471.3 | 38    | -7.0  | 22.62                | -81   | -23.0  | -2.5  | 6.0  | 47      | -1.9149           | 0.0633  |
|                                      |           | 60 mg              | 480.6 | 32    | -11.1 | 16.68                | -43   | -20.5  | -12.5 | -0.5 | 35      | -3.7628           | 0.0007  |
|                                      |           | 120 mg             | 472.0 | 42    | -10.6 | 16.96                | -55   | -19.0  | -9.0  | 0.0  | 32      | -4.0488           | 0.0002  |
|                                      | Week 4    | Placebo            | 487.7 | 96    | -5.2  | 16.96                | -61   | -16.5  | -4.0  | 4.0  | 42      | -3.0263           | 0.0032  |
|                                      |           | 15 mg              | 480.5 | 89    | -6.0  | 15.92                | -48   | -19.0  | -6.0  | 6.0  | 28      | -3.5752           | 0.0006  |
|                                      |           | 30 mg              | 480.1 | 89    | -7.4  | 21.96                | -95   | -20.0  | -5.0  | 7.0  | 36      | -3.1670           | 0.0021  |
|                                      |           | 60 mg              | 482.9 | 89    | -7.4  | 19.17                | -84   | -17.0  | -6.0  | 4.0  | 37      | -3.6381           | 0.0005  |
|                                      |           | 120 mg             | 475.9 | 86    | -10.8 | 18.98                | -55   | -23.0  | -10.0 | 0.0  | 45      | -5.2850           | <0.0001 |
| Week 6                               | Placebo   | 474.0              | 40    | -5.8  | 16.07 | -43                  | -15.5 | -8.5   | 2.0   | 29   | -2.2925 | 0.0274            |         |
|                                      | 15 mg     | 479.4              | 40    | -10.0 | 18.06 | -59                  | -19.5 | -7.5   | 4.0   | 26   | -3.4940 | 0.0012            |         |
|                                      | 30 mg     | 471.6              | 38    | -6.7  | 24.34 | -86                  | -21.0 | -7.0   | 13.0  | 33   | -1.6994 | 0.0976            |         |
|                                      | 60 mg     | 482.1              | 31    | -7.2  | 18.65 | -42                  | -26.0 | -5.0   | 6.0   | 34   | -2.1477 | 0.0399            |         |
|                                      | 120 mg    | 470.8              | 47    | -9.6  | 18.59 | -82                  | -19.0 | -10.0  | -2.0  | 49   | -3.5466 | 0.0009            |         |
| Week 8                               | Placebo   | 487.3              | 94    | -7.2  | 19.08 | -60                  | -19.0 | -8.0   | 3.0   | 46   | -3.6484 | 0.0004            |         |
|                                      | 15 mg     | 481.0              | 86    | -4.6  | 18.60 | -52                  | -17.0 | -4.5   | 9.0   | 41   | -2.2905 | 0.0245            |         |
|                                      | 30 mg     | 480.1              | 89    | -6.1  | 21.60 | -83                  | -18.0 | -5.0   | 9.0   | 39   | -2.6456 | 0.0097            |         |
|                                      | 60 mg     | 481.0              | 85    | -2.2  | 20.57 | -55                  | -14.0 | 1.0    | 12.0  | 36   | -0.9967 | 0.3218            |         |
|                                      | 120 mg    | 475.3              | 83    | -10.2 | 21.00 | -98                  | -24.0 | -7.0   | 4.0   | 37   | -4.4213 | <0.0001           |         |
| Week 12                              | Placebo   | 487.7              | 95    | -7.5  | 20.52 | -55                  | -20.0 | -8.0   | 6.0   | 31   | -3.5405 | 0.0006            |         |
|                                      | 15 mg     | 481.0              | 86    | -8.3  | 18.31 | -58                  | -22.0 | -6.0   | 6.0   | 32   | -4.1817 | <0.0001           |         |
|                                      | 30 mg     | 480.0              | 86    | -7.4  | 20.24 | -78                  | -19.0 | -6.0   | 8.0   | 33   | -3.3826 | 0.0011            |         |
|                                      | 60 mg     | 480.2              | 82    | -2.0  | 19.80 | -57                  | -16.0 | -1.5   | 16.0  | 33   | -0.8924 | 0.3748            |         |
|                                      | 120 mg    | 474.4              | 82    | -8.1  | 22.65 | -105                 | -20.0 | -4.0   | 6.0   | 35   | -3.2427 | 0.0017            |         |
| Week 16                              | Placebo   | 487.7              | 95    | -7.2  | 24.47 | -92                  | -22.0 | -4.0   | 10.0  | 48   | -2.8761 | 0.0050            |         |
|                                      | 15 mg     | 480.0              | 84    | -6.1  | 18.37 | -56                  | -17.0 | -5.0   | 8.0   | 35   | -3.0527 | 0.0030            |         |
|                                      | 30 mg     | 480.4              | 85    | -8.2  | 26.14 | -98                  | -26.0 | -9.0   | 7.0   | 65   | -2.8759 | 0.0051            |         |
|                                      | 60 mg     | 483.6              | 80    | -2.7  | 20.04 | -52                  | -15.0 | -2.0   | 11.0  | 40   | -1.2215 | 0.2255            |         |
|                                      | 120 mg    | 475.0              | 81    | -7.2  | 23.16 | -98                  | -16.0 | -3.0   | 5.0   | 40   | -2.8116 | 0.0062            |         |
| Week 20                              | Placebo   | 487.7              | 95    | -4.4  | 23.95 | -70                  | -19.0 | -3.0   | 10.0  | 53   | -1.7866 | 0.0772            |         |
|                                      | 15 mg     | 480.0              | 84    | -4.6  | 21.43 | -67                  | -17.0 | -4.5   | 11.0  | 38   | -1.9654 | 0.0527            |         |
|                                      | 30 mg     | 480.4              | 85    | -10.0 | 28.33 | -121                 | -25.0 | -9.0   | 6.0   | 59   | -3.2396 | 0.0017            |         |
|                                      | 60 mg     | 482.5              | 79    | -4.2  | 21.91 | -74                  | -20.0 | -5.0   | 11.0  | 42   | -1.6895 | 0.0951            |         |
|                                      | 120 mg    | 475.5              | 79    | -8.5  | 21.94 | -70                  | -23.0 | -11.0  | 7.0   | 46   | -3.4355 | 0.0010            |         |
| Week 24                              | Placebo   | 488.5              | 93    | -10.1 | 26.94 | -135                 | -22.0 | -8.0   | 3.0   | 43   | -3.6028 | 0.0005            |         |
|                                      | 15 mg     | 479.8              | 83    | -6.2  | 21.09 | -55                  | -19.0 | -6.0   | 10.0  | 36   | -2.6905 | 0.0086            |         |
|                                      | 30 mg     | 480.8              | 84    | -8.9  | 27.11 | -93                  | -22.5 | -5.5   | 10.5  | 41   | -2.9938 | 0.0036            |         |
|                                      | 60 mg     | 483.2              | 78    | -7.2  | 22.01 | -78                  | -21.0 | -6.5   | 5.0   | 38   | -2.8860 | 0.0051            |         |
|                                      | 120 mg    | 476.1              | 78    | -10.2 | 22.31 | -79                  | -26.0 | -11.0  | 6.0   | 41   | -4.0343 | 0.0001            |         |
| Week 28                              | Placebo   | 488.2              | 93    | -9.3  | 22.35 | -66                  | -23.0 | -5.0   | 3.0   | 48   | -4.0046 | 0.0001            |         |
|                                      | 15 mg     | 479.4              | 83    | -4.6  | 20.30 | -65                  | -19.0 | -5.0   | 8.0   | 38   | -2.0490 | 0.0437            |         |
|                                      | 30 mg     | 480.8              | 84    | -8.9  | 24.42 | -105                 | -20.0 | -9.5   | 4.0   | 51   | -3.3557 | 0.0012            |         |
|                                      | 60 mg     | 482.9              | 77    | -7.1  | 22.43 | -100                 | -18.0 | -5.0   | 7.0   | 37   | -2.7795 | 0.0069            |         |
|                                      | 120 mg    | 476.2              | 77    | -3.9  | 26.76 | -97                  | -20.0 | -1.0   | 10.0  | 71   | -1.2902 | 0.2009            |         |
| Follow-up                            | Placebo   | 486.7              | 97    | -9.2  | 22.12 | -66                  | -23.0 | -5.0   | 3.0   | 48   | -4.0940 | <0.0001           |         |
|                                      | 15 mg     | 478.6              | 86    | -5.1  | 20.96 | -65                  | -19.0 | -4.5   | 8.0   | 38   | -2.2581 | 0.0265            |         |
|                                      | 30 mg     | 480.2              | 92    | -8.5  | 23.75 | -105                 | -19.0 | -8.5   | 4.0   | 51   | -3.4326 | 0.0009            |         |
|                                      | 60 mg     | 482.6              | 87    | -7.7  | 21.84 | -100                 | -20.0 | -5.0   | 7.0   | 37   | -3.2943 | 0.0014            |         |
|                                      | 120 mg    | 476.4              | 87    | -4.8  | 25.83 | -97                  | -20.0 | -8.0   | 10.0  | 71   | -1.7269 | 0.0878            |         |



表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (4/14)

| Item         | Treatment | Summary Statistics |        |       |         |                      |        |         |        |        | One-sample t-test |         |        |
|--------------|-----------|--------------------|--------|-------|---------|----------------------|--------|---------|--------|--------|-------------------|---------|--------|
|              |           | Baseline Mean      | N      | Mean  | SD      | Change from Baseline |        |         |        |        | t-value           | p-value |        |
|              |           |                    |        |       |         | Min                  | Q1     | Median  | Q3     | Max    |                   |         |        |
| WBC<br>(/μL) | Week 1    | Placebo            | 6641.0 | 39    | -46.2   | 695.04               | -1600  | -400.0  | 0.0    | 400.0  | 1600              | -0.4147 | 0.6807 |
|              |           | 15 mg              | 6478.4 | 37    | 351.4   | 791.84               | -1300  | -200.0  | 600.0  | 800.0  | 1900              | 2.6990  | 0.0105 |
|              |           | 30 mg              | 6694.9 | 39    | 128.2   | 1262.40              | -3000  | -500.0  | -100.0 | 800.0  | 4600              | 0.6342  | 0.5297 |
|              |           | 60 mg              | 6283.8 | 37    | 221.6   | 1158.58              | -3500  | -200.0  | 400.0  | 700.0  | 3300              | 1.1636  | 0.2523 |
|              | 120 mg    | 6412.8             | 47     | -17.0 | 1237.16 | -3400                | -800.0 | 0.0     | 900.0  | 3400   | -0.0943           | 0.9253  |        |
|              | Week 2    | Placebo            | 6650.5 | 97    | -208.2  | 985.06               | -3100  | -800.0  | -200.0 | 300.0  | 2000              | -2.0821 | 0.0400 |
|              |           | 15 mg              | 6542.7 | 89    | 323.6   | 1738.47              | -3100  | -300.0  | 100.0  | 700.0  | 13000             | 1.7560  | 0.0826 |
|              |           | 30 mg              | 6509.9 | 91    | 297.8   | 1313.77              | -3400  | -400.0  | 300.0  | 900.0  | 4100              | 2.1624  | 0.0332 |
|              |           | 60 mg              | 6580.6 | 93    | 180.6   | 916.66               | -2300  | -400.0  | 100.0  | 800.0  | 2700              | 1.9005  | 0.0605 |
|              |           | 120 mg             | 6643.2 | 88    | -168.2  | 1201.20              | -5500  | -900.0  | -250.0 | 550.0  | 3500              | -1.3134 | 0.1925 |
|              | Week 3    | Placebo            | 6637.8 | 37    | -294.6  | 900.91               | -1800  | -900.0  | -400.0 | 100.0  | 2000              | -1.9890 | 0.0543 |
|              |           | 15 mg              | 6447.2 | 36    | 391.7   | 901.23               | -1400  | -100.0  | 300.0  | 700.0  | 2700              | 2.6075  | 0.0133 |
|              |           | 30 mg              | 6760.5 | 38    | -18.4   | 1238.10              | -3400  | -600.0  | -200.0 | 500.0  | 2900              | -0.0917 | 0.9274 |
|              |           | 60 mg              | 6228.1 | 32    | 106.3   | 1035.79              | -2300  | -300.0  | 200.0  | 650.0  | 2500              | 0.5803  | 0.5659 |
|              |           | 120 mg             | 6519.0 | 42    | -307.1  | 1046.81              | -4300  | -900.0  | -250.0 | 400.0  | 2100              | -1.9015 | 0.0643 |
|              | Week 4    | Placebo            | 6675.0 | 96    | -289.6  | 977.16               | -3000  | -900.0  | -250.0 | 300.0  | 2000              | -2.9037 | 0.0046 |
|              |           | 15 mg              | 6542.7 | 89    | 138.2   | 1164.00              | -3500  | -400.0  | 100.0  | 800.0  | 4700              | 1.1201  | 0.2657 |
|              |           | 30 mg              | 6480.9 | 89    | 124.7   | 1237.42              | -4100  | -500.0  | 0.0    | 900.0  | 3900              | 0.9508  | 0.3443 |
|              |           | 60 mg              | 6641.6 | 89    | -68.5   | 1163.42              | -3400  | -600.0  | -100.0 | 600.0  | 4900              | -0.5558 | 0.5798 |
|              |           | 120 mg             | 6703.5 | 86    | -51.2   | 1320.23              | -4800  | -900.0  | -200.0 | 600.0  | 4200              | -0.3594 | 0.7202 |
|              | Week 6    | Placebo            | 6390.0 | 40    | 257.5   | 1059.73              | -2500  | -500.0  | 150.0  | 1150.0 | 2600              | 1.5368  | 0.1324 |
|              |           | 15 mg              | 6255.0 | 40    | 295.0   | 1515.38              | -1700  | -450.0  | 100.0  | 750.0  | 6800              | 1.2312  | 0.2256 |
|              |           | 30 mg              | 6757.9 | 38    | -18.4   | 1179.98              | -3900  | -600.0  | 50.0   | 600.0  | 3100              | -0.0962 | 0.9239 |
|              |           | 60 mg              | 6283.9 | 31    | 212.9   | 1204.92              | -1700  | -600.0  | 0.0    | 1000.0 | 3900              | 0.9838  | 0.3331 |
|              |           | 120 mg             | 6461.7 | 47    | -74.5   | 1353.17              | -4600  | -800.0  | -100.0 | 700.0  | 3900              | -0.3773 | 0.7077 |
|              | Week 8    | Placebo            | 6681.9 | 94    | -402.1  | 989.51               | -4000  | -900.0  | -300.0 | 200.0  | 1700              | -3.9401 | 0.0002 |
|              |           | 15 mg              | 6560.5 | 86    | -22.1   | 973.58               | -2900  | -500.0  | -50.0  | 600.0  | 3100              | -0.2104 | 0.8338 |
|              |           | 30 mg              | 6480.9 | 89    | 142.7   | 1381.85              | -3300  | -700.0  | 0.0    | 900.0  | 4500              | 0.9742  | 0.3326 |
|              |           | 60 mg              | 6591.8 | 85    | 92.9    | 1064.45              | -2200  | -600.0  | 100.0  | 700.0  | 3000              | 0.8050  | 0.4231 |
|              |           | 120 mg             | 6662.7 | 83    | -108.4  | 1336.29              | -5000  | -700.0  | -100.0 | 600.0  | 4300              | -0.7393 | 0.4619 |
|              | Week 12   | Placebo            | 6656.8 | 95    | -210.5  | 1372.25              | -3100  | -1000.0 | -300.0 | 400.0  | 6500              | -1.4953 | 0.1382 |
|              |           | 15 mg              | 6560.5 | 86    | -169.8  | 1147.08              | -3500  | -800.0  | -150.0 | 500.0  | 2700              | -1.3725 | 0.1735 |
|              |           | 30 mg              | 6481.4 | 86    | 108.1   | 1209.30              | -4100  | -600.0  | 100.0  | 800.0  | 4700              | 0.8293  | 0.4093 |
|              |           | 60 mg              | 6567.1 | 82    | -31.7   | 1201.12              | -3400  | -600.0  | 0.0    | 600.0  | 4300              | -0.2390 | 0.8117 |
|              |           | 120 mg             | 6672.0 | 82    | -267.1  | 1145.58              | -4900  | -900.0  | -100.0 | 300.0  | 2500              | -2.1111 | 0.0378 |
|              | Week 16   | Placebo            | 6656.8 | 95    | -312.6  | 1134.68              | -3700  | -1100.0 | -200.0 | 500.0  | 2300              | -2.6855 | 0.0086 |
|              |           | 15 mg              | 6592.9 | 84    | 8.3     | 1150.40              | -3600  | -600.0  | 100.0  | 750.0  | 2900              | 0.0664  | 0.9472 |
|              |           | 30 mg              | 6476.5 | 85    | 278.8   | 1391.14              | -2400  | -600.0  | 200.0  | 900.0  | 6300              | 1.8479  | 0.0681 |
|              |           | 60 mg              | 6552.5 | 80    | -163.8  | 1225.70              | -2900  | -900.0  | -100.0 | 650.0  | 3600              | -1.1949 | 0.2357 |
|              |           | 120 mg             | 6697.5 | 81    | -258.0  | 1148.79              | -4100  | -1000.0 | -100.0 | 400.0  | 2000              | -2.0215 | 0.0466 |
|              | Week 20   | Placebo            | 6656.8 | 95    | -245.3  | 1040.54              | -3100  | -900.0  | -200.0 | 400.0  | 2700              | -2.2974 | 0.0238 |
|              |           | 15 mg              | 6592.9 | 84    | -36.9   | 1077.79              | -3100  | -600.0  | 50.0   | 500.0  | 2500              | -0.3138 | 0.7544 |
|              |           | 30 mg              | 6476.5 | 85    | 85.9    | 1268.97              | -4200  | -600.0  | -100.0 | 800.0  | 3400              | 0.6240  | 0.5343 |
|              |           | 60 mg              | 6550.6 | 79    | -155.7  | 1263.67              | -4200  | -700.0  | 0.0    | 700.0  | 2600              | -1.0951 | 0.2768 |
|              |           | 120 mg             | 6738.0 | 79    | -74.7   | 1267.49              | -3500  | -800.0  | -200.0 | 600.0  | 5400              | -0.5237 | 0.6020 |
|              | Week 24   | Placebo            | 6680.6 | 93    | -349.5  | 1173.68              | -4200  | -1000.0 | -300.0 | 300.0  | 3100              | -2.8714 | 0.0051 |
|              |           | 15 mg              | 6563.9 | 83    | -55.4   | 1175.18              | -3800  | -800.0  | -100.0 | 900.0  | 3200              | -0.4297 | 0.6686 |
|              |           | 30 mg              | 6473.8 | 84    | 78.6    | 1140.29              | -3200  | -550.0  | 100.0  | 850.0  | 2200              | 0.6315  | 0.5294 |
|              |           | 60 mg              | 6575.6 | 78    | -115.4  | 1161.51              | -3000  | -700.0  | -100.0 | 500.0  | 3400              | -0.8773 | 0.3830 |
|              |           | 120 mg             | 6729.5 | 78    | -183.3  | 1247.59              | -4400  | -700.0  | -250.0 | 400.0  | 5200              | -1.2978 | 0.1982 |
|              | Week 28   | Placebo            | 6667.7 | 93    | -341.9  | 1134.18              | -2700  | -1100.0 | -400.0 | 400.0  | 2500              | -2.9074 | 0.0046 |
|              |           | 15 mg              | 6604.8 | 83    | -201.2  | 960.37               | -3100  | -600.0  | -100.0 | 500.0  | 1700              | -1.9087 | 0.0598 |
|              |           | 30 mg              | 6473.8 | 84    | 27.4    | 1287.32              | -4500  | -600.0  | -50.0  | 900.0  | 2400              | 0.1949  | 0.8459 |
|              |           | 60 mg              | 6593.5 | 77    | -139.0  | 1087.51              | -2500  | -800.0  | -300.0 | 600.0  | 4000              | -1.1213 | 0.2657 |
|              |           | 120 mg             | 6741.6 | 77    | -188.3  | 1252.99              | -4400  | -900.0  | -100.0 | 600.0  | 4800              | -1.3188 | 0.1912 |
|              | Follow-up | Placebo            | 6639.2 | 97    | -352.6  | 1125.10              | -2700  | -1100.0 | -400.0 | 400.0  | 2500              | -3.0864 | 0.0026 |
|              |           | 15 mg              | 6564.0 | 86    | -188.4  | 947.12               | -3100  | -600.0  | -150.0 | 500.0  | 1700              | -1.8444 | 0.0686 |
|              |           | 30 mg              | 6494.6 | 92    | 2.2     | 1281.91              | -4500  | -600.0  | -100.0 | 900.0  | 2400              | 0.0163  | 0.9871 |
|              |           | 60 mg              | 6655.2 | 87    | -34.5   | 1099.35              | -2500  | -700.0  | -100.0 | 700.0  | 4000              | -0.2926 | 0.7706 |
|              |           | 120 mg             | 6652.9 | 87    | -157.5  | 1250.27              | -4400  | -900.0  | -100.0 | 600.0  | 4800              | -1.1748 | 0.2433 |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (5/14)

| Item           | Treatment | Summary Statistics |       |       |       |                      |       |        |       |      |         | One-sample t-test |        |
|----------------|-----------|--------------------|-------|-------|-------|----------------------|-------|--------|-------|------|---------|-------------------|--------|
|                |           | Baseline Mean      | N     | Mean  | SD    | Change from Baseline |       |        |       |      | t-value | p-value           |        |
|                |           |                    |       |       |       | Min                  | Q1    | Median | Q3    | Max  |         |                   |        |
| Neutrophil (%) | Week 1    | Placebo            | 58.73 | 39    | -1.58 | 5.404                | -16.0 | -5.10  | -1.10 | 2.90 | 5.7     | -1.8283           | 0.0754 |
|                |           | 15 mg              | 56.63 | 37    | 0.14  | 4.487                | -8.8  | -2.90  | 0.40  | 2.30 | 11.0    | 0.1832            | 0.8557 |
|                |           | 30 mg              | 55.48 | 39    | 0.09  | 6.291                | -12.8 | -4.00  | -0.30 | 4.90 | 15.4    | 0.0865            | 0.9315 |
|                |           | 60 mg              | 56.10 | 35    | 0.77  | 5.767                | -8.6  | -3.60  | 0.20  | 3.90 | 16.4    | 0.7885            | 0.4359 |
|                |           | 120 mg             | 57.65 | 47    | -0.00 | 7.629                | -22.0 | -3.20  | 0.30  | 3.90 | 15.5    | -0.0038           | 0.9970 |
|                | Week 2    | Placebo            | 57.97 | 97    | -1.22 | 6.983                | -20.7 | -5.10  | -1.90 | 2.40 | 23.8    | -1.7229           | 0.0881 |
|                |           | 15 mg              | 57.12 | 88    | 0.50  | 7.843                | -18.3 | -3.20  | -0.10 | 2.95 | 35.9    | 0.5994            | 0.5505 |
|                |           | 30 mg              | 56.77 | 90    | 0.45  | 7.920                | -20.7 | -4.70  | 0.95  | 5.00 | 23.1    | 0.5350            | 0.5940 |
|                |           | 60 mg              | 56.01 | 91    | -0.03 | 5.295                | -12.0 | -3.50  | -0.20 | 4.10 | 11.0    | -0.0515           | 0.9591 |
|                |           | 120 mg             | 56.43 | 87    | 0.20  | 7.269                | -22.2 | -2.80  | -0.20 | 4.10 | 22.0    | 0.2522            | 0.8015 |
|                | Week 3    | Placebo            | 58.88 | 37    | -2.84 | 5.081                | -19.7 | -3.60  | -2.20 | 0.80 | 2.6     | -3.4041           | 0.0016 |
|                |           | 15 mg              | 56.72 | 36    | 0.24  | 5.711                | -16.3 | -3.20  | 1.10  | 4.70 | 11.9    | 0.2539            | 0.8011 |
|                |           | 30 mg              | 55.39 | 38    | -0.71 | 6.066                | -14.5 | -4.00  | -0.65 | 3.80 | 12.5    | -0.7167           | 0.4780 |
|                |           | 60 mg              | 56.71 | 31    | 0.03  | 5.305                | -11.1 | -4.00  | 0.70  | 3.30 | 9.5     | 0.0271            | 0.9786 |
|                |           | 120 mg             | 58.17 | 42    | -1.29 | 6.224                | -13.6 | -5.00  | -0.95 | 1.50 | 18.0    | -1.3388           | 0.1880 |
|                | Week 4    | Placebo            | 58.01 | 96    | -0.97 | 6.286                | -24.9 | -3.55  | -0.65 | 2.30 | 18.2    | -1.5116           | 0.1340 |
|                |           | 15 mg              | 57.12 | 88    | 0.44  | 7.458                | -21.6 | -3.60  | -0.05 | 4.40 | 26.1    | 0.5517            | 0.5826 |
|                |           | 30 mg              | 56.52 | 89    | -0.33 | 7.605                | -24.2 | -3.50  | 0.70  | 4.60 | 17.8    | -0.4084           | 0.6840 |
|                |           | 60 mg              | 55.65 | 88    | -0.58 | 6.356                | -16.8 | -3.90  | -0.65 | 3.05 | 18.6    | -0.8536           | 0.3956 |
|                |           | 120 mg             | 56.45 | 87    | 0.79  | 8.029                | -23.0 | -4.00  | 0.90  | 4.90 | 21.4    | 0.9160            | 0.3622 |
| Week 6         | Placebo   | 58.48              | 40    | -0.98 | 6.720 | -17.1                | -4.65 | 0.90   | 4.05  | 10.1 | -0.9200 | 0.3632            |        |
|                | 15 mg     | 56.59              | 39    | 2.06  | 7.204 | -13.6                | -1.40 | 2.30   | 5.90  | 28.2 | 1.7893  | 0.0815            |        |
|                | 30 mg     | 56.54              | 38    | -2.31 | 6.520 | -17.1                | -6.10 | -1.60  | 1.80  | 14.4 | -2.1796 | 0.0357            |        |
|                | 60 mg     | 55.66              | 30    | -0.05 | 6.850 | -14.2                | -4.80 | 0.60   | 2.90  | 20.0 | -0.0426 | 0.9663            |        |
|                | 120 mg    | 57.80              | 47    | -0.61 | 6.785 | -18.0                | -4.00 | -0.10  | 3.00  | 19.0 | -0.6127 | 0.5431            |        |
| Week 8         | Placebo   | 58.06              | 94    | -1.39 | 6.019 | -17.8                | -4.50 | -1.15  | 2.20  | 15.6 | -2.2363 | 0.0277            |        |
|                | 15 mg     | 57.23              | 85    | -0.18 | 6.358 | -18.6                | -3.10 | -0.30  | 3.60  | 20.3 | -0.2627 | 0.7934            |        |
|                | 30 mg     | 56.52              | 89    | 0.54  | 8.234 | -21.8                | -5.10 | 0.50   | 4.50  | 26.2 | 0.6192  | 0.5374            |        |
|                | 60 mg     | 55.30              | 85    | 0.81  | 6.317 | -13.9                | -3.10 | 0.90   | 4.10  | 18.8 | 1.1847  | 0.2395            |        |
|                | 120 mg    | 56.26              | 84    | 0.25  | 6.760 | -17.2                | -3.80 | 0.25   | 3.55  | 22.0 | 0.3389  | 0.7355            |        |
| Week 12        | Placebo   | 57.99              | 94    | -0.01 | 8.176 | -22.8                | -3.70 | -0.75  | 3.30  | 30.7 | -0.0076 | 0.9940            |        |
|                | 15 mg     | 57.23              | 85    | -0.51 | 7.447 | -22.5                | -5.40 | -0.20  | 4.10  | 16.8 | -0.6307 | 0.5300            |        |
|                | 30 mg     | 56.14              | 86    | 0.33  | 7.726 | -19.8                | -5.30 | 0.40   | 4.80  | 21.4 | 0.3936  | 0.6949            |        |
|                | 60 mg     | 55.32              | 80    | 0.97  | 6.853 | -16.1                | -3.25 | 0.80   | 5.55  | 32.8 | 1.2710  | 0.2075            |        |
|                | 120 mg    | 56.16              | 83    | 1.19  | 7.337 | -18.5                | -3.00 | 0.70   | 5.60  | 28.4 | 1.4737  | 0.1444            |        |
| Week 16        | Placebo   | 57.98              | 95    | -0.54 | 6.797 | -22.0                | -4.60 | -1.20  | 3.00  | 32.4 | -0.7713 | 0.4425            |        |
|                | 15 mg     | 57.14              | 83    | 0.64  | 6.916 | -16.4                | -3.50 | -1.20  | 5.40  | 15.8 | 0.8427  | 0.4018            |        |
|                | 30 mg     | 56.09              | 85    | 1.25  | 7.422 | -17.2                | -3.30 | 1.00   | 5.30  | 24.0 | 1.5536  | 0.1240            |        |
|                | 60 mg     | 55.13              | 79    | 0.55  | 5.960 | -12.8                | -3.30 | 0.50   | 3.00  | 27.0 | 0.8250  | 0.4119            |        |
|                | 120 mg    | 56.04              | 82    | 0.36  | 7.282 | -22.1                | -3.10 | 0.00   | 4.50  | 29.0 | 0.4535  | 0.6514            |        |
| Week 20        | Placebo   | 57.98              | 95    | -0.92 | 6.768 | -29.5                | -4.00 | -0.80  | 3.60  | 23.5 | -1.3280 | 0.1874            |        |
|                | 15 mg     | 57.14              | 83    | -0.46 | 7.084 | -20.2                | -4.60 | 0.30   | 4.60  | 15.2 | -0.5904 | 0.5566            |        |
|                | 30 mg     | 56.09              | 85    | -0.11 | 7.271 | -16.2                | -5.20 | -0.10  | 5.20  | 16.1 | -0.1372 | 0.8912            |        |
|                | 60 mg     | 54.78              | 77    | 0.95  | 6.235 | -14.2                | -3.20 | 1.20   | 5.30  | 15.3 | 1.3342  | 0.1861            |        |
|                | 120 mg    | 56.37              | 78    | 0.44  | 7.267 | -23.5                | -3.40 | 0.50   | 3.30  | 24.1 | 0.5360  | 0.5935            |        |
| Week 24        | Placebo   | 57.98              | 93    | -0.04 | 8.542 | -27.1                | -4.20 | 0.00   | 4.80  | 19.4 | -0.0473 | 0.9623            |        |
|                | 15 mg     | 57.14              | 83    | -0.24 | 6.645 | -22.8                | -4.00 | 0.40   | 3.20  | 15.4 | -0.3287 | 0.7432            |        |
|                | 30 mg     | 56.17              | 84    | 0.95  | 8.073 | -22.0                | -3.40 | 1.25   | 6.45  | 24.7 | 1.0744  | 0.2858            |        |
|                | 60 mg     | 55.08              | 77    | -0.03 | 6.721 | -17.6                | -4.40 | 0.50   | 5.00  | 18.7 | -0.0339 | 0.9730            |        |
|                | 120 mg    | 56.09              | 79    | -0.44 | 7.766 | -24.4                | -4.70 | 0.00   | 4.20  | 20.3 | -0.4984 | 0.6196            |        |
| Week 28        | Placebo   | 58.01              | 94    | -0.80 | 6.526 | -18.0                | -4.40 | -0.50  | 3.00  | 15.9 | -1.1853 | 0.2389            |        |
|                | 15 mg     | 57.09              | 82    | -0.79 | 7.081 | -25.0                | -5.50 | -0.15  | 3.80  | 13.4 | -1.0059 | 0.3175            |        |
|                | 30 mg     | 56.17              | 84    | 0.58  | 6.777 | -20.4                | -3.15 | -0.40  | 5.80  | 14.9 | 0.7873  | 0.4334            |        |
|                | 60 mg     | 55.25              | 76    | 0.10  | 5.723 | -13.3                | -3.55 | -0.10  | 2.80  | 14.0 | 0.1563  | 0.8762            |        |
|                | 120 mg    | 56.08              | 78    | 0.24  | 6.900 | -16.8                | -3.80 | 0.45   | 3.40  | 16.1 | 0.3085  | 0.7585            |        |
| Follow-up      | Placebo   | 57.99              | 98    | -1.08 | 6.747 | -18.0                | -4.50 | -0.60  | 3.00  | 15.9 | -1.5840 | 0.1164            |        |
|                | 15 mg     | 56.94              | 85    | -0.40 | 7.190 | -25.0                | -5.40 | 0.40   | 4.50  | 13.4 | -0.5129 | 0.6094            |        |
|                | 30 mg     | 56.72              | 92    | 0.50  | 6.871 | -20.4                | -3.50 | -0.40  | 5.80  | 14.9 | 0.6949  | 0.4889            |        |
|                | 60 mg     | 56.00              | 86    | 0.22  | 5.802 | -13.3                | -3.50 | 0.10   | 3.00  | 14.0 | 0.3438  | 0.7318            |        |
|                | 120 mg    | 56.56              | 88    | 0.09  | 6.736 | -16.8                | -3.70 | 0.60   | 3.35  | 16.1 | 0.1282  | 0.8983            |        |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (6/14)

| Item           | Treatment | Summary Statistics |                      |       |       |       |       |        |       |      |         | One-sample t-test |        |
|----------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|------|---------|-------------------|--------|
|                |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       |      | t-value | p-value           |        |
|                |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    | Max  |         |                   |        |
| Lymphocyte (%) | Week 1    | Placebo            | 33.03                | 39    | 1.08  | 4.813 | -6.5  | -2.30  | 0.50  | 4.90 | 12.5    | 1.3974            | 0.1704 |
|                |           | 15 mg              | 35.51                | 37    | -0.53 | 4.652 | -12.0 | -2.60  | 0.10  | 2.10 | 8.1     | -0.6962           | 0.4908 |
|                |           | 30 mg              | 35.97                | 39    | -0.27 | 5.900 | -17.9 | -3.10  | 0.10  | 3.50 | 9.2     | -0.2877           | 0.7751 |
|                |           | 60 mg              | 35.89                | 35    | -0.85 | 5.095 | -17.2 | -3.80  | -0.20 | 2.70 | 8.1     | -0.9920           | 0.3282 |
|                |           | 120 mg             | 34.36                | 47    | 0.26  | 6.605 | -14.0 | -2.90  | -0.20 | 3.30 | 18.6    | 0.2672            | 0.7905 |
|                | Week 2    | Placebo            | 33.62                | 97    | 0.89  | 6.143 | -22.9 | -2.60  | 1.60  | 4.10 | 16.1    | 1.4314            | 0.1556 |
|                |           | 15 mg              | 34.89                | 88    | -0.13 | 6.835 | -23.6 | -2.45  | -0.35 | 3.80 | 17.6    | -0.1778           | 0.8593 |
|                |           | 30 mg              | 34.51                | 90    | -0.15 | 7.141 | -19.3 | -4.20  | -0.60 | 3.60 | 20.0    | -0.1993           | 0.8425 |
|                |           | 60 mg              | 35.74                | 91    | 0.11  | 4.795 | -10.1 | -3.50  | 0.20  | 3.00 | 12.7    | 0.2230            | 0.8241 |
|                |           | 120 mg             | 35.47                | 87    | -0.21 | 6.281 | -23.5 | -3.80  | -0.20 | 3.00 | 19.9    | -0.3175           | 0.7516 |
|                | Week 3    | Placebo            | 33.04                | 37    | 2.54  | 4.186 | -3.0  | -0.40  | 1.70  | 4.20 | 15.8    | 3.6959            | 0.0007 |
|                |           | 15 mg              | 35.39                | 36    | -0.23 | 5.482 | -12.5 | -3.30  | -1.00 | 3.85 | 16.1    | -0.2493           | 0.8046 |
|                |           | 30 mg              | 36.14                | 38    | 0.93  | 5.932 | -13.0 | -3.00  | 1.15  | 4.00 | 14.7    | 0.9626            | 0.3420 |
|                |           | 60 mg              | 35.11                | 31    | -0.22 | 4.569 | -9.2  | -3.10  | -1.00 | 2.90 | 10.2    | -0.2712           | 0.7881 |
|                |           | 120 mg             | 33.96                | 42    | 1.17  | 5.360 | -10.2 | -2.30  | 0.95  | 4.60 | 11.4    | 1.4135            | 0.1650 |
|                | Week 4    | Placebo            | 33.58                | 96    | 1.17  | 5.573 | -16.4 | -2.60  | 0.85  | 3.90 | 20.4    | 2.0513            | 0.0430 |
|                |           | 15 mg              | 34.89                | 88    | -0.23 | 7.041 | -22.9 | -4.05  | -0.55 | 3.90 | 18.4    | -0.3119           | 0.7559 |
|                |           | 30 mg              | 34.70                | 89    | 0.68  | 7.341 | -15.6 | -5.00  | -0.60 | 3.60 | 22.6    | 0.8765            | 0.3831 |
|                |           | 60 mg              | 36.09                | 88    | 0.80  | 6.013 | -16.3 | -2.70  | 0.90  | 4.10 | 17.8    | 1.2499            | 0.2147 |
|                |           | 120 mg             | 35.51                | 87    | -0.95 | 7.055 | -20.3 | -4.90  | -1.40 | 3.30 | 19.6    | -1.2552           | 0.2128 |
|                | Week 6    | Placebo            | 33.16                | 40    | 0.75  | 5.911 | -9.2  | -3.55  | -0.20 | 4.25 | 13.8    | 0.8024            | 0.4272 |
|                |           | 15 mg              | 35.33                | 39    | -1.99 | 5.534 | -19.0 | -5.80  | -2.00 | 2.20 | 13.0    | -2.2510           | 0.0303 |
|                |           | 30 mg              | 34.79                | 38    | 2.51  | 6.141 | -11.5 | -1.80  | 1.40  | 8.30 | 16.0    | 2.5201            | 0.0162 |
|                |           | 60 mg              | 36.10                | 30    | -0.09 | 5.746 | -16.3 | -2.10  | -0.50 | 4.00 | 9.9     | -0.0890           | 0.9297 |
|                |           | 120 mg             | 34.40                | 47    | -0.23 | 6.411 | -19.0 | -3.30  | -0.50 | 3.10 | 14.0    | -0.2457           | 0.8070 |
|                | Week 8    | Placebo            | 33.57                | 94    | 1.25  | 5.721 | -15.8 | -1.50  | 1.35  | 4.20 | 16.2    | 2.1167            | 0.0370 |
|                |           | 15 mg              | 34.84                | 85    | 0.37  | 6.098 | -17.0 | -3.10  | 0.10  | 3.50 | 17.1    | 0.5532            | 0.5816 |
|                |           | 30 mg              | 34.70                | 89    | -0.24 | 7.202 | -18.6 | -4.30  | -1.40 | 4.00 | 19.9    | -0.3179           | 0.7513 |
|                |           | 60 mg              | 36.39                | 85    | -0.48 | 6.179 | -19.4 | -4.10  | -0.60 | 3.30 | 15.5    | -0.7232           | 0.4716 |
|                |           | 120 mg             | 35.61                | 84    | -0.70 | 6.318 | -19.6 | -3.65  | -0.60 | 3.10 | 15.5    | -1.0086           | 0.3161 |
|                | Week 12   | Placebo            | 33.63                | 94    | 0.40  | 7.148 | -26.5 | -2.90  | 1.80  | 3.90 | 19.1    | 0.5397            | 0.5907 |
|                |           | 15 mg              | 34.84                | 85    | 0.54  | 7.125 | -16.9 | -3.60  | 0.50  | 4.90 | 20.5    | 0.6957            | 0.4885 |
|                |           | 30 mg              | 35.07                | 86    | 0.04  | 6.956 | -20.2 | -5.00  | -0.55 | 4.60 | 15.8    | 0.0574            | 0.9544 |
|                |           | 60 mg              | 36.34                | 80    | -0.69 | 6.269 | -29.3 | -4.40  | -0.55 | 2.35 | 17.3    | -0.9773           | 0.3314 |
|                |           | 120 mg             | 35.73                | 83    | -1.37 | 6.714 | -24.4 | -4.80  | -1.00 | 2.60 | 16.0    | -1.8540           | 0.0673 |
|                | Week 16   | Placebo            | 33.63                | 95    | 0.52  | 6.250 | -30.0 | -2.80  | 1.10  | 4.90 | 15.9    | 0.8176            | 0.4157 |
|                |           | 15 mg              | 34.92                | 83    | -0.60 | 6.828 | -22.0 | -5.00  | 0.90  | 3.00 | 16.9    | -0.8005           | 0.4257 |
|                |           | 30 mg              | 35.09                | 85    | -0.78 | 6.782 | -22.0 | -3.60  | -0.40 | 2.80 | 16.3    | -1.0636           | 0.2906 |
|                |           | 60 mg              | 36.53                | 79    | -0.57 | 5.924 | -30.0 | -3.20  | -0.40 | 2.60 | 14.1    | -0.8603           | 0.3923 |
|                |           | 120 mg             | 35.87                | 82    | -0.73 | 6.123 | -20.4 | -4.50  | -0.50 | 3.10 | 18.2    | -1.0785           | 0.2840 |
|                | Week 20   | Placebo            | 33.63                | 95    | 0.89  | 6.026 | -21.5 | -3.20  | 1.20  | 4.10 | 23.9    | 1.4352            | 0.1545 |
|                |           | 15 mg              | 34.92                | 83    | 0.70  | 7.230 | -18.4 | -4.00  | 0.10  | 4.10 | 21.5    | 0.8775            | 0.3828 |
|                |           | 30 mg              | 35.09                | 85    | -0.02 | 6.322 | -17.0 | -4.30  | 0.00  | 4.60 | 14.7    | -0.0326           | 0.9741 |
|                |           | 60 mg              | 36.81                | 77    | -0.88 | 5.840 | -14.9 | -4.20  | -0.50 | 3.00 | 13.0    | -1.3288           | 0.1879 |
|                |           | 120 mg             | 35.55                | 78    | -0.57 | 6.783 | -22.8 | -3.10  | -0.10 | 3.40 | 20.1    | -0.7461           | 0.4579 |
|                | Week 24   | Placebo            | 33.58                | 93    | 0.19  | 7.986 | -17.5 | -4.20  | -0.30 | 4.10 | 28.3    | 0.2259            | 0.8217 |
|                |           | 15 mg              | 34.92                | 83    | 0.29  | 6.569 | -14.1 | -2.90  | 0.00  | 3.80 | 21.9    | 0.3977            | 0.6919 |
|                |           | 30 mg              | 35.03                | 84    | -0.43 | 7.197 | -18.6 | -5.20  | -0.80 | 4.15 | 19.6    | -0.5442           | 0.5877 |
| 60 mg          |           | 36.53              | 77                   | 0.30  | 6.260 | -14.4 | -3.90 | -0.20  | 4.10  | 21.6 | 0.4169  | 0.6779            |        |
| 120 mg         |           | 35.76              | 79                   | 0.05  | 6.767 | -18.5 | -4.40 | 0.60   | 4.10  | 21.0 | 0.0698  | 0.9445            |        |
| Week 28        | Placebo   | 33.57              | 94                   | 0.60  | 6.056 | -15.0 | -3.30 | 0.30   | 4.30  | 15.1 | 0.9656  | 0.3367            |        |
|                | 15 mg     | 34.95              | 82                   | 1.03  | 6.853 | -13.0 | -4.10 | 0.80   | 5.50  | 24.8 | 1.3568  | 0.1786            |        |
|                | 30 mg     | 35.03              | 84                   | -0.30 | 5.878 | -13.1 | -4.05 | -0.85  | 2.70  | 17.6 | -0.4641 | 0.6438            |        |
|                | 60 mg     | 36.39              | 76                   | -0.20 | 5.424 | -16.5 | -2.60 | 0.05   | 3.50  | 11.6 | -0.3257 | 0.7456            |        |
|                | 120 mg    | 35.77              | 78                   | -0.54 | 6.655 | -20.1 | -3.30 | -0.65  | 2.50  | 14.3 | -0.7213 | 0.4729            |        |
| Follow-up      | Placebo   | 33.61              | 98                   | 0.84  | 6.170 | -15.0 | -3.30 | 0.35   | 4.50  | 15.1 | 1.3426  | 0.1825            |        |
|                | 15 mg     | 34.99              | 85                   | 0.74  | 6.904 | -13.0 | -4.30 | 0.60   | 5.20  | 24.8 | 0.9834  | 0.3282            |        |
|                | 30 mg     | 34.53              | 92                   | -0.25 | 6.082 | -13.1 | -4.05 | -0.65  | 2.70  | 17.6 | -0.4011 | 0.6893            |        |
|                | 60 mg     | 35.71              | 86                   | -0.21 | 5.499 | -16.5 | -2.80 | 0.05   | 3.50  | 11.6 | -0.3608 | 0.7191            |        |
|                | 120 mg    | 35.39              | 88                   | -0.43 | 6.455 | -20.1 | -3.20 | -0.60  | 2.40  | 14.3 | -0.6210 | 0.5362            |        |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (7/14)

| Item         | Treatment | Summary Statistics |                      |       |       |       |       |        |       |      |         | One-sample t-test |        |
|--------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|------|---------|-------------------|--------|
|              |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       |      | t-value | p-value           |        |
|              |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    | Max  |         |                   |        |
| Monocyte (%) | Week 1    | Placebo            | 4.97                 | 39    | -0.01 | 0.918 | -1.6  | -0.80  | -0.20 | 0.70 | 2.0     | -0.0698           | 0.9448 |
|              |           | 15 mg              | 4.85                 | 37    | 0.21  | 1.722 | -3.3  | -0.70  | 0.30  | 1.00 | 4.9     | 0.7255            | 0.4729 |
|              |           | 30 mg              | 5.25                 | 39    | 0.04  | 1.806 | -4.0  | -0.90  | 0.00  | 0.90 | 4.2     | 0.1241            | 0.9019 |
|              |           | 60 mg              | 4.61                 | 35    | 0.34  | 0.819 | -1.3  | -0.20  | 0.20  | 0.90 | 2.2     | 2.4367            | 0.0202 |
|              |           | 120 mg             | 4.73                 | 47    | 0.19  | 1.149 | -3.6  | -0.30  | 0.10  | 1.00 | 2.3     | 1.1423            | 0.2592 |
|              | Week 2    | Placebo            | 5.28                 | 97    | 0.10  | 1.503 | -3.6  | -0.70  | 0.10  | 0.90 | 5.6     | 0.6757            | 0.5009 |
|              |           | 15 mg              | 5.12                 | 88    | -0.19 | 1.700 | -7.9  | -0.70  | 0.00  | 0.65 | 3.1     | -1.0596           | 0.2923 |
|              |           | 30 mg              | 5.19                 | 90    | 0.08  | 1.506 | -5.1  | -0.70  | 0.00  | 1.00 | 4.3     | 0.5250            | 0.6009 |
|              |           | 60 mg              | 5.05                 | 91    | 0.13  | 1.239 | -5.1  | -0.50  | 0.00  | 0.70 | 4.5     | 1.0154            | 0.3126 |
|              |           | 120 mg             | 4.83                 | 87    | 0.13  | 1.508 | -6.0  | -0.40  | 0.30  | 1.00 | 4.3     | 0.8247            | 0.4118 |
|              | Week 3    | Placebo            | 5.02                 | 37    | -0.04 | 1.055 | -3.5  | -0.40  | 0.00  | 0.50 | 2.4     | -0.2181           | 0.8286 |
|              |           | 15 mg              | 4.86                 | 36    | -0.13 | 1.421 | -4.3  | -0.85  | -0.05 | 0.60 | 3.5     | -0.5395           | 0.5930 |
|              |           | 30 mg              | 5.26                 | 38    | -0.20 | 1.336 | -3.2  | -0.80  | 0.00  | 0.70 | 2.0     | -0.9108           | 0.3683 |
|              |           | 60 mg              | 4.65                 | 31    | 0.19  | 0.735 | -1.6  | -0.30  | 0.30  | 0.60 | 1.6     | 1.4178            | 0.1665 |
|              |           | 120 mg             | 4.70                 | 42    | 0.16  | 1.826 | -7.0  | -0.50  | 0.00  | 1.30 | 4.8     | 0.5830            | 0.5631 |
|              | Week 4    | Placebo            | 5.28                 | 96    | -0.37 | 1.400 | -4.0  | -1.00  | -0.40 | 0.40 | 3.1     | -2.6176           | 0.0103 |
|              |           | 15 mg              | 5.12                 | 88    | -0.21 | 1.631 | -6.0  | -0.80  | 0.00  | 0.80 | 4.0     | -1.1828           | 0.2401 |
|              |           | 30 mg              | 5.22                 | 89    | -0.07 | 1.278 | -2.8  | -0.80  | -0.20 | 0.60 | 4.0     | -0.4893           | 0.6258 |
|              |           | 60 mg              | 5.12                 | 88    | -0.14 | 1.391 | -4.3  | -0.80  | -0.15 | 0.75 | 3.0     | -0.9119           | 0.3643 |
|              |           | 120 mg             | 4.81                 | 87    | 0.15  | 1.530 | -3.8  | -0.60  | 0.10  | 0.90 | 5.1     | 0.9042            | 0.3684 |
|              | Week 6    | Placebo            | 5.03                 | 40    | 0.10  | 1.021 | -1.5  | -0.80  | -0.10 | 0.70 | 2.9     | 0.6350            | 0.5292 |
|              |           | 15 mg              | 4.98                 | 39    | -0.10 | 1.520 | -6.1  | -0.70  | 0.00  | 1.00 | 2.1     | -0.4215           | 0.6758 |
|              |           | 30 mg              | 5.31                 | 38    | -0.28 | 1.270 | -3.4  | -1.00  | -0.30 | 0.60 | 2.9     | -1.3415           | 0.1879 |
|              |           | 60 mg              | 4.67                 | 30    | 0.25  | 0.980 | -2.1  | -0.20  | 0.30  | 0.80 | 2.1     | 1.3785            | 0.1786 |
|              |           | 120 mg             | 4.61                 | 47    | 0.70  | 1.927 | -2.0  | -0.40  | 0.50  | 1.20 | 10.0    | 2.4906            | 0.0164 |
|              | Week 8    | Placebo            | 5.26                 | 94    | -0.16 | 1.351 | -5.3  | -0.80  | -0.15 | 0.60 | 3.4     | -1.1296           | 0.2615 |
|              |           | 15 mg              | 5.06                 | 85    | -0.20 | 1.433 | -6.0  | -0.80  | 0.00  | 0.40 | 5.0     | -1.2720           | 0.2069 |
|              |           | 30 mg              | 5.22                 | 89    | 0.06  | 1.686 | -5.2  | -0.80  | 0.00  | 1.00 | 5.0     | 0.3522            | 0.7255 |
|              |           | 60 mg              | 5.15                 | 85    | -0.12 | 1.258 | -4.6  | -0.80  | 0.10  | 0.70 | 3.0     | -0.9055           | 0.3678 |
|              |           | 120 mg             | 4.82                 | 84    | 0.42  | 1.373 | -3.3  | -0.30  | 0.50  | 1.00 | 4.0     | 2.7897            | 0.0065 |
|              | Week 12   | Placebo            | 5.26                 | 94    | -0.29 | 1.602 | -6.0  | -1.10  | -0.20 | 0.70 | 3.3     | -1.7579           | 0.0821 |
|              |           | 15 mg              | 5.06                 | 85    | -0.08 | 1.115 | -5.0  | -0.50  | -0.10 | 0.70 | 2.1     | -0.6711           | 0.5040 |
|              |           | 30 mg              | 5.20                 | 86    | -0.15 | 1.224 | -4.1  | -0.70  | -0.10 | 0.60 | 2.2     | -1.1277           | 0.2626 |
|              |           | 60 mg              | 5.17                 | 80    | -0.04 | 1.457 | -4.5  | -0.80  | 0.00  | 0.80 | 5.9     | -0.2609           | 0.7948 |
|              |           | 120 mg             | 4.80                 | 83    | 0.10  | 1.396 | -4.6  | -0.50  | 0.10  | 0.80 | 3.2     | 0.6368            | 0.5260 |
|              | Week 16   | Placebo            | 5.27                 | 95    | -0.24 | 1.605 | -5.2  | -1.00  | -0.30 | 0.80 | 3.8     | -1.4444           | 0.1519 |
|              |           | 15 mg              | 5.09                 | 83    | -0.08 | 1.327 | -4.2  | -0.80  | 0.10  | 0.60 | 3.2     | -0.5709           | 0.5697 |
|              |           | 30 mg              | 5.21                 | 85    | -0.13 | 1.397 | -4.0  | -1.00  | 0.00  | 0.70 | 6.0     | -0.8382           | 0.4043 |
|              |           | 60 mg              | 5.18                 | 79    | 0.06  | 1.101 | -2.7  | -0.70  | 0.10  | 0.70 | 3.8     | 0.4598            | 0.6469 |
|              |           | 120 mg             | 4.82                 | 82    | 0.20  | 1.520 | -6.5  | -0.50  | 0.20  | 1.00 | 5.0     | 1.1988            | 0.2341 |
|              | Week 20   | Placebo            | 5.27                 | 95    | -0.10 | 1.511 | -4.0  | -0.90  | 0.00  | 0.70 | 4.3     | -0.6721           | 0.5032 |
|              |           | 15 mg              | 5.09                 | 83    | -0.14 | 1.550 | -7.0  | -0.90  | -0.10 | 0.50 | 5.0     | -0.8425           | 0.4019 |
|              |           | 30 mg              | 5.21                 | 85    | 0.17  | 1.297 | -3.0  | -0.60  | 0.10  | 0.80 | 3.6     | 1.2379            | 0.2192 |
|              |           | 60 mg              | 5.21                 | 77    | 0.01  | 1.207 | -2.9  | -0.60  | 0.00  | 0.70 | 2.8     | 0.0661            | 0.9475 |
|              |           | 120 mg             | 4.84                 | 78    | 0.03  | 1.351 | -4.6  | -0.70  | 0.10  | 1.00 | 2.7     | 0.1844            | 0.8542 |
|              | Week 24   | Placebo            | 5.26                 | 93    | -0.27 | 1.472 | -4.6  | -1.00  | -0.10 | 0.70 | 3.3     | -1.7681           | 0.0804 |
|              |           | 15 mg              | 5.09                 | 83    | -0.14 | 1.491 | -7.0  | -0.80  | 0.00  | 0.50 | 5.0     | -0.8614           | 0.3916 |
|              |           | 30 mg              | 5.21                 | 84    | -0.08 | 1.491 | -5.0  | -0.85  | -0.05 | 0.60 | 5.0     | -0.4682           | 0.6409 |
| 60 mg        |           | 5.22               | 77                   | -0.09 | 1.254 | -3.4  | -0.70 | -0.10  | 0.70  | 5.8  | -0.6362 | 0.5266            |        |
|              | 120 mg    | 4.87               | 79                   | 0.24  | 1.324 | -2.7  | -0.50 | 0.20   | 1.00  | 3.3  | 1.5977  | 0.1142            |        |
| Week 28      | Placebo   | 5.26               | 94                   | -0.02 | 1.483 | -4.1  | -0.90 | 0.10   | 0.80  | 3.7  | -0.1182 | 0.9062            |        |
|              | 15 mg     | 5.10               | 82                   | -0.20 | 1.317 | -5.0  | -0.90 | 0.00   | 0.70  | 3.7  | -1.3998 | 0.1654            |        |
|              | 30 mg     | 5.21               | 84                   | -0.04 | 1.644 | -6.1  | -0.90 | 0.00   | 0.95  | 5.4  | -0.2124 | 0.8323            |        |
|              | 60 mg     | 5.22               | 76                   | 0.21  | 1.288 | -3.0  | -0.50 | 0.30   | 0.80  | 4.9  | 1.4517  | 0.1508            |        |
|              | 120 mg    | 4.87               | 78                   | 0.06  | 1.367 | -4.6  | -0.60 | 0.10   | 0.60  | 3.5  | 0.3645  | 0.7165            |        |
| Follow-up    | Placebo   | 5.29               | 98                   | -0.03 | 1.481 | -4.1  | -0.90 | 0.10   | 0.80  | 3.7  | -0.1979 | 0.8436            |        |
|              | 15 mg     | 5.16               | 85                   | -0.26 | 1.382 | -5.0  | -0.90 | -0.10  | 0.60  | 3.7  | -1.7659 | 0.0810            |        |
|              | 30 mg     | 5.20               | 92                   | 0.01  | 1.703 | -6.1  | -0.90 | 0.00   | 1.00  | 5.4  | 0.0612  | 0.9513            |        |
|              | 60 mg     | 5.11               | 86                   | 0.19  | 1.260 | -3.0  | -0.60 | 0.25   | 0.80  | 4.9  | 1.3697  | 0.1744            |        |
|              | 120 mg    | 4.82               | 88                   | 0.10  | 1.331 | -4.6  | -0.50 | 0.15   | 0.70  | 3.5  | 0.7049  | 0.4827            |        |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (8/14)

| Item           | Treatment | Summary Statistics |                      |       |       |       |       |        |       |      |         | One-sample t-test |        |
|----------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|------|---------|-------------------|--------|
|                |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       |      | t-value | p-value           |        |
|                |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    | Max  |         |                   |        |
| Eosinophil (%) | Week 1    | Placebo            | 2.83                 | 39    | 0.45  | 1.089 | -1.2  | -0.20  | 0.30  | 0.80 | 3.8     | 2.5576            | 0.0147 |
|                |           | 15 mg              | 2.59                 | 37    | 0.14  | 1.117 | -2.1  | -0.30  | -0.10 | 0.50 | 4.0     | 0.7801            | 0.4404 |
|                |           | 30 mg              | 2.79                 | 39    | 0.09  | 1.386 | -3.0  | -0.60  | -0.10 | 0.50 | 6.5     | 0.4158            | 0.6799 |
|                |           | 60 mg              | 2.88                 | 35    | -0.21 | 1.518 | -7.5  | -0.50  | -0.10 | 0.40 | 2.8     | -0.8018           | 0.4283 |
|                |           | 120 mg             | 2.67                 | 47    | -0.33 | 1.028 | -4.3  | -0.70  | -0.20 | 0.10 | 2.0     | -2.1710           | 0.0351 |
|                | Week 2    | Placebo            | 2.65                 | 97    | 0.17  | 1.335 | -3.1  | -0.60  | 0.00  | 0.70 | 4.0     | 1.2849            | 0.2019 |
|                |           | 15 mg              | 2.44                 | 88    | -0.20 | 1.194 | -4.2  | -0.60  | -0.05 | 0.35 | 3.6     | -1.5354           | 0.1283 |
|                |           | 30 mg              | 3.01                 | 90    | -0.36 | 1.603 | -9.0  | -0.70  | -0.10 | 0.30 | 2.7     | -2.1233           | 0.0365 |
|                |           | 60 mg              | 2.65                 | 91    | -0.20 | 1.011 | -3.0  | -0.80  | -0.10 | 0.30 | 3.3     | -1.8766           | 0.0638 |
|                |           | 120 mg             | 2.72                 | 87    | -0.08 | 1.182 | -5.5  | -0.50  | 0.00  | 0.60 | 2.6     | -0.6441           | 0.5212 |
|                | Week 3    | Placebo            | 2.61                 | 37    | 0.31  | 1.174 | -3.7  | -0.10  | 0.20  | 0.90 | 2.8     | 1.5830            | 0.1222 |
|                |           | 15 mg              | 2.60                 | 36    | 0.09  | 1.376 | -4.3  | -0.20  | 0.15  | 0.65 | 4.0     | 0.4117            | 0.6831 |
|                |           | 30 mg              | 2.74                 | 38    | 0.01  | 1.676 | -5.0  | -0.70  | -0.20 | 0.70 | 6.3     | 0.0290            | 0.9770 |
|                |           | 60 mg              | 3.00                 | 31    | 0.02  | 0.994 | -1.7  | -0.60  | -0.30 | 0.60 | 2.6     | 0.1265            | 0.9002 |
|                |           | 120 mg             | 2.60                 | 42    | 0.02  | 0.992 | -2.7  | -0.60  | 0.00  | 0.80 | 2.0     | 0.1244            | 0.9016 |
|                | Week 4    | Placebo            | 2.65                 | 96    | 0.15  | 1.278 | -3.5  | -0.55  | 0.20  | 0.70 | 4.0     | 1.1422            | 0.2563 |
|                |           | 15 mg              | 2.44                 | 88    | -0.03 | 1.302 | -3.8  | -0.60  | 0.00  | 0.50 | 5.0     | -0.2047           | 0.8383 |
|                |           | 30 mg              | 3.04                 | 89    | -0.25 | 1.429 | -5.5  | -0.70  | -0.20 | 0.60 | 4.1     | -1.6318           | 0.1063 |
|                |           | 60 mg              | 2.58                 | 88    | -0.04 | 1.272 | -4.1  | -0.80  | 0.00  | 0.55 | 4.0     | -0.3184           | 0.7509 |
|                |           | 120 mg             | 2.69                 | 87    | -0.02 | 1.207 | -3.7  | -0.50  | -0.10 | 0.60 | 2.8     | -0.1777           | 0.8594 |
|                | Week 6    | Placebo            | 2.86                 | 40    | 0.07  | 1.494 | -3.7  | -0.35  | 0.10  | 0.70 | 3.9     | 0.2751            | 0.7847 |
|                |           | 15 mg              | 2.66                 | 39    | 0.07  | 1.823 | -3.5  | -0.70  | 0.10  | 0.60 | 5.1     | 0.2371            | 0.8138 |
|                |           | 30 mg              | 2.88                 | 38    | 0.02  | 1.525 | -3.1  | -0.70  | -0.05 | 0.70 | 6.5     | 0.0957            | 0.9242 |
|                |           | 60 mg              | 3.00                 | 30    | -0.05 | 1.225 | -3.1  | -0.50  | 0.05  | 0.40 | 3.8     | -0.2385           | 0.8132 |
|                |           | 120 mg             | 2.64                 | 47    | 0.18  | 1.126 | -1.8  | -0.50  | 0.10  | 0.60 | 4.2     | 1.0880            | 0.2823 |
|                | Week 8    | Placebo            | 2.62                 | 94    | 0.25  | 1.299 | -3.3  | -0.20  | 0.15  | 0.70 | 4.3     | 1.8971            | 0.0609 |
|                |           | 15 mg              | 2.45                 | 85    | 0.01  | 1.157 | -4.3  | -0.50  | 0.00  | 0.60 | 3.9     | 0.0750            | 0.9404 |
|                |           | 30 mg              | 3.04                 | 89    | -0.33 | 2.027 | -7.3  | -0.80  | -0.20 | 0.60 | 8.7     | -1.5273           | 0.1303 |
|                |           | 60 mg              | 2.60                 | 85    | -0.13 | 1.228 | -5.5  | -0.60  | -0.20 | 0.40 | 3.3     | -1.0070           | 0.3168 |
|                |           | 120 mg             | 2.77                 | 84    | 0.01  | 1.199 | -3.4  | -0.75  | 0.00  | 0.65 | 2.6     | 0.1001            | 0.9205 |
|                | Week 12   | Placebo            | 2.63                 | 94    | -0.07 | 1.529 | -6.6  | -0.70  | 0.00  | 0.70 | 3.2     | -0.4386           | 0.6620 |
|                |           | 15 mg              | 2.45                 | 85    | -0.01 | 1.138 | -3.2  | -0.70  | 0.00  | 0.50 | 4.0     | -0.1048           | 0.9168 |
|                |           | 30 mg              | 3.06                 | 86    | -0.22 | 1.987 | -6.0  | -0.80  | -0.20 | 0.40 | 8.4     | -1.0367           | 0.3028 |
|                |           | 60 mg              | 2.61                 | 80    | -0.17 | 1.248 | -3.5  | -0.75  | -0.10 | 0.30 | 4.6     | -1.1826           | 0.2405 |
|                |           | 120 mg             | 2.78                 | 83    | 0.05  | 1.177 | -3.3  | -0.70  | 0.10  | 0.70 | 5.1     | 0.3636            | 0.7171 |
|                | Week 16   | Placebo            | 2.64                 | 95    | 0.26  | 1.326 | -5.8  | -0.40  | 0.40  | 0.80 | 4.1     | 1.9344            | 0.0561 |
|                |           | 15 mg              | 2.43                 | 83    | 0.01  | 1.128 | -2.7  | -0.70  | 0.00  | 0.60 | 5.0     | 0.0584            | 0.9536 |
|                |           | 30 mg              | 3.08                 | 85    | -0.30 | 1.837 | -7.2  | -0.60  | -0.10 | 0.30 | 7.7     | -1.5176           | 0.1329 |
|                |           | 60 mg              | 2.59                 | 79    | 0.02  | 1.015 | -3.2  | -0.40  | 0.00  | 0.50 | 3.8     | 0.1330            | 0.8945 |
|                |           | 120 mg             | 2.73                 | 82    | 0.14  | 1.378 | -4.0  | -0.50  | 0.30  | 1.00 | 3.4     | 0.9380            | 0.3511 |
|                | Week 20   | Placebo            | 2.64                 | 95    | 0.09  | 1.292 | -4.3  | -0.40  | 0.10  | 0.70 | 3.6     | 0.7066            | 0.4816 |
|                |           | 15 mg              | 2.43                 | 83    | -0.11 | 1.065 | -3.3  | -0.70  | -0.10 | 0.50 | 2.8     | -0.9172           | 0.3617 |
|                |           | 30 mg              | 3.08                 | 85    | -0.05 | 3.021 | -8.2  | -0.70  | -0.10 | 0.50 | 18.4    | -0.1580           | 0.8749 |
|                |           | 60 mg              | 2.62                 | 77    | 0.03  | 1.286 | -4.3  | -0.50  | -0.10 | 0.60 | 4.2     | 0.1950            | 0.8459 |
|                |           | 120 mg             | 2.70                 | 78    | 0.10  | 1.161 | -4.1  | -0.50  | 0.10  | 0.90 | 2.5     | 0.7510            | 0.4550 |
|                | Week 24   | Placebo            | 2.69                 | 93    | 0.14  | 1.758 | -7.1  | -0.50  | 0.10  | 0.60 | 9.7     | 0.7786            | 0.4382 |
|                |           | 15 mg              | 2.43                 | 83    | 0.05  | 1.121 | -2.2  | -0.70  | -0.10 | 0.80 | 3.6     | 0.3722            | 0.7107 |
|                |           | 30 mg              | 3.06                 | 84    | -0.40 | 1.881 | -8.7  | -1.00  | -0.25 | 0.60 | 5.0     | -1.9547           | 0.0540 |
| 60 mg          |           | 2.59               | 77                   | -0.12 | 1.578 | -4.5  | -0.70 | -0.10  | 0.40  | 5.3  | -0.6501 | 0.5176            |        |
| 120 mg         |           | 2.75               | 79                   | 0.08  | 1.386 | -5.4  | -0.50 | 0.10   | 0.90  | 2.7  | 0.5195  | 0.6049            |        |
| Week 28        | Placebo   | 2.67               | 94                   | 0.19  | 1.465 | -3.7  | -0.70 | 0.30   | 1.00  | 4.1  | 1.2460  | 0.2159            |        |
|                | 15 mg     | 2.44               | 82                   | -0.08 | 1.075 | -3.9  | -0.70 | -0.10  | 0.40  | 2.5  | -0.6578 | 0.5126            |        |
|                | 30 mg     | 3.06               | 84                   | -0.25 | 1.997 | -9.0  | -0.75 | -0.10  | 0.75  | 5.3  | -1.1473 | 0.2546            |        |
|                | 60 mg     | 2.59               | 76                   | -0.06 | 1.273 | -3.9  | -0.55 | -0.10  | 0.60  | 5.2  | -0.4144 | 0.6798            |        |
|                | 120 mg    | 2.75               | 78                   | 0.18  | 1.078 | -3.1  | -0.50 | 0.15   | 1.00  | 2.7  | 1.4597  | 0.1484            |        |
| Follow-up      | Placebo   | 2.63               | 98                   | 0.24  | 1.476 | -3.7  | -0.60 | 0.30   | 1.10  | 4.1  | 1.6086  | 0.1110            |        |
|                | 15 mg     | 2.48               | 85                   | -0.11 | 1.105 | -3.9  | -0.70 | -0.10  | 0.40  | 2.5  | -0.9125 | 0.3641            |        |
|                | 30 mg     | 3.02               | 92                   | -0.26 | 1.960 | -9.0  | -0.80 | -0.10  | 0.75  | 5.3  | -1.2499 | 0.2145            |        |
|                | 60 mg     | 2.64               | 86                   | -0.13 | 1.253 | -3.9  | -0.60 | -0.20  | 0.50  | 5.2  | -0.9981 | 0.3211            |        |
|                | 120 mg    | 2.69               | 88                   | 0.17  | 1.064 | -3.1  | -0.50 | 0.10   | 1.00  | 2.7  | 1.5027  | 0.1365            |        |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (9/14)

| Item         | Treatment | Summary Statistics |                      |       |       |       |       |        |       |      |         | One-sample t-test |        |
|--------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|------|---------|-------------------|--------|
|              |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       |      | t-value | p-value           |        |
|              |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    | Max  |         |                   |        |
| Basophil (%) | Week 1    | Placebo            | 0.45                 | 39    | 0.06  | 0.314 | -0.8  | -0.10  | 0.10  | 0.20 | 0.8     | 1.1205            | 0.2695 |
|              |           | 15 mg              | 0.42                 | 37    | 0.05  | 0.521 | -0.6  | -0.20  | 0.00  | 0.10 | 2.0     | 0.5680            | 0.5736 |
|              |           | 30 mg              | 0.51                 | 39    | 0.03  | 0.398 | -1.3  | -0.20  | 0.00  | 0.20 | 1.0     | 0.4826            | 0.6321 |
|              |           | 60 mg              | 0.52                 | 35    | -0.05 | 0.268 | -0.5  | -0.20  | -0.10 | 0.10 | 0.5     | -1.0081           | 0.3205 |
|              |           | 120 mg             | 0.60                 | 47    | -0.12 | 0.377 | -1.1  | -0.30  | -0.10 | 0.10 | 0.6     | -2.1676           | 0.0354 |
|              | Week 2    | Placebo            | 0.48                 | 97    | 0.05  | 0.486 | -2.0  | -0.20  | 0.10  | 0.30 | 1.5     | 1.0449            | 0.2987 |
|              |           | 15 mg              | 0.42                 | 88    | 0.02  | 0.330 | -0.7  | -0.20  | 0.00  | 0.10 | 2.0     | 0.4517            | 0.6526 |
|              |           | 30 mg              | 0.52                 | 90    | -0.02 | 0.393 | -1.7  | -0.10  | 0.00  | 0.20 | 1.3     | -0.5092           | 0.6119 |
|              |           | 60 mg              | 0.54                 | 91    | -0.02 | 0.331 | -0.8  | -0.20  | 0.00  | 0.20 | 1.0     | -0.6335           | 0.5280 |
|              |           | 120 mg             | 0.55                 | 87    | -0.05 | 0.472 | -1.5  | -0.20  | 0.00  | 0.20 | 1.5     | -0.9082           | 0.3663 |
|              | Week 3    | Placebo            | 0.45                 | 37    | 0.03  | 0.313 | -0.8  | -0.10  | 0.00  | 0.20 | 0.7     | 0.6308            | 0.5322 |
|              |           | 15 mg              | 0.43                 | 36    | 0.02  | 0.235 | -0.4  | -0.10  | 0.00  | 0.10 | 0.7     | 0.4960            | 0.6230 |
|              |           | 30 mg              | 0.47                 | 38    | -0.03 | 0.371 | -1.3  | -0.10  | 0.00  | 0.10 | 0.9     | -0.5253           | 0.6025 |
|              |           | 60 mg              | 0.53                 | 31    | -0.01 | 0.263 | -0.6  | -0.20  | 0.00  | 0.20 | 0.6     | -0.2732           | 0.7866 |
|              |           | 120 mg             | 0.57                 | 42    | -0.07 | 0.342 | -1.0  | -0.20  | 0.00  | 0.20 | 0.7     | -1.2636           | 0.2135 |
|              | Week 4    | Placebo            | 0.48                 | 96    | 0.03  | 0.365 | -1.2  | -0.15  | 0.00  | 0.20 | 1.1     | 0.7550            | 0.4521 |
|              |           | 15 mg              | 0.42                 | 88    | 0.02  | 0.297 | -0.6  | -0.10  | 0.00  | 0.10 | 1.0     | 0.7525            | 0.4538 |
|              |           | 30 mg              | 0.52                 | 89    | -0.04 | 0.423 | -1.4  | -0.20  | 0.00  | 0.10 | 1.1     | -0.8774           | 0.3826 |
|              |           | 60 mg              | 0.55                 | 88    | -0.04 | 0.366 | -1.0  | -0.20  | 0.00  | 0.10 | 1.4     | -0.9897           | 0.3251 |
|              |           | 120 mg             | 0.54                 | 87    | 0.04  | 0.434 | -1.2  | -0.20  | 0.00  | 0.20 | 1.5     | 0.7662            | 0.4456 |
|              | Week 6    | Placebo            | 0.47                 | 40    | 0.06  | 0.336 | -1.0  | -0.05  | 0.10  | 0.20 | 0.7     | 1.1303            | 0.2653 |
|              |           | 15 mg              | 0.44                 | 39    | -0.04 | 0.266 | -0.9  | -0.20  | 0.00  | 0.10 | 0.8     | -0.8426           | 0.4047 |
|              |           | 30 mg              | 0.47                 | 38    | 0.05  | 0.361 | -1.0  | -0.10  | 0.00  | 0.40 | 0.7     | 0.8095            | 0.4234 |
|              |           | 60 mg              | 0.58                 | 30    | -0.05 | 0.276 | -0.8  | -0.20  | -0.05 | 0.10 | 0.7     | -0.9249           | 0.3626 |
|              |           | 120 mg             | 0.56                 | 47    | -0.04 | 0.485 | -1.7  | -0.20  | 0.00  | 0.10 | 2.0     | -0.6011           | 0.5507 |
|              | Week 8    | Placebo            | 0.49                 | 94    | 0.04  | 0.332 | -1.0  | -0.10  | 0.00  | 0.20 | 1.2     | 1.2415            | 0.2175 |
|              |           | 15 mg              | 0.42                 | 85    | 0.00  | 0.314 | -1.1  | -0.10  | 0.00  | 0.10 | 1.0     | 0.1036            | 0.9177 |
|              |           | 30 mg              | 0.52                 | 89    | -0.03 | 0.373 | -1.3  | -0.20  | 0.00  | 0.10 | 1.4     | -0.8230           | 0.4127 |
|              |           | 60 mg              | 0.56                 | 85    | -0.07 | 0.341 | -1.0  | -0.20  | 0.00  | 0.10 | 1.1     | -1.8793           | 0.0637 |
|              |           | 120 mg             | 0.54                 | 84    | 0.01  | 0.376 | -1.0  | -0.20  | 0.00  | 0.20 | 1.5     | 0.3484            | 0.7284 |
|              | Week 12   | Placebo            | 0.48                 | 94    | -0.03 | 0.378 | -2.0  | -0.10  | 0.00  | 0.10 | 1.0     | -0.8193           | 0.4147 |
|              |           | 15 mg              | 0.42                 | 85    | 0.06  | 0.306 | -0.8  | -0.10  | 0.10  | 0.20 | 1.0     | 1.8102            | 0.0738 |
|              |           | 30 mg              | 0.52                 | 86    | -0.01 | 0.445 | -2.0  | -0.20  | 0.00  | 0.20 | 1.4     | -0.1211           | 0.9039 |
|              |           | 60 mg              | 0.56                 | 80    | -0.08 | 0.372 | -1.1  | -0.30  | 0.00  | 0.20 | 1.1     | -1.8051           | 0.0749 |
|              |           | 120 mg             | 0.53                 | 83    | 0.01  | 0.432 | -1.2  | -0.20  | 0.00  | 0.20 | 1.8     | 0.2288            | 0.8196 |
|              | Week 16   | Placebo            | 0.48                 | 95    | -0.01 | 0.403 | -2.0  | -0.10  | 0.00  | 0.20 | 0.9     | -0.2803           | 0.7799 |
|              |           | 15 mg              | 0.42                 | 83    | 0.04  | 0.369 | -0.5  | -0.10  | 0.00  | 0.20 | 2.0     | 0.8913            | 0.3754 |
|              |           | 30 mg              | 0.53                 | 85    | -0.04 | 0.374 | -2.0  | -0.20  | 0.00  | 0.20 | 0.7     | -0.9574           | 0.3411 |
|              |           | 60 mg              | 0.56                 | 79    | -0.05 | 0.274 | -1.0  | -0.20  | -0.10 | 0.10 | 0.8     | -1.4807           | 0.1427 |
|              |           | 120 mg             | 0.53                 | 82    | 0.02  | 0.352 | -1.0  | -0.20  | 0.00  | 0.20 | 1.2     | 0.5339            | 0.5949 |
|              | Week 20   | Placebo            | 0.48                 | 95    | -0.01 | 0.276 | -1.0  | -0.10  | 0.00  | 0.10 | 0.9     | -0.4466           | 0.6562 |
|              |           | 15 mg              | 0.42                 | 83    | 0.01  | 0.319 | -1.0  | -0.20  | 0.00  | 0.20 | 1.0     | -0.3782           | 0.7062 |
|              |           | 30 mg              | 0.53                 | 85    | -0.01 | 0.442 | -2.0  | -0.20  | 0.00  | 0.20 | 1.2     | -0.1962           | 0.8449 |
|              |           | 60 mg              | 0.57                 | 77    | -0.11 | 0.417 | -2.0  | -0.30  | -0.10 | 0.10 | 0.7     | -2.2666           | 0.0263 |
|              |           | 120 mg             | 0.54                 | 78    | 0.01  | 0.445 | -2.4  | -0.10  | 0.00  | 0.20 | 1.2     | 0.1018            | 0.9192 |
|              | Week 24   | Placebo            | 0.49                 | 93    | -0.03 | 0.387 | -2.0  | -0.20  | 0.00  | 0.20 | 1.3     | -0.8304           | 0.4085 |
|              |           | 15 mg              | 0.42                 | 83    | 0.05  | 0.328 | -0.7  | -0.10  | 0.00  | 0.20 | 2.0     | 1.3382            | 0.1845 |
|              |           | 30 mg              | 0.53                 | 84    | -0.05 | 0.448 | -2.0  | -0.20  | 0.00  | 0.10 | 1.0     | -1.0956           | 0.2764 |
| 60 mg        |           | 0.56               | 77                   | -0.06 | 0.400 | -2.0  | -0.20 | 0.00   | 0.10  | 1.4  | -1.2521 | 0.2144            |        |
| 120 mg       |           | 0.53               | 79                   | 0.06  | 0.350 | -0.8  | -0.10 | 0.00   | 0.20  | 1.2  | 1.6082  | 0.1118            |        |
| Week 28      | Placebo   | 0.49               | 94                   | 0.02  | 0.329 | -1.1  | -0.20 | 0.00   | 0.20  | 1.0  | 0.7215  | 0.4724            |        |
|              | 15 mg     | 0.43               | 82                   | 0.04  | 0.376 | -0.8  | -0.10 | 0.00   | 0.20  | 1.0  | 0.8511  | 0.3972            |        |
|              | 30 mg     | 0.53               | 84                   | -0.03 | 0.417 | -1.5  | -0.20 | 0.00   | 0.20  | 0.9  | -0.7057 | 0.4823            |        |
|              | 60 mg     | 0.54               | 76                   | -0.05 | 0.348 | -1.5  | -0.20 | 0.00   | 0.20  | 0.5  | -1.1875 | 0.2388            |        |
|              | 120 mg    | 0.53               | 78                   | 0.07  | 0.351 | -1.1  | -0.10 | 0.10   | 0.20  | 1.4  | 1.7076  | 0.0917            |        |
| Follow-up    | Placebo   | 0.48               | 98                   | 0.03  | 0.326 | -1.1  | -0.20 | 0.00   | 0.20  | 1.0  | 0.9903  | 0.3245            |        |
|              | 15 mg     | 0.43               | 85                   | 0.03  | 0.377 | -0.8  | -0.20 | 0.00   | 0.20  | 1.0  | 0.7766  | 0.4396            |        |
|              | 30 mg     | 0.53               | 92                   | -0.03 | 0.421 | -1.5  | -0.20 | 0.00   | 0.20  | 0.9  | -0.7188 | 0.4741            |        |
|              | 60 mg     | 0.53               | 86                   | -0.05 | 0.344 | -1.5  | -0.20 | 0.00   | 0.20  | 0.8  | -1.2554 | 0.2128            |        |
|              | 120 mg    | 0.54               | 88                   | 0.06  | 0.403 | -1.1  | -0.10 | 0.10   | 0.20  | 1.8  | 1.5093  | 0.1348            |        |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (10/14)

| Item                                       | Treatment | Summary Statistics |       |       |       |                      |       |        |       |      | One-sample t-test |         |         |
|--|-----------|--------------------|-------|-------|-------|----------------------|-------|--------|-------|------|-------------------|---------|---------|
|  |           | Baseline Mean      | N     | Mean  | SD    | Change from Baseline |       |        |       |      | t-value           | p-value |         |
|  |           |                    |       |       |       | Min                  | Q1    | Median | Q3    | Max  |                   |         |         |
| Platelets<br>( $\times 10^3/\mu\text{L}$ ) | Week 1    | Placebo            | 25.04 | 38    | 0.28  | 2.458                | -5.9  | -1.20  | 0.85  | 1.30 | 5.5               | 0.6931  | 0.4926  |
|  |           | 15 mg              | 25.88 | 36    | 0.07  | 2.394                | -4.6  | -1.45  | -0.20 | 1.65 | 4.9               | 0.1740  | 0.8628  |
|  |           | 30 mg              | 23.85 | 39    | 0.48  | 1.885                | -4.4  | -0.60  | 0.80  | 1.70 | 4.2               | 1.6052  | 0.1167  |
|  |           | 60 mg              | 23.91 | 37    | 0.00  | 1.841                | -3.2  | -1.30  | -0.10 | 0.80 | 4.9               | 0.0000  | 1.0000  |
|  |           | 120 mg             | 23.61 | 47    | 0.82  | 2.301                | -4.3  | -0.40  | 0.90  | 1.80 | 8.4               | 2.4341  | 0.0189  |
|  | Week 2    | Placebo            | 24.11 | 97    | 0.25  | 2.988                | -5.3  | -1.30  | -0.10 | 1.80 | 11.8              | 0.8121  | 0.4188  |
|  |           | 15 mg              | 24.83 | 87    | 0.46  | 2.519                | -5.7  | -1.00  | 0.50  | 2.10 | 8.9               | 1.7155  | 0.0899  |
|  |           | 30 mg              | 23.77 | 91    | 1.02  | 2.288                | -4.1  | -0.60  | 1.10  | 2.60 | 6.7               | 4.2373  | <0.0001 |
|  |           | 60 mg              | 23.72 | 91    | 0.99  | 2.674                | -8.4  | -0.40  | 1.00  | 2.50 | 13.2              | 3.5480  | 0.0006  |
|  |           | 120 mg             | 24.26 | 88    | 0.46  | 2.172                | -6.5  | -1.00  | 0.60  | 1.90 | 9.2               | 2.0029  | 0.0483  |
|  | Week 3    | Placebo            | 24.29 | 37    | 0.57  | 2.274                | -3.3  | -1.10  | 0.50  | 2.60 | 4.2               | 1.5328  | 0.1341  |
|  |           | 15 mg              | 25.90 | 35    | 0.24  | 2.370                | -4.7  | -0.80  | -0.20 | 1.20 | 7.9               | 0.5921  | 0.5577  |
|  |           | 30 mg              | 23.69 | 38    | 0.96  | 2.874                | -4.5  | -0.50  | 0.35  | 2.60 | 8.9               | 2.0488  | 0.0476  |
|  |           | 60 mg              | 23.58 | 32    | 0.63  | 1.662                | -4.3  | -0.45  | 0.65  | 1.80 | 3.7               | 2.1586  | 0.0387  |
|  |           | 120 mg             | 23.49 | 42    | 0.38  | 2.361                | -6.0  | -1.10  | 0.30  | 1.90 | 5.0               | 1.0456  | 0.3019  |
|  | Week 4    | Placebo            | 24.05 | 96    | 0.18  | 2.442                | -9.8  | -1.10  | 0.20  | 1.75 | 5.5               | 0.7273  | 0.4688  |
|  |           | 15 mg              | 24.85 | 86    | 0.82  | 2.295                | -4.2  | -1.00  | 0.75  | 1.80 | 12.3              | 3.3032  | 0.0014  |
|  |           | 30 mg              | 23.77 | 89    | 0.90  | 2.290                | -4.6  | -0.70  | 0.90  | 2.20 | 9.1               | 3.6985  | 0.0004  |
|  |           | 60 mg              | 23.68 | 87    | 0.29  | 2.299                | -8.8  | -0.90  | 0.30  | 1.80 | 8.5               | 1.1797  | 0.2414  |
|  |           | 120 mg             | 24.33 | 86    | 0.82  | 2.529                | -6.6  | -0.50  | 0.95  | 1.80 | 8.2               | 3.0145  | 0.0034  |
|  | Week 6    | Placebo            | 23.57 | 40    | 0.71  | 2.064                | -3.6  | -0.65  | 0.55  | 2.25 | 4.9               | 2.1758  | 0.0357  |
|  |           | 15 mg              | 24.84 | 39    | 1.46  | 4.970                | -3.9  | -0.70  | 0.40  | 2.30 | 24.5              | 1.8332  | 0.0746  |
|  |           | 30 mg              | 23.61 | 38    | 0.89  | 2.672                | -4.5  | -0.40  | 0.70  | 2.60 | 9.9               | 2.0462  | 0.0479  |
|  |           | 60 mg              | 23.30 | 31    | 1.36  | 2.536                | -5.1  | 0.30   | 1.00  | 2.80 | 6.0               | 2.9892  | 0.0055  |
|  |           | 120 mg             | 22.73 | 47    | 1.01  | 2.351                | -6.0  | -0.60  | 1.20  | 2.10 | 7.1               | 2.9345  | 0.0052  |
|  | Week 8    | Placebo            | 24.16 | 94    | -0.14 | 2.264                | -5.2  | -1.70  | -0.40 | 1.60 | 5.4               | -0.6012 | 0.5491  |
|  |           | 15 mg              | 24.79 | 84    | 0.81  | 2.752                | -5.6  | -0.75  | 0.75  | 1.90 | 15.1              | 2.7075  | 0.0082  |
|  |           | 30 mg              | 23.77 | 89    | 0.65  | 2.599                | -6.5  | -1.10  | 0.70  | 2.10 | 12.0              | 2.3614  | 0.0204  |
|  |           | 60 mg              | 23.42 | 84    | 0.30  | 2.471                | -8.3  | -1.15  | 0.35  | 1.90 | 7.1               | 1.1041  | 0.2727  |
|  |           | 120 mg             | 24.03 | 83    | 0.90  | 2.546                | -6.4  | -0.70  | 0.80  | 2.40 | 7.4               | 3.2156  | 0.0019  |
|  | Week 12   | Placebo            | 24.10 | 95    | -0.27 | 2.438                | -8.1  | -1.70  | 0.10  | 1.20 | 5.6               | -1.0690 | 0.2878  |
|  |           | 15 mg              | 24.94 | 83    | 0.28  | 2.424                | -9.3  | -1.20  | 0.30  | 1.60 | 7.6               | 1.0415  | 0.3007  |
|  |           | 30 mg              | 23.61 | 86    | 0.72  | 2.958                | -9.0  | -0.90  | 0.70  | 2.50 | 9.4               | 2.2526  | 0.0269  |
|  |           | 60 mg              | 23.37 | 81    | 0.47  | 3.101                | -14.6 | -0.90  | 0.80  | 2.00 | 6.2               | 1.3508  | 0.1806  |
|  |           | 120 mg             | 23.97 | 82    | 0.52  | 2.826                | -7.7  | -1.00  | 0.60  | 2.30 | 11.4              | 1.6569  | 0.1014  |
|  | Week 16   | Placebo            | 24.10 | 95    | -0.11 | 2.589                | -6.8  | -2.00  | 0.00  | 1.80 | 5.3               | -0.4082 | 0.6841  |
|  |           | 15 mg              | 24.88 | 82    | 0.05  | 2.389                | -6.3  | -1.30  | 0.20  | 1.60 | 5.9               | 0.1895  | 0.8502  |
|  |           | 30 mg              | 23.61 | 85    | 0.58  | 2.643                | -6.3  | -0.80  | 0.60  | 2.20 | 7.7               | 2.0273  | 0.0458  |
|  |           | 60 mg              | 23.23 | 79    | 0.10  | 2.177                | -6.1  | -1.10  | -0.10 | 1.40 | 5.3               | 0.4238  | 0.6729  |
|  |           | 120 mg             | 24.26 | 81    | 0.95  | 2.933                | -6.9  | -0.80  | 0.90  | 2.40 | 12.0              | 2.9249  | 0.0045  |
|  | Week 20   | Placebo            | 24.01 | 94    | -0.13 | 2.647                | -6.2  | -1.80  | 0.00  | 1.30 | 6.2               | -0.4910 | 0.6246  |
|  |           | 15 mg              | 25.04 | 81    | 0.44  | 3.374                | -7.4  | -1.60  | 0.40  | 1.70 | 16.7              | 1.1725  | 0.2445  |
|  |           | 30 mg              | 23.61 | 85    | 0.47  | 2.749                | -6.4  | -1.20  | 0.30  | 2.20 | 8.7               | 1.5627  | 0.1219  |
|  |           | 60 mg              | 23.24 | 78    | 0.17  | 3.042                | -12.6 | -1.40  | 0.45  | 2.10 | 6.8               | 0.4802  | 0.6325  |
|  |           | 120 mg             | 24.29 | 79    | 1.05  | 3.215                | -7.4  | -0.90  | 1.00  | 2.40 | 19.3              | 2.9012  | 0.0048  |
|  | Week 24   | Placebo            | 24.18 | 93    | -0.49 | 2.647                | -6.1  | -2.20  | -0.40 | 1.60 | 7.9               | -1.7823 | 0.0780  |
|  |           | 15 mg              | 24.97 | 81    | -0.25 | 2.862                | -13.2 | -1.60  | -0.20 | 1.00 | 10.4              | -0.7997 | 0.4263  |
|  |           | 30 mg              | 23.60 | 84    | 0.35  | 2.963                | -6.6  | -1.55  | 0.20  | 2.05 | 10.1              | 1.0678  | 0.2887  |
| 60 mg                                      |           | 23.34              | 77    | 0.13  | 2.625 | -8.2                 | -1.30 | 0.30   | 1.60  | 6.7  | 0.4254            | 0.6718  |         |
| 120 mg                                     |           | 24.38              | 78    | 0.59  | 2.263 | -5.0                 | -0.70 | 0.55   | 1.70  | 9.1  | 2.3166            | 0.0232  |         |
| Week 28                                    | Placebo   | 24.12              | 93    | -0.14 | 2.741 | -6.8                 | -1.40 | -0.30  | 1.80  | 6.1  | -0.5031           | 0.6161  |         |
|  | 15 mg     | 24.94              | 81    | 0.02  | 2.328 | -5.2                 | -1.50 | -0.10  | 1.40  | 7.9  | 0.0811            | 0.9355  |         |
|  | 30 mg     | 23.60              | 84    | -0.11 | 2.861 | -10.4                | -1.95 | 0.20   | 1.65  | 8.4  | -0.3432           | 0.7323  |         |
|  | 60 mg     | 23.27              | 76    | 0.06  | 2.628 | -7.9                 | -1.60 | 0.15   | 1.70  | 7.0  | 0.1920            | 0.8482  |         |
|  | 120 mg    | 24.48              | 77    | 0.17  | 2.238 | -5.5                 | -1.10 | 0.30   | 1.30  | 6.2  | 0.6722            | 0.5035  |         |
| Follow-up                                  | Placebo   | 24.01              | 97    | -0.06 | 2.868 | -6.8                 | -1.40 | -0.30  | 1.80  | 9.2  | -0.1947           | 0.8460  |         |
|  | 15 mg     | 25.04              | 84    | 0.08  | 2.738 | -6.1                 | -1.50 | -0.15  | 1.40  | 12.3 | 0.2789            | 0.7810  |         |
|  | 30 mg     | 23.66              | 92    | -0.10 | 2.754 | -10.4                | -1.90 | 0.20   | 1.45  | 8.4  | -0.3482           | 0.7285  |         |
|  | 60 mg     | 23.71              | 86    | -0.00 | 2.551 | -7.9                 | -1.70 | 0.10   | 1.70  | 7.0  | -0.0127           | 0.9899  |         |
|  | 120 mg    | 24.35              | 87    | 0.24  | 2.196 | -5.5                 | -1.10 | 0.40   | 1.30  | 6.2  | 1.0352            | 0.3035  |         |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (11/14)

| Item      | Treatment | Summary Statistics |                      |       |       |       |       |        |       |      |         | One-sample t-test |        |
|-----------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|------|---------|-------------------|--------|
|           |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       |      | t-value | p-value           |        |
|           |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    | Max  |         |                   |        |
| MCV (fL)  | Week 1    | Placebo            | 91.97                | 39    | 0.21  | 1.673 | -3.0  | -1.00  | 0.00  | 1.00 | 4.0     | 0.7657            | 0.4486 |
|           |           | 15 mg              | 91.02                | 37    | -0.16 | 1.405 | -5.0  | -1.00  | 0.00  | 1.00 | 3.0     | -0.7022           | 0.4870 |
|           |           | 30 mg              | 91.85                | 39    | -0.05 | 1.731 | -4.0  | -1.00  | 0.00  | 1.00 | 4.0     | -0.1942           | 0.8470 |
|           |           | 60 mg              | 92.98                | 37    | -0.14 | 1.653 | -3.0  | -1.00  | 0.00  | 1.00 | 5.0     | -0.5272           | 0.6013 |
|           |           | 120 mg             | 92.77                | 47    | 0.03  | 1.343 | -3.0  | -1.00  | 0.00  | 1.00 | 4.0     | 0.1303            | 0.8969 |
|           | Week 2    | Placebo            | 92.37                | 97    | -0.17 | 1.631 | -5.4  | -1.00  | 0.00  | 1.00 | 4.0     | -1.0519           | 0.2955 |
|           |           | 15 mg              | 91.42                | 89    | -0.01 | 1.506 | -6.0  | -1.00  | 0.00  | 1.00 | 5.0     | -0.0915           | 0.9273 |
|           |           | 30 mg              | 91.97                | 91    | 0.04  | 1.335 | -3.0  | -1.00  | 0.00  | 1.00 | 3.0     | 0.3062            | 0.7602 |
|           |           | 60 mg              | 92.24                | 93    | -0.05 | 1.462 | -3.0  | -1.00  | 0.00  | 1.00 | 4.0     | -0.3616           | 0.7185 |
|           |           | 120 mg             | 92.37                | 88    | 0.46  | 1.519 | -2.0  | -0.20  | 0.00  | 1.00 | 7.0     | 2.8696            | 0.0052 |
|           | Week 3    | Placebo            | 92.19                | 37    | -0.22 | 1.566 | -3.0  | -1.00  | 0.00  | 1.00 | 3.0     | -0.8399           | 0.4065 |
|           |           | 15 mg              | 90.83                | 36    | 0.46  | 1.592 | -4.0  | 0.00   | 0.85  | 1.00 | 5.0     | 1.7480            | 0.0892 |
|           |           | 30 mg              | 91.95                | 38    | 0.10  | 1.740 | -4.0  | -1.00  | 0.00  | 1.00 | 4.0     | 0.3542            | 0.7252 |
|           |           | 60 mg              | 92.69                | 32    | 0.20  | 1.410 | -3.0  | -1.00  | 0.00  | 1.00 | 3.0     | 0.7897            | 0.4357 |
|           |           | 120 mg             | 92.71                | 42    | 0.48  | 1.615 | -4.0  | -1.00  | 0.00  | 2.00 | 3.0     | 1.9401            | 0.0593 |
|           | Week 4    | Placebo            | 92.39                | 96    | -0.17 | 1.460 | -3.4  | -1.00  | 0.00  | 1.00 | 3.0     | -1.1604           | 0.2488 |
|           |           | 15 mg              | 91.42                | 89    | -0.11 | 1.715 | -6.0  | -1.00  | 0.00  | 1.00 | 5.0     | -0.6304           | 0.5301 |
|           |           | 30 mg              | 92.08                | 89    | -0.03 | 1.377 | -3.0  | -1.00  | 0.00  | 1.00 | 3.8     | -0.2309           | 0.8180 |
|           |           | 60 mg              | 92.25                | 89    | 0.17  | 1.509 | -4.0  | -1.00  | 0.00  | 1.00 | 4.0     | 1.0886            | 0.2793 |
|           |           | 120 mg             | 92.32                | 86    | 0.11  | 1.671 | -4.0  | -1.00  | 0.00  | 1.00 | 3.1     | 0.6000            | 0.5501 |
|           | Week 6    | Placebo            | 92.38                | 40    | 0.45  | 1.600 | -3.0  | -1.00  | 0.50  | 1.50 | 4.0     | 1.7782            | 0.0832 |
|           |           | 15 mg              | 90.97                | 40    | 0.36  | 1.896 | -5.0  | -1.00  | 1.00  | 2.00 | 4.0     | 1.1842            | 0.2435 |
|           |           | 30 mg              | 91.87                | 38    | 0.17  | 1.803 | -4.0  | -1.00  | 0.00  | 1.00 | 4.0     | 0.5758            | 0.5682 |
|           |           | 60 mg              | 93.49                | 31    | -0.08 | 1.571 | -3.0  | -1.00  | 0.00  | 1.00 | 2.0     | -0.2858           | 0.7770 |
|           |           | 120 mg             | 93.34                | 47    | 0.06  | 1.981 | -4.0  | -1.00  | 0.00  | 1.00 | 4.0     | 0.2136            | 0.8318 |
|           | Week 8    | Placebo            | 92.37                | 94    | -0.19 | 1.630 | -5.0  | -1.00  | 0.00  | 1.00 | 3.0     | -1.1076           | 0.2709 |
|           |           | 15 mg              | 91.35                | 86    | 0.03  | 1.980 | -6.0  | -1.00  | 0.00  | 1.00 | 7.0     | 0.1634            | 0.8706 |
|           |           | 30 mg              | 92.08                | 89    | 0.02  | 1.772 | -4.0  | -1.00  | 0.00  | 1.00 | 7.5     | 0.0957            | 0.9240 |
|           |           | 60 mg              | 92.39                | 85    | -0.23 | 1.806 | -4.0  | -1.00  | 0.00  | 1.00 | 4.0     | -1.1828           | 0.2402 |
|           |           | 120 mg             | 92.53                | 83    | 0.30  | 1.794 | -3.0  | -1.00  | 0.00  | 2.00 | 5.0     | 1.5359            | 0.1284 |
|           | Week 12   | Placebo            | 92.38                | 95    | -0.25 | 1.750 | -5.0  | -1.00  | 0.00  | 1.00 | 4.0     | -1.3715           | 0.1735 |
|           |           | 15 mg              | 91.35                | 86    | -0.03 | 2.006 | -6.0  | -1.00  | 0.00  | 1.00 | 8.0     | -0.1398           | 0.8892 |
|           |           | 30 mg              | 92.26                | 86    | -0.24 | 2.004 | -4.0  | -2.00  | 0.00  | 1.00 | 6.0     | -1.1195           | 0.2661 |
|           |           | 60 mg              | 92.48                | 82    | -0.44 | 1.693 | -5.0  | -2.00  | 0.00  | 1.00 | 3.0     | -2.3613           | 0.0206 |
|           |           | 120 mg             | 92.52                | 82    | 0.01  | 1.478 | -3.0  | -1.00  | 0.00  | 1.00 | 4.0     | 0.0448            | 0.9644 |
|           | Week 16   | Placebo            | 92.38                | 95    | -0.39 | 1.679 | -5.0  | -1.00  | 0.00  | 1.00 | 5.0     | -2.2365           | 0.0277 |
|           |           | 15 mg              | 91.38                | 84    | -0.18 | 2.185 | -6.0  | -1.00  | 0.00  | 1.00 | 7.0     | -0.7490           | 0.4560 |
|           |           | 30 mg              | 92.28                | 85    | -0.38 | 1.946 | -4.0  | -2.00  | -1.00 | 1.00 | 5.0     | -1.8171           | 0.0728 |
|           |           | 60 mg              | 92.48                | 80    | -0.38 | 1.772 | -4.0  | -1.80  | 0.00  | 1.00 | 4.0     | -1.9060           | 0.0603 |
|           |           | 120 mg             | 92.51                | 81    | -0.33 | 1.667 | -4.0  | -2.00  | 0.00  | 1.00 | 4.0     | -1.7664           | 0.0812 |
|           | Week 20   | Placebo            | 92.38                | 95    | -0.65 | 1.685 | -5.0  | -2.00  | -1.00 | 0.00 | 3.0     | -3.7636           | 0.0003 |
|           |           | 15 mg              | 91.38                | 84    | -0.38 | 2.081 | -5.0  | -1.00  | 0.00  | 0.95 | 10.0    | -1.6828           | 0.0962 |
|           |           | 30 mg              | 92.28                | 85    | -0.50 | 1.935 | -5.0  | -2.00  | -1.00 | 1.00 | 4.0     | -2.3596           | 0.0206 |
|           |           | 60 mg              | 92.49                | 79    | -0.61 | 1.923 | -5.0  | -2.00  | 0.00  | 1.00 | 3.0     | -2.8374           | 0.0058 |
|           |           | 120 mg             | 92.48                | 79    | -0.18 | 1.810 | -5.0  | -1.00  | 0.00  | 1.00 | 4.0     | -0.8887           | 0.3769 |
|           | Week 24   | Placebo            | 92.34                | 93    | -0.74 | 1.862 | -8.7  | -1.00  | -1.00 | 0.00 | 4.0     | -3.8208           | 0.0002 |
|           |           | 15 mg              | 91.37                | 83    | -0.72 | 2.108 | -6.0  | -2.00  | -1.00 | 0.00 | 6.0     | -3.0986           | 0.0027 |
|           |           | 30 mg              | 92.34                | 84    | -0.81 | 2.208 | -9.0  | -2.00  | -1.00 | 0.00 | 6.0     | -3.3645           | 0.0012 |
| 60 mg     |           | 92.49              | 78                   | -0.86 | 1.864 | -7.0  | -2.00 | -1.00  | 0.00  | 4.0  | -4.0942 | 0.0001            |        |
| 120 mg    |           | 92.48              | 78                   | -0.58 | 1.879 | -6.0  | -2.00 | -1.00  | 0.00  | 4.0  | -2.7481 | 0.0075            |        |
| Week 28   | Placebo   | 92.35              | 93                   | -0.70 | 2.129 | -7.5  | -2.00 | -1.00  | 0.00  | 4.0  | -3.1714 | 0.0021            |        |
|           | 15 mg     | 91.39              | 83                   | -0.66 | 2.147 | -5.0  | -2.00 | -1.00  | 1.00  | 6.0  | -2.7965 | 0.0064            |        |
|           | 30 mg     | 92.34              | 84                   | -0.68 | 2.372 | -8.0  | -2.00 | -1.00  | 0.50  | 7.8  | -2.6401 | 0.0099            |        |
|           | 60 mg     | 92.44              | 77                   | -0.86 | 1.924 | -7.0  | -2.00 | -1.00  | 0.30  | 3.0  | -3.9205 | 0.0002            |        |
|           | 120 mg    | 92.40              | 77                   | -0.89 | 1.863 | -5.0  | -2.00 | -1.00  | 0.00  | 6.0  | -4.2139 | <0.0001           |        |
| Follow-up | Placebo   | 92.35              | 97                   | -0.70 | 2.106 | -7.5  | -2.00 | -1.00  | 0.00  | 4.0  | -3.2839 | 0.0014            |        |
|           | 15 mg     | 91.47              | 86                   | -0.71 | 2.118 | -5.0  | -2.00 | -1.00  | 1.00  | 6.0  | -3.0908 | 0.0027            |        |
|           | 30 mg     | 92.03              | 92                   | -0.68 | 2.365 | -8.0  | -2.00 | -1.00  | 0.50  | 7.8  | -2.7509 | 0.0072            |        |
|           | 60 mg     | 92.12              | 87                   | -0.83 | 1.875 | -7.0  | -2.00 | -1.00  | 1.00  | 3.0  | -4.1278 | <0.0001           |        |
|           | 120 mg    | 92.29              | 87                   | -0.80 | 1.799 | -5.0  | -2.00 | -1.00  | 0.00  | 6.0  | -4.1654 | <0.0001           |        |



表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (12/14)

| Item      | Treatment | Summary Statistics |       |       |       |                      |       |        |       |      |         | One-sample t-test |        |
|-----------|-----------|--------------------|-------|-------|-------|----------------------|-------|--------|-------|------|---------|-------------------|--------|
|           |           | Baseline Mean      | N     | Mean  | SD    | Change from Baseline |       |        |       |      | t-value | p-value           |        |
|           |           |                    |       |       |       | Min                  | Q1    | Median | Q3    | Max  |         |                   |        |
| MCH (pg)  | Week 1    | Placebo            | 30.48 | 39    | -0.26 | 0.625                | -2.0  | -0.70  | -0.30 | 0.30 | 0.8     | -2.6125           | 0.0128 |
|           |           | 15 mg              | 29.67 | 37    | 0.15  | 0.405                | -0.9  | -0.10  | 0.20  | 0.50 | 0.9     | 2.1907            | 0.0350 |
|           |           | 30 mg              | 30.28 | 39    | 0.05  | 0.457                | -1.3  | -0.20  | 0.00  | 0.40 | 0.9     | 0.7358            | 0.4664 |
|           |           | 60 mg              | 30.53 | 37    | -0.05 | 0.428                | -0.8  | -0.40  | 0.00  | 0.20 | 0.7     | -0.6915           | 0.4937 |
|           |           | 120 mg             | 30.74 | 47    | -0.12 | 0.520                | -1.0  | -0.40  | -0.20 | 0.30 | 0.9     | -1.5435           | 0.1296 |
|           | Week 2    | Placebo            | 30.60 | 97    | -0.08 | 0.574                | -2.0  | -0.40  | -0.10 | 0.30 | 0.9     | -1.4506           | 0.1502 |
|           |           | 15 mg              | 30.09 | 89    | 0.03  | 0.501                | -1.1  | -0.30  | 0.10  | 0.30 | 1.1     | 0.4870            | 0.6274 |
|           |           | 30 mg              | 30.35 | 91    | -0.00 | 0.494                | -1.7  | -0.30  | 0.10  | 0.40 | 0.8     | -0.0425           | 0.9662 |
|           |           | 60 mg              | 30.63 | 93    | -0.02 | 0.518                | -1.7  | -0.30  | 0.10  | 0.30 | 1.1     | -0.4400           | 0.6609 |
|           |           | 120 mg             | 30.68 | 88    | -0.01 | 0.549                | -1.2  | -0.35  | -0.05 | 0.35 | 1.6     | -0.0971           | 0.9229 |
|           | Week 3    | Placebo            | 30.52 | 37    | -0.08 | 0.592                | -2.1  | -0.30  | -0.10 | 0.20 | 1.2     | -0.7772           | 0.4421 |
|           |           | 15 mg              | 29.68 | 36    | 0.23  | 0.524                | -1.2  | -0.20  | 0.30  | 0.70 | 1.0     | 2.6704            | 0.0114 |
|           |           | 30 mg              | 30.25 | 38    | -0.03 | 0.510                | -1.2  | -0.30  | -0.05 | 0.30 | 1.2     | -0.3817           | 0.7048 |
|           |           | 60 mg              | 30.50 | 32    | -0.06 | 0.356                | -0.7  | -0.30  | -0.10 | 0.25 | 0.6     | -0.9923           | 0.3287 |
|           |           | 120 mg             | 30.72 | 42    | -0.07 | 0.518                | -1.1  | -0.40  | -0.10 | 0.30 | 1.0     | -0.8635           | 0.3929 |
|           | Week 4    | Placebo            | 30.64 | 96    | -0.05 | 0.569                | -1.5  | -0.45  | -0.05 | 0.30 | 1.3     | -0.9142           | 0.3629 |
|           |           | 15 mg              | 30.09 | 89    | 0.04  | 0.497                | -1.2  | -0.30  | 0.00  | 0.40 | 1.3     | 0.8314            | 0.4080 |
|           |           | 30 mg              | 30.41 | 89    | -0.03 | 0.498                | -1.3  | -0.30  | -0.10 | 0.30 | 1.3     | -0.5109           | 0.6107 |
|           |           | 60 mg              | 30.64 | 89    | 0.02  | 0.481                | -1.1  | -0.30  | 0.00  | 0.30 | 1.4     | 0.4190            | 0.6762 |
|           |           | 120 mg             | 30.65 | 86    | -0.10 | 0.526                | -1.4  | -0.50  | -0.10 | 0.30 | 1.0     | -1.7421           | 0.0851 |
|           | Week 6    | Placebo            | 30.70 | 40    | 0.02  | 0.630                | -1.9  | -0.40  | 0.05  | 0.50 | 1.6     | 0.1756            | 0.8615 |
|           |           | 15 mg              | 29.82 | 40    | 0.14  | 0.644                | -1.7  | -0.30  | 0.10  | 0.50 | 1.7     | 1.3268            | 0.1923 |
|           |           | 30 mg              | 30.38 | 38    | 0.06  | 0.571                | -1.5  | -0.20  | 0.10  | 0.40 | 1.1     | 0.6823            | 0.4993 |
|           |           | 60 mg              | 30.79 | 31    | 0.06  | 0.729                | -0.6  | -0.40  | -0.10 | 0.30 | 3.1     | 0.4682            | 0.6430 |
|           |           | 120 mg             | 30.84 | 47    | -0.05 | 0.641                | -1.4  | -0.50  | 0.00  | 0.40 | 1.2     | -0.5004           | 0.6192 |
|           | Week 8    | Placebo            | 30.64 | 94    | 0.01  | 0.582                | -1.6  | -0.30  | 0.00  | 0.30 | 2.3     | 0.2303            | 0.8184 |
|           |           | 15 mg              | 30.10 | 86    | 0.06  | 0.663                | -1.9  | -0.30  | 0.10  | 0.50 | 3.3     | 0.8138            | 0.4180 |
|           |           | 30 mg              | 30.41 | 89    | 0.02  | 0.498                | -1.0  | -0.30  | 0.10  | 0.40 | 1.2     | 0.4258            | 0.6713 |
|           |           | 60 mg              | 30.69 | 85    | -0.09 | 0.494                | -1.5  | -0.40  | -0.10 | 0.30 | 0.9     | -1.6909           | 0.0946 |
|           |           | 120 mg             | 30.74 | 83    | -0.10 | 0.592                | -1.6  | -0.40  | -0.10 | 0.20 | 1.8     | -1.5018           | 0.1370 |
|           | Week 12   | Placebo            | 30.65 | 95    | -0.08 | 0.589                | -1.8  | -0.50  | -0.10 | 0.30 | 1.3     | -1.3936           | 0.1667 |
|           |           | 15 mg              | 30.10 | 86    | -0.01 | 0.801                | -2.6  | -0.40  | -0.10 | 0.40 | 3.1     | -0.1077           | 0.9145 |
|           |           | 30 mg              | 30.50 | 86    | 0.02  | 0.639                | -1.2  | -0.50  | 0.05  | 0.40 | 1.9     | 0.3207            | 0.7492 |
|           |           | 60 mg              | 30.71 | 82    | -0.14 | 0.642                | -1.5  | -0.70  | -0.10 | 0.30 | 1.2     | -1.9087           | 0.0598 |
|           |           | 120 mg             | 30.71 | 82    | -0.13 | 0.553                | -1.3  | -0.60  | -0.10 | 0.30 | 1.2     | -2.1185           | 0.0372 |
|           | Week 16   | Placebo            | 30.65 | 95    | -0.09 | 0.560                | -1.8  | -0.40  | -0.10 | 0.30 | 1.4     | -1.5559           | 0.1231 |
|           |           | 15 mg              | 30.10 | 84    | 0.05  | 0.694                | -1.8  | -0.30  | 0.00  | 0.40 | 2.9     | 0.6289            | 0.5311 |
|           |           | 30 mg              | 30.49 | 85    | 0.06  | 0.645                | -1.1  | -0.30  | 0.00  | 0.50 | 1.8     | 0.8911            | 0.3754 |
|           |           | 60 mg              | 30.74 | 80    | -0.11 | 0.666                | -1.7  | -0.60  | -0.15 | 0.30 | 1.5     | -1.4276           | 0.1573 |
|           |           | 120 mg             | 30.71 | 81    | -0.17 | 0.631                | -1.7  | -0.50  | -0.10 | 0.20 | 1.8     | -2.4665           | 0.0158 |
|           | Week 20   | Placebo            | 30.65 | 95    | -0.07 | 0.661                | -2.3  | -0.40  | -0.10 | 0.30 | 2.2     | -1.0858           | 0.2803 |
|           |           | 15 mg              | 30.10 | 84    | -0.00 | 0.675                | -1.3  | -0.45  | 0.00  | 0.30 | 3.0     | -0.0162           | 0.9871 |
|           |           | 30 mg              | 30.49 | 85    | 0.03  | 0.666                | -2.6  | -0.40  | 0.10  | 0.40 | 2.1     | 0.4234            | 0.6731 |
|           |           | 60 mg              | 30.73 | 79    | -0.15 | 0.628                | -1.4  | -0.60  | -0.10 | 0.20 | 1.5     | -2.1128           | 0.0378 |
|           |           | 120 mg             | 30.71 | 79    | -0.21 | 0.650                | -2.1  | -0.50  | -0.20 | 0.20 | 1.1     | -2.9261           | 0.0045 |
|           | Week 24   | Placebo            | 30.63 | 93    | -0.13 | 0.738                | -2.7  | -0.50  | -0.10 | 0.30 | 1.9     | -1.6864           | 0.0951 |
|           |           | 15 mg              | 30.07 | 83    | 0.04  | 0.778                | -1.9  | -0.30  | 0.10  | 0.50 | 3.2     | 0.5222            | 0.6030 |
|           |           | 30 mg              | 30.50 | 84    | -0.06 | 0.775                | -3.3  | -0.50  | -0.10 | 0.30 | 2.2     | -0.6898           | 0.4923 |
| 60 mg     |           | 30.72              | 78    | -0.12 | 0.620 | -1.3                 | -0.60 | -0.25  | 0.30  | 1.6  | -1.7520 | 0.0837            |        |
| 120 mg    |           | 30.71              | 78    | -0.15 | 0.848 | -2.5                 | -0.70 | -0.10  | 0.30  | 3.5  | -1.5480 | 0.1257            |        |
| Week 28   | Placebo   | 30.65              | 93    | -0.12 | 0.778 | -3.9                 | -0.50 | 0.00   | 0.30  | 1.4  | -1.4930 | 0.1389            |        |
|           | 15 mg     | 30.08              | 83    | -0.04 | 0.862 | -2.1                 | -0.50 | 0.00   | 0.40  | 3.4  | -0.4200 | 0.6756            |        |
|           | 30 mg     | 30.50              | 84    | 0.05  | 0.784 | -3.7                 | -0.40 | 0.00   | 0.40  | 2.5  | 0.5987  | 0.5510            |        |
|           | 60 mg     | 30.69              | 77    | -0.25 | 0.704 | -1.4                 | -0.80 | -0.30  | 0.20  | 1.7  | -3.1223 | 0.0025            |        |
|           | 120 mg    | 30.65              | 77    | -0.33 | 0.704 | -2.5                 | -0.70 | -0.30  | 0.00  | 2.6  | -4.0927 | 0.0001            |        |
| Follow-up | Placebo   | 30.60              | 97    | -0.12 | 0.763 | -3.9                 | -0.50 | 0.00   | 0.30  | 1.4  | -1.4907 | 0.1393            |        |
|           | 15 mg     | 30.07              | 86    | -0.06 | 0.859 | -2.1                 | -0.50 | 0.00   | 0.40  | 3.4  | -0.6907 | 0.4917            |        |
|           | 30 mg     | 30.38              | 92    | 0.04  | 0.767 | -3.7                 | -0.40 | 0.00   | 0.40  | 2.5  | 0.4485  | 0.6549            |        |
|           | 60 mg     | 30.57              | 87    | -0.24 | 0.678 | -1.4                 | -0.80 | -0.20  | 0.20  | 1.7  | -3.3183 | 0.0013            |        |
|           | 120 mg    | 30.63              | 87    | -0.30 | 0.682 | -2.5                 | -0.70 | -0.30  | 0.00  | 2.6  | -4.1497 | <0.0001           |        |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (13/14)

| Item      | Treatment | Summary Statistics |       |       |       |                      |       |        |       |      |         | One-sample t-test |        |
|-----------|-----------|--------------------|-------|-------|-------|----------------------|-------|--------|-------|------|---------|-------------------|--------|
|           |           | Baseline Mean      | N     | Mean  | SD    | Change from Baseline |       |        |       |      | t-value | p-value           |        |
|           |           |                    |       |       |       | Min                  | Q1    | Median | Q3    | Max  |         |                   |        |
| MCHC (%)  | Week 1    | Placebo            | 33.15 | 39    | -0.38 | 0.940                | -2.4  | -1.10  | -0.40 | 0.40 | 1.2     | -2.5562           | 0.0147 |
|           |           | 15 mg              | 32.59 | 37    | 0.20  | 0.633                | -1.0  | -0.20  | 0.20  | 0.50 | 1.7     | 1.9209            | 0.0627 |
|           |           | 30 mg              | 32.95 | 39    | 0.09  | 0.741                | -1.5  | -0.60  | 0.10  | 0.70 | 2.1     | 0.7348            | 0.4670 |
|           |           | 60 mg              | 32.85 | 37    | 0.02  | 0.758                | -2.1  | -0.40  | -0.10 | 0.70 | 1.3     | 0.1952            | 0.8464 |
|           |           | 120 mg             | 33.14 | 47    | -0.14 | 0.763                | -1.4  | -0.90  | -0.20 | 0.50 | 1.4     | -1.2419           | 0.2206 |
|           | Week 2    | Placebo            | 33.14 | 97    | -0.06 | 0.785                | -2.1  | -0.60  | 0.00  | 0.40 | 2.0     | -0.7110           | 0.4788 |
|           |           | 15 mg              | 32.91 | 89    | 0.00  | 0.756                | -2.3  | -0.50  | 0.00  | 0.50 | 3.2     | 0.0561            | 0.9554 |
|           |           | 30 mg              | 33.00 | 91    | -0.02 | 0.668                | -1.9  | -0.50  | 0.00  | 0.50 | 1.5     | -0.2510           | 0.8024 |
|           |           | 60 mg              | 33.21 | 93    | -0.00 | 0.720                | -1.7  | -0.50  | 0.00  | 0.50 | 2.3     | -0.0576           | 0.9542 |
|           |           | 120 mg             | 33.21 | 88    | -0.16 | 0.833                | -3.3  | -0.60  | -0.15 | 0.40 | 1.7     | -1.7650           | 0.0811 |
|           | Week 3    | Placebo            | 33.12 | 37    | -0.01 | 0.847                | -2.4  | -0.60  | 0.20  | 0.60 | 1.7     | -0.0971           | 0.9232 |
|           |           | 15 mg              | 32.66 | 36    | 0.08  | 0.677                | -1.4  | -0.30  | 0.10  | 0.60 | 1.4     | 0.6650            | 0.5104 |
|           |           | 30 mg              | 32.88 | 38    | -0.06 | 0.833                | -2.4  | -0.70  | -0.15 | 0.60 | 1.5     | -0.4481           | 0.6567 |
|           |           | 60 mg              | 32.92 | 32    | -0.13 | 0.693                | -1.5  | -0.60  | 0.00  | 0.25 | 1.6     | -1.0967           | 0.2812 |
|           |           | 120 mg             | 33.14 | 42    | -0.23 | 0.706                | -1.5  | -0.60  | -0.30 | 0.00 | 1.5     | -2.0987           | 0.0420 |
|           | Week 4    | Placebo            | 33.18 | 96    | -0.04 | 0.776                | -2.2  | -0.60  | 0.00  | 0.50 | 1.7     | -0.4733           | 0.6371 |
|           |           | 15 mg              | 32.91 | 89    | 0.09  | 0.724                | -1.4  | -0.40  | 0.10  | 0.50 | 2.5     | 1.1714            | 0.2446 |
|           |           | 30 mg              | 33.02 | 89    | -0.03 | 0.690                | -1.6  | -0.50  | 0.00  | 0.40 | 1.9     | -0.3535           | 0.7245 |
|           |           | 60 mg              | 33.23 | 89    | -0.06 | 0.772                | -2.1  | -0.60  | -0.10 | 0.50 | 1.7     | -0.6725           | 0.5031 |
|           |           | 120 mg             | 33.21 | 86    | -0.14 | 0.760                | -1.8  | -0.70  | -0.20 | 0.30 | 2.0     | -1.7594           | 0.0821 |
| Week 6    | Placebo   | 33.25              | 40    | -0.16 | 0.968 | -2.8                 | -0.80 | -0.20  | 0.30  | 2.3  | -1.0455 | 0.3022            |        |
|           | 15 mg     | 32.76              | 40    | -0.00 | 0.837 | -1.3                 | -0.75 | -0.05  | 0.60  | 1.9  | -0.0189 | 0.9850            |        |
|           | 30 mg     | 33.06              | 38    | -0.01 | 0.829 | -1.5                 | -0.50 | 0.10   | 0.40  | 1.9  | -0.0979 | 0.9225            |        |
|           | 60 mg     | 32.96              | 31    | 0.03  | 0.831 | -1.4                 | -0.50 | 0.00   | 0.50  | 2.8  | 0.1946  | 0.8470            |        |
|           | 120 mg    | 33.06              | 47    | -0.07 | 0.896 | -2.1                 | -0.70 | -0.10  | 0.60  | 2.0  | -0.5695 | 0.5718            |        |
| Week 8    | Placebo   | 33.18              | 94    | 0.06  | 0.829 | -2.3                 | -0.50 | 0.10   | 0.60  | 2.6  | 0.7468  | 0.4570            |        |
|           | 15 mg     | 32.94              | 86    | 0.04  | 0.814 | -1.7                 | -0.60 | -0.10  | 0.70  | 2.1  | 0.4767  | 0.6348            |        |
|           | 30 mg     | 33.02              | 89    | 0.03  | 0.680 | -2.4                 | -0.30 | 0.10   | 0.50  | 1.3  | 0.4052  | 0.6863            |        |
|           | 60 mg     | 33.24              | 85    | -0.03 | 0.766 | -2.4                 | -0.50 | 0.00   | 0.40  | 2.3  | -0.4107 | 0.6823            |        |
|           | 120 mg    | 33.23              | 83    | -0.20 | 0.825 | -2.2                 | -0.60 | -0.10  | 0.30  | 1.4  | -2.1552 | 0.0341            |        |
| Week 12   | Placebo   | 33.19              | 95    | -0.03 | 0.884 | -2.7                 | -0.50 | 0.00   | 0.40  | 2.1  | -0.3018 | 0.7635            |        |
|           | 15 mg     | 32.94              | 86    | -0.01 | 0.876 | -2.2                 | -0.60 | -0.10  | 0.60  | 2.6  | -0.1108 | 0.9121            |        |
|           | 30 mg     | 33.06              | 86    | 0.11  | 0.769 | -1.7                 | -0.30 | 0.20   | 0.70  | 2.1  | 1.3327  | 0.1862            |        |
|           | 60 mg     | 33.22              | 82    | -0.01 | 0.829 | -1.7                 | -0.60 | 0.05   | 0.60  | 1.8  | -0.0666 | 0.9471            |        |
|           | 120 mg    | 33.19              | 82    | -0.14 | 0.781 | -1.8                 | -0.70 | -0.05  | 0.50  | 1.7  | -1.6259 | 0.1079            |        |
| Week 16   | Placebo   | 33.19              | 95    | 0.03  | 0.823 | -3.0                 | -0.40 | 0.10   | 0.50  | 2.3  | 0.3364  | 0.7373            |        |
|           | 15 mg     | 32.93              | 84    | 0.10  | 0.830 | -2.3                 | -0.50 | 0.15   | 0.75  | 2.4  | 1.1040  | 0.2728            |        |
|           | 30 mg     | 33.05              | 85    | 0.20  | 0.838 | -1.8                 | -0.30 | 0.20   | 0.80  | 2.1  | 2.1998  | 0.0306            |        |
|           | 60 mg     | 33.26              | 80    | 0.02  | 0.853 | -1.9                 | -0.60 | 0.05   | 0.60  | 2.1  | 0.2489  | 0.8041            |        |
|           | 120 mg    | 33.20              | 81    | -0.09 | 0.840 | -2.0                 | -0.50 | -0.10  | 0.50  | 2.6  | -0.9127 | 0.3641            |        |
| Week 20   | Placebo   | 33.19              | 95    | 0.14  | 0.817 | -1.7                 | -0.30 | 0.10   | 0.70  | 1.9  | 1.6573  | 0.1008            |        |
|           | 15 mg     | 32.93              | 84    | 0.12  | 0.706 | -1.5                 | -0.30 | 0.10   | 0.50  | 2.3  | 1.5141  | 0.1338            |        |
|           | 30 mg     | 33.05              | 85    | 0.22  | 0.830 | -2.3                 | -0.30 | 0.20   | 0.90  | 1.7  | 2.4557  | 0.0161            |        |
|           | 60 mg     | 33.24              | 79    | 0.04  | 0.900 | -1.8                 | -0.70 | 0.00   | 0.60  | 2.1  | 0.4378  | 0.6628            |        |
|           | 120 mg    | 33.21              | 79    | -0.15 | 0.789 | -1.7                 | -0.70 | -0.10  | 0.40  | 1.7  | -1.6972 | 0.0936            |        |
| Week 24   | Placebo   | 33.19              | 93    | 0.11  | 0.915 | -1.8                 | -0.40 | 0.00   | 0.70  | 2.4  | 1.1564  | 0.2505            |        |
|           | 15 mg     | 32.91              | 83    | 0.30  | 0.772 | -1.6                 | -0.20 | 0.40   | 0.80  | 2.1  | 3.5546  | 0.0006            |        |
|           | 30 mg     | 33.04              | 84    | 0.22  | 0.861 | -1.9                 | -0.35 | 0.30   | 0.80  | 2.2  | 2.2933  | 0.0244            |        |
|           | 60 mg     | 33.23              | 78    | 0.18  | 0.891 | -2.2                 | -0.30 | 0.10   | 0.80  | 2.1  | 1.7928  | 0.0769            |        |
|           | 120 mg    | 33.21              | 78    | 0.06  | 0.893 | -1.4                 | -0.70 | 0.05   | 0.70  | 2.6  | 0.5577  | 0.5787            |        |
| Week 28   | Placebo   | 33.20              | 93    | 0.12  | 0.985 | -2.2                 | -0.50 | 0.10   | 0.80  | 2.0  | 1.1899  | 0.2372            |        |
|           | 15 mg     | 32.91              | 83    | 0.17  | 0.815 | -1.5                 | -0.50 | 0.10   | 0.80  | 2.0  | 1.8985  | 0.0611            |        |
|           | 30 mg     | 33.04              | 84    | 0.30  | 0.789 | -1.5                 | -0.20 | 0.20   | 0.90  | 2.6  | 3.4993  | 0.0008            |        |
|           | 60 mg     | 33.22              | 77    | 0.04  | 0.937 | -1.9                 | -0.50 | -0.10  | 0.60  | 2.5  | 0.4134  | 0.6804            |        |
|           | 120 mg    | 33.17              | 77    | -0.04 | 0.708 | -1.7                 | -0.50 | -0.10  | 0.40  | 1.6  | -0.4509 | 0.6533            |        |
| Follow-up | Placebo   | 33.15              | 97    | 0.13  | 0.972 | -2.2                 | -0.50 | 0.10   | 0.80  | 2.0  | 1.3156  | 0.1914            |        |
|           | 15 mg     | 32.88              | 86    | 0.16  | 0.807 | -1.5                 | -0.50 | 0.10   | 0.70  | 2.0  | 1.8304  | 0.0707            |        |
|           | 30 mg     | 33.00              | 92    | 0.28  | 0.772 | -1.5                 | -0.20 | 0.20   | 0.90  | 2.6  | 3.4578  | 0.0008            |        |
|           | 60 mg     | 33.20              | 87    | 0.05  | 0.911 | -1.9                 | -0.50 | -0.10  | 0.60  | 2.5  | 0.4826  | 0.6306            |        |
|           | 120 mg    | 33.19              | 87    | -0.04 | 0.696 | -1.7                 | -0.50 | -0.10  | 0.40  | 1.6  | -0.6004 | 0.5498            |        |

表 2.7.4.7-付録-12 血液学的検査の評価時点別の変化量 (CCT-001 試験) (14/14)

| Item                   | Treatment | Summary Statistics |                      |       |       |       |       |       |         |         |         | One-sample t-test |        |
|------------------------|-----------|--------------------|----------------------|-------|-------|-------|-------|-------|---------|---------|---------|-------------------|--------|
|                        |           | Baseline Mean      | Change from Baseline |       |       |       |       | Max   | t-value | p-value |         |                   |        |
|                        |           |                    | N                    | Mean  | SD    | Min   | Q1    |       |         |         | Median  | Q3                |        |
| Prothrombin Time (sec) | Week 1    | Placebo            | 11.08                | 39    | -0.29 | 1.161 | -6.9  | -0.40 | -0.10   | 0.20    | 0.8     | -1.5580           | 0.1275 |
|                        |           | 15 mg              | 10.66                | 37    | -0.01 | 0.532 | -1.9  | -0.30 | -0.10   | 0.40    | 0.9     | -0.1237           | 0.9023 |
|                        |           | 30 mg              | 10.78                | 39    | -0.11 | 0.803 | -3.2  | -0.50 | 0.00    | 0.20    | 2.0     | -0.8371           | 0.4078 |
|                        |           | 60 mg              | 10.74                | 37    | -0.01 | 0.497 | -0.9  | -0.20 | -0.10   | 0.20    | 1.6     | -0.1655           | 0.8695 |
|                        |           | 120 mg             | 10.77                | 47    | 0.00  | 0.727 | -3.5  | -0.20 | 0.10    | 0.30    | 1.9     | 0.0201            | 0.9841 |
|                        | Week 2    | Placebo            | 10.93                | 97    | -0.17 | 1.271 | -8.1  | -0.30 | -0.10   | 0.20    | 4.5     | -1.2860           | 0.2015 |
|                        |           | 15 mg              | 10.70                | 89    | 0.14  | 0.672 | -2.4  | -0.10 | 0.10    | 0.40    | 3.6     | 1.9085            | 0.0596 |
|                        |           | 30 mg              | 10.66                | 91    | 0.05  | 0.746 | -2.8  | -0.30 | 0.10    | 0.40    | 3.4     | 0.5763            | 0.5659 |
|                        |           | 60 mg              | 10.76                | 93    | 0.03  | 0.815 | -3.3  | -0.30 | 0.00    | 0.30    | 5.6     | 0.4071            | 0.6848 |
|                        |           | 120 mg             | 10.68                | 89    | 0.01  | 0.583 | -3.6  | -0.30 | 0.00    | 0.30    | 1.2     | 0.2364            | 0.8136 |
|                        | Week 3    | Placebo            | 11.08                | 36    | -0.08 | 1.248 | -6.8  | -0.30 | 0.05    | 0.35    | 1.1     | -0.4006           | 0.6912 |
|                        |           | 15 mg              | 10.66                | 36    | -0.00 | 0.519 | -1.8  | -0.30 | 0.00    | 0.35    | 1.0     | -0.0321           | 0.9745 |
|                        |           | 30 mg              | 10.77                | 38    | 0.07  | 0.883 | -2.8  | -0.40 | 0.00    | 0.30    | 3.3     | 0.4963            | 0.6226 |
|                        |           | 60 mg              | 10.76                | 32    | -0.03 | 0.420 | -0.8  | -0.30 | -0.05   | 0.20    | 1.0     | -0.4209           | 0.6767 |
|                        |           | 120 mg             | 10.83                | 42    | -0.04 | 0.695 | -3.2  | -0.30 | -0.05   | 0.30    | 0.9     | -0.3996           | 0.6915 |
|                        | Week 4    | Placebo            | 10.91                | 96    | -0.10 | 1.253 | -8.7  | -0.20 | 0.05    | 0.30    | 2.8     | -0.7903           | 0.4313 |
|                        |           | 15 mg              | 10.70                | 89    | 0.16  | 0.885 | -1.3  | -0.20 | 0.10    | 0.30    | 6.5     | 1.7242            | 0.0882 |
|                        |           | 30 mg              | 10.66                | 89    | 0.17  | 1.048 | -3.2  | -0.30 | 0.00    | 0.40    | 4.5     | 1.5368            | 0.1279 |
|                        |           | 60 mg              | 10.76                | 89    | -0.03 | 0.645 | -3.0  | -0.40 | 0.00    | 0.20    | 2.5     | -0.3942           | 0.6944 |
|                        |           | 120 mg             | 10.68                | 87    | 0.08  | 0.761 | -3.6  | -0.30 | 0.00    | 0.50    | 3.4     | 0.9580            | 0.3408 |
|                        | Week 6    | Placebo            | 11.06                | 40    | -0.10 | 1.238 | -7.2  | -0.25 | 0.00    | 0.20    | 1.3     | -0.4982           | 0.6211 |
|                        |           | 15 mg              | 10.71                | 40    | 0.03  | 0.591 | -1.6  | -0.30 | 0.00    | 0.25    | 1.5     | 0.3213            | 0.7497 |
|                        |           | 30 mg              | 10.82                | 38    | -0.02 | 0.684 | -2.6  | -0.30 | 0.05    | 0.40    | 1.2     | -0.2133           | 0.8322 |
|                        |           | 60 mg              | 10.75                | 31    | -0.10 | 0.484 | -0.8  | -0.40 | -0.20   | 0.10    | 1.2     | -1.1131           | 0.2745 |
|                        |           | 120 mg             | 10.83                | 47    | -0.05 | 0.707 | -3.3  | -0.40 | 0.00    | 0.30    | 1.3     | -0.5155           | 0.6087 |
|                        | Week 8    | Placebo            | 10.92                | 94    | -0.07 | 1.276 | -8.8  | -0.30 | 0.10    | 0.40    | 1.7     | -0.5499           | 0.5837 |
|                        |           | 15 mg              | 10.71                | 86    | 0.06  | 0.833 | -2.1  | -0.30 | -0.05   | 0.30    | 6.0     | 0.6735            | 0.5025 |
|                        |           | 30 mg              | 10.66                | 89    | -0.02 | 0.619 | -2.9  | -0.40 | 0.00    | 0.30    | 2.3     | -0.3594           | 0.7202 |
|                        |           | 60 mg              | 10.77                | 86    | 0.02  | 0.716 | -3.2  | -0.30 | 0.00    | 0.30    | 2.8     | 0.3165            | 0.7524 |
|                        |           | 120 mg             | 10.68                | 84    | 0.06  | 0.854 | -3.9  | -0.30 | 0.05    | 0.35    | 3.7     | 0.6512            | 0.5167 |
|                        | Week 12   | Placebo            | 10.91                | 95    | -0.09 | 1.263 | -8.3  | -0.30 | 0.00    | 0.30    | 3.5     | -0.7228           | 0.4716 |
|                        |           | 15 mg              | 10.71                | 86    | 0.07  | 0.883 | -1.9  | -0.30 | 0.00    | 0.30    | 6.6     | 0.7447            | 0.4585 |
|                        |           | 30 mg              | 10.65                | 86    | -0.11 | 0.625 | -3.1  | -0.50 | -0.10   | 0.30    | 1.2     | -1.6898           | 0.0947 |
|                        |           | 60 mg              | 10.76                | 82    | -0.06 | 0.596 | -3.2  | -0.40 | 0.00    | 0.30    | 1.1     | -0.9825           | 0.3288 |
|                        |           | 120 mg             | 10.69                | 83    | -0.05 | 0.645 | -3.2  | -0.30 | 0.00    | 0.40    | 1.4     | -0.7314           | 0.4666 |
|                        | Week 16   | Placebo            | 10.84                | 94    | 0.10  | 1.307 | -7.3  | -0.20 | 0.10    | 0.30    | 8.5     | 0.7262            | 0.4696 |
|                        |           | 15 mg              | 10.71                | 84    | 0.13  | 0.951 | -1.9  | -0.30 | 0.10    | 0.40    | 6.8     | 1.2508            | 0.2145 |
|                        |           | 30 mg              | 10.65                | 85    | 0.11  | 0.807 | -2.1  | -0.40 | 0.00    | 0.50    | 4.0     | 1.2642            | 0.2097 |
|                        |           | 60 mg              | 10.78                | 79    | -0.09 | 0.599 | -3.3  | -0.30 | 0.00    | 0.30    | 1.0     | -1.2969           | 0.1985 |
|                        |           | 120 mg             | 10.69                | 82    | 0.05  | 0.679 | -3.7  | -0.20 | 0.10    | 0.40    | 2.4     | 0.6018            | 0.5490 |
|                        | Week 20   | Placebo            | 10.91                | 95    | 0.07  | 1.656 | -8.3  | -0.30 | 0.10    | 0.40    | 10.0    | 0.4090            | 0.6835 |
|                        |           | 15 mg              | 10.71                | 84    | 0.09  | 0.901 | -2.2  | -0.30 | 0.10    | 0.40    | 5.4     | 0.9562            | 0.3418 |
|                        |           | 30 mg              | 10.65                | 85    | 0.07  | 0.884 | -3.3  | -0.30 | 0.00    | 0.50    | 5.7     | 0.7607            | 0.4490 |
|                        |           | 60 mg              | 10.77                | 78    | 0.05  | 1.090 | -3.4  | -0.40 | -0.10   | 0.40    | 7.8     | 0.4363            | 0.6638 |
|                        |           | 120 mg             | 10.69                | 80    | -0.00 | 0.687 | -4.1  | -0.20 | 0.00    | 0.40    | 1.1     | -0.0163           | 0.9871 |
|                        | Week 24   | Placebo            | 10.91                | 93    | 0.03  | 1.609 | -8.5  | -0.30 | 0.00    | 0.30    | 7.3     | 0.1869            | 0.8522 |
|                        |           | 15 mg              | 10.71                | 84    | 0.21  | 1.218 | -1.3  | -0.25 | 0.10    | 0.40    | 9.9     | 1.5764            | 0.1187 |
|                        |           | 30 mg              | 10.66                | 84    | 0.21  | 0.935 | -3.2  | -0.15 | 0.10    | 0.45    | 4.8     | 2.0539            | 0.0431 |
| 60 mg                  |           | 10.77              | 78                   | 0.03  | 0.915 | -3.7  | -0.30 | 0.00  | 0.30    | 5.9     | 0.2723  | 0.7861            |        |
| 120 mg                 |           | 10.68              | 79                   | -0.01 | 0.701 | -3.9  | -0.30 | 0.10  | 0.40    | 1.2     | -0.1605 | 0.8729            |        |
| Week 28                | Placebo   | 10.91              | 94                   | -0.05 | 1.453 | -8.7  | -0.40 | 0.00  | 0.40    | 5.0     | -0.3265 | 0.7447            |        |
|                        | 15 mg     | 10.72              | 83                   | 0.10  | 0.923 | -2.3  | -0.20 | 0.10  | 0.40    | 6.4     | 1.0349  | 0.3038            |        |
|                        | 30 mg     | 10.66              | 84                   | 0.06  | 0.809 | -3.6  | -0.20 | 0.00  | 0.35    | 3.5     | 0.7285  | 0.4684            |        |
|                        | 60 mg     | 10.76              | 77                   | 0.06  | 1.005 | -3.0  | -0.40 | 0.00  | 0.20    | 5.4     | 0.5558  | 0.5800            |        |
|                        | 120 mg    | 10.68              | 78                   | -0.04 | 0.602 | -3.3  | -0.30 | 0.00  | 0.40    | 0.9     | -0.6211 | 0.5364            |        |
| Follow-up              | Placebo   | 10.92              | 98                   | -0.06 | 1.425 | -8.7  | -0.40 | 0.00  | 0.30    | 5.0     | -0.3827 | 0.7028            |        |
|                        | 15 mg     | 10.72              | 86                   | 0.13  | 0.921 | -2.3  | -0.20 | 0.10  | 0.40    | 6.4     | 1.3354  | 0.1853            |        |
|                        | 30 mg     | 10.67              | 92                   | 0.06  | 0.781 | -3.6  | -0.20 | 0.00  | 0.35    | 3.5     | 0.7079  | 0.4808            |        |
|                        | 60 mg     | 10.74              | 87                   | 0.07  | 0.972 | -3.0  | -0.40 | 0.10  | 0.40    | 5.4     | 0.6950  | 0.4890            |        |
|                        | 120 mg    | 10.68              | 88                   | -0.03 | 0.583 | -3.3  | -0.30 | 0.00  | 0.40    | 0.9     | -0.5481 | 0.5850            |        |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (1/14)

|            |           | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|------------|-----------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|            |           | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|            |           | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Hemoglobin | Week 1    | Lower   | -      | -     | -     | 3      | 1      | -     | 4     | 2      | 1      | -     | 3     | 1      | -      | 1     | 2     | 1      | 2      | -     | 3     |    |
|            |           | Normal  | -      | 34    | 1     | 35     | 1      | 28    | -     | 29     | -      | 30    | 3     | 33     | -      | 32    | 2     | 34     | -      | 41    | 1     | 42 |
|            |           | Upper   | -      | 2     | 2     | 4      | -      | 1     | 3     | 4      | -      | 2     | 1     | 3      | -      | -     | 2     | 2      | -      | -     | 2     | 2  |
|            |           | Total   | -      | 36    | 3     | 39     | 4      | 30    | 3     | 37     | 2      | 33    | 4     | 39     | 1      | 32    | 4     | 37     | 1      | 43    | 3     | 47 |
|            | Week 2    | Lower   | 2      | 4     | -     | 6      | 4      | 4     | -     | 8      | 3      | 3     | -     | 6      | 2      | 2     | -     | 4      | 1      | 4     | -     | 5  |
|            |           | Normal  | 1      | 77    | 7     | 85     | 1      | 71    | 3     | 75     | -      | 76    | 3     | 79     | 1      | 81    | 2     | 84     | -      | 77    | 5     | 82 |
|            |           | Upper   | -      | 1     | 5     | 6      | -      | -     | 6     | 6      | -      | 3     | 3     | 6      | -      | 3     | 2     | 5      | -      | 1     | -     | 1  |
|            |           | Total   | 3      | 82    | 12    | 97     | 5      | 75    | 9     | 89     | 3      | 82    | 6     | 91     | 3      | 86    | 4     | 93     | 1      | 82    | 5     | 88 |
|            | Week 3    | Lower   | -      | -     | -     | -      | 3      | -     | -     | 3      | 2      | 1     | -     | 3      | 1      | -     | -     | 1      | 1      | 1     | -     | 2  |
|            |           | Normal  | -      | 33    | 1     | 34     | -      | 31    | -     | 31     | -      | 30    | 2     | 32     | -      | 27    | 3     | 30     | -      | 37    | 3     | 40 |
|            |           | Upper   | -      | 1     | 2     | 3      | -      | -     | 2     | 2      | -      | 2     | 1     | 3      | -      | 1     | -     | 1      | -      | -     | -     | -  |
|            |           | Total   | -      | 34    | 3     | 37     | 3      | 31    | 2     | 36     | 2      | 33    | 3     | 38     | 1      | 28    | 3     | 32     | 1      | 38    | 3     | 42 |
|            | Week 4    | Lower   | 3      | 2     | -     | 5      | 4      | 3     | -     | 7      | 3      | 2     | -     | 5      | 2      | 3     | -     | 5      | 1      | 3     | -     | 4  |
|            |           | Normal  | -      | 74    | 7     | 81     | 1      | 69    | 5     | 75     | -      | 75    | 3     | 78     | 1      | 77    | 3     | 81     | -      | 77    | 5     | 82 |
|            |           | Upper   | -      | 5     | 5     | 10     | -      | 3     | 4     | 7      | -      | 3     | 3     | 6      | -      | 1     | 2     | 3      | -      | -     | -     | -  |
|            |           | Total   | 3      | 81    | 12    | 96     | 5      | 75    | 9     | 89     | 3      | 80    | 6     | 89     | 3      | 81    | 5     | 89     | 1      | 80    | 5     | 86 |
|            | Week 6    | Lower   | -      | -     | -     | -      | 3      | 2     | -     | 5      | 2      | 2     | -     | 4      | 1      | -     | -     | 1      | 1      | 2     | -     | 3  |
|            |           | Normal  | -      | 36    | -     | 36     | -      | 32    | -     | 32     | -      | 30    | 2     | 32     | -      | 27    | 2     | 29     | -      | 41    | 2     | 43 |
|            |           | Upper   | -      | 1     | 3     | 4      | -      | -     | 3     | 3      | -      | 1     | 1     | 2      | -      | -     | 1     | 1      | -      | -     | 1     | 1  |
|            |           | Total   | -      | 37    | 3     | 40     | 3      | 34    | 3     | 40     | 2      | 33    | 3     | 38     | 1      | 27    | 3     | 31     | 1      | 43    | 3     | 47 |
|            | Week 8    | Lower   | 2      | 3     | -     | 5      | 4      | 3     | -     | 7      | 3      | 2     | -     | 5      | 2      | 3     | -     | 5      | -      | 4     | -     | 4  |
|            |           | Normal  | 1      | 74    | 6     | 81     | 1      | 67    | 6     | 74     | -      | 74    | 2     | 76     | 1      | 75    | 2     | 78     | 1      | 74    | 2     | 77 |
|            |           | Upper   | -      | 2     | 6     | 8      | -      | 2     | 3     | 5      | -      | 4     | 4     | 8      | -      | 1     | 1     | 2      | -      | -     | 2     | 2  |
|            |           | Total   | 3      | 79    | 12    | 94     | 5      | 72    | 9     | 86     | 3      | 80    | 6     | 89     | 3      | 79    | 3     | 85     | 1      | 78    | 4     | 83 |
|            | Week 12   | Lower   | 2      | 3     | -     | 5      | 3      | 5     | -     | 8      | 2      | -     | -     | 2      | 2      | 1     | -     | 3      | 1      | 3     | -     | 4  |
|            |           | Normal  | 1      | 74    | 8     | 83     | 2      | 65    | 4     | 71     | -      | 77    | 3     | 80     | 1      | 74    | 1     | 76     | -      | 74    | 3     | 77 |
|            |           | Upper   | -      | 3     | 4     | 7      | -      | 2     | 5     | 7      | -      | 1     | 3     | 4      | -      | 2     | 1     | 3      | -      | -     | 1     | 1  |
|            |           | Total   | 3      | 80    | 12    | 95     | 5      | 72    | 9     | 86     | 2      | 78    | 6     | 86     | 3      | 77    | 2     | 82     | 1      | 77    | 4     | 82 |
|            | Week 16   | Lower   | 2      | 4     | -     | 6      | 4      | 4     | -     | 8      | 2      | 4     | -     | 6      | 3      | 1     | -     | 4      | -      | 3     | -     | 3  |
|            |           | Normal  | 1      | 74    | 8     | 83     | 1      | 67    | 4     | 72     | -      | 72    | 3     | 75     | -      | 72    | 2     | 74     | 1      | 73    | 2     | 76 |
|            |           | Upper   | -      | 2     | 4     | 6      | -      | -     | 4     | 4      | -      | 1     | 3     | 4      | -      | 1     | 1     | 2      | -      | -     | 2     | 2  |
|            |           | Total   | 3      | 80    | 12    | 95     | 5      | 71    | 8     | 84     | 2      | 77    | 6     | 85     | 3      | 74    | 3     | 80     | 1      | 76    | 4     | 81 |
|            | Week 20   | Lower   | 2      | 3     | -     | 5      | 4      | 5     | -     | 9      | 1      | 4     | -     | 5      | 2      | 2     | -     | 4      | -      | 3     | -     | 3  |
|            |           | Normal  | 1      | 73    | 6     | 80     | 1      | 63    | 5     | 69     | 1      | 72    | 3     | 76     | 1      | 71    | 1     | 73     | 1      | 70    | 2     | 73 |
|            |           | Upper   | -      | 4     | 6     | 10     | -      | 3     | 3     | 6      | -      | 1     | 3     | 4      | -      | 1     | 1     | 2      | -      | 1     | 2     | 3  |
|            |           | Total   | 3      | 80    | 12    | 95     | 5      | 71    | 8     | 84     | 2      | 77    | 6     | 85     | 3      | 74    | 2     | 79     | 1      | 74    | 4     | 79 |
|            | Week 24   | Lower   | 2      | 3     | -     | 5      | 3      | 6     | -     | 9      | 1      | 5     | -     | 6      | -      | 2     | -     | 2      | 1      | 5     | -     | 6  |
|            |           | Normal  | 1      | 74    | 10    | 85     | 2      | 62    | 4     | 68     | 1      | 68    | 4     | 73     | 3      | 70    | 2     | 75     | -      | 67    | 3     | 70 |
|            |           | Upper   | -      | 1     | 2     | 3      | -      | 2     | 4     | 6      | -      | 3     | 2     | 5      | -      | 1     | -     | 1      | -      | 1     | 1     | 2  |
|            |           | Total   | 3      | 78    | 12    | 93     | 5      | 70    | 8     | 83     | 2      | 76    | 6     | 84     | 3      | 73    | 2     | 78     | 1      | 73    | 4     | 78 |
|            | Week 28   | Lower   | 2      | 4     | -     | 6      | 3      | 4     | -     | 7      | 1      | 4     | -     | 5      | 2      | 1     | -     | 3      | -      | 4     | -     | 4  |
|            |           | Normal  | 1      | 71    | 7     | 79     | 2      | 63    | 6     | 71     | 1      | 70    | 4     | 75     | 1      | 70    | 2     | 73     | 1      | 65    | 2     | 68 |
|            |           | Upper   | -      | 4     | 4     | 8      | -      | 3     | 2     | 5      | -      | 2     | 2     | 4      | -      | 1     | -     | 1      | -      | 3     | 2     | 5  |
|            |           | Total   | 3      | 79    | 11    | 93     | 5      | 70    | 8     | 83     | 2      | 76    | 6     | 84     | 3      | 72    | 2     | 77     | 1      | 72    | 4     | 77 |
|            | Follow-up | Lower   | 2      | 4     | -     | 6      | 3      | 5     | -     | 8      | 2      | 4     | -     | 6      | 2      | 3     | -     | 5      | -      | 5     | -     | 5  |
|            |           | Normal  | 1      | 75    | 7     | 83     | 2      | 65    | 6     | 73     | 1      | 76    | 4     | 81     | 1      | 76    | 2     | 79     | 1      | 73    | 3     | 77 |
|            |           | Upper   | -      | 4     | 4     | 8      | -      | 3     | 2     | 5      | -      | 2     | 3     | 5      | -      | 1     | 2     | 3      | -      | 3     | 2     | 5  |
|            |           | Total   | 3      | 83    | 11    | 97     | 5      | 73    | 8     | 86     | 3      | 82    | 7     | 92     | 3      | 80    | 4     | 87     | 1      | 81    | 5     | 87 |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (2/14)

|            |           | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|------------|-----------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|            |           | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|            |           | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Hematocrit | Week 1    | Lower   | -      | -     | -     | -      | -      | -     | -     | 1      | 1      | -     | -     | 2      | 1      | -     | -     | 1      | -      | -     | -     | 1  |
|            |           | Normal  | -      | 31    | 2     | 33     | 1      | 31    | -     | 32     | 1      | 31    | 2     | 34     | -      | 34    | -     | 34     | -      | 43    | 2     | 45 |
|            |           | Upper   | -      | 3     | 3     | 6      | -      | 2     | 2     | 4      | -      | 1     | 2     | 3      | -      | -     | 2     | 2      | -      | -     | 1     | 1  |
|            |           | Total   | -      | 34    | 5     | 39     | 1      | 34    | 2     | 37     | 2      | 33    | 4     | 39     | 1      | 34    | 2     | 37     | 1      | 43    | 3     | 47 |
|            | Week 2    | Lower   | 1      | 1     | -     | 2      | 2      | 4     | -     | 6      | 2      | 2     | -     | 4      | 2      | 2     | -     | 4      | 1      | 3     | -     | 4  |
|            |           | Normal  | -      | 81    | 7     | 88     | -      | 73    | -     | 73     | 1      | 74    | 4     | 79     | -      | 85    | 1     | 86     | -      | 78    | 2     | 80 |
|            |           | Upper   | -      | 2     | 5     | 7      | -      | 5     | 5     | 10     | -      | 2     | 6     | 8      | -      | 2     | 1     | 3      | -      | -     | 4     | 4  |
|            |           | Total   | 1      | 84    | 12    | 97     | 2      | 82    | 5     | 89     | 3      | 78    | 10    | 91     | 2      | 89    | 2     | 93     | 1      | 81    | 6     | 88 |
|            | Week 3    | Lower   | -      | -     | -     | -      | 1      | 1     | -     | 2      | 2      | -     | -     | 2      | 1      | -     | -     | 1      | 1      | 1     | -     | 2  |
|            |           | Normal  | -      | 31    | 2     | 33     | -      | 31    | -     | 31     | -      | 31    | 3     | 34     | -      | 29    | 1     | 30     | -      | 37    | 2     | 39 |
|            |           | Upper   | -      | 1     | 3     | 4      | -      | 2     | 1     | 3      | -      | 1     | 1     | 2      | -      | 1     | -     | 1      | -      | -     | 1     | 1  |
|            |           | Total   | -      | 32    | 5     | 37     | 1      | 34    | 1     | 36     | 2      | 32    | 4     | 38     | 1      | 30    | 1     | 32     | 1      | 38    | 3     | 42 |
|            | Week 4    | Lower   | 1      | 1     | -     | 2      | 2      | 2     | -     | 4      | 3      | -     | -     | 3      | 2      | 3     | -     | 5      | 1      | 4     | -     | 5  |
|            |           | Normal  | -      | 81    | 6     | 87     | -      | 77    | -     | 77     | -      | 72    | 7     | 79     | -      | 80    | 1     | 81     | -      | 74    | 6     | 80 |
|            |           | Upper   | -      | 1     | 6     | 7      | -      | 3     | 5     | 8      | -      | 4     | 3     | 7      | -      | 2     | 1     | 3      | -      | 1     | -     | 1  |
|            |           | Total   | 1      | 83    | 12    | 96     | 2      | 82    | 5     | 89     | 3      | 76    | 10    | 89     | 2      | 85    | 2     | 89     | 1      | 79    | 6     | 86 |
|            | Week 6    | Lower   | -      | -     | -     | -      | 1      | -     | -     | 1      | 2      | -     | -     | 2      | 1      | -     | -     | 1      | 1      | 1     | -     | 2  |
|            |           | Normal  | -      | 35    | -     | 35     | -      | 35    | -     | 35     | -      | 32    | 4     | 36     | -      | 27    | -     | 27     | -      | 42    | 2     | 44 |
|            |           | Upper   | -      | 1     | 4     | 5      | -      | 2     | 2     | 4      | -      | -     | -     | -      | -      | 2     | 1     | 3      | -      | -     | 1     | 1  |
|            |           | Total   | -      | 36    | 4     | 40     | 1      | 37    | 2     | 40     | 2      | 32    | 4     | 38     | 1      | 29    | 1     | 31     | 1      | 43    | 3     | 47 |
|            | Week 8    | Lower   | 1      | 1     | -     | 2      | 1      | 2     | -     | 3      | 2      | 1     | -     | 3      | 2      | 2     | -     | 4      | -      | -     | -     | -  |
|            |           | Normal  | -      | 80    | 7     | 87     | 1      | 73    | -     | 74     | 1      | 68    | 6     | 75     | -      | 79    | -     | 79     | 1      | 74    | 4     | 79 |
|            |           | Upper   | -      | -     | 5     | 5      | -      | 4     | 5     | 9      | -      | 7     | 4     | 11     | -      | 1     | 1     | 2      | -      | 3     | 1     | 4  |
|            |           | Total   | 1      | 81    | 12    | 94     | 2      | 79    | 5     | 86     | 3      | 76    | 10    | 89     | 2      | 82    | 1     | 85     | 1      | 77    | 5     | 83 |
|            | Week 12   | Lower   | 1      | 1     | -     | 2      | 1      | 2     | -     | 3      | 2      | 1     | -     | 3      | 2      | 2     | -     | 4      | -      | 2     | -     | 2  |
|            |           | Normal  | -      | 80    | 8     | 88     | 1      | 74    | -     | 75     | -      | 71    | 5     | 76     | -      | 74    | -     | 74     | 1      | 73    | 5     | 79 |
|            |           | Upper   | -      | 1     | 4     | 5      | -      | 3     | 5     | 8      | -      | 2     | 5     | 7      | -      | 3     | 1     | 4      | -      | 1     | -     | 1  |
|            |           | Total   | 1      | 82    | 12    | 95     | 2      | 79    | 5     | 86     | 2      | 74    | 10    | 86     | 2      | 79    | 1     | 82     | 1      | 76    | 5     | 82 |
|            | Week 16   | Lower   | 1      | 3     | -     | 4      | 2      | 3     | -     | 5      | 2      | 1     | -     | 3      | 2      | 2     | -     | 4      | -      | 2     | -     | 2  |
|            |           | Normal  | -      | 75    | 8     | 83     | -      | 73    | 1     | 74     | -      | 71    | 7     | 78     | -      | 72    | 1     | 73     | 1      | 72    | 3     | 76 |
|            |           | Upper   | -      | 4     | 4     | 8      | -      | 2     | 3     | 5      | -      | 1     | 3     | 4      | -      | 3     | -     | 3      | -      | 1     | 2     | 3  |
|            |           | Total   | 1      | 82    | 12    | 95     | 2      | 78    | 4     | 84     | 2      | 73    | 10    | 85     | 2      | 77    | 1     | 80     | 1      | 75    | 5     | 81 |
|            | Week 20   | Lower   | 1      | 2     | -     | 3      | 2      | 6     | -     | 8      | 1      | 2     | -     | 3      | 1      | 2     | -     | 3      | -      | 1     | -     | 1  |
|            |           | Normal  | -      | 75    | 6     | 81     | -      | 68    | -     | 68     | 1      | 70    | 8     | 79     | 1      | 72    | -     | 73     | 1      | 72    | 3     | 76 |
|            |           | Upper   | -      | 5     | 6     | 11     | -      | 4     | 4     | 8      | -      | 1     | 2     | 3      | -      | 2     | 1     | 3      | -      | -     | 2     | 2  |
|            |           | Total   | 1      | 82    | 12    | 95     | 2      | 78    | 4     | 84     | 2      | 73    | 10    | 85     | 2      | 76    | 1     | 79     | 1      | 73    | 5     | 79 |
|            | Week 24   | Lower   | 1      | 3     | -     | 4      | 1      | 6     | -     | 7      | 1      | 2     | -     | 3      | -      | 1     | -     | 1      | 1      | 3     | -     | 4  |
|            |           | Normal  | -      | 75    | 7     | 82     | 1      | 70    | 1     | 72     | 1      | 68    | 6     | 75     | 2      | 74    | 1     | 77     | -      | 67    | 3     | 70 |
|            |           | Upper   | -      | 2     | 5     | 7      | -      | 1     | 3     | 4      | -      | 2     | 4     | 6      | -      | -     | -     | -      | -      | 2     | 2     | 4  |
|            |           | Total   | 1      | 80    | 12    | 93     | 2      | 77    | 4     | 83     | 2      | 72    | 10    | 84     | 2      | 75    | 1     | 78     | 1      | 72    | 5     | 78 |
|            | Week 28   | Lower   | 1      | 2     | -     | 3      | 1      | 3     | -     | 4      | 1      | 3     | -     | 4      | 2      | 1     | -     | 3      | -      | 2     | -     | 2  |
|            |           | Normal  | -      | 76    | 6     | 82     | 1      | 72    | 1     | 74     | 1      | 67    | 8     | 76     | -      | 71    | 1     | 72     | 1      | 66    | 3     | 70 |
|            |           | Upper   | -      | 3     | 5     | 8      | -      | 2     | 3     | 5      | -      | 2     | 2     | 4      | -      | 2     | -     | 2      | -      | 3     | 2     | 5  |
|            |           | Total   | 1      | 81    | 11    | 93     | 2      | 77    | 4     | 83     | 2      | 72    | 10    | 84     | 2      | 74    | 1     | 77     | 1      | 71    | 5     | 77 |
|            | Follow-up | Lower   | 1      | 2     | -     | 3      | 1      | 3     | -     | 4      | 2      | 3     | -     | 5      | 2      | 3     | -     | 5      | -      | 3     | -     | 3  |
|            |           | Normal  | -      | 80    | 6     | 86     | 1      | 75    | 1     | 77     | 1      | 74    | 8     | 83     | -      | 77    | 2     | 79     | 1      | 74    | 3     | 78 |
|            |           | Upper   | -      | 3     | 5     | 8      | -      | 2     | 3     | 5      | -      | 2     | 2     | 4      | -      | 2     | 1     | 3      | -      | 3     | 3     | 6  |
|            |           | Total   | 1      | 85    | 11    | 97     | 2      | 80    | 4     | 86     | 3      | 79    | 10    | 92     | 2      | 82    | 3     | 87     | 1      | 80    | 6     | 87 |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (3/14)

|           |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| RBC       | Week 1  | Lower   | -      | 1     | -     | 1      | -      | 1     | -     | 2      | -      | 2     | -     | 2      | 1      | -     | -     | 1      | 2      | -     | -     | 2  |
|           |         | Normal  | -      | 34    | 1     | 35     | -      | 28    | 2     | 30     | 1      | 33    | 1     | 35     | -      | 31    | 2     | 33     | -      | 41    | -     | 42 |
|           |         | Upper   | -      | 1     | 2     | 3      | -      | 1     | 4     | 5      | -      | 1     | 1     | 2      | -      | -     | 3     | 3      | -      | 1     | 2     | 3  |
|           |         | Total   | -      | 36    | 3     | 39     | 1      | 30    | 6     | 37     | 1      | 36    | 2     | 39     | 1      | 31    | 5     | 37     | 2      | 42    | 3     | 47 |
|           | Week 2  | Lower   | 4      | 2     | -     | 6      | 3      | 2     | -     | 5      | 2      | 1     | -     | 3      | 2      | 3     | -     | 5      | 2      | 2     | -     | 4  |
|           |         | Normal  | -      | 79    | 6     | 85     | -      | 70    | 5     | 75     | -      | 76    | 4     | 80     | 1      | 80    | 5     | 86     | -      | 77    | 2     | 79 |
|           |         | Upper   | -      | 2     | 4     | 6      | -      | 1     | 8     | 9      | -      | 3     | 5     | 8      | -      | -     | 2     | 2      | -      | 1     | 4     | 5  |
|           |         | Total   | 4      | 83    | 10    | 97     | 3      | 73    | 13    | 89     | 2      | 80    | 9     | 91     | 3      | 83    | 7     | 93     | 2      | 80    | 6     | 88 |
|           | Week 3  | Lower   | -      | -     | -     | -      | 1      | -     | -     | 1      | 1      | -     | -     | 1      | 1      | -     | -     | 1      | 1      | 1     | -     | 2  |
|           |         | Normal  | -      | 34    | 1     | 35     | -      | 29    | 2     | 31     | -      | 34    | 1     | 35     | -      | 27    | 4     | 31     | 1      | 34    | 2     | 37 |
|           |         | Upper   | -      | 1     | 1     | 2      | -      | -     | 4     | 4      | -      | 1     | 1     | 2      | -      | -     | -     | -      | -      | 2     | 1     | 3  |
|           |         | Total   | -      | 35    | 2     | 37     | 1      | 29    | 6     | 36     | 1      | 35    | 2     | 38     | 1      | 27    | 4     | 32     | 2      | 37    | 3     | 42 |
|           | Week 4  | Lower   | 4      | -     | -     | 4      | 2      | 1     | -     | 3      | 2      | 2     | -     | 4      | 3      | 1     | -     | 4      | 2      | -     | -     | 2  |
|           |         | Normal  | -      | 81    | 6     | 87     | 1      | 72    | 6     | 79     | -      | 73    | 6     | 79     | -      | 77    | 5     | 82     | -      | 78    | 2     | 80 |
|           |         | Upper   | -      | 1     | 4     | 5      | -      | -     | 7     | 7      | -      | 3     | 3     | 6      | -      | 1     | 2     | 3      | -      | -     | 4     | 4  |
|           |         | Total   | 4      | 82    | 10    | 96     | 3      | 73    | 13    | 89     | 2      | 78    | 9     | 89     | 3      | 79    | 7     | 89     | 2      | 78    | 6     | 86 |
|           | Week 6  | Lower   | -      | -     | -     | -      | 1      | 1     | -     | 2      | 1      | 2     | -     | 3      | 1      | -     | -     | 1      | 2      | 1     | -     | 3  |
|           |         | Normal  | -      | 37    | 2     | 39     | -      | 32    | 1     | 33     | -      | 32    | 2     | 34     | -      | 26    | 3     | 29     | -      | 42    | 1     | 43 |
|           |         | Upper   | -      | -     | 1     | 1      | -      | -     | 5     | 5      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | -     | 1     | 1  |
|           |         | Total   | -      | 37    | 3     | 40     | 1      | 33    | 6     | 40     | 1      | 35    | 2     | 38     | 1      | 26    | 4     | 31     | 2      | 43    | 2     | 47 |
|           | Week 8  | Lower   | 4      | 1     | -     | 5      | 2      | 1     | -     | 3      | 1      | 1     | -     | 2      | 3      | 1     | -     | 4      | 1      | 1     | -     | 2  |
|           |         | Normal  | -      | 77    | 5     | 82     | 1      | 68    | 6     | 75     | 1      | 73    | 6     | 80     | -      | 74    | 4     | 78     | 1      | 74    | 2     | 77 |
|           |         | Upper   | -      | 2     | 5     | 7      | -      | 1     | 7     | 8      | -      | 4     | 3     | 7      | -      | 2     | 1     | 3      | -      | 1     | 3     | 4  |
|           |         | Total   | 4      | 80    | 10    | 94     | 3      | 70    | 13    | 86     | 2      | 78    | 9     | 89     | 3      | 77    | 5     | 85     | 2      | 76    | 5     | 83 |
|           | Week 12 | Lower   | 2      | -     | -     | 2      | 3      | 2     | -     | 5      | 2      | -     | -     | 2      | 3      | 1     | -     | 4      | 1      | -     | -     | 1  |
|           |         | Normal  | 2      | 78    | 8     | 88     | -      | 67    | 5     | 72     | -      | 73    | 5     | 78     | -      | 72    | 1     | 73     | 1      | 74    | 3     | 78 |
|           |         | Upper   | -      | 3     | 2     | 5      | -      | 1     | 8     | 9      | -      | 2     | 4     | 6      | -      | 2     | 3     | 5      | -      | 1     | 2     | 3  |
|           |         | Total   | 4      | 81    | 10    | 95     | 3      | 70    | 13    | 86     | 2      | 75    | 9     | 86     | 3      | 75    | 4     | 82     | 2      | 75    | 5     | 82 |
|           | Week 16 | Lower   | 3      | 3     | -     | 6      | 2      | 2     | -     | 4      | 2      | 3     | -     | 5      | 2      | 1     | -     | 3      | 1      | 2     | -     | 3  |
|           |         | Normal  | 1      | 75    | 7     | 83     | 1      | 67    | 3     | 71     | -      | 68    | 7     | 75     | -      | 71    | 5     | 76     | 1      | 72    | 3     | 76 |
|           |         | Upper   | -      | 3     | 3     | 6      | -      | -     | 9     | 9      | -      | 3     | 2     | 5      | -      | 1     | -     | 1      | -      | -     | 2     | 2  |
|           |         | Total   | 4      | 81    | 10    | 95     | 3      | 69    | 12    | 84     | 2      | 74    | 9     | 85     | 2      | 73    | 5     | 80     | 2      | 74    | 5     | 81 |
|           | Week 20 | Lower   | 3      | -     | -     | 3      | 3      | 3     | -     | 6      | 1      | 4     | -     | 5      | 1      | 2     | -     | 3      | -      | 2     | -     | 2  |
|           |         | Normal  | 1      | 77    | 7     | 85     | -      | 63    | 4     | 67     | 1      | 67    | 6     | 74     | 1      | 69    | 1     | 71     | 2      | 69    | 2     | 73 |
|           |         | Upper   | -      | 4     | 3     | 7      | -      | 3     | 8     | 11     | -      | 3     | 3     | 6      | -      | 2     | 3     | 5      | -      | 1     | 3     | 4  |
|           |         | Total   | 4      | 81    | 10    | 95     | 3      | 69    | 12    | 84     | 2      | 74    | 9     | 85     | 2      | 73    | 4     | 79     | 2      | 72    | 5     | 79 |
|           | Week 24 | Lower   | 3      | 4     | -     | 7      | 2      | 4     | -     | 6      | 1      | 4     | -     | 5      | 1      | 1     | -     | 2      | 2      | 2     | -     | 4  |
|           |         | Normal  | 1      | 75    | 8     | 84     | 1      | 61    | 4     | 66     | 1      | 63    | 7     | 71     | 1      | 71    | 4     | 76     | -      | 69    | 3     | 72 |
|           |         | Upper   | -      | -     | 2     | 2      | -      | 3     | 8     | 11     | -      | 6     | 2     | 8      | -      | -     | -     | -      | -      | -     | 2     | 2  |
|           |         | Total   | 4      | 79    | 10    | 93     | 3      | 68    | 12    | 83     | 2      | 73    | 9     | 84     | 2      | 72    | 4     | 78     | 2      | 71    | 5     | 78 |
| Week 28   | Lower   | 4       | 2      | -     | 6     | 2      | 2      | -     | 4     | 1      | 5      | -     | 6     | 1      | -      | -     | 1     | 2      | 1      | -     | 3     |    |
|           | Normal  | -       | 77     | 6     | 83    | 1      | 63     | 6     | 70    | 1      | 65     | 6     | 72    | 1      | 69     | 4     | 74    | -      | 66     | 2     | 68    |    |
|           | Upper   | -       | 1      | 3     | 4     | -      | 3      | 6     | 9     | -      | 3      | 3     | 6     | -      | 2      | -     | 2     | -      | 3      | 3     | 6     |    |
|           | Total   | 4       | 80     | 9     | 93    | 3      | 68     | 12    | 83    | 2      | 73     | 9     | 84    | 2      | 71     | 4     | 77    | 2      | 70     | 5     | 77    |    |
| Follow-up | Lower   | 4       | 2      | -     | 6     | 2      | 2      | -     | 4     | 1      | 5      | -     | 6     | 2      | -      | -     | 2     | 2      | 1      | -     | 3     |    |
|           | Normal  | -       | 81     | 6     | 87    | 1      | 66     | 6     | 73    | 1      | 73     | 6     | 80    | 1      | 75     | 6     | 82    | -      | 75     | 2     | 77    |    |
|           | Upper   | -       | 1      | 3     | 4     | -      | 3      | 6     | 9     | -      | 3      | 3     | 6     | -      | 2      | 1     | 3     | -      | 3      | 4     | 7     |    |
|           | Total   | 4       | 84     | 9     | 97    | 3      | 71     | 12    | 86    | 2      | 81     | 9     | 92    | 3      | 77     | 7     | 87    | 2      | 79     | 6     | 87    |    |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (4/14)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| WBC       | Week 1 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           |        | Normal  | 1      | 31    | 3     | 35     | -      | 32    | -     | 32     | -      | 36    | -     | 36     | -      | 31    | -     | 31     | -      | 41    | 1     | 42 |
|           |        | Upper   | -      | 1     | 3     | 4      | -      | -     | 5     | 5      | -      | 2     | 1     | 3      | -      | 1     | 4     | 5      | -      | 3     | 2     | 5  |
|           |        | Total   | 1      | 32    | 6     | 39     | -      | 32    | 5     | 37     | -      | 38    | 1     | 39     | -      | 33    | 4     | 37     | -      | 44    | 3     | 47 |
|           | Week 2 | Lower   | 2      | -     | -     | 2      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | 1      | -     | 2     |    |
|           |        | Normal  | 1      | 79    | 7     | 87     | -      | 78    | 3     | 81     | -      | 78    | 4     | 82     | -      | 79    | 2     | 81     | -      | 74    | 5     | 79 |
|           |        | Upper   | -      | 4     | 4     | 8      | -      | 4     | 4     | 8      | -      | 7     | 2     | 9      | -      | 3     | 9     | 12     | -      | 1     | 6     | 7  |
|           |        | Total   | 3      | 83    | 11    | 97     | -      | 82    | 7     | 89     | -      | 85    | 6     | 91     | -      | 82    | 11    | 93     | 1      | 76    | 11    | 88 |
|           | Week 3 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 30    | 4     | 34     | -      | 30    | -     | 30     | -      | 34    | -     | 34     | -      | 28    | 1     | 29     | -      | 39    | 2     | 41 |
|           |        | Upper   | -      | -     | 2     | 2      | -      | 1     | 5     | 6      | -      | 3     | 1     | 4      | -      | 1     | 2     | 3      | -      | -     | 1     | 1  |
|           |        | Total   | 1      | 30    | 6     | 37     | -      | 31    | 5     | 36     | -      | 37    | 1     | 38     | -      | 29    | 3     | 32     | -      | 39    | 3     | 42 |
|           | Week 4 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | -      | -     | -     |    |
|           |        | Normal  | 1      | 81    | 6     | 88     | -      | 79    | 2     | 81     | -      | 78    | 2     | 80     | -      | 74    | 4     | 78     | -      | 74    | 4     | 78 |
|           |        | Upper   | -      | 2     | 5     | 7      | -      | 3     | 5     | 8      | -      | 5     | 4     | 9      | -      | 2     | 7     | 9      | -      | 1     | 7     | 8  |
|           |        | Total   | 2      | 83    | 11    | 96     | -      | 82    | 7     | 89     | -      | 83    | 6     | 89     | -      | 78    | 11    | 89     | -      | 75    | 11    | 86 |
|           | Week 6 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 2      | -     | 2     |    |
|           |        | Normal  | -      | 34    | 2     | 36     | -      | 35    | 1     | 36     | -      | 36    | -     | 36     | -      | 27    | 2     | 29     | 1      | 39    | 1     | 41 |
|           |        | Upper   | -      | 1     | 2     | 3      | -      | 2     | 2     | 4      | -      | 1     | 1     | 2      | -      | 1     | 1     | 2      | -      | 2     | 2     | 4  |
|           |        | Total   | 1      | 35    | 4     | 40     | -      | 37    | 3     | 40     | -      | 37    | 1     | 38     | -      | 28    | 3     | 31     | 1      | 43    | 3     | 47 |
| Week 8    | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | 1       | 81     | 6     | 88    | -      | 77     | 3     | 80    | -      | 79     | 2     | 81    | -      | 73     | 3     | 76    | 1      | 65     | 4     | 70    |    |
|           | Upper  | -       | -      | 5     | 5     | -      | 2      | 4     | 6     | -      | 4      | 4     | 8     | -      | 2      | 7     | 9     | -      | 6      | 7     | 13    |    |
|           | Total  | 2       | 81     | 11    | 94    | -      | 79     | 7     | 86    | -      | 83     | 6     | 89    | -      | 75     | 10    | 85    | 1      | 71     | 11    | 83    |    |
| Week 12   | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | 1       | 78     | 8     | 87    | -      | 77     | 5     | 82    | -      | 77     | 3     | 80    | -      | 70     | 4     | 74    | 1      | 68     | 5     | 74    |    |
|           | Upper  | -       | 4      | 3     | 7     | -      | 2      | 2     | 4     | -      | 3      | 3     | 6     | -      | 3      | 5     | 8     | -      | 2      | 6     | 8     |    |
|           | Total  | 2       | 82     | 11    | 95    | -      | 79     | 7     | 86    | -      | 80     | 6     | 86    | -      | 73     | 9     | 82    | 1      | 70     | 11    | 82    |    |
| Week 16   | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | 1      | -      | -     | 1     |    |
|           | Normal | 1       | 81     | 5     | 87    | -      | 74     | 2     | 76    | -      | 75     | 2     | 77    | -      | 68     | 6     | 74    | -      | 67     | 5     | 72    |    |
|           | Upper  | -       | 1      | 6     | 7     | -      | 3      | 5     | 8     | -      | 3      | 4     | 7     | -      | 3      | 3     | 6     | -      | 2      | 6     | 8     |    |
|           | Total  | 2       | 82     | 11    | 95    | -      | 77     | 7     | 84    | -      | 79     | 6     | 85    | -      | 71     | 9     | 80    | 1      | 69     | 11    | 81    |    |
| Week 20   | Lower  | -       | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     | 1      | -      | -     | 1     |    |
|           | Normal | 2       | 81     | 7     | 90    | -      | 74     | 2     | 76    | -      | 75     | 3     | 78    | -      | 67     | 6     | 73    | -      | 64     | 4     | 68    |    |
|           | Upper  | -       | 1      | 4     | 5     | -      | 1      | 5     | 6     | -      | 3      | 3     | 6     | -      | 3      | 3     | 6     | -      | 3      | 7     | 10    |    |
|           | Total  | 2       | 82     | 11    | 95    | -      | 77     | 7     | 84    | -      | 79     | 6     | 85    | -      | 70     | 9     | 79    | 1      | 67     | 11    | 79    |    |
| Week 24   | Lower  | 1       | -      | -     | 1     | -      | 1      | -     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     | 1      | -      | -     | 1     |    |
|           | Normal | 1       | 78     | 6     | 85    | -      | 74     | 2     | 76    | -      | 73     | 2     | 75    | -      | 67     | 4     | 71    | -      | 64     | 5     | 69    |    |
|           | Upper  | -       | 2      | 5     | 7     | -      | 1      | 5     | 6     | -      | 4      | 4     | 8     | -      | 2      | 5     | 7     | -      | 2      | 6     | 8     |    |
|           | Total  | 2       | 80     | 11    | 93    | -      | 76     | 7     | 83    | -      | 78     | 6     | 84    | -      | 69     | 9     | 78    | 1      | 66     | 11    | 78    |    |
| Week 28   | Lower  | 1       | -      | -     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     |    |
|           | Normal | 1       | 80     | 5     | 86    | -      | 75     | 3     | 78    | -      | 75     | 4     | 79    | -      | 66     | 3     | 69    | -      | 63     | 3     | 66    |    |
|           | Upper  | -       | -      | 6     | 6     | -      | -      | 4     | 4     | -      | 3      | 2     | 5     | -      | 2      | 6     | 8     | -      | 2      | 8     | 10    |    |
|           | Total  | 2       | 80     | 11    | 93    | -      | 76     | 7     | 83    | -      | 78     | 6     | 84    | -      | 68     | 9     | 77    | 1      | 65     | 11    | 77    |    |
| Follow-up | Lower  | 2       | -      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | 1      | -     | 2     |    |
|           | Normal | 1       | 82     | 5     | 88    | -      | 78     | 3     | 81    | -      | 83     | 4     | 87    | -      | 74     | 3     | 77    | -      | 72     | 3     | 75    |    |
|           | Upper  | -       | 1      | 6     | 7     | -      | -      | 4     | 4     | -      | 3      | 2     | 5     | -      | 3      | 7     | 10    | -      | 2      | 8     | 10    |    |
|           | Total  | 3       | 83     | 11    | 97    | -      | 79     | 7     | 86    | -      | 86     | 6     | 92    | -      | 77     | 10    | 87    | 1      | 75     | 11    | 87    |    |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (5/14)

|            |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|------------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|            |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|            |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Neutrophil | Week 1  | Lower   | -      | 1     | -     | 1      | -      | -     | -     | 1      | 1      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 38    | -     | 38     | 1      | 35    | -     | 36     | 1      | 36    | -     | 37     | 1      | 34    | -     | 35     | 1      | 43    | 1     | 45 |
|            |         | Upper   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 2     | -     | 2  |
|            |         | Total   | -      | 39    | -     | 39     | 1      | 36    | -     | 37     | 2      | 37    | -     | 39     | 1      | 34    | -     | 35     | 1      | 45    | 1     | 47 |
|            | Week 2  | Lower   | -      | 1     | -     | 1      | 2      | -     | -     | 2      | 2      | 1     | -     | 3      | 1      | 1     | -     | 2      | 1      | -     | -     | 1  |
|            |         | Normal  | 1      | 92    | 3     | 96     | 1      | 82    | -     | 83     | 1      | 84    | -     | 85     | 2      | 83    | -     | 85     | 2      | 83    | -     | 85 |
|            |         | Upper   | -      | -     | -     | -      | -      | 3     | -     | 3      | -      | 2     | -     | 2      | -      | 2     | 2     | 4      | -      | 1     | -     | 1  |
|            |         | Total   | 1      | 93    | 3     | 97     | 3      | 85    | -     | 88     | 3      | 87    | -     | 90     | 3      | 86    | 2     | 91     | 3      | 84    | -     | 87 |
|            | Week 3  | Lower   | -      | 1     | -     | 1      | -      | 1     | -     | 1      | 1      | 1     | -     | 2      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|            |         | Normal  | -      | 36    | -     | 36     | 1      | 32    | -     | 33     | 1      | 34    | -     | 35     | 1      | 30    | -     | 31     | 1      | 39    | 1     | 41 |
|            |         | Upper   | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -  |
|            |         | Total   | -      | 37    | -     | 37     | 1      | 35    | -     | 36     | 2      | 36    | -     | 38     | 1      | 30    | -     | 31     | 1      | 40    | 1     | 42 |
|            | Week 4  | Lower   | -      | -     | -     | -      | 2      | -     | -     | 2      | 3      | 1     | -     | 4      | 1      | 4     | -     | 5      | -      | 1     | -     | 1  |
|            |         | Normal  | 1      | 92    | 3     | 96     | 1      | 83    | -     | 84     | -      | 84    | -     | 84     | 2      | 80    | 1     | 83     | 3      | 80    | 1     | 84 |
|            |         | Upper   | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 2     | -     | 2  |
|            |         | Total   | 1      | 92    | 3     | 96     | 3      | 85    | -     | 88     | 3      | 86    | -     | 89     | 3      | 84    | 1     | 88     | 3      | 83    | 1     | 87 |
|            | Week 6  | Lower   | -      | 1     | -     | 1      | 1      | -     | -     | 1      | 1      | 1     | -     | 2      | -      | -     | -     | -      | -      | -     | -     | -  |
|            |         | Normal  | -      | 38    | -     | 38     | -      | 37    | -     | 37     | -      | 36    | -     | 36     | 1      | 28    | -     | 29     | -      | 46    | 1     | 47 |
|            |         | Upper   | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -  |
|            |         | Total   | -      | 40    | -     | 40     | 1      | 38    | -     | 39     | 1      | 37    | -     | 38     | 1      | 29    | -     | 30     | -      | 46    | 1     | 47 |
|            | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | 2      | 2     | -     | 4      | 1      | -     | -     | 1  |
|            |         | Normal  | 1      | 90    | 3     | 94     | 3      | 80    | -     | 83     | 3      | 82    | -     | 85     | 1      | 79    | 1     | 81     | 2      | 78    | 1     | 81 |
|            |         | Upper   | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | 3     | -     | 3      | -      | -     | -     | -      | -      | 2     | -     | 2  |
|            |         | Total   | 1      | 90    | 3     | 94     | 3      | 82    | -     | 85     | 3      | 86    | -     | 89     | 3      | 81    | 1     | 85     | 3      | 80    | 1     | 84 |
|            | Week 12 | Lower   | -      | -     | -     | -      | 1      | -     | -     | 1      | 2      | 2     | -     | 4      | 2      | -     | 2     | -      | 1      | -     | -     | 1  |
|            |         | Normal  | 1      | 87    | 3     | 91     | 2      | 78    | -     | 80     | 1      | 79    | -     | 80     | 1      | 74    | 1     | 76     | 3      | 77    | 1     | 81 |
|            |         | Upper   | -      | 3     | -     | 3      | -      | 4     | -     | 4      | -      | 2     | -     | 2      | -      | 2     | -     | 2      | -      | 1     | -     | 1  |
|            |         | Total   | 1      | 90    | 3     | 94     | 3      | 82    | -     | 85     | 3      | 83    | -     | 86     | 3      | 76    | 1     | 80     | 3      | 79    | 1     | 83 |
|            | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | 1      | -     | 2     | -      | -      | -     | -     | 1  |
|            |         | Normal  | 1      | 90    | 2     | 93     | 3      | 78    | -     | 81     | 3      | 78    | -     | 81     | 2      | 74    | 1     | 77     | 3      | 77    | 1     | 81 |
|            |         | Upper   | -      | 1     | 1     | 2      | -      | 2     | -     | 2      | -      | 3     | -     | 3      | -      | -     | -     | -      | -      | -     | -     | -  |
|            |         | Total   | 1      | 91    | 3     | 95     | 3      | 80    | -     | 83     | 3      | 82    | -     | 85     | 3      | 75    | 1     | 79     | 3      | 78    | 1     | 82 |
|            | Week 20 | Lower   | -      | -     | -     | -      | 1      | 2     | -     | 3      | 2      | -     | -     | 2      | -      | -     | -     | -      | -      | -     | -     | -  |
|            |         | Normal  | 1      | 91    | 3     | 95     | 2      | 78    | -     | 80     | 1      | 79    | -     | 80     | 3      | 73    | 1     | 77     | 2      | 74    | 1     | 77 |
|            |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 3     | -     | 3      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|            |         | Total   | 1      | 91    | 3     | 95     | 3      | 80    | -     | 83     | 3      | 82    | -     | 85     | 3      | 73    | 1     | 77     | 2      | 75    | 1     | 78 |
|            | Week 24 | Lower   | -      | -     | -     | -      | 1      | 1     | -     | 2      | 1      | -     | -     | 1      | 1      | 4     | -     | 5      | 1      | 1     | -     | 2  |
|            |         | Normal  | 1      | 87    | 3     | 91     | 2      | 78    | -     | 80     | 2      | 80    | -     | 82     | 2      | 68    | 1     | 71     | 2      | 73    | 1     | 76 |
|            |         | Upper   | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1  |
|            |         | Total   | 1      | 89    | 3     | 93     | 3      | 80    | -     | 83     | 3      | 81    | -     | 84     | 3      | 73    | 1     | 77     | 3      | 75    | 1     | 79 |
|            | Week 28 | Lower   | -      | -     | -     | -      | 1      | 1     | -     | 2      | 2      | 1     | -     | 3      | 2      | 1     | -     | 3      | -      | 1     | -     | 1  |
|            |         | Normal  | 1      | 89    | 3     | 93     | 2      | 78    | -     | 80     | 1      | 78    | -     | 79     | 1      | 70    | 1     | 72     | 3      | 73    | 1     | 77 |
|            |         | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | -     | -     | -  |
|            |         | Total   | 1      | 90    | 3     | 94     | 3      | 79    | -     | 82     | 3      | 81    | -     | 84     | 3      | 72    | 1     | 76     | 3      | 74    | 1     | 78 |
| Follow-up  | Lower   | -       | 1      | -     | 1     | 1      | 1      | -     | 2     | 2      | 1      | -     | 3     | 2      | 1      | -     | 3     | -      | 1      | -     | 1     |    |
|            | Normal  | 1       | 92     | 3     | 96    | 2      | 81     | -     | 83    | 1      | 86     | -     | 87    | 1      | 80     | 1     | 82    | 3      | 83     | 1     | 87    |    |
|            | Upper   | -       | 1      | -     | 1     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|            | Total   | 1       | 94     | 3     | 98    | 3      | 82     | -     | 85    | 3      | 89     | -     | 92    | 3      | 82     | 1     | 86    | 3      | 84     | 1     | 88    |    |



表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (6/14)

|            |           | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|------------|-----------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|            |           | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|            |           | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Lymphocyte | Week 1    | Lower   | -      | -     | -     | -      | 2      | -     | 2     | -      | -      | -     | -     | 1      | -      | -     | 1     | -      | 2      | -     | 2     |    |
|            |           | Normal  | 1      | 37    | -     | 38     | 1      | 32    | -     | 33     | -      | 35    | 1     | 36     | -      | 29    | 3     | 32     | 1      | 43    | 1     | 45 |
|            |           | Upper   | -      | 1     | -     | 1      | -      | -     | 2     | 2      | -      | 1     | 2     | 3      | -      | -     | 2     | 2      | -      | -     | -     | -  |
|            |           | Total   | 1      | 38    | -     | 39     | 1      | 34    | 2     | 37     | -      | 36    | 3     | 39     | 1      | 29    | 5     | 35     | 1      | 45    | 1     | 47 |
|            | Week 2    | Lower   | 1      | -     | -     | 1      | -      | 3     | -     | 3      | 1      | 2     | -     | 3      | 3      | 2     | -     | 5      | 1      | -     | -     | 1  |
|            |           | Normal  | 2      | 92    | 1     | 95     | 1      | 78    | 2     | 81     | -      | 82    | -     | 82     | -      | 75    | 5     | 80     | -      | 82    | 2     | 84 |
|            |           | Upper   | -      | 1     | -     | 1      | -      | 2     | 2     | 4      | -      | 1     | 4     | 5      | -      | 3     | 3     | 6      | -      | -     | 2     | 2  |
|            |           | Total   | 3      | 93    | 1     | 97     | 1      | 83    | 4     | 88     | 1      | 85    | 4     | 90     | 3      | 80    | 8     | 91     | 1      | 82    | 4     | 87 |
|            | Week 3    | Lower   | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | -     | 1     | -      | -      | -     | -     |    |
|            |           | Normal  | -      | 36    | -     | 36     | 1      | 29    | 1     | 31     | -      | 33    | 1     | 34     | -      | 26    | 2     | 28     | 1      | 39    | 1     | 41 |
|            |           | Upper   | -      | 1     | -     | 1      | -      | 2     | 1     | 3      | -      | 1     | 2     | 3      | -      | -     | 2     | 2      | -      | 1     | -     | 1  |
|            |           | Total   | -      | 37    | -     | 37     | 1      | 33    | 2     | 36     | -      | 35    | 3     | 38     | 1      | 26    | 4     | 31     | 1      | 40    | 1     | 42 |
|            | Week 4    | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | 1      | 3     | -     | 4      | 1      | 1     | -     | 2      | 1      | 2     | -     | 3  |
|            |           | Normal  | 3      | 91    | 1     | 95     | 1      | 80    | 2     | 83     | -      | 77    | 2     | 79     | 1      | 74    | 3     | 78     | 1      | 78    | 2     | 81 |
|            |           | Upper   | -      | 1     | -     | 1      | -      | 2     | 2     | 4      | -      | 4     | 2     | 6      | -      | 3     | 5     | 8      | -      | 1     | 2     | 3  |
|            |           | Total   | 3      | 92    | 1     | 96     | 1      | 83    | 4     | 88     | 1      | 84    | 4     | 89     | 2      | 78    | 8     | 88     | 2      | 81    | 4     | 87 |
|            | Week 6    | Lower   | -      | 1     | -     | 1      | -      | 3     | -     | 3      | -      | 1     | -     | 1      | 1      | -     | -     | 1      | -      | 1     | -     | 1  |
|            |           | Normal  | 1      | 36    | -     | 37     | 1      | 33    | 1     | 35     | -      | 34    | -     | 34     | -      | 25    | 2     | 27     | 1      | 45    | -     | 46 |
|            |           | Upper   | -      | 2     | -     | 2      | -      | -     | 1     | 1      | -      | 1     | 2     | 3      | -      | -     | 2     | 2      | -      | -     | -     | -  |
|            |           | Total   | 1      | 39    | -     | 40     | 1      | 36    | 2     | 39     | -      | 36    | 2     | 38     | 1      | 25    | 4     | 30     | 1      | 46    | -     | 47 |
|            | Week 8    | Lower   | -      | -     | -     | -      | -      | 2     | -     | 2      | 1      | 2     | -     | 3      | 1      | -     | -     | 1      | -      | 2     | -     | 2  |
|            |           | Normal  | 3      | 89    | 1     | 93     | 1      | 76    | 3     | 80     | -      | 80    | 4     | 84     | 1      | 73    | 4     | 78     | 2      | 76    | 3     | 81 |
|            |           | Upper   | -      | 1     | -     | 1      | -      | 2     | 1     | 3      | -      | 2     | -     | 2      | -      | 2     | 4     | 6      | -      | -     | 1     | 1  |
|            |           | Total   | 3      | 90    | 1     | 94     | 1      | 80    | 4     | 85     | 1      | 84    | 4     | 89     | 2      | 75    | 8     | 85     | 2      | 78    | 4     | 84 |
|            | Week 12   | Lower   | -      | 3     | -     | 3      | -      | 3     | -     | 3      | 1      | 2     | -     | 3      | 1      | 2     | -     | 3      | 1      | 1     | -     | 2  |
|            |           | Normal  | 3      | 86    | 1     | 90     | 1      | 76    | 2     | 79     | -      | 76    | 2     | 78     | 1      | 67    | 5     | 73     | 1      | 72    | 4     | 77 |
|            |           | Upper   | -      | 1     | -     | 1      | -      | 1     | 2     | 3      | -      | 3     | 2     | 5      | -      | 1     | 3     | 4      | -      | 4     | -     | 4  |
|            |           | Total   | 3      | 90    | 1     | 94     | 1      | 80    | 4     | 85     | 1      | 81    | 4     | 86     | 2      | 70    | 8     | 80     | 2      | 77    | 4     | 83 |
|            | Week 16   | Lower   | 1      | 1     | -     | 2      | -      | 2     | -     | 2      | 1      | 2     | -     | 3      | 2      | -     | -     | 2      | 1      | 1     | -     | 2  |
|            |           | Normal  | 2      | 89    | 1     | 92     | 1      | 73    | 2     | 76     | -      | 75    | 2     | 77     | -      | 68    | 6     | 74     | 1      | 74    | 3     | 78 |
|            |           | Upper   | -      | 1     | -     | 1      | -      | 3     | 2     | 5      | -      | 3     | 2     | 5      | -      | 1     | 2     | 3      | -      | 1     | 1     | 2  |
|            |           | Total   | 3      | 91    | 1     | 95     | 1      | 78    | 4     | 83     | 1      | 80    | 4     | 85     | 2      | 69    | 8     | 79     | 2      | 76    | 4     | 82 |
|            | Week 20   | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | 1      | 3     | -     | 4      | 2      | 1     | -     | 3      | -      | 2     | -     | 2  |
|            |           | Normal  | 3      | 91    | 1     | 95     | 1      | 74    | 2     | 77     | -      | 75    | 2     | 77     | -      | 64    | 7     | 71     | 2      | 71    | 3     | 76 |
|            |           | Upper   | -      | -     | -     | -      | -      | 3     | 2     | 5      | -      | 2     | 2     | 4      | -      | 2     | 1     | 3      | -      | -     | -     | -  |
|            |           | Total   | 3      | 91    | 1     | 95     | 1      | 78    | 4     | 83     | 1      | 80    | 4     | 85     | 2      | 67    | 8     | 77     | 2      | 73    | 3     | 78 |
|            | Week 24   | Lower   | -      | 2     | -     | 2      | -      | 1     | -     | 1      | 1      | 2     | -     | 3      | 1      | -     | -     | 1      | 1      | 2     | -     | 3  |
|            |           | Normal  | 3      | 86    | 1     | 90     | 1      | 74    | 3     | 78     | -      | 76    | 2     | 78     | 1      | 63    | 3     | 67     | 1      | 70    | 3     | 74 |
|            |           | Upper   | -      | 1     | -     | 1      | -      | 3     | 1     | 4      | -      | 1     | 2     | 3      | -      | 4     | 5     | 9      | -      | 1     | 1     | 2  |
|            |           | Total   | 3      | 89    | 1     | 93     | 1      | 78    | 4     | 83     | 1      | 79    | 4     | 84     | 2      | 67    | 8     | 77     | 2      | 73    | 4     | 79 |
|            | Week 28   | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | 1      | 1     | -     | 2      | 2      | 1     | -     | 3      | 1      | -     | -     | 1  |
|            |           | Normal  | 2      | 89    | 1     | 92     | 1      | 73    | 2     | 76     | -      | 76    | 2     | 78     | -      | 64    | 6     | 70     | 1      | 71    | 3     | 75 |
|            |           | Upper   | -      | 1     | -     | 1      | -      | 4     | 2     | 6      | -      | 2     | 2     | 4      | -      | 1     | 2     | 3      | -      | 1     | 1     | 2  |
|            |           | Total   | 3      | 90    | 1     | 94     | 1      | 77    | 4     | 82     | 1      | 79    | 4     | 84     | 2      | 66    | 8     | 76     | 2      | 72    | 4     | 78 |
|            | Follow-up | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | 1      | 1     | -     | 2      | 2      | 1     | -     | 3      | 1      | -     | -     | 1  |
|            |           | Normal  | 2      | 92    | 1     | 95     | 1      | 76    | 2     | 79     | -      | 84    | 2     | 86     | -      | 74    | 6     | 80     | 1      | 81    | 3     | 85 |
|            |           | Upper   | -      | 2     | -     | 2      | -      | 4     | 2     | 6      | -      | 2     | 2     | 4      | -      | 1     | 2     | 3      | -      | 1     | 1     | 2  |
|            |           | Total   | 3      | 94    | 1     | 98     | 1      | 80    | 4     | 85     | 1      | 87    | 4     | 92     | 2      | 76    | 8     | 86     | 2      | 82    | 4     | 88 |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (7/14)

|           |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Monocyte  | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 39    | -     | 39     | -      | 36    | -     | 36     | -      | 39    | -     | 39     | -      | 35    | -     | 35     | -      | 47    | -     | 47 |
|           |         | Upper   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 39    | -     | 39     | -      | 35    | -     | 35     | -      | 47    | -     | 47 |
|           | Week 2  | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | 1      | 94    | 1     | 96     | -      | 87    | -     | 87     | -      | 90    | -     | 90     | -      | 91    | -     | 91     | -      | 87    | -     | 87 |
|           |         | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | 1      | 95    | 1     | 97     | -      | 88    | -     | 88     | -      | 90    | -     | 90     | -      | 91    | -     | 91     | -      | 87    | -     | 87 |
|           | Week 3  | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |         | Normal  | -      | 37    | -     | 37     | -      | 35    | -     | 35     | -      | 37    | -     | 37     | -      | 31    | -     | 31     | -      | 41    | -     | 41 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | -      | 37    | -     | 37     | -      | 36    | -     | 36     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 42    | -     | 42 |
|           | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           |         | Normal  | 1      | 94    | 1     | 96     | -      | 88    | -     | 88     | -      | 89    | -     | 89     | -      | 87    | -     | 87     | -      | 87    | -     | 87 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | 1      | 94    | 1     | 96     | -      | 88    | -     | 88     | -      | 89    | -     | 89     | -      | 88    | -     | 88     | -      | 87    | -     | 87 |
|           | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 40    | -     | 40     | -      | 39    | -     | 39     | -      | 38    | -     | 38     | -      | 30    | -     | 30     | -      | 46    | -     | 46 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |         | Total   | -      | 40    | -     | 40     | -      | 39    | -     | 39     | -      | 38    | -     | 38     | -      | 30    | -     | 30     | -      | 47    | -     | 47 |
|           | Week 8  | Lower   | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | 1      | 91    | 1     | 93     | -      | 84    | -     | 84     | -      | 89    | -     | 89     | -      | 85    | -     | 85     | -      | 84    | -     | 84 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | 1      | 92    | 1     | 94     | -      | 85    | -     | 85     | -      | 89    | -     | 89     | -      | 85    | -     | 85     | -      | 84    | -     | 84 |
|           | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | 1      | 92    | 1     | 94     | -      | 85    | -     | 85     | -      | 86    | -     | 86     | -      | 80    | -     | 80     | -      | 83    | -     | 83 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | 1      | 92    | 1     | 94     | -      | 85    | -     | 85     | -      | 86    | -     | 86     | -      | 80    | -     | 80     | -      | 83    | -     | 83 |
|           | Week 16 | Lower   | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |         | Normal  | 1      | 92    | 1     | 94     | -      | 82    | -     | 82     | -      | 85    | -     | 85     | -      | 79    | -     | 79     | -      | 81    | -     | 81 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | 1      | 93    | 1     | 95     | -      | 83    | -     | 83     | -      | 85    | -     | 85     | -      | 79    | -     | 79     | -      | 82    | -     | 82 |
|           | Week 20 | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | 1      | 92    | 1     | 94     | -      | 82    | -     | 82     | -      | 85    | -     | 85     | -      | 77    | -     | 77     | -      | 78    | -     | 78 |
|           |         | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | 1      | 93    | 1     | 95     | -      | 83    | -     | 83     | -      | 85    | -     | 85     | -      | 77    | -     | 77     | -      | 78    | -     | 78 |
|           | Week 24 | Lower   | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | 1      | 90    | 1     | 92     | -      | 82    | -     | 82     | -      | 84    | -     | 84     | -      | 77    | -     | 77     | -      | 79    | -     | 79 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | 1      | 91    | 1     | 93     | -      | 83    | -     | 83     | -      | 84    | -     | 84     | -      | 77    | -     | 77     | -      | 79    | -     | 79 |
| Week 28   | Lower   | -       | 2      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | 1       | 90     | 1     | 92    | -      | 81     | -     | 81    | -      | 84     | -     | 84    | -      | 76     | -     | 76    | -      | 78     | -     | 78    |    |
|           | Upper   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total   | 1       | 92     | 1     | 94    | -      | 82     | -     | 82    | -      | 84     | -     | 84    | -      | 76     | -     | 76    | -      | 78     | -     | 78    |    |
| Follow-up | Lower   | -       | 2      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | 1       | 94     | 1     | 96    | -      | 84     | -     | 84    | -      | 92     | -     | 92    | -      | 86     | -     | 86    | -      | 88     | -     | 88    |    |
|           | Upper   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total   | 1       | 96     | 1     | 98    | -      | 85     | -     | 85    | -      | 92     | -     | 92    | -      | 86     | -     | 86    | -      | 88     | -     | 88    |    |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (8/14)

|            |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|------------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|            |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|            |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Eosinophil | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 37    | -     | 37     | -      | 36    | -     | 36     | -      | 37    | 1     | 38     | -      | 34    | -     | 34     | -      | 47    | -     | 47 |
|            |         | Upper   | -      | -     | 2     | 2      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | -     | -     | -  |
|            |         | Total   | -      | 37    | 2     | 39     | -      | 37    | -     | 37     | -      | 38    | 1     | 39     | -      | 34    | 1     | 35     | -      | 47    | -     | 47 |
|            | Week 2  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 94    | 1     | 95     | -      | 87    | -     | 87     | -      | 84    | 5     | 89     | -      | 89    | 1     | 90     | -      | 87    | -     | 87 |
|            |         | Upper   | -      | -     | 2     | 2      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | -     | -     | -  |
|            |         | Total   | -      | 94    | 3     | 97     | -      | 88    | -     | 88     | -      | 84    | 6     | 90     | -      | 89    | 2     | 91     | -      | 87    | -     | 87 |
|            | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 35    | -     | 35     | -      | 35    | -     | 35     | -      | 36    | 1     | 37     | -      | 30    | -     | 30     | -      | 41    | -     | 41 |
|            |         | Upper   | -      | 1     | 1     | 2      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | 1     | -     | 1  |
|            |         | Total   | -      | 36    | 1     | 37     | -      | 36    | -     | 36     | -      | 37    | 1     | 38     | -      | 30    | 1     | 31     | -      | 42    | -     | 42 |
|            | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 93    | 1     | 94     | -      | 87    | -     | 87     | -      | 83    | 5     | 88     | -      | 86    | 1     | 87     | -      | 83    | -     | 83 |
|            |         | Upper   | -      | -     | 2     | 2      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | 4     | -     | 4  |
|            |         | Total   | -      | 93    | 3     | 96     | -      | 88    | -     | 88     | -      | 83    | 6     | 89     | -      | 86    | 2     | 88     | -      | 87    | -     | 87 |
|            | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 38    | 1     | 39     | -      | 38    | -     | 38     | -      | 36    | 1     | 37     | -      | 29    | -     | 29     | -      | 46    | -     | 46 |
|            |         | Upper   | -      | -     | 1     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | 1     | -     | 1  |
|            |         | Total   | -      | 38    | 2     | 40     | -      | 39    | -     | 39     | -      | 37    | 1     | 38     | -      | 29    | 1     | 30     | -      | 47    | -     | 47 |
|            | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 91    | 1     | 92     | -      | 85    | -     | 85     | -      | 82    | 5     | 87     | -      | 83    | 1     | 84     | -      | 82    | -     | 82 |
|            |         | Upper   | -      | -     | 2     | 2      | -      | -     | -     | -      | -      | 1     | 1     | 2      | -      | -     | 1     | 1      | -      | 2     | -     | 2  |
|            |         | Total   | -      | 91    | 3     | 94     | -      | 85    | -     | 85     | -      | 83    | 6     | 89     | -      | 83    | 2     | 85     | -      | 84    | -     | 84 |
|            | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 90    | 3     | 93     | -      | 85    | -     | 85     | -      | 79    | 5     | 84     | -      | 78    | 1     | 79     | -      | 82    | -     | 82 |
|            |         | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | 1     | 2      | -      | -     | 1     | 1      | -      | 1     | -     | 1  |
|            |         | Total   | -      | 91    | 3     | 94     | -      | 85    | -     | 85     | -      | 80    | 6     | 86     | -      | 78    | 2     | 80     | -      | 83    | -     | 83 |
|            | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 91    | 2     | 93     | -      | 83    | -     | 83     | -      | 77    | 6     | 83     | -      | 76    | 1     | 77     | -      | 79    | -     | 79 |
|            |         | Upper   | -      | 1     | 1     | 2      | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | 1     | 1     | 2      | -      | 3     | -     | 3  |
|            |         | Total   | -      | 92    | 3     | 95     | -      | 83    | -     | 83     | -      | 79    | 6     | 85     | -      | 77    | 2     | 79     | -      | 82    | -     | 82 |
|            | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 91    | 2     | 93     | -      | 83    | -     | 83     | -      | 77    | 5     | 82     | -      | 74    | 1     | 75     | -      | 76    | -     | 76 |
|            |         | Upper   | -      | 1     | 1     | 2      | -      | -     | -     | -      | -      | 2     | 1     | 3      | -      | 1     | 1     | 2      | -      | 2     | -     | 2  |
|            |         | Total   | -      | 92    | 3     | 95     | -      | 83    | -     | 83     | -      | 79    | 6     | 85     | -      | 75    | 2     | 77     | -      | 78    | -     | 78 |
|            | Week 24 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 89    | 2     | 91     | -      | 82    | -     | 82     | -      | 78    | 5     | 83     | -      | 75    | 1     | 76     | -      | 78    | -     | 78 |
|            |         | Upper   | -      | 1     | 1     | 2      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | 1     | -     | 1  |
|            |         | Total   | -      | 90    | 3     | 93     | -      | 83    | -     | 83     | -      | 78    | 6     | 84     | -      | 75    | 2     | 77     | -      | 79    | -     | 79 |
| Week 28    | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|            | Normal  | -       | 91     | 2     | 93    | -      | 82     | -     | 82    | -      | 77     | 6     | 83    | -      | 74     | 1     | 75    | -      | 76     | -     | 76    |    |
|            | Upper   | -       | -      | 1     | 1     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | 1     | 1     | -      | 2      | -     | 2     |    |
|            | Total   | -       | 91     | 3     | 94    | -      | 82     | -     | 82    | -      | 78     | 6     | 84    | -      | 74     | 2     | 76    | -      | 78     | -     | 78    |    |
| Follow-up  | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|            | Normal  | -       | 95     | 2     | 97    | -      | 85     | -     | 85    | -      | 85     | 6     | 91    | -      | 84     | 1     | 85    | -      | 86     | -     | 86    |    |
|            | Upper   | -       | -      | 1     | 1     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | 1     | 1     | -      | 2      | -     | 2     |    |
|            | Total   | -       | 95     | 3     | 98    | -      | 85     | -     | 85    | -      | 86     | 6     | 92    | -      | 84     | 2     | 86    | -      | 88     | -     | 88    |    |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (9/14)

|           |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Basophil  | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 39    | -     | 39     | -      | 35    | -     | 35     | -      | 46    | 1     | 47 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 39    | -     | 39     | -      | 35    | -     | 35     | -      | 46    | 1     | 47 |
|           | Week 2  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 97    | -     | 97     | -      | 88    | -     | 88     | -      | 90    | -     | 90     | -      | 90    | 1     | 91     | -      | 86    | 1     | 87 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 97    | -     | 97     | -      | 88    | -     | 88     | -      | 90    | -     | 90     | -      | 90    | 1     | 91     | -      | 86    | 1     | 87 |
|           | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 37    | -     | 37     | -      | 36    | -     | 36     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 42    | -     | 42 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 37    | -     | 37     | -      | 36    | -     | 36     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 42    | -     | 42 |
|           | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 96    | -     | 96     | -      | 88    | -     | 88     | -      | 88    | -     | 88     | -      | 87    | -     | 87     | -      | 86    | 1     | 87 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     |    |
|           |         | Total   | -      | 96    | -     | 96     | -      | 88    | -     | 88     | -      | 89    | -     | 89     | -      | 87    | 1     | 88     | -      | 86    | 1     | 87 |
|           | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 40    | -     | 40     | -      | 39    | -     | 39     | -      | 38    | -     | 38     | -      | 30    | -     | 30     | -      | 45    | 1     | 46 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |         | Total   | -      | 40    | -     | 40     | -      | 39    | -     | 39     | -      | 38    | -     | 38     | -      | 30    | -     | 30     | -      | 46    | 1     | 47 |
|           | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 93    | -     | 93     | -      | 85    | -     | 85     | -      | 89    | -     | 89     | -      | 84    | 1     | 85     | -      | 83    | 1     | 84 |
|           |         | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 94    | -     | 94     | -      | 85    | -     | 85     | -      | 89    | -     | 89     | -      | 84    | 1     | 85     | -      | 83    | 1     | 84 |
|           | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 94    | -     | 94     | -      | 85    | -     | 85     | -      | 86    | -     | 86     | -      | 79    | 1     | 80     | -      | 82    | 1     | 83 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 94    | -     | 94     | -      | 85    | -     | 85     | -      | 86    | -     | 86     | -      | 79    | 1     | 80     | -      | 82    | 1     | 83 |
|           | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 95    | -     | 95     | -      | 83    | -     | 83     | -      | 85    | -     | 85     | -      | 78    | -     | 78     | -      | 81    | 1     | 82 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | 1     | -      | -      | -     |       |    |
|           |         | Total   | -      | 95    | -     | 95     | -      | 83    | -     | 83     | -      | 85    | -     | 85     | -      | 78    | 1     | 79     | -      | 81    | 1     | 82 |
|           | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 95    | -     | 95     | -      | 83    | -     | 83     | -      | 85    | -     | 85     | -      | 76    | 1     | 77     | -      | 76    | 1     | 77 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |         | Total   | -      | 95    | -     | 95     | -      | 83    | -     | 83     | -      | 85    | -     | 85     | -      | 76    | 1     | 77     | -      | 77    | 1     | 78 |
|           | Week 24 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 93    | -     | 93     | -      | 83    | -     | 83     | -      | 84    | -     | 84     | -      | 76    | -     | 76     | -      | 78    | -     | 78 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | 1     | -      | -      | 1     | 1     |    |
|           |         | Total   | -      | 93    | -     | 93     | -      | 83    | -     | 83     | -      | 84    | -     | 84     | -      | 76    | 1     | 77     | -      | 78    | 1     | 79 |
| Week 28   | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 94     | -     | 94    | -      | 82     | -     | 82    | -      | 84     | -     | 84    | -      | 76     | -     | 76    | -      | 76     | 1     | 77    |    |
|           | Upper   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Total   | -       | 94     | -     | 94    | -      | 82     | -     | 82    | -      | 84     | -     | 84    | -      | 76     | -     | 76    | -      | 77     | 1     | 78    |    |
| Follow-up | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 98     | -     | 98    | -      | 85     | -     | 85    | -      | 92     | -     | 92    | -      | 86     | -     | 86    | -      | 86     | 1     | 87    |    |
|           | Upper   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Total   | -       | 98     | -     | 98    | -      | 85     | -     | 85    | -      | 92     | -     | 92    | -      | 86     | -     | 86    | -      | 87     | 1     | 88    |    |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (10/14)

|           |           | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|-----------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |           | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |           | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Platelets | Week 1    | Lower   | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 33    | 1     | 34     | -      | 33    | -     | 33     | -      | 36    | 1     | 37     | -      | 35    | -     | 35     | -      | 45    | 1     | 46 |
|           |           | Upper   | -      | 1     | 3     | 4      | -      | 1     | 1     | 2      | -      | -     | 2     | 2      | -      | -     | 2     | 2      | -      | -     | 1     | 1  |
|           |           | Total   | -      | 34    | 4     | 38     | 1      | 34    | 1     | 36     | -      | 36    | 3     | 39     | -      | 35    | 2     | 37     | -      | 45    | 2     | 47 |
|           | Week 2    | Lower   | 3      | -     | -     | 3      | 1      | -     | -     | 1      | -      | -     | -     | -      | 1      | 2     | -     | 3      | -      | 1     | -     | 1  |
|           |           | Normal  | -      | 84    | 2     | 86     | -      | 79    | -     | 79     | -      | 80    | 1     | 81     | -      | 81    | -     | 81     | -      | 81    | 2     | 83 |
|           |           | Upper   | -      | 3     | 5     | 8      | -      | 2     | 5     | 7      | -      | 5     | 5     | 10     | -      | 5     | 2     | 7      | -      | 2     | 2     | 4  |
|           |           | Total   | 3      | 87    | 7     | 97     | 1      | 81    | 5     | 87     | -      | 85    | 6     | 91     | 1      | 88    | 2     | 91     | -      | 84    | 4     | 88 |
|           | Week 3    | Lower   | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | 2      | -     | 2     |    |
|           |           | Normal  | -      | 33    | 2     | 35     | -      | 32    | -     | 32     | -      | 36    | -     | 36     | -      | 31    | -     | 31     | -      | 38    | 1     | 39 |
|           |           | Upper   | -      | 1     | 1     | 2      | -      | 1     | 1     | 2      | -      | -     | 2     | 2      | -      | -     | 1     | 1      | -      | -     | 1     | 1  |
|           |           | Total   | -      | 34    | 3     | 37     | 1      | 33    | 1     | 35     | -      | 36    | 2     | 38     | -      | 31    | 1     | 32     | -      | 40    | 2     | 42 |
|           | Week 4    | Lower   | 2      | -     | -     | 2      | 1      | -     | -     | 1      | -      | 1     | -     | 1      | 1      | 1     | -     | 2      | -      | 2     | -     | 2  |
|           |           | Normal  | 1      | 82    | 1     | 84     | -      | 76    | -     | 76     | -      | 80    | 2     | 82     | -      | 83    | -     | 83     | -      | 79    | -     | 79 |
|           |           | Upper   | -      | 4     | 6     | 10     | -      | 4     | 5     | 9      | -      | 2     | 4     | 6      | -      | -     | 2     | 2      | -      | 1     | 4     | 5  |
|           |           | Total   | 3      | 86    | 7     | 96     | 1      | 80    | 5     | 86     | -      | 83    | 6     | 89     | 1      | 84    | 2     | 87     | -      | 82    | 4     | 86 |
|           | Week 6    | Lower   | 1      | -     | -     | 1      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |           | Normal  | -      | 35    | -     | 35     | -      | 33    | -     | 33     | -      | 36    | 1     | 37     | -      | 31    | -     | 31     | -      | 45    | -     | 45 |
|           |           | Upper   | -      | 2     | 2     | 4      | -      | 4     | 1     | 5      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | 1     | 1  |
|           |           | Total   | 1      | 37    | 2     | 40     | 1      | 37    | 1     | 39     | -      | 36    | 2     | 38     | -      | 31    | -     | 31     | -      | 46    | 1     | 47 |
|           | Week 8    | Lower   | 2      | -     | -     | 2      | 1      | -     | -     | 1      | -      | 1     | -     | 1      | 1      | 1     | -     | 2      | -      | 1     | -     | 1  |
|           |           | Normal  | 1      | 83    | 1     | 85     | -      | 76    | 1     | 77     | -      | 80    | 1     | 81     | -      | 78    | -     | 78     | -      | 75    | 1     | 76 |
|           |           | Upper   | -      | 1     | 6     | 7      | -      | 2     | 4     | 6      | -      | 2     | 5     | 7      | -      | 3     | 1     | 4      | -      | 3     | 3     | 6  |
|           |           | Total   | 3      | 84    | 7     | 94     | 1      | 78    | 5     | 84     | -      | 83    | 6     | 89     | 1      | 82    | 1     | 84     | -      | 79    | 4     | 83 |
|           | Week 12   | Lower   | 3      | 1     | -     | 4      | -      | -     | -     | -      | -      | 1     | -     | 1      | 1      | 1     | -     | 3      | -      | 1     | -     | 1  |
|           |           | Normal  | -      | 82    | 2     | 84     | -      | 76    | 1     | 77     | -      | 77    | 2     | 79     | -      | 76    | -     | 76     | -      | 76    | 1     | 77 |
|           |           | Upper   | -      | 2     | 5     | 7      | -      | 2     | 4     | 6      | -      | 3     | 3     | 6      | -      | 2     | -     | 2      | -      | 2     | 2     | 4  |
|           |           | Total   | 3      | 85    | 7     | 95     | -      | 78    | 5     | 83     | -      | 81    | 5     | 86     | 1      | 80    | -     | 81     | -      | 79    | 3     | 82 |
|           | Week 16   | Lower   | 3      | -     | -     | 3      | 1      | -     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | 1      | -      | 2     | -     | 2  |
|           |           | Normal  | -      | 84    | 1     | 85     | -      | 74    | 1     | 75     | -      | 77    | 3     | 80     | -      | 78    | -     | 78     | -      | 74    | 1     | 75 |
|           |           | Upper   | -      | 1     | 6     | 7      | -      | 2     | 4     | 6      | -      | 2     | 2     | 4      | -      | -     | -     | -      | -      | 1     | 3     | 4  |
|           |           | Total   | 3      | 85    | 7     | 95     | 1      | 76    | 5     | 82     | -      | 80    | 5     | 85     | 1      | 78    | -     | 79     | -      | 77    | 4     | 81 |
|           | Week 20   | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | 1     | -     | 2      | -      | 1     | -     | 1  |
|           |           | Normal  | 2      | 83    | 1     | 86     | -      | 73    | 3     | 76     | -      | 79    | 2     | 81     | -      | 76    | -     | 76     | -      | 73    | 1     | 74 |
|           |           | Upper   | -      | 1     | 6     | 7      | -      | 3     | 2     | 5      | -      | 1     | 3     | 4      | -      | -     | -     | -      | -      | 1     | 3     | 4  |
|           |           | Total   | 3      | 84    | 7     | 94     | -      | 76    | 5     | 81     | -      | 80    | 5     | 85     | 1      | 77    | -     | 78     | -      | 75    | 4     | 79 |
|           | Week 24   | Lower   | 3      | 1     | -     | 4      | 1      | -     | -     | 1      | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     | -  |
|           |           | Normal  | -      | 81    | 1     | 82     | -      | 73    | 1     | 74     | -      | 77    | 2     | 79     | -      | 76    | -     | 76     | -      | 72    | 1     | 73 |
|           |           | Upper   | -      | 1     | 6     | 7      | -      | 2     | 4     | 6      | -      | 2     | 3     | 5      | -      | -     | -     | -      | -      | 2     | 3     | 5  |
|           |           | Total   | 3      | 83    | 7     | 93     | 1      | 75    | 5     | 81     | -      | 79    | 5     | 84     | 1      | 76    | -     | 77     | -      | 74    | 4     | 78 |
|           | Week 28   | Lower   | 1      | -     | -     | 1      | 1      | -     | -     | 1      | -      | -     | -     | -      | 1      | 1     | -     | 2      | -      | -     | -     | -  |
|           |           | Normal  | 2      | 82    | -     | 84     | -      | 73    | 1     | 74     | -      | 77    | 2     | 79     | -      | 74    | -     | 74     | -      | 73    | 1     | 74 |
|           |           | Upper   | -      | 1     | 7     | 8      | -      | 2     | 4     | 6      | -      | 2     | 3     | 5      | -      | -     | -     | -      | -      | 3     | 3     | 3  |
|           |           | Total   | 3      | 83    | 7     | 93     | 1      | 75    | 5     | 81     | -      | 79    | 5     | 84     | 1      | 75    | -     | 76     | -      | 73    | 4     | 77 |
|           | Follow-up | Lower   | 1      | -     | -     | 1      | 1      | -     | -     | 1      | -      | -     | -     | -      | 1      | 1     | -     | 2      | -      | -     | -     | -  |
|           |           | Normal  | 2      | 86    | -     | 88     | -      | 75    | 1     | 76     | -      | 84    | 3     | 87     | -      | 81    | -     | 81     | -      | 83    | 1     | 84 |
|           |           | Upper   | -      | 1     | 7     | 8      | -      | 3     | 4     | 7      | -      | 2     | 3     | 5      | -      | 1     | 2     | 3      | -      | -     | 3     | 3  |
|           |           | Total   | 3      | 87    | 7     | 97     | 1      | 78    | 5     | 84     | -      | 86    | 6     | 92     | 1      | 83    | 2     | 86     | -      | 83    | 4     | 87 |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (11/14)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| MCV       | Week 1 | Lower   | -      | -     | 1     | 2      | 2      | -     | 4     | 1      | -      | -     | 1     | 1      | 1      | -     | 2     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 38    | -     | 38     | -      | 32    | -     | 32     | 1      | 37    | -     | 38     | -      | 33    | -     | 33     | -      | 47    | -     | 47 |
|           |        | Upper   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | 2     | 2      | -      | -     | -     | -  |
|           |        | Total   | 1      | 38    | -     | 39     | 2      | 35    | -     | 37     | 2      | 37    | -     | 39     | 1      | 34    | 2     | 37     | -      | 47    | -     | 47 |
|           | Week 2 | Lower   | 2      | -     | -     | 2      | 4      | 1     | -     | 5      | 2      | -     | -     | 2      | 1      | -     | -     | 1      | 2      | -     | 2     |    |
|           |        | Normal  | 2      | 93    | -     | 95     | -      | 83    | -     | 83     | 3      | 86    | -     | 89     | 2      | 88    | 1     | 91     | 1      | 83    | -     | 84 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | 1     | 1     | -      | 2      | -     | 2     |    |
|           |        | Total   | 4      | 93    | -     | 97     | 4      | 84    | 1     | 89     | 5      | 86    | -     | 91     | 3      | 88    | 2     | 93     | 3      | 85    | -     | 88 |
|           | Week 3 | Lower   | 1      | 1     | -     | 2      | 2      | 1     | -     | 3      | 1      | -     | -     | 1      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 35    | -     | 35     | -      | 33    | -     | 33     | 1      | 36    | -     | 37     | -      | 30    | -     | 30     | -      | 41    | -     | 41 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | 1     | -      | 1      | -     | 1     |    |
|           |        | Total   | 1      | 36    | -     | 37     | 2      | 34    | -     | 36     | 2      | 36    | -     | 38     | 1      | 30    | 1     | 32     | -      | 42    | -     | 42 |
|           | Week 4 | Lower   | 3      | -     | -     | 3      | 4      | 2     | -     | 6      | 4      | -     | -     | 4      | 1      | -     | -     | 1      | 3      | -     | 3     |    |
|           |        | Normal  | 1      | 91    | -     | 92     | -      | 81    | -     | 81     | 1      | 84    | -     | 85     | 2      | 84    | -     | 86     | -      | 82    | -     | 82 |
|           |        | Upper   | -      | 1     | -     | 1      | -      | 1     | 1     | 2      | -      | -     | -     | -      | -      | 2     | 2     | -      | 1      | -     | 1     |    |
|           |        | Total   | 4      | 92    | -     | 96     | 4      | 84    | 1     | 89     | 5      | 84    | -     | 89     | 3      | 84    | 2     | 89     | 3      | 83    | -     | 86 |
|           | Week 6 | Lower   | -      | 1     | -     | 1      | 2      | -     | -     | 2      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |        | Normal  | 1      | 38    | -     | 39     | -      | 37    | -     | 37     | 1      | 36    | -     | 37     | 1      | 28    | -     | 29     | -      | 46    | -     | 46 |
|           |        | Upper   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 2     | 2     | -      | -      | -     | -     |    |
|           |        | Total   | 1      | 39    | -     | 40     | 2      | 38    | -     | 40     | 2      | 36    | -     | 38     | 1      | 28    | 2     | 31     | -      | 47    | -     | 47 |
| Week 8    | Lower  | 3       | 2      | -     | 5     | 3      | 2      | -     | 5     | 4      | -      | -     | 4     | -      | 1      | -     | 1     | 2      | 1      | -     | 3     |    |
|           | Normal | 1       | 88     | -     | 89    | 1      | 79     | -     | 80    | 1      | 83     | -     | 84    | 2      | 80     | -     | 82    | -      | 79     | -     | 79    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | 1     | 1     | -      | 1      | -     | 1     | -      | -      | 2     | 2     | -      | 1      | -     | 1     |    |
|           | Total  | 4       | 90     | -     | 94    | 4      | 81     | 1     | 86    | 5      | 84     | -     | 89    | 2      | 81     | 2     | 85    | 2      | 81     | -     | 83    |    |
| Week 12   | Lower  | 3       | 1      | -     | 4     | 3      | 1      | -     | 4     | 3      | -      | -     | 3     | 1      | 1      | -     | 2     | 2      | 1      | -     | 3     |    |
|           | Normal | 1       | 90     | -     | 91    | 1      | 80     | -     | 81    | 1      | 81     | -     | 82    | 1      | 77     | -     | 78    | -      | 78     | -     | 78    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | 1     | 1     | -      | 1      | -     | 1     | -      | -      | 2     | 2     | -      | 1      | -     | 1     |    |
|           | Total  | 4       | 91     | -     | 95    | 4      | 81     | 1     | 86    | 4      | 82     | -     | 86    | 2      | 78     | 2     | 82    | 2      | 80     | -     | 82    |    |
| Week 16   | Lower  | 3       | 3      | -     | 6     | 3      | 2      | -     | 5     | 3      | -      | -     | 3     | 2      | 1      | -     | 3     | 2      | 1      | -     | 3     |    |
|           | Normal | 1       | 88     | -     | 89    | 1      | 77     | -     | 78    | 1      | 81     | -     | 82    | -      | 75     | -     | 75    | -      | 78     | -     | 78    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | 1     | 1     | -      | -      | -     | -     | -      | 2      | 2     | -     | -      | -      | -     |       |    |
|           | Total  | 4       | 91     | -     | 95    | 4      | 79     | 1     | 84    | 4      | 81     | -     | 85    | 2      | 76     | 2     | 80    | 2      | 79     | -     | 81    |    |
| Week 20   | Lower  | 3       | 1      | -     | 4     | 3      | 3      | -     | 6     | 2      | -      | -     | 2     | -      | -      | -     | 2     | 1      | 1      | -     | 3     |    |
|           | Normal | 1       | 90     | -     | 91    | 1      | 76     | -     | 77    | 2      | 81     | -     | 83    | 2      | 75     | 1     | 78    | -      | 76     | -     | 76    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | 1     | 1     | -      | -      | -     | -     | -      | 1      | 1     | -     | -      | -      | -     |       |    |
|           | Total  | 4       | 91     | -     | 95    | 4      | 79     | 1     | 84    | 4      | 81     | -     | 85    | 2      | 75     | 2     | 79    | 2      | 77     | -     | 79    |    |
| Week 24   | Lower  | 4       | 1      | -     | 5     | 3      | 2      | -     | 5     | 3      | -      | -     | 3     | 2      | -      | -     | 2     | 1      | 1      | -     | 2     |    |
|           | Normal | -       | 87     | -     | 87    | 1      | 76     | -     | 77    | 1      | 79     | -     | 80    | -      | 74     | 2     | 76    | 1      | 75     | -     | 76    |    |
|           | Upper  | -       | 1      | -     | 1     | -      | -      | 1     | 1     | -      | 1      | -     | 1     | -      | -      | 1     | -     | -      | -      | -     |       |    |
|           | Total  | 4       | 89     | -     | 93    | 4      | 78     | 1     | 83    | 4      | 80     | -     | 84    | 2      | 74     | 2     | 78    | 2      | 76     | -     | 78    |    |
| Week 28   | Lower  | 2       | 1      | -     | 3     | 3      | 3      | -     | 6     | 3      | 1      | -     | 4     | 2      | -      | -     | 2     | 1      | 2      | -     | 3     |    |
|           | Normal | 2       | 88     | -     | 90    | 1      | 75     | -     | 76    | 1      | 77     | -     | 78    | -      | 73     | -     | 73    | 1      | 73     | -     | 74    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | 1     | 1     | -      | 2      | -     | 2     | -      | 2      | 2     | -     | -      | -      | -     |       |    |
|           | Total  | 4       | 89     | -     | 93    | 4      | 78     | 1     | 83    | 4      | 80     | -     | 84    | 2      | 73     | 2     | 77    | 2      | 75     | -     | 77    |    |
| Follow-up | Lower  | 2       | 1      | -     | 3     | 3      | 3      | -     | 6     | 4      | 2      | -     | 6     | 3      | -      | -     | 3     | 2      | 2      | -     | 4     |    |
|           | Normal | 2       | 92     | -     | 94    | 1      | 78     | -     | 79    | 1      | 83     | -     | 84    | -      | 82     | -     | 82    | 1      | 82     | -     | 83    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | 1     | 1     | -      | 2      | -     | 2     | -      | -      | 2     | 2     | -      | -      | -     |       |    |
|           | Total  | 4       | 93     | -     | 97    | 4      | 81     | 1     | 86    | 5      | 87     | -     | 92    | 3      | 82     | 2     | 87    | 3      | 84     | -     | 87    |    |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (12/14)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |    |
| MCH       | Week 1 | Lower   | -      | 1     | -     | 1      | 4      | -     | -     | 4      | 3      | 1     | -     | 4      | 2      | -     | -     | 2      | -      | -     | -     | -  |    |
|           |        | Normal  | 2      | 36    | -     | 38     | 3      | 30    | -     | 33     | -      | 35    | -     | 35     | -      | 33    | -     | 1      | 34     | 1     | 44    | 1  | 46 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | 1  | 1  |
|           |        | Total   | 2      | 37    | -     | 39     | 7      | 30    | -     | 37     | 3      | 36    | -     | 39     | 2      | 34    | 1     | 37     | 1      | 44    | 2     | 47 | 47 |
|           | Week 2 | Lower   | 3      | 2     | -     | 5      | 9      | 1     | -     | 10     | 7      | 1     | -     | 8      | 3      | -     | -     | 3      | 3      | 2     | -     | 5  |    |
|           |        | Normal  | 1      | 89    | 1     | 91     | 3      | 75    | -     | 78     | -      | 83    | -     | 83     | -      | 87    | 1     | 88     | 1      | 78    | 2     | 81 |    |
|           |        | Upper   | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | 1     | 1     | 2  |    |
|           |        | Total   | 4      | 91    | 2     | 97     | 12     | 76    | 1     | 89     | 7      | 84    | -     | 91     | 3      | 89    | 1     | 93     | 4      | 81    | 3     | 88 |    |
|           | Week 3 | Lower   | 2      | 1     | -     | 3      | 3      | -     | -     | 3      | 3      | -     | -     | 3      | 2      | -     | -     | 2      | 1      | 1     | -     | 2  |    |
|           |        | Normal  | -      | 34    | -     | 34     | 3      | 30    | -     | 33     | -      | 35    | -     | 35     | -      | 29    | 1     | 30     | -      | 38    | -     | 38 |    |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | 2  |    |
|           |        | Total   | 2      | 35    | -     | 37     | 6      | 30    | -     | 36     | 3      | 35    | -     | 38     | 2      | 29    | 1     | 32     | 1      | 39    | 2     | 42 |    |
|           | Week 4 | Lower   | 3      | 2     | -     | 5      | 9      | -     | -     | 9      | 5      | -     | -     | 5      | 2      | -     | -     | 2      | 3      | -     | -     | 3  |    |
|           |        | Normal  | 1      | 87    | 1     | 89     | 3      | 76    | -     | 79     | -      | 84    | -     | 84     | -      | 85    | 1     | 86     | 1      | 78    | 1     | 80 |    |
|           |        | Upper   | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | 2     | 3  |    |
|           |        | Total   | 4      | 90    | 2     | 96     | 12     | 76    | 1     | 89     | 5      | 84    | -     | 89     | 2      | 86    | 1     | 89     | 4      | 79    | 3     | 86 |    |
|           | Week 6 | Lower   | 2      | -     | -     | 2      | 3      | 2     | -     | 5      | 2      | -     | -     | 2      | -      | -     | -     | -      | 1      | -     | -     | 1  |    |
|           |        | Normal  | -      | 37    | -     | 37     | 3      | 31    | -     | 34     | -      | 34    | -     | 34     | -      | 29    | 1     | 30     | -      | 43    | 1     | 44 |    |
|           |        | Upper   | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | -     | 2     | 2  |    |
|           |        | Total   | 2      | 38    | -     | 40     | 6      | 34    | -     | 40     | 2      | 36    | -     | 38     | -      | 30    | 1     | 31     | 1      | 43    | 3     | 47 |    |
| Week 8    | Lower  | 4       | 1      | -     | 5     | 10     | -      | -     | 10    | 5      | 1      | -     | 6     | 1      | -      | 1     | 4     | 1      | -      | 5     |       |    |    |
|           | Normal | -       | 87     | 1     | 88    | 2      | 72     | -     | 74    | -      | 83     | -     | 83    | -      | 82     | 1     | 83    | -      | 74     | 1     | 75    |    |    |
|           | Upper  | -       | -      | 1     | 1     | -      | 1      | 1     | 2     | -      | -      | -     | -     | 1      | -      | 1     | -     | 1      | 2      | 3     |       |    |    |
|           | Total  | 4       | 88     | 2     | 94    | 12     | 73     | 1     | 86    | 5      | 84     | -     | 89    | 1      | 83     | 1     | 85    | 4      | 76     | 3     | 83    |    |    |
| Week 12   | Lower  | 4       | 1      | -     | 5     | 8      | 1      | -     | 9     | 4      | 1      | -     | 5     | 1      | -      | 2     | 4     | -      | -      | 4     |       |    |    |
|           | Normal | -       | 88     | 1     | 89    | 4      | 72     | -     | 76    | -      | 80     | -     | 80    | -      | 79     | 1     | 80    | -      | 74     | 2     | 76    |    |    |
|           | Upper  | -       | -      | 1     | 1     | -      | -      | 1     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | 1      | 1     | 2     |    |    |
|           | Total  | 4       | 89     | 2     | 95    | 12     | 73     | 1     | 86    | 4      | 82     | -     | 86    | 1      | 80     | 1     | 82    | 4      | 75     | 3     | 82    |    |    |
| Week 16   | Lower  | 4       | 1      | -     | 5     | 8      | 1      | -     | 9     | 4      | 1      | -     | 5     | 1      | -      | 2     | 4     | -      | -      | 4     |       |    |    |
|           | Normal | -       | 88     | 1     | 89    | 4      | 70     | -     | 74    | -      | 79     | -     | 79    | -      | 76     | 1     | 77    | -      | 73     | 2     | 75    |    |    |
|           | Upper  | -       | -      | 1     | 1     | -      | -      | 1     | 1     | -      | 1      | -     | 1     | -      | 1      | -     | 1     | -      | 1      | 1     | 2     |    |    |
|           | Total  | 4       | 89     | 2     | 95    | 12     | 71     | 1     | 84    | 4      | 81     | -     | 85    | 1      | 78     | 1     | 80    | 4      | 74     | 3     | 81    |    |    |
| Week 20   | Lower  | 4       | 2      | -     | 6     | 9      | 1      | -     | 10    | 4      | 1      | -     | 5     | 1      | -      | 1     | 4     | 1      | -      | 5     |       |    |    |
|           | Normal | -       | 87     | 1     | 88    | 3      | 70     | -     | 73    | -      | 79     | -     | 79    | -      | 77     | 1     | 78    | -      | 71     | 1     | 72    |    |    |
|           | Upper  | -       | -      | 1     | 1     | -      | -      | 1     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | 2     | 2     |    |    |
|           | Total  | 4       | 89     | 2     | 95    | 12     | 71     | 1     | 84    | 4      | 81     | -     | 85    | 1      | 77     | 1     | 79    | 4      | 72     | 3     | 79    |    |    |
| Week 24   | Lower  | 3       | 1      | -     | 4     | 6      | 1      | -     | 7     | 4      | -      | -     | 4     | -      | -      | -     | 3     | 1      | -      | 4     |       |    |    |
|           | Normal | 1       | 85     | 1     | 87    | 6      | 69     | -     | 75    | -      | 79     | -     | 79    | 1      | 75     | 1     | 77    | 1      | 69     | 1     | 71    |    |    |
|           | Upper  | -       | 1      | 1     | 2     | -      | -      | 1     | 1     | -      | 1      | -     | 1     | -      | 1      | -     | 1     | -      | 1      | 2     | 3     |    |    |
|           | Total  | 4       | 87     | 2     | 93    | 12     | 70     | 1     | 83    | 4      | 80     | -     | 84    | 1      | 76     | 1     | 78    | 4      | 71     | 3     | 78    |    |    |
| Week 28   | Lower  | 3       | 1      | -     | 4     | 9      | 1      | -     | 10    | 3      | -      | -     | 3     | 1      | 1      | -     | 2     | 3      | 1      | -     | 4     |    |    |
|           | Normal | 1       | 86     | 1     | 88    | 3      | 69     | -     | 72    | 1      | 79     | -     | 80    | -      | 74     | 1     | 75    | 1      | 70     | 2     | 73    |    |    |
|           | Upper  | -       | -      | 1     | 1     | -      | -      | 1     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     |    |    |
|           | Total  | 4       | 87     | 2     | 93    | 12     | 70     | 1     | 83    | 4      | 80     | -     | 84    | 1      | 75     | 1     | 77    | 4      | 71     | 2     | 77    |    |    |
| Follow-up | Lower  | 3       | 1      | -     | 4     | 9      | 2      | -     | 11    | 5      | -      | -     | 5     | 3      | 2      | -     | 5     | 3      | 2      | -     | 5     |    |    |
|           | Normal | 1       | 90     | 1     | 92    | 3      | 71     | -     | 74    | 2      | 84     | -     | 86    | -      | 81     | 1     | 82    | 1      | 79     | 2     | 82    |    |    |
|           | Upper  | -       | -      | 1     | 1     | -      | -      | 1     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     |    |    |
|           | Total  | 4       | 91     | 2     | 97    | 12     | 73     | 1     | 86    | 7      | 85     | -     | 92    | 3      | 83     | 1     | 87    | 4      | 81     | 2     | 87    |    |    |

表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (13/14)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| MCHC      | Week 1 | Lower   | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 38    | 1     | 39     | -      | 36    | -     | 36     | 1      | 36    | -     | 38     | -      | 37    | -     | 37     | -      | 44    | 1     | 45 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 2     | -     | 2  |
|           |        | Total   | -      | 38    | 1     | 39     | 1      | 36    | -     | 37     | 1      | 37    | 1     | 39     | -      | 37    | -     | 37     | -      | 46    | 1     | 47 |
|           | Week 2 | Lower   | 1      | -     | -     | 1      | 2      | -     | -     | 2      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 90    | 2     | 92     | -      | 83    | 1     | 84     | -      | 88    | 1     | 89     | -      | 90    | 1     | 91     | -      | 83    | -     | 83 |
|           |        | Upper   | -      | 2     | 2     | 4      | -      | 3     | -     | 3      | -      | 1     | -     | 1      | -      | 1     | 1     | 2      | -      | 3     | 2     | 5  |
|           |        | Total   | 1      | 92    | 4     | 97     | 2      | 86    | 1     | 89     | 1      | 89    | 1     | 91     | -      | 91    | 2     | 93     | -      | 86    | 2     | 88 |
|           | Week 3 | Lower   | -      | -     | -     | -      | 1      | -     | -     | 1      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 36    | -     | 36     | -      | 34    | -     | 34     | -      | 36    | -     | 36     | -      | 32    | -     | 32     | -      | 41    | -     | 41 |
|           |        | Upper   | -      | -     | 1     | 1      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | 1     | 1  |
|           |        | Total   | -      | 36    | 1     | 37     | 1      | 35    | -     | 36     | 1      | 36    | 1     | 38     | -      | 32    | -     | 32     | -      | 41    | 1     | 42 |
|           | Week 4 | Lower   | 1      | -     | -     | 1      | 2      | -     | -     | 2      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 91    | 2     | 93     | -      | 85    | 1     | 86     | -      | 85    | 1     | 86     | -      | 86    | 2     | 88     | -      | 83    | -     | 83 |
|           |        | Upper   | -      | -     | 2     | 2      | -      | 1     | -     | 1      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | 1     | 2     | 3  |
|           |        | Total   | 1      | 91    | 4     | 96     | 2      | 86    | 1     | 89     | 1      | 87    | 1     | 89     | -      | 87    | 2     | 89     | -      | 84    | 2     | 86 |
|           | Week 6 | Lower   | -      | -     | -     | -      | 1      | -     | -     | 1      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 36    | 1     | 37     | -      | 39    | -     | 39     | -      | 34    | -     | 34     | -      | 30    | -     | 30     | -      | 46    | 1     | 47 |
|           |        | Upper   | -      | 3     | -     | 3      | -      | -     | -     | -      | -      | 2     | 1     | 3      | -      | 1     | -     | 1      | -      | -     | -     |    |
|           |        | Total   | -      | 39    | 1     | 40     | 1      | 39    | -     | 40     | 1      | 36    | 1     | 38     | -      | 31    | -     | 31     | -      | 46    | 1     | 47 |
| Week 8    | Lower  | 1       | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 88     | 1     | 89    | 1      | 82     | -     | 83    | -      | 86     | -     | 86    | -      | 82     | 2     | 84    | -      | 80     | -     | 80    |    |
|           | Upper  | -       | 1      | 3     | 4     | -      | 1      | 1     | 2     | -      | 1      | 1     | 2     | -      | 1      | -     | 1     | -      | 1      | 2     | 3     |    |
|           | Total  | 1       | 89     | 4     | 94    | 2      | 83     | 1     | 86    | 1      | 87     | 1     | 89    | -      | 83     | 2     | 85    | -      | 81     | 2     | 83    |    |
| Week 12   | Lower  | 1       | -      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 90     | 3     | 93    | 1      | 80     | 1     | 82    | -      | 84     | 1     | 85    | -      | 79     | 2     | 81    | -      | 80     | 1     | 81    |    |
|           | Upper  | -       | -      | 1     | 1     | -      | 3      | -     | 3     | -      | 1      | -     | 1     | -      | 1      | -     | 1     | -      | -      | 1     | 1     |    |
|           | Total  | 1       | 90     | 4     | 95    | 2      | 83     | 1     | 86    | -      | 85     | 1     | 86    | -      | 80     | 2     | 82    | -      | 80     | 2     | 82    |    |
| Week 16   | Lower  | 1       | -      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 87     | 1     | 88    | 1      | 78     | -     | 79    | -      | 79     | 1     | 80    | -      | 78     | 2     | 80    | -      | 78     | -     | 78    |    |
|           | Upper  | -       | 3      | 3     | 6     | -      | 3      | 1     | 4     | -      | 5      | -     | 5     | -      | -      | -     | -     | -      | 1      | 2     | 3     |    |
|           | Total  | 1       | 90     | 4     | 95    | 2      | 81     | 1     | 84    | -      | 84     | 1     | 85    | -      | 78     | 2     | 80    | -      | 79     | 2     | 81    |    |
| Week 20   | Lower  | 1       | -      | -     | 1     | 2      | -      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 88     | 2     | 90    | -      | 81     | 1     | 82    | -      | 82     | 1     | 83    | -      | 73     | 2     | 75    | -      | 77     | -     | 77    |    |
|           | Upper  | -       | 2      | 2     | 4     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | 4      | -     | 4     | -      | -      | 2     | 2     |    |
|           | Total  | 1       | 90     | 4     | 95    | 2      | 81     | 1     | 84    | -      | 84     | 1     | 85    | -      | 77     | 2     | 79    | -      | 77     | 2     | 79    |    |
| Week 24   | Lower  | 1       | -      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 84     | 3     | 87    | 1      | 79     | 1     | 81    | -      | 81     | 1     | 82    | -      | 74     | 1     | 75    | -      | 74     | 1     | 75    |    |
|           | Upper  | -       | 4      | 1     | 5     | -      | 1      | -     | 1     | -      | 2      | -     | 2     | -      | 2      | 1     | 3     | -      | 2      | 1     | 3     |    |
|           | Total  | 1       | 88     | 4     | 93    | 2      | 80     | 1     | 83    | -      | 83     | 1     | 84    | -      | 76     | 2     | 78    | -      | 76     | 2     | 78    |    |
| Week 28   | Lower  | 1       | -      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 82     | 2     | 84    | 1      | 79     | 1     | 81    | -      | 79     | 1     | 80    | -      | 72     | 2     | 74    | -      | 75     | -     | 75    |    |
|           | Upper  | -       | 6      | 2     | 8     | -      | 1      | -     | 1     | -      | 4      | -     | 4     | -      | 3      | -     | 3     | -      | 1      | 1     | 2     |    |
|           | Total  | 1       | 88     | 4     | 93    | 2      | 80     | 1     | 83    | -      | 83     | 1     | 84    | -      | 75     | 2     | 77    | -      | 76     | 1     | 77    |    |
| Follow-up | Lower  | 1       | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 86     | 2     | 88    | 1      | 82     | 1     | 84    | -      | 86     | 1     | 87    | -      | 81     | 2     | 83    | -      | 85     | -     | 85    |    |
|           | Upper  | -       | 6      | 2     | 8     | -      | 1      | -     | 1     | -      | 4      | -     | 4     | -      | 4      | -     | 4     | -      | 1      | 1     | 2     |    |
|           | Total  | 1       | 92     | 4     | 97    | 2      | 83     | 1     | 86    | 1      | 90     | 1     | 92    | -      | 85     | 2     | 87    | -      | 86     | 1     | 87    |    |



表 2.7.4.7-付録-13 血液学的検査の評価時点別正異判定クロス表 (CCT-001 試験) (14/14)

|                  |           | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|------------------|-----------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|                  |           | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|                  |           | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Prothrombin Time | Week 1    | Lower   | -      | 1     | -     | 1      | -      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                  |           | Normal  | -      | 36    | 1     | 37     | -      | 34    | 1     | 35     | -      | 35    | 1     | 36     | -      | 36    | -     | 36     | 1      | 43    | 1     | 45 |
|                  |           | Upper   | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | -     | 2     | 2  |
|                  |           | Total   | -      | 37    | 2     | 39     | 1      | 35    | 1     | 37     | -      | 38    | 1     | 39     | -      | 37    | -     | 37     | 1      | 43    | 3     | 47 |
|                  | Week 2    | Lower   | -      | 1     | -     | 1      | -      | -     | -     | -      | 1      | 1     | -     | 2      | -      | 1     | -     | 1      | 1      | 1     | -     | 2  |
|                  |           | Normal  | 1      | 89    | 2     | 92     | 2      | 80    | 1     | 83     | -      | 83    | 2     | 85     | -      | 87    | 1     | 88     | -      | 82    | 2     | 84 |
|                  |           | Upper   | -      | 1     | 3     | 4      | -      | 3     | 3     | 6      | -      | 4     | -     | 4      | -      | 3     | 1     | 4      | -      | 1     | 2     | 3  |
|                  |           | Total   | 1      | 91    | 5     | 97     | 2      | 83    | 4     | 89     | 1      | 88    | 2     | 91     | -      | 91    | 2     | 93     | 1      | 84    | 4     | 89 |
|                  | Week 3    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                  |           | Normal  | -      | 34    | 1     | 35     | 1      | 34    | 1     | 36     | -      | 35    | 1     | 36     | -      | 32    | -     | 32     | -      | 39    | 1     | 40 |
|                  |           | Upper   | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | -     | -     | -      | -      | -     | 2     | 2  |
|                  |           | Total   | -      | 34    | 2     | 36     | 1      | 34    | 1     | 36     | -      | 37    | 1     | 38     | -      | 32    | -     | 32     | -      | 39    | 3     | 42 |
|                  | Week 4    | Lower   | -      | 1     | -     | 1      | 1      | 3     | -     | 4      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|                  |           | Normal  | 1      | 88    | 3     | 92     | 1      | 78    | 1     | 80     | 1      | 80    | 1     | 82     | -      | 82    | 1     | 83     | 1      | 78    | 3     | 82 |
|                  |           | Upper   | -      | 1     | 2     | 3      | -      | 2     | 3     | 5      | -      | 5     | 1     | 6      | -      | 5     | 1     | 6      | -      | 4     | 1     | 5  |
|                  |           | Total   | 1      | 90    | 5     | 96     | 2      | 83    | 4     | 89     | 1      | 86    | 2     | 89     | -      | 87    | 2     | 89     | 1      | 82    | 4     | 87 |
|                  | Week 6    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                  |           | Normal  | -      | 38    | 1     | 39     | 1      | 38    | 1     | 40     | -      | 36    | 1     | 37     | -      | 31    | -     | 31     | -      | 41    | 2     | 43 |
|                  |           | Upper   | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 2     | 2     | 4  |
|                  |           | Total   | -      | 38    | 2     | 40     | 1      | 38    | 1     | 40     | -      | 37    | 1     | 38     | -      | 31    | -     | 31     | -      | 43    | 4     | 47 |
|                  | Week 8    | Lower   | -      | -     | -     | -      | -      | 2     | -     | 2      | 1      | -     | -     | 1      | -      | 1     | -     | 1      | 1      | 1     | -     | 2  |
|                  |           | Normal  | 1      | 86    | 2     | 89     | 2      | 78    | 1     | 81     | -      | 84    | 2     | 86     | -      | 80    | 1     | 81     | -      | 74    | 3     | 77 |
|                  |           | Upper   | -      | 2     | 3     | 5      | -      | -     | 3     | 3      | -      | 2     | -     | 2      | -      | 3     | 1     | 4      | -      | 4     | 1     | 5  |
|                  |           | Total   | 1      | 88    | 5     | 94     | 2      | 80    | 4     | 86     | 1      | 86    | 2     | 89     | -      | 84    | 2     | 86     | 1      | 79    | 4     | 84 |
|                  | Week 12   | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | 1      | 3     | -     | 4      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|                  |           | Normal  | 1      | 88    | 2     | 91     | 2      | 79    | 2     | 83     | -      | 77    | 2     | 79     | -      | 79    | 1     | 80     | 1      | 76    | 3     | 80 |
|                  |           | Upper   | -      | 1     | 3     | 4      | -      | -     | 2     | 2      | -      | 3     | -     | 3      | -      | 1     | 1     | 2      | -      | 1     | 1     | 2  |
|                  |           | Total   | 1      | 89    | 5     | 95     | 2      | 80    | 4     | 86     | 1      | 83    | 2     | 86     | -      | 80    | 2     | 82     | 1      | 78    | 4     | 83 |
|                  | Week 16   | Lower   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 2     | -     | 2      | -      | 1     | -     | 1  |
|                  |           | Normal  | 1      | 87    | 1     | 89     | 2      | 77    | 2     | 81     | 1      | 79    | 1     | 81     | -      | 75    | 1     | 76     | 1      | 73    | 3     | 77 |
|                  |           | Upper   | -      | 1     | 3     | 4      | -      | 1     | 2     | 3      | -      | 2     | 1     | 3      | -      | -     | 1     | 1      | -      | 3     | 1     | 4  |
|                  |           | Total   | 1      | 89    | 4     | 94     | 2      | 78    | 4     | 84     | 1      | 82    | 2     | 85     | -      | 77    | 2     | 79     | 1      | 77    | 4     | 82 |
|                  | Week 20   | Lower   | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | -     | -     |    |
|                  |           | Normal  | 1      | 87    | 2     | 90     | 2      | 76    | 2     | 80     | 1      | 77    | 2     | 80     | -      | 72    | 1     | 73     | 1      | 73    | 3     | 77 |
|                  |           | Upper   | -      | 1     | 3     | 4      | -      | 1     | 2     | 3      | -      | 3     | -     | 3      | -      | 3     | 1     | 4      | -      | 2     | 1     | 3  |
|                  |           | Total   | 1      | 89    | 5     | 95     | 2      | 78    | 4     | 84     | 1      | 82    | 2     | 85     | -      | 76    | 2     | 78     | 1      | 75    | 4     | 80 |
|                  | Week 24   | Lower   | 1      | 1     | -     | 2      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|                  |           | Normal  | -      | 83    | 2     | 85     | 2      | 78    | 2     | 82     | 1      | 77    | 2     | 80     | -      | 74    | 1     | 75     | 1      | 72    | 3     | 76 |
|                  |           | Upper   | -      | 3     | 3     | 6      | -      | -     | 2     | 2      | -      | 4     | -     | 4      | -      | 2     | 1     | 3      | -      | 1     | 1     | 2  |
|                  |           | Total   | 1      | 87    | 5     | 93     | 2      | 78    | 4     | 84     | 1      | 81    | 2     | 84     | -      | 76    | 2     | 78     | 1      | 74    | 4     | 79 |
|                  | Week 28   | Lower   | 1      | 2     | -     | 3      | 1      | 1     | -     | 2      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | 2     | -     | 2  |
|                  |           | Normal  | -      | 83    | 2     | 85     | 1      | 75    | 3     | 79     | -      | 77    | 2     | 79     | -      | 73    | 2     | 75     | 1      | 71    | 2     | 74 |
|                  |           | Upper   | -      | 3     | 3     | 6      | -      | 1     | 1     | 2      | -      | 4     | -     | 4      | -      | 2     | -     | 2      | -      | -     | 2     | 2  |
|                  |           | Total   | 1      | 88    | 5     | 94     | 2      | 77    | 4     | 83     | 1      | 81    | 2     | 84     | -      | 75    | 2     | 77     | 1      | 73    | 4     | 78 |
|                  | Follow-up | Lower   | 1      | 2     | -     | 3      | 1      | 1     | -     | 2      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | 2     | -     | 2  |
|                  |           | Normal  | -      | 87    | 2     | 89     | 1      | 77    | 3     | 81     | -      | 85    | 2     | 87     | -      | 82    | 2     | 84     | 1      | 81    | 2     | 84 |
|                  |           | Upper   | -      | 3     | 3     | 6      | -      | 2     | 1     | 3      | -      | 4     | -     | 4      | -      | 3     | -     | 3      | -      | -     | 2     | 2  |
|                  |           | Total   | 1      | 92    | 5     | 98     | 2      | 80    | 4     | 86     | 1      | 89    | 2     | 92     | -      | 85    | 2     | 87     | 1      | 83    | 4     | 88 |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (1/24)

| Item                    | Treatment | Summary Statistics |                      |       |       |       |       |        |       |         | One-sample t-test |         |         |
|-------------------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|---------|-------------------|---------|---------|
|                         |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       | t-value | p-value           |         |         |
|                         |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    |         |                   | Max     |         |
| Total Protein<br>(g/dL) | Week 1    | Placebo            | 7.55                 | 39    | -0.09 | 0.337 | -0.9  | -0.30  | -0.10 | 0.20    | 0.6               | -1.6140 | 0.1148  |
|                         |           | 15 mg              | 7.60                 | 37    | -0.09 | 0.322 | -1.0  | -0.30  | -0.10 | 0.10    | 0.6               | -1.6841 | 0.1008  |
|                         |           | 30 mg              | 7.44                 | 39    | 0.02  | 0.291 | -0.4  | -0.20  | 0.00  | 0.20    | 0.6               | 0.4398  | 0.6626  |
|                         |           | 60 mg              | 7.51                 | 37    | -0.16 | 0.277 | -0.9  | -0.30  | -0.20 | 0.10    | 0.2               | -3.5579 | 0.0011  |
|                         |           | 120 mg             | 7.52                 | 47    | -0.08 | 0.259 | -0.6  | -0.30  | -0.10 | 0.10    | 0.4               | -2.1962 | 0.0332  |
|                         | Week 2    | Placebo            | 7.63                 | 96    | -0.09 | 0.329 | -1.2  | -0.30  | -0.10 | 0.10    | 0.7               | -2.6350 | 0.0098  |
|                         |           | 15 mg              | 7.63                 | 89    | -0.04 | 0.305 | -0.9  | -0.20  | 0.00  | 0.20    | 0.6               | -1.1454 | 0.2552  |
|                         |           | 30 mg              | 7.51                 | 91    | -0.06 | 0.314 | -1.0  | -0.30  | -0.10 | 0.20    | 0.5               | -1.9704 | 0.0519  |
|                         |           | 60 mg              | 7.53                 | 93    | -0.09 | 0.314 | -1.1  | -0.30  | -0.10 | 0.10    | 0.7               | -2.8067 | 0.0061  |
|                         |           | 120 mg             | 7.56                 | 89    | -0.13 | 0.295 | -0.8  | -0.40  | -0.10 | 0.10    | 0.7               | -4.2352 | <0.0001 |
|                         | Week 3    | Placebo            | 7.59                 | 36    | -0.03 | 0.327 | -0.8  | -0.20  | 0.00  | 0.20    | 0.8               | -0.5104 | 0.6130  |
|                         |           | 15 mg              | 7.56                 | 36    | -0.03 | 0.330 | -0.8  | -0.20  | 0.00  | 0.10    | 0.9               | -0.6054 | 0.5488  |
|                         |           | 30 mg              | 7.44                 | 38    | -0.05 | 0.327 | -0.9  | -0.30  | 0.00  | 0.10    | 1.0               | -0.9427 | 0.3519  |
|                         |           | 60 mg              | 7.53                 | 32    | -0.11 | 0.296 | -0.8  | -0.30  | -0.15 | 0.10    | 0.4               | -2.1504 | 0.0394  |
|                         |           | 120 mg             | 7.54                 | 42    | -0.13 | 0.268 | -0.6  | -0.30  | -0.10 | 0.10    | 0.4               | -3.0534 | 0.0040  |
|                         | Week 4    | Placebo            | 7.63                 | 95    | -0.03 | 0.309 | -0.8  | -0.20  | 0.00  | 0.20    | 0.7               | -0.9636 | 0.3377  |
|                         |           | 15 mg              | 7.63                 | 89    | 0.01  | 0.344 | -1.0  | -0.20  | 0.00  | 0.20    | 0.9               | 0.1849  | 0.8538  |
|                         |           | 30 mg              | 7.51                 | 89    | -0.03 | 0.317 | -0.6  | -0.30  | -0.10 | 0.20    | 1.0               | -1.0023 | 0.3190  |
|                         |           | 60 mg              | 7.54                 | 89    | -0.06 | 0.330 | -0.8  | -0.30  | -0.10 | 0.10    | 0.9               | -1.7679 | 0.0805  |
|                         |           | 120 mg             | 7.56                 | 87    | -0.05 | 0.340 | -1.0  | -0.30  | -0.10 | 0.20    | 0.7               | -1.4494 | 0.1509  |
| Week 6                  | Placebo   | 7.59               | 40                   | -0.01 | 0.324 | -0.9  | -0.20 | 0.00   | 0.20  | 0.6     | -0.1952           | 0.8463  |         |
|                         | 15 mg     | 7.62               | 40                   | -0.05 | 0.307 | -0.9  | -0.20 | 0.00   | 0.20  | 0.4     | -1.0295           | 0.3096  |         |
|                         | 30 mg     | 7.46               | 38                   | -0.05 | 0.269 | -0.6  | -0.20 | -0.05  | 0.10  | 0.8     | -1.1463           | 0.2590  |         |
|                         | 60 mg     | 7.53               | 31                   | -0.13 | 0.302 | -0.8  | -0.30 | -0.10  | 0.10  | 0.6     | -2.3755           | 0.0241  |         |
|                         | 120 mg    | 7.55               | 47                   | -0.08 | 0.300 | -0.6  | -0.30 | -0.10  | 0.10  | 0.9     | -1.7993           | 0.0785  |         |
| Week 8                  | Placebo   | 7.63               | 94                   | -0.08 | 0.343 | -0.9  | -0.30 | -0.10  | 0.20  | 0.8     | -2.2584           | 0.0263  |         |
|                         | 15 mg     | 7.63               | 86                   | -0.02 | 0.326 | -0.8  | -0.20 | 0.00   | 0.20  | 0.9     | -0.4625           | 0.6449  |         |
|                         | 30 mg     | 7.51               | 89                   | -0.03 | 0.364 | -0.9  | -0.20 | -0.10  | 0.10  | 0.9     | -0.8744           | 0.3843  |         |
|                         | 60 mg     | 7.54               | 86                   | -0.03 | 0.360 | -0.8  | -0.30 | -0.10  | 0.20  | 1.0     | -0.8696           | 0.3870  |         |
|                         | 120 mg    | 7.55               | 84                   | -0.06 | 0.337 | -0.9  | -0.30 | -0.10  | 0.10  | 0.8     | -1.7151           | 0.0901  |         |
| Week 12                 | Placebo   | 7.63               | 94                   | -0.06 | 0.361 | -1.2  | -0.30 | 0.00   | 0.20  | 1.2     | -1.4862           | 0.1406  |         |
|                         | 15 mg     | 7.63               | 86                   | -0.05 | 0.316 | -0.8  | -0.30 | 0.00   | 0.20  | 0.7     | -1.3645           | 0.1760  |         |
|                         | 30 mg     | 7.53               | 86                   | -0.08 | 0.359 | -0.9  | -0.30 | -0.10  | 0.20  | 0.8     | -2.0131           | 0.0473  |         |
|                         | 60 mg     | 7.54               | 82                   | -0.08 | 0.363 | -1.0  | -0.30 | -0.10  | 0.20  | 0.9     | -1.9445           | 0.0553  |         |
|                         | 120 mg    | 7.55               | 83                   | -0.06 | 0.379 | -0.8  | -0.30 | 0.00   | 0.20  | 1.0     | -1.4468           | 0.1518  |         |
| Week 16                 | Placebo   | 7.63               | 94                   | -0.08 | 0.322 | -0.8  | -0.30 | -0.10  | 0.10  | 0.7     | -2.3988           | 0.0184  |         |
|                         | 15 mg     | 7.62               | 84                   | -0.03 | 0.316 | -0.8  | -0.25 | 0.00   | 0.15  | 1.1     | -0.8275           | 0.4103  |         |
|                         | 30 mg     | 7.52               | 85                   | -0.07 | 0.373 | -1.2  | -0.30 | -0.10  | 0.20  | 1.1     | -1.6560           | 0.1014  |         |
|                         | 60 mg     | 7.56               | 80                   | -0.07 | 0.398 | -1.1  | -0.30 | -0.10  | 0.20  | 0.9     | -1.5722           | 0.1199  |         |
|                         | 120 mg    | 7.56               | 82                   | -0.04 | 0.387 | -0.9  | -0.30 | 0.00   | 0.20  | 1.3     | -1.0000           | 0.3203  |         |
| Week 20                 | Placebo   | 7.63               | 94                   | -0.07 | 0.366 | -1.1  | -0.30 | -0.05  | 0.20  | 1.1     | -1.9140           | 0.0587  |         |
|                         | 15 mg     | 7.62               | 84                   | -0.03 | 0.366 | -0.8  | -0.20 | -0.05  | 0.20  | 1.0     | -0.6551           | 0.5142  |         |
|                         | 30 mg     | 7.52               | 85                   | -0.09 | 0.401 | -1.1  | -0.30 | -0.10  | 0.20  | 1.0     | -1.9995           | 0.0488  |         |
|                         | 60 mg     | 7.56               | 79                   | -0.07 | 0.355 | -0.9  | -0.30 | -0.10  | 0.20  | 0.8     | -1.6776           | 0.0974  |         |
|                         | 120 mg    | 7.56               | 80                   | -0.06 | 0.384 | -0.8  | -0.30 | -0.10  | 0.15  | 1.1     | -1.4839           | 0.1418  |         |
| Week 24                 | Placebo   | 7.63               | 93                   | -0.13 | 0.364 | -1.1  | -0.40 | -0.10  | 0.20  | 0.7     | -3.5049           | 0.0007  |         |
|                         | 15 mg     | 7.62               | 84                   | -0.02 | 0.369 | -0.7  | -0.30 | 0.00   | 0.20  | 1.2     | -0.3846           | 0.7015  |         |
|                         | 30 mg     | 7.53               | 84                   | -0.11 | 0.324 | -1.0  | -0.30 | -0.10  | 0.10  | 0.7     | -3.2024           | 0.0019  |         |
|                         | 60 mg     | 7.57               | 78                   | -0.12 | 0.374 | -1.0  | -0.40 | -0.10  | 0.10  | 0.8     | -2.8168           | 0.0062  |         |
|                         | 120 mg    | 7.56               | 79                   | -0.09 | 0.363 | -1.0  | -0.30 | -0.10  | 0.10  | 0.9     | -2.1715           | 0.0329  |         |
| Week 28                 | Placebo   | 7.63               | 93                   | -0.07 | 0.376 | -1.0  | -0.30 | -0.10  | 0.20  | 1.1     | -1.7363           | 0.0859  |         |
|                         | 15 mg     | 7.63               | 83                   | -0.03 | 0.361 | -1.2  | -0.30 | 0.00   | 0.20  | 0.7     | -0.6394           | 0.5243  |         |
|                         | 30 mg     | 7.53               | 84                   | -0.10 | 0.323 | -1.1  | -0.30 | -0.10  | 0.10  | 0.7     | -2.9728           | 0.0039  |         |
|                         | 60 mg     | 7.57               | 77                   | -0.13 | 0.371 | -0.9  | -0.40 | -0.10  | 0.10  | 0.6     | -3.0415           | 0.0032  |         |
|                         | 120 mg    | 7.55               | 78                   | -0.03 | 0.366 | -0.8  | -0.30 | -0.05  | 0.20  | 0.9     | -0.7120           | 0.4786  |         |
| Follow-up               | Placebo   | 7.63               | 97                   | -0.06 | 0.374 | -1.0  | -0.30 | -0.10  | 0.20  | 1.1     | -1.6289           | 0.1066  |         |
|                         | 15 mg     | 7.63               | 86                   | -0.03 | 0.356 | -1.2  | -0.30 | 0.00   | 0.20  | 0.7     | -0.6673           | 0.5064  |         |
|                         | 30 mg     | 7.51               | 92                   | -0.09 | 0.315 | -1.1  | -0.30 | -0.10  | 0.10  | 0.7     | -2.7766           | 0.0067  |         |
|                         | 60 mg     | 7.55               | 87                   | -0.13 | 0.363 | -0.9  | -0.40 | -0.10  | 0.10  | 0.6     | -3.2752           | 0.0015  |         |
|                         | 120 mg    | 7.55               | 88                   | -0.06 | 0.357 | -0.8  | -0.30 | -0.10  | 0.20  | 0.9     | -1.5238           | 0.1312  |         |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (2/24)

| Item              | Treatment | Summary Statistics |                      |       |       |       |       |        |       |         |         | One-sample t-test |        |
|-------------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|---------|---------|-------------------|--------|
|                   |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       |         | t-value | p-value           |        |
|                   |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    | Max     |         |                   |        |
| Albumin<br>(g/dL) | Week 1    | Placebo            | 4.54                 | 39    | -0.04 | 0.189 | -0.5  | -0.20  | -0.10 | 0.10    | 0.3     | -1.3571           | 0.1828 |
|                   | 15 mg     | 4.67               | 37                   | -0.06 | 0.203 | -0.6  | -0.20 | -0.10  | 0.10  | 0.3     | -1.7783 | 0.0838            |        |
|                   | 30 mg     | 4.57               | 39                   | 0.03  | 0.227 | -0.4  | -0.10 | 0.00   | 0.20  | 0.5     | 0.8454  | 0.4032            |        |
|                   | 60 mg     | 4.69               | 37                   | -0.10 | 0.195 | -0.5  | -0.20 | -0.10  | 0.00  | 0.3     | -3.2027 | 0.0028            |        |
|                   | 120 mg    | 4.62               | 47                   | -0.03 | 0.176 | -0.4  | -0.20 | 0.00   | 0.10  | 0.3     | -1.1630 | 0.2508            |        |
|                   | Week 2    | Placebo            | 4.63                 | 96    | -0.05 | 0.221 | -0.6  | -0.20  | -0.10 | 0.10    | 0.5     | -2.0784           | 0.0404 |
|                   | 15 mg     | 4.66               | 89                   | -0.02 | 0.201 | -0.5  | -0.20 | 0.00   | 0.10  | 0.5     | -0.9508 | 0.3443            |        |
|                   | 30 mg     | 4.62               | 91                   | -0.02 | 0.218 | -0.4  | -0.20 | 0.00   | 0.10  | 0.6     | -1.0592 | 0.2924            |        |
|                   | 60 mg     | 4.64               | 93                   | -0.06 | 0.203 | -0.9  | -0.20 | -0.10  | 0.00  | 0.5     | -3.0723 | 0.0028            |        |
|                   | 120 mg    | 4.62               | 89                   | -0.08 | 0.197 | -0.5  | -0.20 | 0.00   | 0.10  | 0.3     | -3.8156 | 0.0003            |        |
|                   | Week 3    | Placebo            | 4.56                 | 36    | -0.02 | 0.195 | -0.5  | -0.10  | 0.00  | 0.10    | 0.3     | -0.5971           | 0.5543 |
|                   | 15 mg     | 4.65               | 36                   | -0.01 | 0.222 | -0.6  | -0.20 | 0.00   | 0.10  | 0.4     | -0.2251 | 0.8232            |        |
|                   | 30 mg     | 4.58               | 38                   | 0.01  | 0.260 | -0.5  | -0.20 | 0.00   | 0.20  | 0.7     | 0.2498  | 0.8041            |        |
|                   | 60 mg     | 4.68               | 32                   | -0.07 | 0.206 | -0.4  | -0.20 | -0.10  | 0.10  | 0.4     | -1.8044 | 0.0809            |        |
|                   | 120 mg    | 4.63               | 42                   | -0.06 | 0.219 | -0.6  | -0.20 | 0.00   | 0.10  | 0.3     | -1.7638 | 0.0852            |        |
|                   | Week 4    | Placebo            | 4.64                 | 95    | -0.01 | 0.196 | -0.5  | -0.10  | 0.00  | 0.10    | 0.5     | -0.7340           | 0.4648 |
|                   | 15 mg     | 4.66               | 89                   | -0.02 | 0.219 | -0.6  | -0.20 | 0.00   | 0.10  | 0.6     | -0.8709 | 0.3862            |        |
|                   | 30 mg     | 4.62               | 89                   | -0.01 | 0.224 | -0.5  | -0.10 | -0.10  | 0.10  | 0.7     | -0.5686 | 0.5711            |        |
|                   | 60 mg     | 4.66               | 89                   | -0.05 | 0.220 | -0.4  | -0.20 | -0.10  | 0.10  | 0.8     | -2.1242 | 0.0365            |        |
|                   | 120 mg    | 4.62               | 87                   | -0.03 | 0.213 | -0.6  | -0.20 | 0.00   | 0.10  | 0.4     | -1.3582 | 0.1779            |        |
| Week 6            | Placebo   | 4.56               | 40                   | -0.03 | 0.233 | -0.5  | -0.20 | -0.10  | 0.15  | 0.5     | -0.6790 | 0.5012            |        |
| 15 mg             | 4.66      | 40                 | -0.07                | 0.199 | -0.5  | -0.20 | 0.00  | 0.10   | 0.3   | -2.2250 | 0.0319  |                   |        |
| 30 mg             | 4.58      | 38                 | -0.01                | 0.230 | -0.5  | -0.20 | 0.00  | 0.10   | 0.7   | -0.1410 | 0.8886  |                   |        |
| 60 mg             | 4.72      | 31                 | -0.09                | 0.229 | -0.6  | -0.20 | -0.10 | 0.00   | 0.5   | -2.2705 | 0.0305  |                   |        |
| 120 mg            | 4.63      | 47                 | -0.05                | 0.200 | -0.4  | -0.20 | -0.10 | 0.10   | 0.4   | -1.7516 | 0.0865  |                   |        |
| Week 8            | Placebo   | 4.64               | 94                   | -0.07 | 0.215 | -0.6  | -0.20 | -0.05  | 0.10  | 0.5     | -2.9771 | 0.0037            |        |
| 15 mg             | 4.65      | 86                 | -0.02                | 0.226 | -0.7  | -0.20 | 0.00  | 0.10   | 0.6   | -0.9551 | 0.3422  |                   |        |
| 30 mg             | 4.62      | 89                 | -0.04                | 0.249 | -0.7  | -0.20 | 0.00  | 0.10   | 0.7   | -1.4482 | 0.1511  |                   |        |
| 60 mg             | 4.66      | 86                 | -0.05                | 0.231 | -0.6  | -0.20 | -0.10 | 0.10   | 0.6   | -2.1946 | 0.0309  |                   |        |
| 120 mg            | 4.62      | 84                 | -0.05                | 0.219 | -0.5  | -0.20 | 0.00  | 0.10   | 0.5   | -2.2453 | 0.0274  |                   |        |
| Week 12           | Placebo   | 4.64               | 94                   | -0.06 | 0.216 | -0.6  | -0.20 | -0.05  | 0.10  | 0.5     | -2.7750 | 0.0067            |        |
| 15 mg             | 4.65      | 86                 | -0.06                | 0.193 | -0.5  | -0.20 | -0.10 | 0.10   | 0.4   | -2.7444 | 0.0074  |                   |        |
| 30 mg             | 4.63      | 86                 | -0.06                | 0.251 | -0.6  | -0.20 | -0.10 | 0.10   | 0.8   | -2.2794 | 0.0251  |                   |        |
| 60 mg             | 4.66      | 82                 | -0.08                | 0.211 | -0.6  | -0.20 | -0.10 | 0.10   | 0.4   | -3.4987 | 0.0008  |                   |        |
| 120 mg            | 4.63      | 83                 | -0.09                | 0.231 | -0.7  | -0.20 | -0.10 | 0.10   | 0.5   | -3.4639 | 0.0008  |                   |        |
| Week 16           | Placebo   | 4.64               | 94                   | -0.05 | 0.220 | -0.7  | -0.20 | 0.00   | 0.10  | 0.4     | -2.1581 | 0.0335            |        |
| 15 mg             | 4.65      | 84                 | -0.06                | 0.199 | -0.6  | -0.20 | -0.10 | 0.10   | 0.5   | -2.7905 | 0.0065  |                   |        |
| 30 mg             | 4.63      | 85                 | -0.08                | 0.244 | -0.6  | -0.20 | -0.10 | 0.10   | 0.8   | -2.9320 | 0.0043  |                   |        |
| 60 mg             | 4.69      | 80                 | -0.07                | 0.247 | -0.6  | -0.30 | -0.10 | 0.10   | 0.6   | -2.4864 | 0.0150  |                   |        |
| 120 mg            | 4.62      | 82                 | -0.08                | 0.233 | -0.7  | -0.20 | -0.05 | 0.00   | 0.8   | -2.9898 | 0.0037  |                   |        |
| Week 20           | Placebo   | 4.64               | 94                   | -0.07 | 0.225 | -0.7  | -0.20 | -0.10  | 0.10  | 0.5     | -3.1238 | 0.0024            |        |
| 15 mg             | 4.65      | 84                 | -0.08                | 0.239 | -0.7  | -0.20 | -0.10 | 0.10   | 0.5   | -2.9655 | 0.0039  |                   |        |
| 30 mg             | 4.63      | 85                 | -0.07                | 0.267 | -0.6  | -0.20 | -0.10 | 0.10   | 0.8   | -2.3570 | 0.0208  |                   |        |
| 60 mg             | 4.68      | 79                 | -0.08                | 0.223 | -0.5  | -0.20 | -0.10 | 0.10   | 0.5   | -3.3343 | 0.0013  |                   |        |
| 120 mg            | 4.62      | 80                 | -0.10                | 0.259 | -0.5  | -0.30 | -0.20 | 0.00   | 0.7   | -3.5797 | 0.0006  |                   |        |
| Week 24           | Placebo   | 4.64               | 93                   | -0.12 | 0.240 | -0.7  | -0.30 | -0.10  | 0.00  | 0.5     | -4.7528 | <0.0001           |        |
| 15 mg             | 4.65      | 84                 | -0.08                | 0.223 | -0.5  | -0.20 | -0.10 | 0.00   | 0.7   | -3.3696 | 0.0011  |                   |        |
| 30 mg             | 4.63      | 84                 | -0.10                | 0.220 | -0.5  | -0.30 | -0.10 | 0.00   | 0.4   | -4.1645 | <0.0001 |                   |        |
| 60 mg             | 4.68      | 78                 | -0.12                | 0.224 | -0.7  | -0.20 | -0.10 | 0.10   | 0.5   | -4.5505 | <0.0001 |                   |        |
| 120 mg            | 4.62      | 79                 | -0.11                | 0.235 | -0.6  | -0.20 | -0.10 | 0.00   | 0.6   | -4.1728 | <0.0001 |                   |        |
| Week 28           | Placebo   | 4.64               | 93                   | -0.12 | 0.222 | -0.6  | -0.30 | -0.10  | 0.00  | 0.4     | -5.1353 | <0.0001           |        |
| 15 mg             | 4.65      | 83                 | -0.10                | 0.234 | -0.8  | -0.20 | -0.10 | 0.10   | 0.4   | -3.8465 | 0.0002  |                   |        |
| 30 mg             | 4.63      | 84                 | -0.12                | 0.209 | -0.6  | -0.30 | -0.10 | 0.00   | 0.4   | -5.4725 | <0.0001 |                   |        |
| 60 mg             | 4.68      | 77                 | -0.12                | 0.238 | -0.8  | -0.30 | -0.10 | 0.00   | 0.4   | -4.5420 | <0.0001 |                   |        |
| 120 mg            | 4.62      | 78                 | -0.08                | 0.240 | -0.7  | -0.20 | -0.10 | 0.10   | 0.5   | -2.7808 | 0.0068  |                   |        |
| Follow-up         | Placebo   | 4.63               | 97                   | -0.11 | 0.221 | -0.6  | -0.30 | -0.10  | 0.00  | 0.4     | -5.0625 | <0.0001           |        |
| 15 mg             | 4.65      | 86                 | -0.10                | 0.247 | -0.9  | -0.20 | -0.10 | 0.10   | 0.4   | -3.9320 | 0.0002  |                   |        |
| 30 mg             | 4.61      | 92                 | -0.11                | 0.208 | -0.6  | -0.30 | -0.10 | 0.00   | 0.4   | -5.2213 | <0.0001 |                   |        |
| 60 mg             | 4.66      | 87                 | -0.12                | 0.242 | -0.8  | -0.30 | -0.10 | 0.00   | 0.4   | -4.7415 | <0.0001 |                   |        |
| 120 mg            | 4.62      | 88                 | -0.09                | 0.234 | -0.7  | -0.20 | -0.10 | 0.05   | 0.5   | -3.4623 | 0.0008  |                   |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (3/24)

| Item               | Treatment | Summary Statistics |                      |       |       |       |      |        |     |         | One-sample t-test |         |        |
|--------------------|-----------|--------------------|----------------------|-------|-------|-------|------|--------|-----|---------|-------------------|---------|--------|
|                    |           | Baseline Mean      | Change from Baseline |       |       |       |      |        |     | t-value | p-value           |         |        |
|                    |           |                    | N                    | Mean  | SD    | Min   | Q1   | Median | Q3  |         |                   | Max     |        |
| AST(GOT)<br>(IU/L) | Week 1    | Placebo            | 24.4                 | 39    | -0.5  | 4.98  | -19  | -3.0   | 0.0 | 2.0     | 8                 | -0.5784 | 0.5664 |
|                    | 15 mg     | 23.5               | 37                   | -0.3  | 4.45  | -11   | -2.0 | -1.0   | 2.0 | 10      | -0.3698           | 0.7137  |        |
|                    | 30 mg     | 23.1               | 39                   | 0.4   | 3.44  | -10   | -1.0 | 1.0    | 3.0 | 9       | 0.6507            | 0.5191  |        |
|                    | 60 mg     | 24.5               | 37                   | -0.1  | 4.97  | -15   | -2.0 | 0.0    | 3.0 | 15      | -0.0662           | 0.9476  |        |
|                    | 120 mg    | 24.6               | 47                   | 2.1   | 8.47  | -22   | -1.0 | 1.0    | 4.0 | 33      | 1.7394            | 0.0887  |        |
|                    | Week 2    | Placebo            | 27.1                 | 96    | -0.2  | 6.79  | -22  | -3.5   | 0.0 | 3.0     | 33                | -0.2555 | 0.7989 |
|                    | 15 mg     | 26.1               | 89                   | 0.6   | 6.68  | -28   | -2.0 | 0.0    | 2.0 | 33      | 0.8723            | 0.3854  |        |
|                    | 30 mg     | 23.1               | 91                   | 0.9   | 6.00  | -22   | -2.0 | 1.0    | 3.0 | 18      | 1.3810            | 0.1707  |        |
|                    | 60 mg     | 27.2               | 93                   | -0.4  | 7.78  | -42   | -3.0 | 0.0    | 2.0 | 29      | -0.4933           | 0.6230  |        |
|                    | 120 mg    | 24.2               | 89                   | 0.4   | 5.22  | -18   | -2.0 | 0.0    | 2.0 | 17      | 0.8116            | 0.4192  |        |
|                    | Week 3    | Placebo            | 25.4                 | 37    | 1.8   | 13.24 | -14  | -3.0   | 0.0 | 3.0     | 70                | 0.8194  | 0.4179 |
|                    | 15 mg     | 23.5               | 36                   | 1.0   | 5.43  | -11   | -1.0 | 1.0    | 3.5 | 13      | 1.1351            | 0.2640  |        |
|                    | 30 mg     | 23.7               | 38                   | 0.3   | 6.93  | -21   | -2.0 | 0.5    | 3.0 | 27      | 0.2810            | 0.7803  |        |
|                    | 60 mg     | 22.5               | 32                   | -0.1  | 5.09  | -10   | -3.0 | 0.0    | 3.0 | 15      | -0.1389           | 0.8904  |        |
|                    | 120 mg    | 24.4               | 42                   | 1.2   | 4.96  | -11   | -2.0 | 1.0    | 4.0 | 18      | 1.5556            | 0.1275  |        |
|                    | Week 4    | Placebo            | 26.8                 | 95    | -0.0  | 8.49  | -30  | -2.0   | 0.0 | 2.0     | 52                | -0.0121 | 0.9904 |
|                    | 15 mg     | 26.1               | 89                   | 0.9   | 6.06  | -18   | -1.0 | 1.0    | 3.0 | 19      | 1.3467            | 0.1815  |        |
|                    | 30 mg     | 23.2               | 89                   | 0.6   | 5.22  | -20   | -2.0 | 1.0    | 3.0 | 18      | 1.1573            | 0.2503  |        |
|                    | 60 mg     | 27.8               | 89                   | -0.4  | 9.44  | -44   | -3.0 | 0.0    | 2.0 | 46      | -0.4378           | 0.6626  |        |
|                    | 120 mg    | 24.1               | 87                   | 0.8   | 6.21  | -13   | -2.0 | 1.0    | 4.0 | 31      | 1.2087            | 0.2301  |        |
| Week 6             | Placebo   | 26.0               | 40                   | 0.8   | 6.68  | -18   | -2.0 | 0.0    | 3.5 | 30      | 0.7817            | 0.4391  |        |
| 15 mg              | 24.7      | 40                 | -0.2                 | 7.62  | -27   | -2.0  | 0.5  | 3.5    | 25  | -0.1659 | 0.8691            |         |        |
| 30 mg              | 24.5      | 38                 | 0.4                  | 4.77  | -15   | -2.0  | 1.0  | 3.0    | 10  | 0.5104  | 0.6128            |         |        |
| 60 mg              | 22.9      | 31                 | 0.6                  | 3.25  | -6    | -1.0  | 1.0  | 4.0    | 5   | 1.1049  | 0.2780            |         |        |
| 120 mg             | 25.9      | 47                 | 0.1                  | 6.90  | -35   | -2.0  | 1.0  | 3.0    | 12  | 0.0846  | 0.9330            |         |        |
| Week 8             | Placebo   | 26.8               | 94                   | -1.4  | 7.38  | -25   | -5.0 | -1.0   | 2.0 | 22      | -1.8585           | 0.0663  |        |
| 15 mg              | 25.9      | 86                 | 0.9                  | 8.25  | -22   | -2.0  | 0.0  | 3.0    | 44  | 1.0460  | 0.2985            |         |        |
| 30 mg              | 23.2      | 89                 | 0.8                  | 7.52  | -29   | -2.0  | 1.0  | 4.0    | 25  | 1.0566  | 0.2936            |         |        |
| 60 mg              | 27.3      | 86                 | -0.4                 | 7.24  | -36   | -3.0  | 0.0  | 4.0    | 22  | -0.5659 | 0.5730            |         |        |
| 120 mg             | 24.0      | 84                 | -0.4                 | 5.96  | -35   | -3.0  | 0.0  | 3.0    | 9   | -0.6771 | 0.5002            |         |        |
| Week 12            | Placebo   | 26.8               | 94                   | 0.1   | 9.68  | -31   | -3.0 | -1.0   | 3.0 | 42      | 0.1491            | 0.8818  |        |
| 15 mg              | 25.9      | 86                 | 1.8                  | 13.48 | -21   | -3.0  | 0.0  | 3.0    | 107 | 1.2395  | 0.2186            |         |        |
| 30 mg              | 23.3      | 86                 | -0.3                 | 6.72  | -25   | -2.0  | 0.0  | 2.0    | 18  | -0.4012 | 0.6893            |         |        |
| 60 mg              | 27.2      | 82                 | -2.0                 | 9.11  | -48   | -4.0  | 0.0  | 2.0    | 13  | -1.9770 | 0.0514            |         |        |
| 120 mg             | 24.3      | 83                 | 0.4                  | 8.98  | -40   | -3.0  | 0.0  | 3.0    | 42  | 0.4154  | 0.6789            |         |        |
| Week 16            | Placebo   | 26.8               | 94                   | -0.0  | 10.97 | -32   | -4.0 | -1.0   | 3.0 | 61      | -0.0094           | 0.9925  |        |
| 15 mg              | 25.9      | 84                 | -0.1                 | 7.51  | -15   | -4.5  | 0.0  | 2.0    | 31  | -0.1017 | 0.9193            |         |        |
| 30 mg              | 23.4      | 85                 | 0.6                  | 7.67  | -21   | -3.0  | 0.0  | 4.0    | 34  | 0.7495  | 0.4557            |         |        |
| 60 mg              | 26.7      | 80                 | -1.8                 | 8.53  | -42   | -3.5  | 0.0  | 2.0    | 12  | -1.8995 | 0.0611            |         |        |
| 120 mg             | 24.0      | 82                 | -0.1                 | 8.62  | -44   | -3.0  | 0.0  | 3.0    | 38  | -0.1538 | 0.8782            |         |        |
| Week 20            | Placebo   | 26.8               | 94                   | 0.8   | 9.18  | -26   | -4.0 | 0.0    | 4.0 | 34      | 0.8093            | 0.4204  |        |
| 15 mg              | 25.9      | 84                 | 1.5                  | 8.87  | -15   | -3.0  | 0.5  | 3.5    | 46  | 1.4999  | 0.1374            |         |        |
| 30 mg              | 23.4      | 85                 | 0.4                  | 6.90  | -26   | -2.0  | 1.0  | 3.0    | 20  | 0.5189  | 0.6052            |         |        |
| 60 mg              | 26.9      | 79                 | -2.2                 | 9.06  | -47   | -5.0  | -1.0 | 2.0    | 12  | -2.1225 | 0.0370            |         |        |
| 120 mg             | 24.2      | 80                 | -0.7                 | 9.95  | -50   | -4.0  | -1.0 | 3.0    | 45  | -0.5845 | 0.5606            |         |        |
| Week 24            | Placebo   | 26.8               | 93                   | 1.4   | 10.99 | -43   | -3.0 | 0.0    | 7.0 | 39      | 1.2072            | 0.2305  |        |
| 15 mg              | 25.9      | 84                 | 0.4                  | 7.59  | -18   | -3.0  | 0.0  | 3.0    | 23  | 0.4602  | 0.6466            |         |        |
| 30 mg              | 23.3      | 84                 | 0.4                  | 8.47  | -25   | -3.0  | 1.0  | 4.0    | 44  | 0.4768  | 0.6348            |         |        |
| 60 mg              | 26.1      | 78                 | -2.8                 | 7.77  | -48   | -6.0  | -1.0 | 2.0    | 9   | -3.1345 | 0.0024            |         |        |
| 120 mg             | 23.9      | 79                 | -1.5                 | 9.13  | -50   | -4.0  | -1.0 | 3.0    | 26  | -1.4904 | 0.1401            |         |        |
| Week 28            | Placebo   | 26.8               | 93                   | -0.0  | 9.45  | -44   | -4.0 | 0.0    | 3.0 | 35      | -0.0329           | 0.9738  |        |
| 15 mg              | 26.0      | 83                 | -0.5                 | 7.42  | -16   | -4.0  | -1.0 | 2.0    | 32  | -0.5917 | 0.5557            |         |        |
| 30 mg              | 23.3      | 84                 | -0.4                 | 8.17  | -25   | -3.0  | -1.0 | 3.0    | 27  | -0.4272 | 0.6703            |         |        |
| 60 mg              | 26.1      | 77                 | -3.3                 | 10.58 | -44   | -7.0  | -3.0 | 0.0    | 55  | -2.7150 | 0.0082            |         |        |
| 120 mg             | 24.2      | 79                 | -0.9                 | 9.51  | -47   | -4.0  | 0.0  | 2.0    | 50  | -0.8403 | 0.4033            |         |        |
| Follow-up          | Placebo   | 27.1               | 97                   | -0.2  | 9.33  | -44   | -4.0 | 0.0    | 3.0 | 35      | -0.1740           | 0.8622  |        |
| 15 mg              | 26.3      | 86                 | -0.8                 | 7.88  | -28   | -4.0  | -1.0 | 2.0    | 32  | -0.8900 | 0.3760            |         |        |
| 30 mg              | 23.0      | 92                 | -0.5                 | 7.90  | -25   | -3.0  | -1.0 | 3.0    | 27  | -0.5672 | 0.5720            |         |        |
| 60 mg              | 26.6      | 87                 | -3.0                 | 10.43 | -44   | -7.0  | -2.0 | 0.0    | 55  | -2.6935 | 0.0085            |         |        |
| 120 mg             | 24.2      | 88                 | -0.7                 | 9.20  | -47   | -4.0  | 0.0  | 3.0    | 50  | -0.6834 | 0.4962            |         |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (4/24)

| Item               | Treatment | Summary Statistics |                      |      |       |       |       |        |      |         | One-sample t-test |         |        |
|--------------------|-----------|--------------------|----------------------|------|-------|-------|-------|--------|------|---------|-------------------|---------|--------|
|                    |           | Baseline Mean      | Change from Baseline |      |       |       |       |        |      | t-value | p-value           |         |        |
|                    |           |                    | N                    | Mean | SD    | Min   | Q1    | Median | Q3   |         |                   | Max     |        |
| ALT(GPT)<br>(IU/L) | Week 1    | Placebo            | 30.4                 | 39   | -1.9  | 8.17  | -36   | -4.0   | -1.0 | 2.0     | 13                | -1.4506 | 0.1551 |
|                    |           | 15 mg              | 29.8                 | 37   | -0.4  | 5.32  | -15   | -2.0   | 0.0  | 2.0     | 14                | -0.4949 | 0.6237 |
|                    |           | 30 mg              | 25.9                 | 39   | -0.1  | 4.23  | -16   | -2.0   | 1.0  | 2.0     | 9                 | -0.1515 | 0.8804 |
|                    |           | 60 mg              | 32.8                 | 37   | 0.1   | 8.88  | -28   | -4.0   | 1.0  | 3.0     | 23                | 0.0555  | 0.9560 |
|                    |           | 120 mg             | 30.6                 | 47   | 3.4   | 7.49  | -13   | -1.0   | 2.0  | 5.0     | 24                | 3.0962  | 0.0033 |
|                    | Week 2    | Placebo            | 33.9                 | 96   | -0.9  | 9.08  | -42   | -5.0   | -1.0 | 3.0     | 33                | -0.9437 | 0.3477 |
|                    |           | 15 mg              | 32.0                 | 89   | 1.2   | 9.24  | -39   | -2.0   | 1.0  | 4.0     | 54                | 1.1816  | 0.2406 |
|                    |           | 30 mg              | 28.2                 | 91   | 2.6   | 8.38  | -22   | -2.0   | 2.0  | 5.0     | 39                | 2.9137  | 0.0045 |
|                    |           | 60 mg              | 36.5                 | 93   | 0.3   | 12.01 | -59   | -3.0   | 1.0  | 3.0     | 43                | 0.2763  | 0.7829 |
|                    |           | 120 mg             | 30.3                 | 89   | 1.8   | 8.64  | -29   | -2.0   | 2.0  | 5.0     | 41                | 1.9251  | 0.0574 |
|                    | Week 3    | Placebo            | 30.3                 | 36   | -0.3  | 13.03 | -46   | -3.0   | 0.0  | 3.0     | 48                | -0.1152 | 0.9090 |
|                    |           | 15 mg              | 30.5                 | 36   | 0.9   | 5.80  | -15   | -2.5   | 1.0  | 4.5     | 11                | 0.8905  | 0.3793 |
|                    |           | 30 mg              | 28.1                 | 38   | 1.3   | 11.39 | -31   | -3.0   | 0.0  | 4.0     | 50                | 0.6976  | 0.4898 |
|                    |           | 60 mg              | 28.2                 | 32   | 0.5   | 7.86  | -16   | -3.5   | 1.0  | 4.0     | 21                | 0.3598  | 0.7215 |
|                    |           | 120 mg             | 29.8                 | 42   | 2.3   | 8.80  | -24   | -3.0   | 2.0  | 7.0     | 25                | 1.6660  | 0.1033 |
|                    | Week 4    | Placebo            | 33.9                 | 95   | 0.6   | 10.27 | -40   | -3.0   | 0.0  | 3.0     | 49                | 0.5393  | 0.5910 |
|                    |           | 15 mg              | 32.0                 | 89   | 1.1   | 8.65  | -37   | -3.0   | 1.0  | 5.0     | 31                | 1.2135  | 0.2282 |
|                    |           | 30 mg              | 28.3                 | 89   | 1.6   | 8.74  | -23   | -3.0   | 0.0  | 5.0     | 33                | 1.7228  | 0.0884 |
|                    |           | 60 mg              | 38.2                 | 89   | 0.7   | 14.48 | -79   | -3.0   | 0.0  | 5.0     | 57                | 0.4320  | 0.6668 |
|                    |           | 120 mg             | 30.2                 | 87   | 2.2   | 13.66 | -24   | -4.0   | 1.0  | 5.0     | 102               | 1.4759  | 0.1436 |
| Week 6             | Placebo   | 32.5               | 40                   | 0.6  | 12.07 | -43   | -3.5  | 0.5    | 5.5  | 35      | 0.3276            | 0.7450  |        |
|                    | 15 mg     | 31.4               | 40                   | -0.2 | 10.06 | -48   | -2.5  | 0.0    | 6.0  | 15      | -0.1257           | 0.9006  |        |
|                    | 30 mg     | 29.8               | 38                   | 0.2  | 7.23  | -12   | -5.0  | -0.5   | 3.0  | 26      | 0.2019            | 0.8411  |        |
|                    | 60 mg     | 29.9               | 32                   | 2.6  | 10.19 | -13   | -2.0  | 2.0    | 5.0  | 42      | 1.4402            | 0.1598  |        |
|                    | 120 mg    | 31.9               | 47                   | -0.1 | 9.36  | -34   | -4.0  | 0.0    | 6.0  | 15      | -0.1090           | 0.9136  |        |
| Week 8             | Placebo   | 33.9               | 94                   | -1.6 | 10.61 | -40   | -4.0  | -1.0   | 2.0  | 36      | -1.4390           | 0.1535  |        |
|                    | 15 mg     | 31.5               | 86                   | 1.9  | 8.92  | -16   | -4.0  | 1.0    | 6.0  | 36      | 1.9460            | 0.0550  |        |
|                    | 30 mg     | 28.3               | 89                   | 1.1  | 10.05 | -41   | -3.0  | 1.0    | 7.0  | 26      | 1.0759            | 0.2849  |        |
|                    | 60 mg     | 37.1               | 86                   | 0.4  | 11.04 | -33   | -5.0  | 0.5    | 6.0  | 44      | 0.3517            | 0.7259  |        |
|                    | 120 mg    | 30.1               | 84                   | -1.1 | 8.75  | -35   | -5.0  | 0.0    | 4.0  | 19      | -1.1217           | 0.2652  |        |
| Week 12            | Placebo   | 33.9               | 94                   | -0.7 | 11.13 | -38   | -5.0  | 0.0    | 3.0  | 48      | -0.6210           | 0.5361  |        |
|                    | 15 mg     | 31.5               | 86                   | 1.7  | 12.03 | -20   | -5.0  | -0.5   | 5.0  | 67      | 1.2730            | 0.2065  |        |
|                    | 30 mg     | 28.5               | 86                   | 0.2  | 10.91 | -48   | -5.0  | 0.0    | 4.0  | 33      | 0.1778            | 0.8593  |        |
|                    | 60 mg     | 37.4               | 82                   | -2.4 | 15.86 | -98   | -7.0  | -1.0   | 5.0  | 39      | -1.3929           | 0.1674  |        |
|                    | 120 mg    | 30.2               | 83                   | 0.1  | 12.69 | -52   | -6.0  | 0.0    | 5.0  | 44      | 0.0865            | 0.9313  |        |
| Week 16            | Placebo   | 33.9               | 94                   | 0.4  | 12.67 | -47   | -5.0  | -1.0   | 5.0  | 42      | 0.3174            | 0.7517  |        |
|                    | 15 mg     | 31.5               | 84                   | -0.9 | 9.23  | -20   | -6.0  | -2.0   | 4.5  | 29      | -0.8861           | 0.3781  |        |
|                    | 30 mg     | 28.7               | 85                   | 2.4  | 16.96 | -48   | -3.0  | 0.0    | 6.0  | 104     | 1.3173            | 0.1913  |        |
|                    | 60 mg     | 36.9               | 80                   | -2.9 | 14.60 | -95   | -8.0  | -0.5   | 4.0  | 26      | -1.7839           | 0.0783  |        |
|                    | 120 mg    | 30.0               | 82                   | -0.4 | 13.58 | -60   | -4.0  | 0.0    | 4.0  | 52      | -0.2521           | 0.8016  |        |
| Week 20            | Placebo   | 33.9               | 94                   | 0.9  | 11.93 | -32   | -4.0  | 0.5    | 4.0  | 53      | 0.7608            | 0.4487  |        |
|                    | 15 mg     | 31.5               | 84                   | 2.6  | 16.96 | -24   | -4.0  | 0.0    | 5.0  | 98      | 1.4155            | 0.1607  |        |
|                    | 30 mg     | 28.7               | 85                   | 0.9  | 10.90 | -57   | -4.0  | 1.0    | 5.0  | 22      | 0.7565            | 0.4515  |        |
|                    | 60 mg     | 37.2               | 79                   | -4.2 | 14.89 | -93   | -11.0 | -1.0   | 4.0  | 18      | -2.5088           | 0.0142  |        |
|                    | 120 mg    | 30.1               | 80                   | -1.3 | 14.90 | -62   | -6.5  | -1.0   | 4.5  | 65      | -0.7652           | 0.4465  |        |
| Week 24            | Placebo   | 34.0               | 93                   | 2.5  | 17.63 | -58   | -3.0  | 1.0    | 8.0  | 73      | 1.3766            | 0.1720  |        |
|                    | 15 mg     | 31.5               | 84                   | 0.3  | 13.07 | -26   | -6.0  | -1.5   | 6.0  | 66      | 0.2337            | 0.8158  |        |
|                    | 30 mg     | 28.5               | 84                   | 1.3  | 12.51 | -54   | -4.0  | 2.0    | 6.5  | 59      | 0.9333            | 0.3534  |        |
|                    | 60 mg     | 37.0               | 78                   | -6.0 | 15.77 | -110  | -12.0 | -3.0   | 3.0  | 13      | -3.3811           | 0.0011  |        |
|                    | 120 mg    | 29.6               | 79                   | -1.8 | 14.15 | -67   | -7.0  | -1.0   | 4.0  | 45      | -1.1049           | 0.2726  |        |
| Week 28            | Placebo   | 34.0               | 93                   | 0.7  | 15.07 | -68   | -4.0  | 0.0    | 6.0  | 59      | 0.4680            | 0.6409  |        |
|                    | 15 mg     | 31.6               | 83                   | -1.6 | 10.15 | -27   | -8.0  | -1.0   | 2.0  | 31      | -1.4163           | 0.1605  |        |
|                    | 30 mg     | 28.5               | 84                   | -0.6 | 14.37 | -68   | -6.0  | -2.0   | 4.0  | 44      | -0.3873           | 0.6995  |        |
|                    | 60 mg     | 37.1               | 77                   | -7.6 | 20.07 | -94   | -13.0 | -6.0   | 0.0  | 87      | -3.3053           | 0.0014  |        |
|                    | 120 mg    | 30.0               | 79                   | -3.6 | 15.43 | -72   | -9.0  | -2.0   | 2.0  | 77      | -2.0922           | 0.0397  |        |
| Follow-up          | Placebo   | 33.9               | 97                   | 0.6  | 14.76 | -68   | -4.0  | 0.0    | 6.0  | 59      | 0.4332            | 0.6658  |        |
|                    | 15 mg     | 32.4               | 86                   | -2.2 | 11.35 | -52   | -8.0  | -1.0   | 2.0  | 31      | -1.8338           | 0.0702  |        |
|                    | 30 mg     | 27.8               | 92                   | -0.8 | 13.80 | -68   | -6.0  | -2.0   | 4.0  | 44      | -0.5288           | 0.5982  |        |
|                    | 60 mg     | 37.6               | 87                   | -6.8 | 19.71 | -94   | -13.0 | -6.0   | 0.0  | 87      | -3.2034           | 0.0019  |        |
|                    | 120 mg    | 30.2               | 88                   | -3.3 | 14.75 | -72   | -9.0  | -2.0   | 2.0  | 77      | -2.1248           | 0.0364  |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (5/24)

| Item            | Treatment | Summary Statistics |                      |       |       |       |      |        |      |         |         | One-sample t-test |        |
|-----------------|-----------|--------------------|----------------------|-------|-------|-------|------|--------|------|---------|---------|-------------------|--------|
|                 |           | Baseline Mean      | Change from Baseline |       |       |       |      |        |      |         | t-value | p-value           |        |
|                 |           |                    | N                    | Mean  | SD    | Min   | Q1   | Median | Q3   | Max     |         |                   |        |
| γ-GTP<br>(IU/L) | Week 1    | Placebo            | 55.2                 | 39    | 0.6   | 12.10 | -38  | -3.0   | -1.0 | 1.0     | 43      | 0.2912            | 0.7724 |
|                 | 15 mg     | 40.2               | 37                   | -1.4  | 5.20  | -12   | -4.0 | -2.0   | 0.0  | 22      | -1.6130 | 0.1155            |        |
|                 | 30 mg     | 43.7               | 39                   | -1.4  | 7.99  | -45   | -2.0 | 0.0    | 1.0  | 6       | -1.1224 | 0.2687            |        |
|                 | 60 mg     | 45.6               | 37                   | -2.0  | 6.12  | -21   | -4.0 | -1.0   | 1.0  | 10      | -2.0142 | 0.0515            |        |
|                 | 120 mg    | 46.4               | 47                   | -0.9  | 7.55  | -25   | -3.0 | -1.0   | 1.0  | 20      | -0.8506 | 0.3994            |        |
|                 | Week 2    | Placebo            | 60.6                 | 96    | -2.1  | 25.96 | -210 | -6.0   | -1.0 | 2.0     | 83      | -0.8061           | 0.4222 |
|                 | 15 mg     | 52.6               | 89                   | 1.3   | 31.29 | -58   | -5.0 | -1.0   | 1.0  | 273     | 0.4065  | 0.6853            |        |
|                 | 30 mg     | 47.5               | 91                   | -0.1  | 11.26 | -48   | -4.0 | 0.0    | 2.0  | 54      | -0.1210 | 0.9039            |        |
|                 | 60 mg     | 58.8               | 93                   | -6.9  | 27.53 | -212  | -5.0 | -2.0   | 1.0  | 21      | -2.4299 | 0.0170            |        |
|                 | 120 mg    | 48.4               | 89                   | -2.2  | 10.15 | -64   | -5.0 | -1.0   | 1.0  | 44      | -2.0160 | 0.0469            |        |
|                 | Week 3    | Placebo            | 57.0                 | 36    | -8.4  | 56.51 | -326 | -3.5   | 0.0  | 3.0     | 51      | -0.8907           | 0.3792 |
|                 | 15 mg     | 40.7               | 36                   | -1.0  | 7.76  | -16   | -4.0 | -1.0   | 1.0  | 24      | -0.7728 | 0.4449            |        |
|                 | 30 mg     | 44.8               | 38                   | 1.3   | 27.63 | -58   | -5.0 | -1.0   | 3.0  | 151     | 0.2876  | 0.7752            |        |
|                 | 60 mg     | 43.4               | 32                   | -4.2  | 8.56  | -34   | -6.5 | -2.0   | 0.5  | 8       | -2.7880 | 0.0090            |        |
|                 | 120 mg    | 47.6               | 42                   | 0.3   | 14.49 | -38   | -6.0 | -1.0   | 1.0  | 54      | 0.1172  | 0.9073            |        |
|                 | Week 4    | Placebo            | 60.6                 | 95    | -2.6  | 44.21 | -357 | -4.0   | -1.0 | 3.0     | 214     | -0.5801           | 0.5632 |
|                 | 15 mg     | 52.6               | 89                   | -1.3  | 17.31 | -74   | -6.0 | -2.0   | 2.0  | 79      | -0.7226 | 0.4718            |        |
|                 | 30 mg     | 47.9               | 89                   | -1.9  | 17.81 | -50   | -7.0 | -2.0   | 2.0  | 123     | -1.0239 | 0.3087            |        |
|                 | 60 mg     | 60.7               | 89                   | -8.9  | 36.29 | -223  | -8.0 | -1.0   | 2.0  | 35      | -2.3106 | 0.0232            |        |
|                 | 120 mg    | 48.1               | 87                   | -1.5  | 9.19  | -46   | -6.0 | -2.0   | 1.0  | 30      | -1.5636 | 0.1216            |        |
| Week 6          | Placebo   | 60.7               | 40                   | -3.3  | 54.47 | -278  | -5.0 | -1.0   | 2.0  | 159     | -0.3860 | 0.7016            |        |
| 15 mg           | 42.3      | 40                 | -0.7                 | 11.15 | -34   | -6.0  | -2.0 | 1.0    | 30   | -0.3830 | 0.7038  |                   |        |
| 30 mg           | 49.1      | 38                 | -3.0                 | 11.63 | -26   | -8.0  | -3.0 | 0.0    | 49   | -1.6046 | 0.1171  |                   |        |
| 60 mg           | 47.0      | 31                 | -3.9                 | 15.31 | -47   | -9.0  | -4.0 | 1.0    | 40   | -1.4308 | 0.1628  |                   |        |
| 120 mg          | 46.7      | 47                 | -1.2                 | 10.04 | -28   | -7.0  | -2.0 | 1.0    | 30   | -0.8423 | 0.4040  |                   |        |
| Week 8          | Placebo   | 60.6               | 94                   | -4.4  | 37.64 | -319  | -7.0 | -1.0   | 4.0  | 116     | -1.1263 | 0.2629            |        |
| 15 mg           | 52.7      | 86                 | -1.1                 | 18.70 | -96   | -7.0  | -2.5 | 3.0    | 73   | -0.5537 | 0.5813  |                   |        |
| 30 mg           | 47.9      | 89                 | -1.2                 | 15.18 | -38   | -7.0  | -2.0 | 3.0    | 74   | -0.7679 | 0.4446  |                   |        |
| 60 mg           | 58.7      | 86                 | -8.5                 | 38.53 | -263  | -8.0  | -1.0 | 2.0    | 58   | -2.0347 | 0.0450  |                   |        |
| 120 mg          | 49.2      | 84                 | -2.2                 | 11.37 | -27   | -7.0  | -2.0 | 1.5    | 38   | -1.7663 | 0.0810  |                   |        |
| Week 12         | Placebo   | 60.6               | 94                   | 0.6   | 33.65 | -172  | -6.0 | -1.0   | 3.0  | 225     | 0.1686  | 0.8665            |        |
| 15 mg           | 52.7      | 86                 | -3.0                 | 15.21 | -65   | -7.0  | -2.0 | 2.0    | 54   | -1.8006 | 0.0753  |                   |        |
| 30 mg           | 48.6      | 86                 | -4.3                 | 15.87 | -80   | -11.0 | -2.0 | 2.0    | 69   | -2.5142 | 0.0138  |                   |        |
| 60 mg           | 57.7      | 82                 | -11.5                | 45.51 | -252  | -10.0 | -2.0 | 2.0    | 38   | -2.2785 | 0.0253  |                   |        |
| 120 mg          | 48.2      | 83                 | -1.1                 | 15.39 | -39   | -9.0  | -2.0 | 2.0    | 81   | -0.6777 | 0.4999  |                   |        |
| Week 16         | Placebo   | 60.6               | 94                   | -1.9  | 36.65 | -276  | -7.0 | -1.0   | 4.0  | 141     | -0.5151 | 0.6077            |        |
| 15 mg           | 52.6      | 84                 | -3.4                 | 12.84 | -49   | -8.0  | -2.0 | 1.5    | 39   | -2.4388 | 0.0169  |                   |        |
| 30 mg           | 48.9      | 85                 | -4.2                 | 17.60 | -105  | -9.0  | -2.0 | 2.0    | 61   | -2.2063 | 0.0301  |                   |        |
| 60 mg           | 53.5      | 80                 | -10.3                | 41.76 | -296  | -8.0  | -1.5 | 2.5    | 35   | -2.1956 | 0.0311  |                   |        |
| 120 mg          | 47.5      | 82                 | -0.3                 | 15.26 | -29   | -7.0  | -1.5 | 3.0    | 71   | -0.1664 | 0.8682  |                   |        |
| Week 20         | Placebo   | 60.6               | 94                   | 2.2   | 24.61 | -50   | -6.0 | -0.5   | 5.0  | 134     | 0.8675  | 0.3879            |        |
| 15 mg           | 52.6      | 84                 | -3.8                 | 15.68 | -56   | -10.0 | -4.0 | 1.5    | 59   | -2.2197 | 0.0292  |                   |        |
| 30 mg           | 48.9      | 85                 | -1.9                 | 22.46 | -63   | -8.0  | -4.0 | 3.0    | 150  | -0.7822 | 0.4363  |                   |        |
| 60 mg           | 53.9      | 79                 | -11.1                | 40.19 | -279  | -12.0 | -2.0 | 2.0    | 40   | -2.4577 | 0.0162  |                   |        |
| 120 mg          | 47.8      | 80                 | -2.4                 | 17.54 | -68   | -8.0  | -3.0 | 2.5    | 53   | -1.2431 | 0.2175  |                   |        |
| Week 24         | Placebo   | 61.1               | 93                   | -0.2  | 31.71 | -220  | -6.0 | -1.0   | 2.0  | 110     | -0.0621 | 0.9506            |        |
| 15 mg           | 52.6      | 84                 | -2.2                 | 17.02 | -58   | -10.0 | -3.0 | 3.0    | 62   | -1.1665 | 0.2468  |                   |        |
| 30 mg           | 47.7      | 84                 | 0.7                  | 29.62 | -45   | -10.0 | -2.0 | 4.0    | 225  | 0.2284  | 0.8199  |                   |        |
| 60 mg           | 53.3      | 78                 | -12.6                | 44.82 | -296  | -13.0 | -3.0 | 1.0    | 37   | -2.4783 | 0.0154  |                   |        |
| 120 mg          | 47.8      | 79                 | -4.3                 | 17.40 | -55   | -10.0 | -3.0 | 2.0    | 65   | -2.1921 | 0.0314  |                   |        |
| Week 28         | Placebo   | 61.1               | 93                   | 1.0   | 24.64 | -113  | -6.0 | -1.0   | 4.0  | 113     | 0.3872  | 0.6995            |        |
| 15 mg           | 53.0      | 83                 | -4.1                 | 16.37 | -48   | -12.0 | -3.0 | 4.0    | 57   | -2.2670 | 0.0260  |                   |        |
| 30 mg           | 47.7      | 84                 | 0.0                  | 27.44 | -49   | -9.0  | -2.0 | 3.0    | 182  | 0.0040  | 0.9968  |                   |        |
| 60 mg           | 53.1      | 77                 | -12.3                | 39.66 | -245  | -15.0 | -3.0 | 2.0    | 18   | -2.7182 | 0.0081  |                   |        |
| 120 mg          | 47.9      | 79                 | -4.5                 | 19.17 | -71   | -11.0 | -3.0 | 2.0    | 62   | -2.0830 | 0.0405  |                   |        |
| Follow-up       | Placebo   | 60.3               | 97                   | 1.0   | 24.25 | -113  | -6.0 | -1.0   | 5.0  | 113     | 0.4187  | 0.6764            |        |
| 15 mg           | 53.7      | 86                 | -3.7                 | 16.34 | -48   | -12.0 | -2.5 | 4.0    | 57   | -2.1179 | 0.0371  |                   |        |
| 30 mg           | 45.9      | 92                 | -0.1                 | 26.24 | -49   | -7.5  | -2.0 | 3.0    | 182  | -0.0318 | 0.9747  |                   |        |
| 60 mg           | 58.4      | 87                 | -14.8                | 46.81 | -270  | -15.0 | -3.0 | 2.0    | 20   | -2.9502 | 0.0041  |                   |        |
| 120 mg          | 48.5      | 88                 | -4.7                 | 19.77 | -71   | -10.0 | -2.5 | 2.0    | 62   | -2.2165 | 0.0293  |                   |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (6/24)

| Item          | Treatment | Summary Statistics |                      |       |       |       |       |        |      |         | One-sample t-test |         |        |
|---------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|------|---------|-------------------|---------|--------|
|               |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |      | t-value | p-value           |         |        |
|               |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3   |         |                   | Max     |        |
| ALP<br>(IU/L) | Week 1    | Placebo            | 248.0                | 39    | -4.4  | 23.17 | -90   | -12.0  | -6.0 | 11.0    | 37                | -1.1819 | 0.2446 |
|               | 15 mg     | 229.8              | 37                   | -3.9  | 15.44 | -51   | -12.0 | -4.0   | 7.0  | 21      | -1.5440           | 0.1313  |        |
|               | 30 mg     | 223.2              | 39                   | 6.6   | 27.89 | -54   | -6.0  | 5.0    | 11.0 | 135     | 1.4699            | 0.1498  |        |
|               | 60 mg     | 241.4              | 37                   | 0.4   | 24.26 | -80   | -7.0  | 4.0    | 15.0 | 56      | 0.1084            | 0.9143  |        |
|               | 120 mg    | 238.6              | 47                   | -5.7  | 24.93 | -79   | -15.0 | -5.0   | 5.0  | 63      | -1.5678           | 0.1238  |        |
|               | Week 2    | Placebo            | 235.7                | 96    | -5.8  | 28.57 | -153  | -17.0  | -5.0 | 10.5    | 56                | -1.9896 | 0.0495 |
|               | 15 mg     | 237.9              | 89                   | -0.5  | 31.68 | -107  | -12.0 | 0.0    | 11.0 | 182     | -0.1472           | 0.8833  |        |
|               | 30 mg     | 237.1              | 91                   | -3.9  | 24.74 | -103  | -15.0 | -2.0   | 11.0 | 70      | -1.5087           | 0.1349  |        |
|               | 60 mg     | 254.8              | 93                   | 0.7   | 25.76 | -117  | -12.0 | 3.0    | 13.0 | 68      | 0.2617            | 0.7942  |        |
|               | 120 mg    | 239.5              | 89                   | -4.8  | 23.34 | -80   | -16.0 | -2.0   | 8.0  | 52      | -1.9255           | 0.0574  |        |
|               | Week 3    | Placebo            | 253.2                | 37    | -8.4  | 48.81 | -212  | -21.0  | -4.0 | 7.0     | 112               | -1.0475 | 0.3019 |
|               | 15 mg     | 234.3              | 36                   | -2.1  | 20.16 | -58   | -14.0 | 3.0    | 11.0 | 30      | -0.6283           | 0.5339  |        |
|               | 30 mg     | 222.1              | 38                   | 4.8   | 33.12 | -45   | -8.0  | 1.5    | 7.0  | 180     | 0.8914            | 0.3785  |        |
|               | 60 mg     | 245.6              | 32                   | -2.7  | 23.91 | -64   | -15.5 | 0.5    | 11.0 | 43      | -0.6358           | 0.5296  |        |
|               | 120 mg    | 240.6              | 42                   | -3.5  | 26.43 | -67   | -16.0 | -5.5   | 5.0  | 59      | -0.8639           | 0.3927  |        |
|               | Week 4    | Placebo            | 237.6                | 95    | -4.4  | 39.53 | -167  | -18.0  | -5.0 | 8.0     | 274               | -1.0900 | 0.2785 |
|               | 15 mg     | 237.9              | 89                   | 2.2   | 23.51 | -64   | -6.0  | 5.0    | 15.0 | 61      | 0.8971            | 0.3721  |        |
|               | 30 mg     | 235.4              | 89                   | -0.8  | 29.10 | -100  | -12.0 | -2.0   | 10.0 | 103     | -0.2550           | 0.7993  |        |
|               | 60 mg     | 252.8              | 89                   | 3.7   | 35.55 | -154  | -10.0 | 2.0    | 20.0 | 122     | 0.9691            | 0.3352  |        |
|               | 120 mg    | 239.6              | 87                   | -1.9  | 29.14 | -112  | -15.0 | -2.0   | 15.0 | 62      | -0.5960           | 0.5527  |        |
| Week 6        | Placebo   | 247.0              | 40                   | -6.1  | 34.04 | -118  | -23.5 | 0.0    | 15.5 | 54      | -1.1286           | 0.2659  |        |
| 15 mg         | 244.5     | 40                 | 0.4                  | 27.68 | -47   | -17.0 | -1.0  | 15.5   | 82   | 0.0857  | 0.9322            |         |        |
| 30 mg         | 238.0     | 38                 | 3.3                  | 28.35 | -51   | -17.0 | -2.0  | 18.0   | 117  | 0.7210  | 0.4754            |         |        |
| 60 mg         | 233.6     | 32                 | 9.4                  | 26.81 | -50   | -7.0  | 10.0  | 23.0   | 72   | 1.9909  | 0.0554            |         |        |
| 120 mg        | 241.4     | 47                 | 0.5                  | 27.36 | -79   | -14.0 | 3.0   | 11.0   | 89   | 0.1173  | 0.9071            |         |        |
| Week 8        | Placebo   | 238.2              | 94                   | -3.4  | 32.86 | -138  | -20.0 | -3.5   | 6.0  | 143     | -0.9952           | 0.3222  |        |
| 15 mg         | 238.8     | 86                 | 2.6                  | 28.12 | -80   | -12.0 | 2.5   | 15.0   | 79   | 0.8551  | 0.3949            |         |        |
| 30 mg         | 235.4     | 89                 | 1.7                  | 30.97 | -121  | -10.0 | 0.0   | 19.0   | 85   | 0.5305  | 0.5971            |         |        |
| 60 mg         | 252.0     | 86                 | 0.8                  | 39.71 | -174  | -10.0 | 2.5   | 20.0   | 94   | 0.1819  | 0.8561            |         |        |
| 120 mg        | 239.5     | 84                 | 0.3                  | 30.49 | -84   | -17.0 | 1.0   | 16.0   | 80   | 0.0751  | 0.9403            |         |        |
| Week 12       | Placebo   | 238.2              | 94                   | -8.1  | 25.59 | -85   | -21.0 | -5.0   | 3.0  | 69      | -3.0714           | 0.0028  |        |
| 15 mg         | 238.8     | 86                 | 1.0                  | 31.24 | -104  | -13.0 | 6.0   | 20.0   | 90   | 0.2899  | 0.7726            |         |        |
| 30 mg         | 233.7     | 86                 | -2.0                 | 31.01 | -92   | -18.0 | 0.0   | 16.0   | 89   | -0.5877 | 0.5583            |         |        |
| 60 mg         | 252.5     | 82                 | -0.8                 | 42.69 | -190  | -15.0 | -2.0  | 22.0   | 75   | -0.1630 | 0.8709            |         |        |
| 120 mg        | 240.0     | 83                 | 3.2                  | 35.48 | -113  | -19.0 | 1.0   | 27.0   | 81   | 0.8321  | 0.4078            |         |        |
| Week 16       | Placebo   | 238.2              | 94                   | -7.2  | 27.82 | -95   | -26.0 | -7.5   | 8.0  | 65      | -2.5176           | 0.0135  |        |
| 15 mg         | 238.5     | 84                 | -1.6                 | 32.03 | -172  | -17.0 | 4.0   | 18.0   | 53   | -0.4530 | 0.6517            |         |        |
| 30 mg         | 233.8     | 85                 | 1.9                  | 34.03 | -163  | -13.0 | 1.0   | 19.0   | 92   | 0.5228  | 0.6025            |         |        |
| 60 mg         | 246.7     | 80                 | 2.3                  | 40.76 | -178  | -17.5 | 5.5   | 25.0   | 115  | 0.5102  | 0.6114            |         |        |
| 120 mg        | 238.7     | 82                 | 8.3                  | 44.32 | -120  | -13.0 | 3.5   | 22.0   | 239  | 1.6894  | 0.0950            |         |        |
| Week 20       | Placebo   | 238.2              | 94                   | -5.5  | 32.11 | -86   | -25.0 | -2.0   | 15.0 | 79      | -1.6735           | 0.0976  |        |
| 15 mg         | 238.5     | 84                 | 4.6                  | 36.12 | -122  | -14.0 | 4.5   | 24.5   | 136  | 1.1568  | 0.2507            |         |        |
| 30 mg         | 233.8     | 85                 | 0.4                  | 36.48 | -88   | -18.0 | 1.0   | 20.0   | 110  | 0.1130  | 0.9103            |         |        |
| 60 mg         | 247.3     | 79                 | -0.5                 | 44.98 | -180  | -19.0 | 0.0   | 28.0   | 108  | -0.1001 | 0.9206            |         |        |
| 120 mg        | 238.4     | 80                 | 1.4                  | 33.41 | -118  | -15.0 | 5.0   | 18.5   | 100  | 0.3715  | 0.7113            |         |        |
| Week 24       | Placebo   | 238.8              | 93                   | -6.1  | 33.28 | -71   | -30.0 | -7.0   | 18.0 | 87      | -1.7634           | 0.0812  |        |
| 15 mg         | 238.5     | 84                 | -0.1                 | 43.79 | -205  | -15.5 | 8.0   | 26.0   | 127  | -0.0199 | 0.9841            |         |        |
| 30 mg         | 230.8     | 84                 | -1.0                 | 32.63 | -127  | -18.0 | -4.5  | 12.0   | 97   | -0.2876 | 0.7744            |         |        |
| 60 mg         | 245.2     | 78                 | -1.5                 | 46.78 | -203  | -17.0 | 3.5   | 25.0   | 90   | -0.2759 | 0.7834            |         |        |
| 120 mg        | 239.1     | 79                 | -1.0                 | 31.55 | -119  | -16.0 | -5.0  | 13.0   | 106  | -0.2817 | 0.7789            |         |        |
| Week 28       | Placebo   | 238.8              | 93                   | -0.7  | 32.42 | -83   | -20.0 | -1.0   | 17.0 | 110     | -0.1983           | 0.8433  |        |
| 15 mg         | 237.7     | 83                 | -1.4                 | 33.51 | -151  | -15.0 | 5.0   | 16.0   | 67   | -0.3767 | 0.7074            |         |        |
| 30 mg         | 230.8     | 84                 | -3.6                 | 32.09 | -110  | -23.0 | -3.5  | 12.5   | 71   | -1.0335 | 0.3044            |         |        |
| 60 mg         | 244.4     | 77                 | -5.0                 | 39.73 | -195  | -17.0 | 1.0   | 13.0   | 68   | -1.1014 | 0.2742            |         |        |
| 120 mg        | 238.4     | 78                 | -2.5                 | 35.11 | -103  | -23.0 | -3.5  | 13.0   | 106  | -0.6256 | 0.5334            |         |        |
| Follow-up     | Placebo   | 236.6              | 97                   | -0.2  | 31.87 | -83   | -19.0 | 0.0    | 17.0 | 110     | -0.0605           | 0.9519  |        |
| 15 mg         | 236.0     | 86                 | -1.9                 | 33.84 | -151  | -19.0 | 5.0   | 16.0   | 67   | -0.5258 | 0.6004            |         |        |
| 30 mg         | 234.6     | 92                 | -3.6                 | 31.07 | -110  | -21.0 | -3.5  | 12.5   | 71   | -1.0971 | 0.2755            |         |        |
| 60 mg         | 252.5     | 87                 | -5.6                 | 42.54 | -195  | -20.0 | 1.0   | 16.0   | 69   | -1.2247 | 0.2240            |         |        |
| 120 mg        | 239.0     | 88                 | 0.8                  | 44.46 | -103  | -23.5 | -2.5  | 18.5   | 253  | 0.1702  | 0.8652            |         |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (7/24)

| Item          | Treatment | Summary Statistics |                      |       |      |       |       |        |       |      |      | One-sample t-test |         |        |
|---------------|-----------|--------------------|----------------------|-------|------|-------|-------|--------|-------|------|------|-------------------|---------|--------|
|               |           | Baseline Mean      | Change from Baseline |       |      |       |       |        |       |      | Max  | t-value           | p-value |        |
|               |           |                    | N                    | Mean  | SD   | Min   | Q1    | Median | Q3    |      |      |                   |         |        |
| LDH<br>(IU/L) | Week 1    | Placebo            | 202.8                | 39    | -2.8 | 14.31 | -27   | -15.0  | -2.0  | 8.0  | 30   | -1.2312           | 0.2258  |        |
|               |           | 15 mg              | 197.8                | 37    | -1.4 | 13.37 | -23   | -13.0  | 0.0   | 8.0  | 35   | -0.6269           | 0.5347  |        |
|               |           | 30 mg              | 186.4                | 39    | 0.5  | 17.32 | -41   | -9.0   | 0.0   | 12.0 | 46   | 0.1941            | 0.8471  |        |
|               |           | 60 mg              | 188.9                | 37    | -1.8 | 17.74 | -56   | -11.0  | -3.0  | 11.0 | 36   | -0.6302           | 0.5325  |        |
|               |           | 120 mg             | 198.9                | 47    | -3.0 | 18.04 | -37   | -13.0  | -3.0  | 7.0  | 47   | -1.1241           | 0.2668  |        |
|               |           | Week 2             | Placebo              | 199.8 | 96   | -2.8  | 20.40 | -61    | -15.5 | -3.0 | 9.0  | 79                | -1.3507 | 0.1800 |
|               |           |                    | 15 mg                | 197.5 | 89   | -1.0  | 16.56 | -37    | -11.0 | 0.0  | 9.0  | 41                | -0.5570 | 0.5789 |
|               |           |                    | 30 mg                | 190.0 | 91   | -0.2  | 15.59 | -45    | -8.0  | 0.0  | 9.0  | 37                | -0.1277 | 0.8987 |
|               |           |                    | 60 mg                | 192.1 | 93   | -2.1  | 15.02 | -51    | -10.0 | -2.0 | 7.0  | 45                | -1.3461 | 0.1816 |
|               |           |                    | 120 mg               | 192.3 | 89   | -1.2  | 17.66 | -33    | -10.0 | -3.0 | 7.0  | 92                | -0.6302 | 0.5302 |
|               |           | Week 3             | Placebo              | 200.9 | 36   | 8.7   | 48.63 | -58    | -8.5  | 4.5  | 12.5 | 269               | 1.0727  | 0.2907 |
|               |           |                    | 15 mg                | 194.6 | 36   | 3.4   | 20.58 | -38    | -8.5  | 2.5  | 13.0 | 62                | 0.9960  | 0.3261 |
|               |           |                    | 30 mg                | 183.9 | 38   | 3.8   | 18.64 | -42    | -8.0  | 4.0  | 15.0 | 50                | 1.2531  | 0.2180 |
|               |           |                    | 60 mg                | 181.0 | 32   | -1.5  | 15.65 | -30    | -11.0 | -5.0 | 6.5  | 40                | -0.5421 | 0.5916 |
|               |           |                    | 120 mg               | 199.8 | 42   | -3.3  | 14.40 | -32    | -13.0 | -4.0 | 4.0  | 33                | -1.4685 | 0.1496 |
|               |           | Week 4             | Placebo              | 199.3 | 95   | -1.9  | 22.04 | -52    | -14.0 | -2.0 | 8.0  | 115               | -0.8517 | 0.3965 |
|               |           |                    | 15 mg                | 197.5 | 89   | 1.7   | 22.60 | -39    | -13.0 | 1.0  | 12.0 | 120               | 0.7082  | 0.4807 |
|               |           |                    | 30 mg                | 190.9 | 89   | 1.2   | 19.15 | -48    | -9.0  | 1.0  | 11.0 | 45                | 0.5758  | 0.5662 |
|               |           |                    | 60 mg                | 191.6 | 89   | -2.0  | 20.76 | -45    | -13.0 | -4.0 | 11.0 | 91                | -0.8987 | 0.3713 |
|               |           |                    | 120 mg               | 192.5 | 87   | 2.1   | 18.60 | -43    | -8.0  | 0.0  | 12.0 | 55                | 1.0318  | 0.3051 |
|               |           | Week 6             | Placebo              | 204.1 | 40   | -5.5  | 16.29 | -40    | -17.5 | -1.0 | 4.0  | 28                | -2.1456 | 0.0382 |
|               |           |                    | 15 mg                | 197.5 | 40   | 5.0   | 36.82 | -41    | -18.0 | -1.0 | 10.5 | 160               | 0.8546  | 0.3980 |
|               |           |                    | 30 mg                | 186.7 | 38   | -1.4  | 20.02 | -46    | -10.0 | 0.0  | 10.0 | 56                | -0.4376 | 0.6642 |
|               |           |                    | 60 mg                | 182.4 | 31   | -1.7  | 14.22 | -42    | -10.0 | -1.0 | 5.0  | 32                | -0.6569 | 0.5163 |
|               |           |                    | 120 mg               | 199.5 | 47   | -1.1  | 20.80 | -52    | -15.0 | -2.0 | 6.0  | 93                | -0.3646 | 0.7171 |
|               |           | Week 8             | Placebo              | 199.5 | 94   | -1.9  | 24.90 | -83    | -12.0 | -2.0 | 8.0  | 103               | -0.7416 | 0.4602 |
|               |           |                    | 15 mg                | 197.8 | 86   | 0.2   | 19.06 | -47    | -10.0 | 0.5  | 10.0 | 52                | 0.0905  | 0.9281 |
|               |           |                    | 30 mg                | 190.9 | 89   | 1.8   | 18.70 | -40    | -10.0 | 2.0  | 11.0 | 55                | 0.8900  | 0.3759 |
|               |           |                    | 60 mg                | 190.8 | 86   | -1.4  | 19.59 | -44    | -11.0 | -3.0 | 9.0  | 49                | -0.6439 | 0.5214 |
|               |           |                    | 120 mg               | 193.1 | 84   | -0.6  | 17.37 | -49    | -9.5  | -2.5 | 11.0 | 52                | -0.3141 | 0.7543 |
|               |           | Week 12            | Placebo              | 199.5 | 94   | -2.9  | 19.06 | -95    | -12.0 | -1.0 | 7.0  | 55                | -1.4554 | 0.1489 |
|               |           |                    | 15 mg                | 197.8 | 86   | 2.9   | 22.30 | -53    | -9.0  | 0.0  | 11.0 | 97                | 1.2136  | 0.2283 |
|               |           |                    | 30 mg                | 189.9 | 85   | -0.6  | 20.04 | -55    | -12.0 | 0.0  | 11.0 | 49                | -0.2760 | 0.7832 |
|               |           |                    | 60 mg                | 191.4 | 82   | -3.1  | 19.24 | -45    | -12.0 | -3.0 | 7.0  | 59                | -1.4690 | 0.1457 |
|               |           |                    | 120 mg               | 193.1 | 83   | -1.5  | 17.51 | -44    | -11.0 | -2.0 | 8.0  | 47                | -0.8024 | 0.4247 |
|               |           | Week 16            | Placebo              | 199.5 | 94   | -1.2  | 21.70 | -80    | -16.0 | 0.0  | 11.0 | 61                | -0.5324 | 0.5957 |
|               |           |                    | 15 mg                | 198.3 | 84   | -0.2  | 19.06 | -53    | -15.0 | 0.0  | 11.0 | 54                | -0.0744 | 0.9409 |
|               |           |                    | 30 mg                | 190.1 | 85   | 2.4   | 25.75 | -48    | -13.0 | 0.0  | 13.0 | 162               | 0.8425  | 0.4019 |
|               |           |                    | 60 mg                | 189.6 | 80   | -0.8  | 20.22 | -51    | -11.0 | -3.5 | 8.5  | 77                | -0.3650 | 0.7161 |
|               |           |                    | 120 mg               | 192.1 | 82   | 0.8   | 21.86 | -47    | -16.0 | 0.5  | 17.0 | 64                | 0.3132  | 0.7549 |
|               |           | Week 20            | Placebo              | 199.5 | 94   | -3.3  | 20.34 | -67    | -11.0 | -3.0 | 7.0  | 58                | -1.5767 | 0.1183 |
|               |           |                    | 15 mg                | 198.3 | 84   | 4.4   | 23.75 | -46    | -10.0 | 1.0  | 13.0 | 100               | 1.6999  | 0.0929 |
|               |           |                    | 30 mg                | 190.1 | 85   | 1.7   | 19.66 | -45    | -10.0 | 1.0  | 13.0 | 49                | 0.8164  | 0.4166 |
|               |           |                    | 60 mg                | 190.4 | 79   | -2.2  | 19.24 | -42    | -14.0 | -1.0 | 9.0  | 67                | -1.0175 | 0.3121 |
|               |           |                    | 120 mg               | 192.5 | 80   | -1.5  | 20.12 | -51    | -15.5 | -2.5 | 12.5 | 66                | -0.6557 | 0.5139 |
|               |           | Week 24            | Placebo              | 199.4 | 93   | -2.7  | 22.58 | -82    | -21.0 | 2.0  | 14.0 | 39                | -1.1663 | 0.2465 |
|               |           |                    | 15 mg                | 198.3 | 84   | 1.6   | 18.45 | -33    | -8.5  | -1.0 | 8.5  | 73                | 0.7805  | 0.4373 |
|               |           |                    | 30 mg                | 190.7 | 84   | 0.3   | 18.82 | -51    | -13.5 | 1.0  | 10.5 | 51                | 0.1565  | 0.8760 |
|               |           | 60 mg              | 189.9                | 78    | -4.3 | 18.54 | -45   | -18.0  | -5.0  | 9.0  | 69   | -2.0394           | 0.0448  |        |
|               |           | 120 mg             | 192.5                | 79    | -1.3 | 22.88 | -60   | -14.0  | -2.0  | 13.0 | 73   | -0.4966           | 0.6209  |        |
|               | Week 28   | Placebo            | 199.4                | 93    | -5.0 | 22.03 | -68   | -19.0  | -4.0  | 7.0  | 62   | -2.1797           | 0.0318  |        |
|               |           | 15 mg              | 199.0                | 83    | -4.4 | 19.28 | -46   | -17.0  | -7.0  | 9.0  | 36   | -2.1009           | 0.0387  |        |
|               |           | 30 mg              | 190.7                | 84    | -0.2 | 21.08 | -52   | -13.0  | 1.5   | 10.5 | 84   | -0.0932           | 0.9260  |        |
|               |           | 60 mg              | 189.9                | 77    | -5.9 | 20.55 | -50   | -19.0  | -8.0  | 3.0  | 70   | -2.5287           | 0.0135  |        |
|               |           | 120 mg             | 193.2                | 78    | 2.2  | 19.50 | -44   | -11.0  | 2.0   | 14.0 | 47   | 0.9988            | 0.3210  |        |
|               | Follow-up | Placebo            | 199.4                | 97    | -4.7 | 21.68 | -68   | -19.0  | -4.0  | 7.0  | 62   | -2.1496           | 0.0341  |        |
|               |           | 15 mg              | 198.5                | 86    | -5.5 | 20.33 | -72   | -18.0  | -7.0  | 8.0  | 36   | -2.4934           | 0.0146  |        |
|               |           | 30 mg              | 190.8                | 92    | -0.9 | 20.62 | -52   | -14.5  | 1.0   | 10.0 | 84   | -0.4297           | 0.6684  |        |
|               |           | 60 mg              | 190.6                | 87    | -6.0 | 19.81 | -50   | -18.0  | -9.0  | 3.0  | 70   | -2.8415           | 0.0056  |        |
|               |           | 120 mg             | 192.9                | 88    | 2.0  | 19.38 | -44   | -11.0  | 2.0   | 15.5 | 47   | 0.9515            | 0.3440  |        |



表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (8/24)

| Item          | Treatment | Summary Statistics |                      |        |        |        |       |        |      |         |         | One-sample t-test |         |
|---------------|-----------|--------------------|----------------------|--------|--------|--------|-------|--------|------|---------|---------|-------------------|---------|
|               |           | Baseline Mean      | Change from Baseline |        |        |        |       |        |      |         | Max     | t-value           | p-value |
|               |           |                    | N                    | Mean   | SD     | Min    | Q1    | Median | Q3   |         |         |                   |         |
| CPK<br>(IU/L) | Week 1    | Placebo            | 110.8                | 39     | 4.9    | 45.05  | -110  | -11.0  | 4.0  | 14.0    | 210     | 0.6825            | 0.4991  |
|               | 15 mg     | 137.4              | 37                   | -1.3   | 44.40  | -228   | -10.0 | 3.0    | 13.0 | 90      | -0.1777 | 0.8599            |         |
|               | 30 mg     | 106.4              | 39                   | -1.3   | 29.97  | -75    | -21.0 | -4.0   | 22.0 | 52      | -0.2778 | 0.7827            |         |
|               | 60 mg     | 123.4              | 37                   | 9.9    | 77.88  | -197   | -11.0 | -3.0   | 25.0 | 386     | 0.7726  | 0.4448            |         |
|               | 120 mg    | 131.0              | 47                   | 33.0   | 213.27 | -88    | -24.0 | -6.0   | 13.0 | 1382    | 1.0615  | 0.2940            |         |
|               | Week 2    | Placebo            | 130.9                | 96     | 14.8   | 228.23 | -596  | -11.5  | 3.0  | 14.5    | 2021    | 0.6355            | 0.5267  |
|               | 15 mg     | 148.1              | 89                   | -7.4   | 70.94  | -363   | -16.0 | 1.0    | 11.0 | 169     | -0.9831 | 0.3282            |         |
|               | 30 mg     | 119.7              | 91                   | 10.6   | 93.71  | -445   | -13.0 | 0.0    | 22.0 | 640     | 1.0807  | 0.2827            |         |
|               | 60 mg     | 125.4              | 93                   | -5.6   | 51.09  | -239   | -16.0 | -2.0   | 12.0 | 148     | -1.0513 | 0.2959            |         |
|               | 120 mg    | 120.4              | 89                   | 0.7    | 33.85  | -68    | -17.0 | -1.0   | 13.0 | 109     | 0.2035  | 0.8392            |         |
|               | Week 3    | Placebo            | 109.6                | 37     | 11.5   | 57.32  | -138  | -10.0  | 5.0  | 22.0    | 211     | 1.2190            | 0.2308  |
|               | 15 mg     | 140.9              | 36                   | -1.8   | 53.52  | -181   | -23.0 | 2.5    | 11.5 | 191     | -0.2024 | 0.8408            |         |
|               | 30 mg     | 106.1              | 38                   | 27.7   | 114.51 | -82    | -21.0 | 0.0    | 18.0 | 451     | 1.4932  | 0.1439            |         |
|               | 60 mg     | 112.4              | 32                   | 8.8    | 26.64  | -37    | -7.5  | 2.0    | 23.0 | 79      | 1.8776  | 0.0699            |         |
|               | 120 mg    | 119.0              | 42                   | -1.0   | 39.41  | -58    | -13.0 | -0.5   | 8.0  | 201     | -0.1644 | 0.8702            |         |
|               | Week 4    | Placebo            | 132.5                | 95     | -4.6   | 77.06  | -526  | -12.0  | 1.0  | 17.0    | 205     | -0.5778           | 0.5648  |
|               | 15 mg     | 148.1              | 89                   | -6.2   | 80.87  | -325   | -18.0 | 0.0    | 12.0 | 442     | -0.7275 | 0.4689            |         |
|               | 30 mg     | 120.8              | 89                   | 7.2    | 77.93  | -471   | -13.0 | 0.0    | 23.0 | 340     | 0.8665  | 0.3886            |         |
|               | 60 mg     | 124.6              | 89                   | 53.7   | 478.99 | -213   | -15.0 | 4.0    | 20.0 | 4496    | 1.0578  | 0.2930            |         |
|               | 120 mg    | 119.8              | 87                   | 2.0    | 62.62  | -230   | -14.0 | 2.0    | 15.0 | 460     | 0.2979  | 0.7665            |         |
| Week 6        | Placebo   | 112.7              | 40                   | -5.9   | 30.90  | -108   | -15.0 | -3.0   | 8.0  | 76      | -1.2077 | 0.2344            |         |
| 15 mg         | 141.0     | 40                 | -10.3                | 67.33  | -285   | -29.5  | -4.0  | 9.0    | 175  | -0.9629 | 0.3415  |                   |         |
| 30 mg         | 102.1     | 38                 | 3.2                  | 48.66  | -103   | -18.0  | 4.0   | 15.0   | 164  | 0.4101  | 0.6841  |                   |         |
| 60 mg         | 115.8     | 31                 | 2.5                  | 48.77  | -92    | -20.0  | -3.0  | 18.0   | 202  | 0.2873  | 0.7759  |                   |         |
| 120 mg        | 121.9     | 47                 | 14.7                 | 87.56  | -132   | -15.0  | 2.0   | 23.0   | 546  | 1.1511  | 0.2556  |                   |         |
| Week 8        | Placebo   | 133.4              | 94                   | -7.9   | 76.80  | -537   | -12.0 | 0.5    | 18.0 | 147     | -0.9938 | 0.3229            |         |
| 15 mg         | 148.2     | 86                 | -12.5                | 75.95  | -359   | -24.0  | -2.0  | 18.0   | 186  | -1.5206 | 0.1321  |                   |         |
| 30 mg         | 120.8     | 89                 | 5.1                  | 63.29  | -464   | -9.0   | 3.0   | 20.0   | 136  | 0.7570  | 0.4511  |                   |         |
| 60 mg         | 126.5     | 86                 | 1.8                  | 63.37  | -187   | -22.0  | -0.5  | 22.0   | 350  | 0.2637  | 0.7926  |                   |         |
| 120 mg        | 121.0     | 84                 | 9.4                  | 37.01  | -106   | -9.0   | 2.0   | 22.5   | 147  | 2.3260  | 0.0225  |                   |         |
| Week 12       | Placebo   | 133.4              | 94                   | -1.0   | 112.41 | -661   | -14.0 | 4.5    | 24.0 | 493     | -0.0890 | 0.9293            |         |
| 15 mg         | 148.2     | 86                 | -7.3                 | 78.30  | -409   | -22.0  | -1.0  | 16.0   | 225  | -0.8677 | 0.3880  |                   |         |
| 30 mg         | 120.6     | 85                 | -1.6                 | 61.01  | -474   | -9.0   | 3.0   | 21.0   | 154  | -0.2489 | 0.8041  |                   |         |
| 60 mg         | 129.3     | 82                 | -0.7                 | 48.49  | -157   | -22.0  | 1.0   | 26.0   | 129  | -0.1389 | 0.8898  |                   |         |
| 120 mg        | 120.3     | 83                 | 6.3                  | 61.93  | -83    | -17.0  | 0.0   | 13.0   | 412  | 0.9322  | 0.3539  |                   |         |
| Week 16       | Placebo   | 133.4              | 94                   | -0.2   | 140.43 | -702   | -21.0 | 4.5    | 31.0 | 855     | -0.0140 | 0.9889            |         |
| 15 mg         | 150.3     | 84                 | 4.0                  | 98.81  | -393   | -12.0  | 5.0   | 24.0   | 632  | 0.3699  | 0.7124  |                   |         |
| 30 mg         | 120.9     | 85                 | 14.0                 | 108.08 | -469   | -10.0  | 5.0   | 24.0   | 640  | 1.1932  | 0.2362  |                   |         |
| 60 mg         | 128.4     | 80                 | 5.3                  | 65.85  | -200   | -18.0  | -1.0  | 18.5   | 418  | 0.7216  | 0.4727  |                   |         |
| 120 mg        | 120.1     | 82                 | 9.6                  | 69.99  | -165   | -14.0  | -1.5  | 19.0   | 502  | 1.2464  | 0.2162  |                   |         |
| Week 20       | Placebo   | 133.4              | 94                   | -14.6  | 103.84 | -661   | -19.0 | -1.5   | 15.0 | 320     | -1.3608 | 0.1769            |         |
| 15 mg         | 150.3     | 84                 | 23.6                 | 181.41 | -397   | -14.5  | 1.5   | 33.0   | 1505 | 1.1914  | 0.2369  |                   |         |
| 30 mg         | 120.9     | 85                 | 2.1                  | 73.39  | -446   | -17.0  | -1.0  | 20.0   | 283  | 0.2675  | 0.7897  |                   |         |
| 60 mg         | 128.9     | 79                 | 14.6                 | 63.75  | -185   | -13.0  | 7.0   | 26.0   | 228  | 2.0384  | 0.0449  |                   |         |
| 120 mg        | 121.5     | 80                 | 7.1                  | 45.41  | -142   | -14.0  | 1.5   | 27.0   | 169  | 1.3935  | 0.1674  |                   |         |
| Week 24       | Placebo   | 133.8              | 93                   | -7.8   | 96.51  | -693   | -22.0 | 6.0    | 18.0 | 182     | -0.7790 | 0.4380            |         |
| 15 mg         | 150.3     | 84                 | -6.4                 | 73.39  | -364   | -28.0  | 0.0   | 21.0   | 191  | -1.7939 | 0.4295  |                   |         |
| 30 mg         | 121.7     | 84                 | 6.7                  | 64.33  | -308   | -14.5  | 4.0   | 20.0   | 268  | 0.9549  | 0.3424  |                   |         |
| 60 mg         | 129.7     | 78                 | 0.1                  | 39.45  | -153   | -9.0   | 2.0   | 16.0   | 132  | 0.0316  | 0.9749  |                   |         |
| 120 mg        | 122.5     | 79                 | 13.4                 | 103.94 | -187   | -19.0  | -1.0  | 22.0   | 807  | 1.1495  | 0.2539  |                   |         |
| Week 28       | Placebo   | 133.8              | 93                   | -16.2  | 111.77 | -635   | -16.0 | -1.0   | 20.0 | 196     | -1.3944 | 0.1666            |         |
| 15 mg         | 151.1     | 83                 | -18.1                | 87.16  | -384   | -29.0  | -4.0  | 13.0   | 248  | -1.8878 | 0.0626  |                   |         |
| 30 mg         | 121.7     | 84                 | -2.5                 | 66.61  | -469   | -20.0  | -1.0  | 20.5   | 198  | -0.3456 | 0.7305  |                   |         |
| 60 mg         | 129.0     | 77                 | 2.9                  | 60.84  | -218   | -20.0  | 4.0   | 21.0   | 305  | 0.4233  | 0.6733  |                   |         |
| 120 mg        | 122.8     | 78                 | -0.6                 | 46.75  | -229   | -20.0  | 1.0   | 17.0   | 176  | -0.1041 | 0.9173  |                   |         |
| Follow-up     | Placebo   | 130.9              | 97                   | -15.1  | 109.61 | -635   | -16.0 | -1.0   | 20.0 | 196     | -1.3608 | 0.1768            |         |
| 15 mg         | 150.1     | 86                 | -18.8                | 86.44  | -384   | -27.0  | -4.0  | 12.0   | 248  | -2.0210 | 0.0464  |                   |         |
| 30 mg         | 120.3     | 92                 | -2.6                 | 63.75  | -469   | -19.5  | -2.0  | 18.5   | 198  | -0.3925 | 0.6956  |                   |         |
| 60 mg         | 124.2     | 87                 | 0.9                  | 59.08  | -218   | -20.0  | 3.0   | 19.0   | 305  | 0.1379  | 0.8906  |                   |         |
| 120 mg        | 120.7     | 88                 | -0.9                 | 46.98  | -229   | -19.5  | -1.0  | 16.5   | 176  | -0.1747 | 0.8617  |                   |         |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (9/24)

| Item                 | Treatment | Summary Statistics |                      |      |       |       |       |        |       |         | One-sample t-test |         |         |        |
|----------------------|-----------|--------------------|----------------------|------|-------|-------|-------|--------|-------|---------|-------------------|---------|---------|--------|
|                      |           | Baseline Mean      | Change from Baseline |      |       |       |       |        | Max   | t-value | p-value           |         |         |        |
|                      |           |                    | N                    | Mean | SD    | Min   | Q1    | Median |       |         |                   | Q3      |         |        |
| Uric Acid<br>(mg/dL) | Week 1    | Placebo            | 5.62                 | 39   | -0.11 | 0.510 | -1.3  | -0.40  | 0.00  | 0.20    | 1.1               | -1.3493 | 0.1852  |        |
|                      |           | 15 mg              | 5.74                 | 37   | -0.02 | 0.659 | -2.6  | -0.20  | 0.00  | 0.30    | 1.5               | -0.1995 | 0.8430  |        |
|                      |           | 30 mg              | 5.67                 | 39   | -0.15 | 0.548 | -1.7  | -0.50  | -0.10 | 0.30    | 0.7               | -1.6958 | 0.0981  |        |
|                      |           | 60 mg              | 5.86                 | 37   | 0.00  | 0.757 | -3.0  | -0.40  | 0.00  | 0.40    | 1.5               | 0.0217  | 0.9828  |        |
|                      |           | 120 mg             | 5.45                 | 47   | 0.07  | 0.604 | -1.7  | -0.30  | 0.00  | 0.50    | 1.6               | 0.8456  | 0.4022  |        |
|                      |           | Week 2             | Placebo              | 5.69 | 96    | 0.00  | 0.693 | -1.8   | -0.40 | 0.10    | 0.40              | 2.9     | 0.0442  | 0.9649 |
|                      |           |                    | 15 mg                | 5.68 | 89    | 0.09  | 0.575 | -2.4   | -0.30 | 0.20    | 0.50              | 1.4     | 1.5301  | 0.1296 |
|                      |           |                    | 30 mg                | 5.61 | 91    | 0.04  | 0.614 | -2.0   | -0.30 | 0.00    | 0.50              | 1.7     | 0.5631  | 0.5748 |
|                      |           |                    | 60 mg                | 5.78 | 93    | 0.07  | 0.826 | -3.5   | -0.30 | 0.10    | 0.40              | 4.3     | 0.8030  | 0.4240 |
|                      |           |                    | 120 mg               | 5.34 | 89    | 0.18  | 0.625 | -2.1   | -0.20 | 0.20    | 0.50              | 2.2     | 2.7312  | 0.0076 |
|                      |           | Week 3             | Placebo              | 5.67 | 36    | -0.15 | 0.623 | -1.3   | -0.70 | -0.15   | 0.30              | 1.3     | -1.4176 | 0.1651 |
|                      |           |                    | 15 mg                | 5.72 | 36    | 0.02  | 0.671 | -1.8   | -0.40 | 0.00    | 0.50              | 1.3     | 0.1739  | 0.8630 |
|                      |           |                    | 30 mg                | 5.74 | 38    | 0.06  | 0.872 | -2.8   | -0.30 | 0.10    | 0.60              | 1.9     | 0.3907  | 0.6983 |
|                      |           |                    | 60 mg                | 6.02 | 32    | -0.19 | 0.681 | -2.7   | -0.50 | -0.05   | 0.20              | 1.3     | -1.5576 | 0.1295 |
|                      |           |                    | 120 mg               | 5.48 | 42    | 0.10  | 0.694 | -1.8   | -0.30 | 0.10    | 0.50              | 2.0     | 0.9340  | 0.3558 |
|                      |           | Week 4             | Placebo              | 5.73 | 95    | -0.01 | 0.619 | -2.1   | -0.30 | 0.00    | 0.30              | 2.5     | -0.1492 | 0.8817 |
|                      |           |                    | 15 mg                | 5.68 | 89    | 0.02  | 0.665 | -2.1   | -0.40 | 0.10    | 0.40              | 1.7     | 0.3506  | 0.7267 |
|                      |           |                    | 30 mg                | 5.61 | 89    | 0.01  | 0.662 | -2.2   | -0.30 | 0.00    | 0.40              | 2.4     | 0.1762  | 0.8605 |
|                      |           |                    | 60 mg                | 5.81 | 89    | 0.03  | 0.844 | -2.8   | -0.30 | 0.00    | 0.30              | 4.9     | 0.3264  | 0.7449 |
|                      |           |                    | 120 mg               | 5.31 | 87    | 0.17  | 0.575 | -1.7   | -0.20 | 0.20    | 0.50              | 1.4     | 2.8147  | 0.0061 |
|                      |           | Week 6             | Placebo              | 5.78 | 40    | -0.25 | 0.773 | -2.3   | -0.55 | -0.20   | 0.20              | 1.3     | -2.0652 | 0.0456 |
|                      |           |                    | 15 mg                | 5.68 | 40    | -0.13 | 0.673 | -2.7   | -0.30 | -0.10   | 0.25              | 1.3     | -1.2458 | 0.2203 |
|                      |           |                    | 30 mg                | 5.59 | 38    | -0.08 | 0.819 | -2.5   | -0.60 | 0.00    | 0.40              | 2.4     | -0.6143 | 0.5428 |
|                      |           |                    | 60 mg                | 5.89 | 31    | 0.01  | 0.851 | -2.9   | -0.40 | 0.00    | 0.80              | 1.1     | 0.0844  | 0.9333 |
|                      |           |                    | 120 mg               | 5.52 | 47    | -0.01 | 0.717 | -2.1   | -0.40 | 0.10    | 0.40              | 1.5     | -0.0610 | 0.9516 |
|                      |           | Week 8             | Placebo              | 5.72 | 94    | -0.17 | 0.700 | -2.8   | -0.60 | -0.10   | 0.30              | 1.5     | -2.3563 | 0.0206 |
|                      |           |                    | 15 mg                | 5.63 | 86    | -0.01 | 0.649 | -2.4   | -0.40 | 0.00    | 0.50              | 1.3     | -0.1828 | 0.8554 |
|                      |           |                    | 30 mg                | 5.61 | 89    | -0.01 | 0.765 | -1.9   | -0.40 | 0.00    | 0.40              | 2.7     | -0.1108 | 0.9120 |
|                      |           |                    | 60 mg                | 5.80 | 86    | -0.07 | 0.686 | -3.1   | -0.50 | -0.10   | 0.40              | 2.1     | -0.9594 | 0.3401 |
|                      |           |                    | 120 mg               | 5.34 | 84    | 0.10  | 0.553 | -1.4   | -0.30 | 0.15    | 0.45              | 1.6     | 1.7163  | 0.0898 |
|                      |           | Week 12            | Placebo              | 5.72 | 94    | -0.09 | 0.753 | -2.1   | -0.60 | -0.10   | 0.30              | 2.7     | -1.1504 | 0.2529 |
|                      |           |                    | 15 mg                | 5.63 | 86    | 0.05  | 0.690 | -2.1   | -0.40 | 0.10    | 0.40              | 1.6     | 0.7030  | 0.4840 |
|                      |           |                    | 30 mg                | 5.65 | 86    | -0.02 | 0.753 | -2.1   | -0.50 | 0.00    | 0.50              | 2.3     | -0.2004 | 0.8416 |
|                      |           |                    | 60 mg                | 5.84 | 82    | -0.13 | 0.739 | -2.9   | -0.50 | -0.05   | 0.40              | 1.9     | -1.6298 | 0.1070 |
|                      |           |                    | 120 mg               | 5.38 | 83    | 0.01  | 0.621 | -1.9   | -0.40 | 0.00    | 0.40              | 1.6     | 0.1238  | 0.9018 |
|                      |           | Week 16            | Placebo              | 5.72 | 94    | -0.14 | 0.799 | -3.3   | -0.60 | -0.10   | 0.40              | 1.7     | -1.6660 | 0.0991 |
|                      |           |                    | 15 mg                | 5.61 | 84    | 0.05  | 0.731 | -1.8   | -0.40 | 0.10    | 0.50              | 2.2     | 0.6270  | 0.5323 |
|                      |           |                    | 30 mg                | 5.64 | 85    | -0.06 | 0.849 | -3.8   | -0.50 | 0.00    | 0.30              | 2.1     | -0.6903 | 0.4919 |
|                      |           |                    | 60 mg                | 5.83 | 80    | -0.15 | 0.895 | -3.8   | -0.60 | -0.10   | 0.30              | 2.2     | -1.5373 | 0.1282 |
|                      |           |                    | 120 mg               | 5.34 | 82    | 0.05  | 0.719 | -1.6   | -0.40 | 0.10    | 0.50              | 3.2     | 0.6607  | 0.5107 |
|                      |           | Week 20            | Placebo              | 5.72 | 94    | -0.16 | 0.851 | -3.5   | -0.60 | 0.00    | 0.30              | 1.7     | -1.8067 | 0.0740 |
|                      |           |                    | 15 mg                | 5.61 | 84    | 0.03  | 0.769 | -2.3   | -0.40 | 0.00    | 0.60              | 1.8     | 0.2981  | 0.7664 |
|                      |           |                    | 30 mg                | 5.64 | 85    | 0.04  | 0.873 | -2.9   | -0.40 | 0.00    | 0.50              | 2.9     | 0.3976  | 0.6919 |
|                      |           |                    | 60 mg                | 5.84 | 79    | -0.19 | 0.957 | -4.3   | -0.70 | -0.20   | 0.40              | 2.1     | -1.7512 | 0.0838 |
|                      |           |                    | 120 mg               | 5.35 | 80    | 0.09  | 0.634 | -1.6   | -0.40 | 0.10    | 0.60              | 1.5     | 1.3228  | 0.1897 |
|                      |           | Week 24            | Placebo              | 5.74 | 93    | -0.14 | 0.707 | -2.4   | -0.50 | -0.10   | 0.20              | 1.5     | -1.9494 | 0.0543 |
|                      |           |                    | 15 mg                | 5.61 | 84    | 0.08  | 0.712 | -1.9   | -0.40 | 0.00    | 0.65              | 1.4     | 1.0109  | 0.3150 |
|                      |           |                    | 30 mg                | 5.66 | 83    | -0.11 | 0.816 | -2.7   | -0.50 | -0.10   | 0.40              | 2.4     | -1.2101 | 0.2297 |
|                      |           | 60 mg              | 5.83                 | 78   | -0.15 | 0.853 | -3.1  | -0.60  | -0.10 | 0.40    | 1.8               | -1.5527 | 0.1246  |        |
|                      |           | 120 mg             | 5.39                 | 79   | -0.05 | 0.730 | -2.2  | -0.60  | 0.00  | 0.40    | 1.5               | -0.5703 | 0.5701  |        |
|                      | Week 28   | Placebo            | 5.74                 | 93   | -0.23 | 0.767 | -3.7  | -0.60  | -0.20 | 0.20    | 1.2               | -2.9084 | 0.0046  |        |
|                      |           | 15 mg              | 5.61                 | 83   | -0.20 | 0.927 | -3.6  | -0.80  | -0.20 | 0.20    | 2.3               | -1.9648 | 0.0528  |        |
|                      |           | 30 mg              | 5.66                 | 84   | -0.21 | 0.741 | -1.9  | -0.65  | -0.10 | 0.40    | 1.6               | -2.5464 | 0.0127  |        |
|                      |           | 60 mg              | 5.83                 | 77   | -0.22 | 0.797 | -3.8  | -0.70  | -0.10 | 0.30    | 1.2               | -2.4295 | 0.0175  |        |
|                      |           | 120 mg             | 5.40                 | 78   | 0.02  | 0.736 | -2.6  | -0.30  | 0.10  | 0.50    | 2.4               | 0.2307  | 0.8182  |        |
|                      | Follow-up | Placebo            | 5.70                 | 97   | -0.21 | 0.762 | -3.7  | -0.60  | -0.20 | 0.30    | 1.2               | -2.7044 | 0.0081  |        |
|                      |           | 15 mg              | 5.68                 | 86   | -0.19 | 0.921 | -3.6  | -0.70  | -0.15 | 0.20    | 2.3               | -1.9087 | 0.0597  |        |
|                      |           | 30 mg              | 5.61                 | 92   | -0.20 | 0.725 | -1.9  | -0.60  | -0.10 | 0.40    | 1.6               | -2.6621 | 0.0092  |        |
|                      |           | 60 mg              | 5.80                 | 87   | -0.18 | 0.785 | -3.8  | -0.60  | -0.10 | 0.40    | 1.3               | -2.1174 | 0.0371  |        |
|                      |           | 120 mg             | 5.34                 | 88   | -0.02 | 0.722 | -2.6  | -0.35  | 0.00  | 0.45    | 2.4               | -0.2067 | 0.8367  |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (10/24)

| Item                       | Treatment | Summary Statistics |                      |       |       |       |       |        |      |         | One-sample t-test |         |        |
|----------------------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|------|---------|-------------------|---------|--------|
|                            |           | Baseline Mean      | Change from Baseline |       |       |       |       |        | Max  | t-value | p-value           |         |        |
|                            |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median |      |         |                   | Q3      |        |
| Total Bilirubin<br>(mg/dL) | Week 1    | Placebo            | 0.54                 | 39    | -0.02 | 0.144 | -0.4  | -0.10  | 0.00 | 0.10    | 0.2               | -1.0000 | 0.3236 |
|                            |           | 15 mg              | 0.52                 | 37    | -0.04 | 0.112 | -0.3  | -0.10  | 0.00 | 0.00    | 0.2               | -2.2076 | 0.0337 |
|                            |           | 30 mg              | 0.55                 | 39    | -0.02 | 0.221 | -0.7  | -0.10  | 0.00 | 0.10    | 0.5               | -0.5785 | 0.5664 |
|                            |           | 60 mg              | 0.52                 | 37    | 0.04  | 0.155 | -0.3  | -0.10  | 0.10 | 0.10    | 0.4               | 1.4831  | 0.1468 |
|                            |           | 120 mg             | 0.54                 | 47    | -0.01 | 0.166 | -0.4  | -0.10  | 0.00 | 0.10    | 0.3               | -0.5262 | 0.6013 |
|                            | Week 2    | Placebo            | 0.59                 | 96    | -0.01 | 0.226 | -1.1  | -0.10  | 0.00 | 0.10    | 0.6               | -0.5871 | 0.5585 |
|                            |           | 15 mg              | 0.55                 | 89    | -0.00 | 0.165 | -0.4  | -0.10  | 0.00 | 0.10    | 0.5               | -0.1284 | 0.8981 |
|                            |           | 30 mg              | 0.58                 | 91    | -0.01 | 0.235 | -0.8  | -0.10  | 0.00 | 0.10    | 1.0               | -0.4018 | 0.6888 |
|                            |           | 60 mg              | 0.56                 | 93    | -0.01 | 0.173 | -0.5  | -0.10  | 0.00 | 0.10    | 0.5               | -0.7178 | 0.4747 |
|                            |           | 120 mg             | 0.55                 | 89    | 0.01  | 0.150 | -0.4  | -0.10  | 0.00 | 0.10    | 0.5               | 0.3545  | 0.7238 |
|                            | Week 3    | Placebo            | 0.54                 | 36    | -0.00 | 0.193 | -0.5  | -0.10  | 0.00 | 0.10    | 0.4               | -0.0862 | 0.9318 |
|                            |           | 15 mg              | 0.54                 | 36    | -0.02 | 0.149 | -0.3  | -0.10  | 0.00 | 0.10    | 0.4               | -0.8919 | 0.3785 |
|                            |           | 30 mg              | 0.54                 | 38    | -0.06 | 0.211 | -0.8  | -0.10  | 0.00 | 0.00    | 0.6               | -1.6113 | 0.1156 |
|                            |           | 60 mg              | 0.51                 | 32    | -0.01 | 0.176 | -0.4  | -0.10  | 0.00 | 0.10    | 0.3               | -0.2010 | 0.8420 |
|                            |           | 120 mg             | 0.55                 | 42    | -0.02 | 0.140 | -0.4  | -0.10  | 0.00 | 0.10    | 0.3               | -0.8811 | 0.3834 |
|                            | Week 4    | Placebo            | 0.59                 | 95    | 0.05  | 0.237 | -1.2  | -0.10  | 0.00 | 0.20    | 0.5               | 1.8623  | 0.0657 |
|                            |           | 15 mg              | 0.55                 | 89    | -0.01 | 0.158 | -0.4  | -0.10  | 0.00 | 0.10    | 0.5               | -0.8038 | 0.4237 |
|                            |           | 30 mg              | 0.58                 | 89    | -0.01 | 0.219 | -0.7  | -0.10  | 0.00 | 0.10    | 0.7               | -0.6284 | 0.5313 |
|                            |           | 60 mg              | 0.57                 | 89    | -0.02 | 0.174 | -0.6  | -0.10  | 0.00 | 0.10    | 0.3               | -0.9734 | 0.3330 |
|                            |           | 120 mg             | 0.54                 | 87    | 0.02  | 0.168 | -0.5  | -0.10  | 0.00 | 0.10    | 0.5               | 0.8916  | 0.3751 |
| Week 6                     | Placebo   | 0.55               | 40                   | 0.04  | 0.166 | -0.4  | 0.00  | 0.10   | 0.10 | 0.4     | 1.4294            | 0.1609  |        |
|                            | 15 mg     | 0.54               | 40                   | -0.02 | 0.136 | -0.5  | -0.10 | 0.00   | 0.10 | 0.2     | -0.8159           | 0.4195  |        |
|                            | 30 mg     | 0.57               | 38                   | 0.02  | 0.246 | -0.6  | -0.10 | 0.00   | 0.10 | 0.9     | 0.4618            | 0.6469  |        |
|                            | 60 mg     | 0.51               | 31                   | 0.02  | 0.175 | -0.3  | -0.10 | 0.00   | 0.20 | 0.3     | 0.7202            | 0.4770  |        |
|                            | 120 mg    | 0.52               | 47                   | 0.01  | 0.183 | -0.4  | -0.10 | 0.00   | 0.10 | 0.5     | 0.3980            | 0.6925  |        |
| Week 8                     | Placebo   | 0.59               | 94                   | 0.00  | 0.216 | -1.0  | -0.10 | 0.00   | 0.10 | 0.5     | 0.1434            | 0.8863  |        |
|                            | 15 mg     | 0.55               | 86                   | 0.00  | 0.175 | -0.6  | -0.10 | 0.00   | 0.10 | 0.4     | 0.0615            | 0.9511  |        |
|                            | 30 mg     | 0.58               | 89                   | -0.02 | 0.235 | -0.8  | -0.20 | 0.00   | 0.10 | 0.7     | -0.6763           | 0.5007  |        |
|                            | 60 mg     | 0.58               | 86                   | 0.04  | 0.202 | -0.5  | -0.10 | 0.00   | 0.10 | 0.9     | 1.6525            | 0.1021  |        |
|                            | 120 mg    | 0.55               | 84                   | 0.02  | 0.171 | -0.4  | -0.10 | 0.00   | 0.10 | 0.6     | 1.2123            | 0.2288  |        |
| Week 12                    | Placebo   | 0.59               | 94                   | 0.06  | 0.268 | -1.0  | -0.10 | 0.10   | 0.20 | 1.4     | 2.0380            | 0.0444  |        |
|                            | 15 mg     | 0.55               | 86                   | -0.01 | 0.184 | -0.5  | -0.10 | 0.00   | 0.10 | 0.4     | -0.4693           | 0.6401  |        |
|                            | 30 mg     | 0.59               | 86                   | -0.01 | 0.240 | -0.6  | -0.10 | 0.00   | 0.10 | 0.9     | -0.2690           | 0.7886  |        |
|                            | 60 mg     | 0.58               | 82                   | 0.02  | 0.206 | -0.7  | -0.10 | 0.00   | 0.10 | 0.7     | 1.0173            | 0.3121  |        |
|                            | 120 mg    | 0.54               | 83                   | -0.00 | 0.198 | -0.5  | -0.10 | 0.00   | 0.10 | 0.7     | -0.1111           | 0.9118  |        |
| Week 16                    | Placebo   | 0.59               | 94                   | 0.05  | 0.232 | -0.7  | -0.10 | 0.10   | 0.20 | 0.7     | 2.1808            | 0.0317  |        |
|                            | 15 mg     | 0.55               | 84                   | 0.03  | 0.176 | -0.5  | 0.00  | 0.00   | 0.10 | 0.8     | 1.3662            | 0.1756  |        |
|                            | 30 mg     | 0.59               | 85                   | 0.02  | 0.264 | -0.6  | -0.10 | 0.00   | 0.20 | 1.0     | 0.7811            | 0.4369  |        |
|                            | 60 mg     | 0.59               | 80                   | 0.02  | 0.192 | -0.5  | -0.10 | 0.00   | 0.10 | 0.7     | 1.0505            | 0.2967  |        |
|                            | 120 mg    | 0.55               | 82                   | 0.03  | 0.182 | -0.3  | -0.10 | 0.00   | 0.10 | 0.8     | 1.5792            | 0.1182  |        |
| Week 20                    | Placebo   | 0.59               | 94                   | 0.07  | 0.290 | -0.9  | -0.10 | 0.10   | 0.20 | 1.3     | 2.2787            | 0.0250  |        |
|                            | 15 mg     | 0.55               | 84                   | 0.02  | 0.175 | -0.5  | -0.10 | 0.00   | 0.10 | 0.6     | 1.0000            | 0.3202  |        |
|                            | 30 mg     | 0.59               | 85                   | 0.03  | 0.262 | -0.8  | -0.10 | 0.00   | 0.20 | 0.8     | 1.0340            | 0.3041  |        |
|                            | 60 mg     | 0.59               | 79                   | 0.03  | 0.182 | -0.3  | -0.10 | 0.00   | 0.10 | 0.6     | 1.6646            | 0.1000  |        |
|                            | 120 mg    | 0.55               | 80                   | 0.03  | 0.207 | -0.5  | -0.10 | 0.00   | 0.10 | 1.0     | 1.0826            | 0.2823  |        |
| Week 24                    | Placebo   | 0.59               | 93                   | 0.07  | 0.252 | -1.1  | -0.10 | 0.10   | 0.20 | 0.6     | 2.5966            | 0.0110  |        |
|                            | 15 mg     | 0.55               | 84                   | 0.04  | 0.198 | -0.5  | -0.05 | 0.00   | 0.20 | 0.5     | 2.0411            | 0.0444  |        |
|                            | 30 mg     | 0.59               | 84                   | 0.07  | 0.224 | -0.5  | -0.10 | 0.10   | 0.20 | 0.8     | 3.0208            | 0.0034  |        |
|                            | 60 mg     | 0.58               | 78                   | 0.07  | 0.213 | -0.5  | -0.10 | 0.10   | 0.20 | 0.7     | 2.7099            | 0.0083  |        |
|                            | 120 mg    | 0.55               | 79                   | 0.03  | 0.189 | -0.5  | -0.10 | 0.00   | 0.10 | 0.9     | 1.2483            | 0.2156  |        |
| Week 28                    | Placebo   | 0.59               | 93                   | 0.05  | 0.272 | -1.5  | 0.00  | 0.10   | 0.20 | 0.9     | 1.7148            | 0.0897  |        |
|                            | 15 mg     | 0.56               | 83                   | 0.06  | 0.237 | -0.4  | -0.10 | 0.00   | 0.20 | 1.2     | 2.2191            | 0.0292  |        |
|                            | 30 mg     | 0.59               | 84                   | 0.07  | 0.246 | -0.7  | -0.10 | 0.00   | 0.20 | 1.0     | 2.7446            | 0.0074  |        |
|                            | 60 mg     | 0.58               | 77                   | 0.08  | 0.197 | -0.4  | 0.00  | 0.10   | 0.20 | 0.9     | 3.3475            | 0.0013  |        |
|                            | 120 mg    | 0.55               | 78                   | 0.08  | 0.203 | -0.4  | 0.00  | 0.10   | 0.20 | 0.8     | 3.4505            | 0.0009  |        |
| Follow-up                  | Placebo   | 0.59               | 97                   | 0.05  | 0.268 | -1.5  | 0.00  | 0.10   | 0.20 | 0.9     | 1.7066            | 0.0911  |        |
|                            | 15 mg     | 0.55               | 86                   | 0.06  | 0.234 | -0.4  | -0.10 | 0.00   | 0.20 | 1.2     | 2.2077            | 0.0300  |        |
|                            | 30 mg     | 0.57               | 92                   | 0.08  | 0.245 | -0.7  | -0.10 | 0.00   | 0.20 | 1.0     | 3.1868            | 0.0020  |        |
|                            | 60 mg     | 0.56               | 87                   | 0.06  | 0.207 | -0.6  | 0.00  | 0.10   | 0.20 | 0.9     | 2.7442            | 0.0074  |        |
|                            | 120 mg    | 0.55               | 88                   | 0.07  | 0.203 | -0.4  | -0.10 | 0.10   | 0.20 | 0.8     | 3.0404            | 0.0031  |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (11/24)

| Item               | Treatment | Summary Statistics |                      |       |       |       |       |        |       |         | One-sample t-test |         |         |
|--------------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|---------|-------------------|---------|---------|
|                    |           | Baseline Mean      | Change from Baseline |       |       |       |       |        | Max   | t-value | p-value           |         |         |
|                    |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median |       |         |                   | Q3      |         |
| ChE<br>(IU/L/37°C) | Week 1    | Placebo            | 392.7                | 39    | -9.4  | 24.71 | -71   | -27.0  | -9.0  | 8.0     | 40                | -2.3783 | 0.0225  |
|                    |           | 15 mg              | 414.8                | 37    | -9.4  | 19.10 | -49   | -23.0  | -6.0  | 1.0     | 35                | -3.0033 | 0.0048  |
|                    |           | 30 mg              | 395.3                | 39    | -3.7  | 23.62 | -47   | -20.0  | -8.0  | 8.0     | 55                | -0.9829 | 0.3319  |
|                    |           | 60 mg              | 397.7                | 37    | -11.5 | 18.97 | -64   | -24.0  | -8.0  | 4.0     | 17                | -3.6921 | 0.0007  |
|                    |           | 120 mg             | 395.3                | 47    | -9.3  | 21.74 | -65   | -23.0  | -14.0 | 6.0     | 46                | -2.9456 | 0.0050  |
|                    | Week 2    | Placebo            | 409.6                | 96    | -10.9 | 23.83 | -63   | -26.0  | -13.0 | 2.5     | 66                | -4.4848 | <0.0001 |
|                    |           | 15 mg              | 414.0                | 89    | -8.6  | 24.33 | -61   | -29.0  | -9.0  | 9.0     | 44                | -3.3323 | 0.0013  |
|                    |           | 30 mg              | 407.5                | 91    | -11.1 | 21.41 | -60   | -26.0  | -11.0 | 5.0     | 34                | -4.9362 | <0.0001 |
|                    |           | 60 mg              | 406.3                | 93    | -14.6 | 25.68 | -116  | -30.0  | -14.0 | 4.0     | 36                | -5.4760 | <0.0001 |
|                    |           | 120 mg             | 406.1                | 89    | -16.5 | 25.86 | -113  | -31.0  | -12.0 | 1.0     | 46                | -6.0089 | <0.0001 |
|                    | Week 3    | Placebo            | 394.5                | 36    | -9.5  | 27.51 | -74   | -30.0  | -5.5  | 8.0     | 36                | -2.0662 | 0.0463  |
|                    |           | 15 mg              | 412.3                | 36    | -11.1 | 27.04 | -63   | -34.0  | -3.5  | 8.5     | 36                | -2.4715 | 0.0185  |
|                    |           | 30 mg              | 397.6                | 38    | -13.1 | 33.37 | -113  | -29.0  | -12.0 | 17.0    | 38                | -2.4206 | 0.0205  |
|                    |           | 60 mg              | 395.0                | 32    | -15.8 | 23.22 | -61   | -25.0  | -10.0 | 0.5     | 24                | -3.8449 | 0.0006  |
|                    |           | 120 mg             | 397.7                | 42    | -17.5 | 25.05 | -69   | -31.0  | -16.5 | -3.0    | 29                | -4.5144 | <0.0001 |
|                    | Week 4    | Placebo            | 410.3                | 95    | -7.8  | 24.97 | -80   | -24.0  | -5.0  | 8.0     | 71                | -3.0488 | 0.0030  |
|                    |           | 15 mg              | 414.0                | 89    | -9.2  | 28.96 | -80   | -31.0  | -8.0  | 10.0    | 101               | -2.9869 | 0.0037  |
|                    |           | 30 mg              | 408.1                | 89    | -13.0 | 28.16 | -86   | -31.0  | -14.0 | 3.0     | 75                | -4.3698 | <0.0001 |
|                    |           | 60 mg              | 406.3                | 89    | -11.5 | 26.99 | -84   | -28.0  | -11.0 | 5.0     | 55                | -4.0332 | 0.0001  |
|                    |           | 120 mg             | 406.1                | 87    | -17.0 | 25.03 | -67   | -36.0  | -17.0 | 1.0     | 46                | -6.3265 | <0.0001 |
| Week 6             | Placebo   | 395.0              | 40                   | -11.1 | 23.28 | -63   | -24.0 | -13.5  | -1.0  | 47      | -3.0022           | 0.0047  |         |
|                    | 15 mg     | 414.5              | 40                   | -19.3 | 30.87 | -125  | -36.5 | -21.0  | -1.0  | 48      | -3.9535           | 0.0003  |         |
|                    | 30 mg     | 400.9              | 38                   | -18.7 | 35.51 | -99   | -44.0 | -18.5  | 10.0  | 53      | -3.2479           | 0.0025  |         |
|                    | 60 mg     | 396.9              | 31                   | -22.3 | 29.16 | -85   | -39.0 | -16.0  | 3.0   | 23      | -4.2497           | 0.0002  |         |
|                    | 120 mg    | 388.7              | 47                   | -20.2 | 32.07 | -113  | -37.0 | -20.0  | 5.0   | 41      | -4.3117           | <0.0001 |         |
| Week 8             | Placebo   | 410.6              | 94                   | -12.4 | 24.04 | -116  | -26.0 | -12.0  | 3.0   | 37      | -4.9904           | <0.0001 |         |
|                    | 15 mg     | 413.1              | 86                   | -9.6  | 31.99 | -92   | -29.0 | -11.5  | 10.0  | 71      | -2.7879           | 0.0065  |         |
|                    | 30 mg     | 408.1              | 89                   | -15.5 | 34.05 | -119  | -41.0 | -17.0  | 6.0   | 64      | -4.2990           | <0.0001 |         |
|                    | 60 mg     | 403.5              | 86                   | -13.4 | 34.66 | -85   | -35.0 | -15.0  | 7.0   | 99      | -3.5869           | 0.0006  |         |
|                    | 120 mg    | 404.3              | 84                   | -22.2 | 34.30 | -120  | -46.0 | -24.0  | -3.0  | 88      | -5.9258           | <0.0001 |         |
| Week 12            | Placebo   | 410.6              | 94                   | -13.6 | 32.63 | -121  | -32.0 | -9.5   | 7.0   | 72      | -4.0528           | 0.0001  |         |
|                    | 15 mg     | 413.1              | 86                   | -13.4 | 31.24 | -89   | -38.0 | -13.0  | 9.0   | 61      | -3.9724           | 0.0001  |         |
|                    | 30 mg     | 408.7              | 86                   | -18.2 | 33.43 | -120  | -44.0 | -18.0  | 3.0   | 46      | -5.0578           | <0.0001 |         |
|                    | 60 mg     | 404.6              | 82                   | -16.9 | 34.22 | -88   | -44.0 | -17.0  | 8.0   | 55      | -4.4734           | <0.0001 |         |
|                    | 120 mg    | 403.6              | 83                   | -20.7 | 36.09 | -98   | -45.0 | -18.0  | 0.0   | 75      | -5.2191           | <0.0001 |         |
| Week 16            | Placebo   | 410.6              | 94                   | -13.6 | 32.53 | -93   | -35.0 | -14.5  | 8.0   | 65      | -4.0581           | 0.0001  |         |
|                    | 15 mg     | 413.5              | 84                   | -12.6 | 32.03 | -110  | -33.0 | -15.5  | 2.5   | 80      | -3.6044           | 0.0005  |         |
|                    | 30 mg     | 409.1              | 85                   | -18.6 | 37.61 | -164  | -38.0 | -18.0  | 6.0   | 62      | -4.5657           | <0.0001 |         |
|                    | 60 mg     | 403.3              | 80                   | -17.9 | 36.77 | -103  | -42.0 | -19.5  | 8.5   | 68      | -4.3636           | <0.0001 |         |
|                    | 120 mg    | 405.4              | 82                   | -20.6 | 38.15 | -109  | -46.0 | -19.0  | 3.0   | 68      | -4.8972           | <0.0001 |         |
| Week 20            | Placebo   | 410.6              | 94                   | -11.1 | 33.11 | -97   | -30.0 | -11.0  | 11.0  | 56      | -3.2619           | 0.0015  |         |
|                    | 15 mg     | 413.5              | 84                   | -13.3 | 30.70 | -66   | -36.0 | -16.0  | 7.5   | 58      | -3.9804           | 0.0001  |         |
|                    | 30 mg     | 409.1              | 85                   | -23.5 | 39.38 | -152  | -45.0 | -22.0  | 2.0   | 69      | -5.4943           | <0.0001 |         |
|                    | 60 mg     | 403.4              | 79                   | -20.9 | 34.39 | -90   | -52.0 | -23.0  | 4.0   | 67      | -5.4044           | <0.0001 |         |
|                    | 120 mg    | 401.4              | 80                   | -22.3 | 38.70 | -119  | -45.5 | -20.5  | 4.0   | 63      | -5.1511           | <0.0001 |         |
| Week 24            | Placebo   | 412.1              | 93                   | -18.5 | 42.27 | -265  | -41.0 | -13.0  | 6.0   | 58      | -4.2246           | <0.0001 |         |
|                    | 15 mg     | 413.5              | 84                   | -13.4 | 31.74 | -78   | -35.5 | -15.0  | 4.5   | 90      | -3.8602           | 0.0002  |         |
|                    | 30 mg     | 407.8              | 84                   | -20.0 | 30.41 | -103  | -41.0 | -15.5  | 3.0   | 58      | -6.0269           | <0.0001 |         |
|                    | 60 mg     | 403.9              | 78                   | -24.0 | 35.08 | -122  | -45.0 | -20.0  | 0.0   | 66      | -6.0489           | <0.0001 |         |
|                    | 120 mg    | 401.5              | 79                   | -26.0 | 41.60 | -160  | -53.0 | -19.0  | 3.0   | 59      | -5.5471           | <0.0001 |         |
| Week 28            | Placebo   | 412.1              | 93                   | -12.5 | 37.87 | -105  | -39.0 | -14.0  | 18.0  | 86      | -3.1816           | 0.0020  |         |
|                    | 15 mg     | 413.0              | 83                   | -10.9 | 34.75 | -98   | -35.0 | -14.0  | 15.0  | 74      | -2.8457           | 0.0056  |         |
|                    | 30 mg     | 407.8              | 84                   | -16.4 | 29.38 | -103  | -38.5 | -15.0  | 3.5   | 40      | -5.1055           | <0.0001 |         |
|                    | 60 mg     | 405.0              | 77                   | -22.1 | 34.35 | -103  | -45.0 | -17.0  | 4.0   | 46      | -5.6360           | <0.0001 |         |
|                    | 120 mg    | 402.4              | 78                   | -19.7 | 43.01 | -191  | -44.0 | -15.5  | 8.0   | 71      | -4.0441           | 0.0001  |         |
| Follow-up          | Placebo   | 409.9              | 97                   | -13.4 | 41.06 | -175  | -39.0 | -14.0  | 18.0  | 86      | -3.2120           | 0.0018  |         |
|                    | 15 mg     | 415.0              | 86                   | -12.6 | 38.51 | -145  | -36.0 | -14.0  | 15.0  | 74      | -3.0271           | 0.0033  |         |
|                    | 30 mg     | 406.5              | 92                   | -16.8 | 29.28 | -103  | -38.5 | -16.5  | 4.5   | 40      | -5.5012           | <0.0001 |         |
|                    | 60 mg     | 407.4              | 87                   | -22.2 | 33.26 | -103  | -44.0 | -17.0  | 4.0   | 46      | -6.2216           | <0.0001 |         |
|                    | 120 mg    | 407.0              | 88                   | -21.1 | 41.67 | -191  | -46.5 | -16.0  | 7.0   | 71      | -4.7426           | <0.0001 |         |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (12/24)

| Item                  | Treatment | Summary Statistics |                      |       |        |        |        |        |        |        |         | One-sample t-test |         |        |
|-----------------------|-----------|--------------------|----------------------|-------|--------|--------|--------|--------|--------|--------|---------|-------------------|---------|--------|
|                       |           | Baseline Mean      | Change from Baseline |       |        |        |        |        |        |        | t-value | p-value           |         |        |
|                       |           |                    | N                    | Mean  | SD     | Min    | Q1     | Median | Q3     | Max    |         |                   |         |        |
| Creatinine<br>(mg/dL) | Week 1    | Placebo            | 0.728                | 39    | 0.003  | 0.0608 | -0.16  | -0.030 | 0.000  | 0.040  | 0.12    | 0.3421            | 0.7341  |        |
|                       |           | 15 mg              | 0.709                | 37    | 0.005  | 0.0684 | -0.17  | -0.030 | 0.000  | 0.040  | 0.16    | 0.4806            | 0.6337  |        |
|                       |           | 30 mg              | 0.728                | 39    | 0.010  | 0.0582 | -0.14  | -0.020 | 0.020  | 0.050  | 0.14    | 1.1006            | 0.2780  |        |
|                       |           | 60 mg              | 0.714                | 37    | -0.008 | 0.0594 | -0.24  | -0.020 | 0.000  | 0.030  | 0.11    | -0.7748           | 0.4435  |        |
|                       |           | 120 mg             | 0.712                | 47    | 0.004  | 0.0652 | -0.15  | -0.030 | 0.000  | 0.050  | 0.15    | 0.4696            | 0.6409  |        |
|                       |           | Week 2             | Placebo              | 0.772 | 96     | -0.009 | 0.0750 | -0.30  | -0.040 | 0.000  | 0.030   | 0.22              | -1.1565 | 0.2504 |
|                       |           |                    | 15 mg                | 0.710 | 89     | 0.015  | 0.0777 | -0.13  | -0.030 | 0.010  | 0.050   | 0.51              | 1.8548  | 0.0670 |
|                       |           |                    | 30 mg                | 0.754 | 91     | 0.004  | 0.0656 | -0.18  | -0.030 | 0.000  | 0.030   | 0.21              | 0.5116  | 0.6102 |
|                       |           |                    | 60 mg                | 0.738 | 93     | 0.009  | 0.0735 | -0.20  | -0.030 | 0.000  | 0.040   | 0.29              | 1.1423  | 0.2563 |
|                       |           |                    | 120 mg               | 0.714 | 89     | 0.006  | 0.0558 | -0.10  | -0.030 | 0.010  | 0.040   | 0.17              | 1.0633  | 0.2906 |
|                       |           | Week 3             | Placebo              | 0.734 | 36     | 0.007  | 0.0565 | -0.14  | -0.025 | 0.010  | 0.055   | 0.09              | 0.7373  | 0.4658 |
|                       |           |                    | 15 mg                | 0.696 | 36     | 0.006  | 0.0675 | -0.20  | -0.030 | 0.010  | 0.045   | 0.13              | 0.5680  | 0.5737 |
|                       |           |                    | 30 mg                | 0.736 | 38     | 0.008  | 0.0683 | -0.19  | -0.020 | 0.020  | 0.040   | 0.15              | 0.7359  | 0.4664 |
|                       |           |                    | 60 mg                | 0.736 | 32     | 0.003  | 0.0959 | -0.26  | -0.040 | -0.010 | 0.040   | 0.37              | 0.1843  | 0.8550 |
|                       |           |                    | 120 mg               | 0.718 | 42     | 0.002  | 0.0557 | -0.17  | -0.030 | 0.000  | 0.030   | 0.11              | 0.2218  | 0.8256 |
|                       |           | Week 4             | Placebo              | 0.770 | 95     | -0.008 | 0.0663 | -0.30  | -0.040 | -0.010 | 0.030   | 0.15              | -1.1923 | 0.2361 |
|                       |           |                    | 15 mg                | 0.710 | 89     | 0.011  | 0.0560 | -0.14  | -0.030 | 0.000  | 0.050   | 0.22              | 1.8156  | 0.0728 |
|                       |           |                    | 30 mg                | 0.757 | 89     | -0.012 | 0.0678 | -0.34  | -0.040 | 0.000  | 0.020   | 0.17              | -1.6730 | 0.0979 |
|                       |           |                    | 60 mg                | 0.734 | 89     | 0.010  | 0.0807 | -0.23  | -0.030 | 0.000  | 0.030   | 0.49              | 1.1296  | 0.2617 |
|                       |           |                    | 120 mg               | 0.712 | 87     | 0.006  | 0.0544 | -0.19  | -0.020 | 0.000  | 0.030   | 0.15              | 1.0643  | 0.2902 |
|                       |           | Week 6             | Placebo              | 0.732 | 40     | 0.003  | 0.0852 | -0.17  | -0.030 | -0.010 | 0.030   | 0.33              | 0.2228  | 0.8249 |
|                       |           |                    | 15 mg                | 0.686 | 40     | -0.005 | 0.0894 | -0.20  | -0.060 | 0.000  | 0.025   | 0.37              | -0.3362 | 0.7385 |
|                       |           |                    | 30 mg                | 0.729 | 38     | 0.002  | 0.0720 | -0.27  | -0.020 | 0.005  | 0.040   | 0.11              | 0.1577  | 0.8755 |
|                       |           |                    | 60 mg                | 0.734 | 32     | 0.019  | 0.0903 | -0.10  | -0.030 | 0.000  | 0.040   | 0.40              | 1.2137  | 0.2340 |
|                       |           |                    | 120 mg               | 0.719 | 47     | -0.002 | 0.0621 | -0.17  | -0.040 | 0.000  | 0.040   | 0.11              | -0.1879 | 0.8518 |
|                       |           | Week 8             | Placebo              | 0.769 | 94     | -0.017 | 0.0905 | -0.41  | -0.060 | -0.020 | 0.030   | 0.26              | -1.8227 | 0.0716 |
|                       |           |                    | 15 mg                | 0.710 | 86     | -0.002 | 0.0762 | -0.43  | -0.040 | 0.010  | 0.040   | 0.15              | -0.2832 | 0.7777 |
|                       |           |                    | 30 mg                | 0.757 | 89     | -0.012 | 0.0776 | -0.35  | -0.050 | -0.010 | 0.020   | 0.25              | -1.4747 | 0.1439 |
|                       |           |                    | 60 mg                | 0.734 | 86     | 0.006  | 0.0621 | -0.17  | -0.030 | 0.000  | 0.040   | 0.23              | 0.9548  | 0.3424 |
|                       |           |                    | 120 mg               | 0.719 | 84     | -0.003 | 0.0520 | -0.12  | -0.030 | 0.000  | 0.030   | 0.11              | -0.5247 | 0.6012 |
|                       |           | Week 12            | Placebo              | 0.769 | 94     | -0.015 | 0.0731 | -0.33  | -0.050 | -0.020 | 0.030   | 0.13              | -2.0024 | 0.0482 |
|                       |           |                    | 15 mg                | 0.710 | 86     | 0.008  | 0.0829 | -0.19  | -0.040 | 0.005  | 0.050   | 0.32              | 0.8711  | 0.3861 |
|                       |           |                    | 30 mg                | 0.764 | 86     | -0.005 | 0.0749 | -0.33  | -0.050 | -0.005 | 0.030   | 0.20              | -0.6189 | 0.5377 |
|                       |           |                    | 60 mg                | 0.741 | 82     | 0.003  | 0.0684 | -0.13  | -0.030 | 0.000  | 0.040   | 0.32              | 0.3874  | 0.6995 |
|                       |           |                    | 120 mg               | 0.717 | 83     | -0.010 | 0.0653 | -0.19  | -0.050 | -0.010 | 0.030   | 0.17              | -1.3617 | 0.1770 |
|                       |           | Week 16            | Placebo              | 0.769 | 94     | -0.026 | 0.0855 | -0.33  | -0.060 | -0.030 | 0.020   | 0.20              | -2.9668 | 0.0038 |
|                       |           |                    | 15 mg                | 0.709 | 84     | 0.009  | 0.0781 | -0.24  | -0.045 | 0.010  | 0.055   | 0.28              | 1.0483  | 0.2976 |
|                       |           |                    | 30 mg                | 0.761 | 85     | 0.002  | 0.0871 | -0.18  | -0.050 | -0.010 | 0.050   | 0.37              | 0.2242  | 0.8231 |
|                       |           |                    | 60 mg                | 0.735 | 80     | -0.005 | 0.0939 | -0.31  | -0.060 | 0.000  | 0.035   | 0.55              | -0.5001 | 0.6184 |
|                       |           |                    | 120 mg               | 0.716 | 82     | 0.000  | 0.0564 | -0.11  | -0.040 | 0.000  | 0.040   | 0.14              | 0.0196  | 0.9844 |
|                       |           | Week 20            | Placebo              | 0.769 | 94     | -0.024 | 0.1035 | -0.46  | -0.060 | -0.010 | 0.030   | 0.22              | -2.2022 | 0.0301 |
|                       |           |                    | 15 mg                | 0.709 | 84     | 0.002  | 0.0672 | -0.18  | -0.035 | 0.010  | 0.040   | 0.16              | 0.2924  | 0.7707 |
|                       |           |                    | 30 mg                | 0.761 | 85     | -0.007 | 0.0890 | -0.32  | -0.060 | -0.010 | 0.050   | 0.22              | -0.7189 | 0.4742 |
|                       |           |                    | 60 mg                | 0.735 | 79     | -0.011 | 0.0745 | -0.30  | -0.040 | -0.010 | 0.030   | 0.19              | -1.2686 | 0.2083 |
|                       |           |                    | 120 mg               | 0.721 | 80     | 0.002  | 0.0627 | -0.16  | -0.040 | 0.000  | 0.040   | 0.16              | 0.2675  | 0.7898 |
|                       |           | Week 24            | Placebo              | 0.773 | 93     | -0.028 | 0.1016 | -0.36  | -0.070 | -0.010 | 0.020   | 0.33              | -2.6524 | 0.0094 |
|                       |           |                    | 15 mg                | 0.709 | 84     | 0.002  | 0.0653 | -0.22  | -0.040 | -0.005 | 0.050   | 0.14              | 0.2338  | 0.8157 |
|                       |           |                    | 30 mg                | 0.766 | 84     | -0.007 | 0.0676 | -0.26  | -0.040 | -0.010 | 0.025   | 0.25              | -0.9525 | 0.3436 |
|                       |           | 60 mg              | 0.736                | 78    | -0.006 | 0.0848 | -0.31  | -0.060 | -0.010 | 0.040  | 0.27    | -0.6011           | 0.5496  |        |
|                       |           | 120 mg             | 0.723                | 79    | -0.004 | 0.0706 | -0.14  | -0.060 | -0.010 | 0.030  | 0.23    | -0.5420           | 0.5894  |        |
|                       | Week 28   | Placebo            | 0.773                | 93    | -0.030 | 0.0900 | -0.44  | -0.060 | -0.020 | 0.020  | 0.22    | -3.1702           | 0.0021  |        |
|                       |           | 15 mg              | 0.707                | 83    | 0.000  | 0.0759 | -0.25  | -0.040 | -0.010 | 0.050  | 0.21    | 0.0000            | 1.0000  |        |
|                       |           | 30 mg              | 0.766                | 84    | -0.020 | 0.0640 | -0.21  | -0.045 | -0.015 | 0.030  | 0.18    | -2.7975           | 0.0064  |        |
|                       |           | 60 mg              | 0.736                | 77    | 0.001  | 0.0701 | -0.17  | -0.040 | -0.010 | 0.030  | 0.29    | 0.1789            | 0.8585  |        |
|                       |           | 120 mg             | 0.723                | 78    | -0.005 | 0.0730 | -0.20  | -0.050 | -0.005 | 0.040  | 0.17    | -0.5587           | 0.5780  |        |
|                       | Follow-up | Placebo            | 0.770                | 97    | -0.027 | 0.0916 | -0.44  | -0.070 | -0.020 | 0.020  | 0.22    | -2.9262           | 0.0043  |        |
|                       |           | 15 mg              | 0.709                | 86    | 0.005  | 0.0932 | -0.25  | -0.040 | -0.010 | 0.050  | 0.51    | 0.5324            | 0.5959  |        |
|                       |           | 30 mg              | 0.756                | 92    | -0.018 | 0.0650 | -0.21  | -0.045 | -0.015 | 0.030  | 0.18    | -2.7256           | 0.0077  |        |
|                       |           | 60 mg              | 0.731                | 87    | 0.010  | 0.0891 | -0.17  | -0.040 | -0.010 | 0.040  | 0.42    | 1.0831            | 0.2818  |        |
|                       |           | 120 mg             | 0.715                | 88    | -0.001 | 0.0724 | -0.20  | -0.050 | 0.000  | 0.045  | 0.17    | -0.0883           | 0.9298  |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (13/24)

| Item           | Treatment | Summary Statistics |                      |       |       |       |       |        |       |         | One-sample t-test |         |        |
|----------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|---------|-------------------|---------|--------|
|                |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       | t-value | p-value           |         |        |
|                |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    |         |                   | Max     |        |
| BUN<br>(mg/dL) | Week 1    | Placebo            | 14.87                | 39    | -1.23 | 2.832 | -7.0  | -3.00  | -1.00 | 1.00    | 4.0               | -2.7200 | 0.0098 |
|                |           | 15 mg              | 13.99                | 37    | 0.19  | 2.597 | -6.0  | -1.00  | 1.00  | 2.00    | 5.0               | 0.4431  | 0.6603 |
|                |           | 30 mg              | 14.84                | 39    | -0.42 | 3.562 | -8.0  | -3.00  | -1.00 | 2.00    | 9.0               | -0.7416 | 0.4629 |
|                |           | 60 mg              | 13.62                | 37    | -0.28 | 2.980 | -8.0  | -2.00  | -1.00 | 2.00    | 5.0               | -0.5738 | 0.5697 |
|                |           | 120 mg             | 14.40                | 47    | -0.78 | 3.647 | -12.0 | -3.00  | 0.00  | 1.00    | 9.0               | -1.4640 | 0.1500 |
|                | Week 2    | Placebo            | 14.72                | 96    | -0.82 | 3.292 | -8.0  | -3.00  | -1.00 | 1.00    | 6.0               | -2.4399 | 0.0165 |
|                |           | 15 mg              | 13.62                | 89    | 0.12  | 2.216 | -6.0  | -1.00  | 0.00  | 2.00    | 4.0               | 0.5165  | 0.6068 |
|                |           | 30 mg              | 14.94                | 91    | 0.06  | 3.101 | -7.0  | -2.00  | 0.00  | 2.00    | 8.0               | 0.1961  | 0.8450 |
|                |           | 60 mg              | 13.71                | 93    | 0.50  | 3.836 | -10.0 | -2.00  | 0.00  | 2.00    | 16.0              | 1.2516  | 0.2139 |
|                |           | 120 mg             | 14.14                | 89    | -0.41 | 2.812 | -12.0 | -2.00  | 0.00  | 1.00    | 6.0               | -1.3682 | 0.1747 |
|                | Week 3    | Placebo            | 15.45                | 36    | -1.91 | 2.867 | -12.0 | -3.50  | -1.70 | 0.00    | 3.0               | -3.9999 | 0.0003 |
|                |           | 15 mg              | 13.99                | 36    | 0.34  | 3.692 | -6.0  | -1.00  | 0.00  | 1.50    | 15.0              | 0.5598  | 0.5792 |
|                |           | 30 mg              | 15.02                | 38    | 0.33  | 3.621 | -11.0 | -1.80  | 1.00  | 3.00    | 8.0               | 0.5601  | 0.5788 |
|                |           | 60 mg              | 13.91                | 32    | 0.02  | 3.969 | -14.0 | -2.00  | -0.85 | 3.00    | 7.0               | 0.0267  | 0.9789 |
|                |           | 120 mg             | 14.38                | 42    | -0.33 | 3.451 | -14.0 | -1.00  | 0.00  | 1.00    | 6.0               | -0.6216 | 0.5377 |
|                | Week 4    | Placebo            | 14.82                | 95    | -1.00 | 2.512 | -9.0  | -3.00  | -1.00 | 1.00    | 6.0               | -3.8763 | 0.0002 |
|                |           | 15 mg              | 13.62                | 89    | 0.04  | 2.639 | -6.0  | -2.00  | 0.00  | 2.00    | 7.0               | 0.1406  | 0.8885 |
|                |           | 30 mg              | 15.00                | 89    | -0.86 | 2.851 | -13.0 | -2.00  | -1.00 | 1.00    | 4.0               | -2.8299 | 0.0058 |
|                |           | 60 mg              | 13.76                | 89    | -0.43 | 2.868 | -11.0 | -2.00  | 0.00  | 1.00    | 8.0               | -1.4269 | 0.1572 |
|                |           | 120 mg             | 14.14                | 87    | -0.27 | 3.216 | -15.0 | -2.00  | -0.20 | 2.00    | 7.0               | -0.7802 | 0.4374 |
| Week 6         | Placebo   | 15.10              | 40                   | -0.85 | 3.480 | -9.0  | -2.10 | -1.00  | 1.00  | 6.0     | -1.5355           | 0.1327  |        |
|                | 15 mg     | 14.08              | 40                   | -0.59 | 2.177 | -5.0  | -2.00 | -1.00  | 1.45  | 4.0     | -1.7213           | 0.0931  |        |
|                | 30 mg     | 15.18              | 38                   | -0.40 | 2.834 | -6.0  | -2.00 | -1.00  | 1.00  | 5.0     | -0.8757           | 0.3868  |        |
|                | 60 mg     | 13.27              | 31                   | 0.08  | 3.578 | -11.0 | -2.00 | 0.50   | 2.00  | 9.0     | 0.1255            | 0.9010  |        |
|                | 120 mg    | 14.74              | 47                   | -0.65 | 3.327 | -11.0 | -2.00 | -1.00  | 1.00  | 7.0     | -1.3373           | 0.1877  |        |
| Week 8         | Placebo   | 14.74              | 94                   | -0.59 | 3.019 | -8.0  | -3.00 | -1.00  | 1.00  | 6.0     | -1.8789           | 0.0634  |        |
|                | 15 mg     | 13.60              | 86                   | 0.05  | 3.018 | -7.0  | -2.00 | 0.00   | 2.00  | 7.0     | 0.1429            | 0.8867  |        |
|                | 30 mg     | 15.00              | 89                   | -0.37 | 2.931 | -7.0  | -2.00 | -1.00  | 2.00  | 7.8     | -1.1936           | 0.2359  |        |
|                | 60 mg     | 13.75              | 86                   | 0.07  | 3.090 | -8.0  | -2.00 | 0.00   | 2.00  | 8.0     | 0.1954            | 0.8455  |        |
|                | 120 mg    | 14.12              | 84                   | -0.29 | 3.074 | -10.0 | -2.00 | 0.00   | 2.00  | 6.0     | -0.8624           | 0.3909  |        |
| Week 12        | Placebo   | 14.74              | 94                   | -0.60 | 2.832 | -8.0  | -2.00 | -1.00  | 1.00  | 6.0     | -2.0506           | 0.0431  |        |
|                | 15 mg     | 13.60              | 86                   | -0.01 | 3.227 | -8.0  | -2.00 | 0.00   | 2.00  | 10.0    | -0.0234           | 0.9814  |        |
|                | 30 mg     | 15.05              | 86                   | -0.32 | 3.147 | -9.0  | -2.00 | 0.00   | 2.00  | 7.0     | -0.9527           | 0.3435  |        |
|                | 60 mg     | 13.85              | 82                   | -0.73 | 3.586 | -16.0 | -3.00 | 0.00   | 1.30  | 8.0     | -1.8353           | 0.0701  |        |
|                | 120 mg    | 14.17              | 83                   | -0.08 | 3.537 | -13.0 | -2.00 | 0.00   | 2.00  | 8.0     | -0.2017           | 0.8407  |        |
| Week 16        | Placebo   | 14.74              | 94                   | -1.05 | 3.599 | -15.0 | -3.00 | -1.00  | 1.00  | 8.0     | -2.8402           | 0.0055  |        |
|                | 15 mg     | 13.63              | 84                   | 0.26  | 2.651 | -6.0  | -2.00 | 0.00   | 2.00  | 6.5     | 0.8849            | 0.3787  |        |
|                | 30 mg     | 14.98              | 85                   | -0.23 | 3.419 | -10.0 | -2.00 | 0.00   | 2.00  | 12.0    | -0.6187           | 0.5378  |        |
|                | 60 mg     | 13.92              | 80                   | -0.46 | 3.333 | -13.0 | -3.00 | 0.00   | 2.00  | 5.0     | -1.2309           | 0.2220  |        |
|                | 120 mg    | 14.01              | 82                   | 0.09  | 3.593 | -8.0  | -3.00 | 0.00   | 2.00  | 12.0    | 0.2182            | 0.8278  |        |
| Week 20        | Placebo   | 14.74              | 94                   | -0.89 | 3.290 | -14.0 | -3.00 | -1.00  | 1.00  | 9.0     | -2.6369           | 0.0098  |        |
|                | 15 mg     | 13.63              | 84                   | -0.26 | 2.617 | -11.0 | -2.00 | 0.00   | 1.50  | 5.6     | -0.9213           | 0.3596  |        |
|                | 30 mg     | 14.98              | 85                   | -0.40 | 3.879 | -13.0 | -3.00 | 0.00   | 2.00  | 9.0     | -0.9395           | 0.3502  |        |
|                | 60 mg     | 13.87              | 79                   | -0.14 | 3.860 | -14.0 | -2.00 | 0.00   | 2.00  | 11.0    | -0.3177           | 0.7516  |        |
|                | 120 mg    | 13.99              | 80                   | -0.08 | 3.310 | -7.0  | -2.00 | 0.00   | 2.00  | 8.0     | -0.2162           | 0.8294  |        |
| Week 24        | Placebo   | 14.73              | 93                   | -0.45 | 3.663 | -15.0 | -2.20 | 0.00   | 2.00  | 7.0     | -1.1748           | 0.2431  |        |
|                | 15 mg     | 13.63              | 84                   | -0.13 | 3.015 | -7.0  | -2.00 | -0.50  | 2.00  | 7.0     | -0.3981           | 0.6916  |        |
|                | 30 mg     | 14.96              | 84                   | -0.21 | 3.692 | -8.0  | -3.00 | 0.00   | 2.00  | 7.0     | -0.5290           | 0.5982  |        |
|                | 60 mg     | 13.85              | 78                   | -0.42 | 3.833 | -13.0 | -2.00 | -0.85  | 1.00  | 10.0    | -0.9570           | 0.3416  |        |
|                | 120 mg    | 14.05              | 79                   | -0.22 | 2.697 | -7.0  | -2.00 | 0.00   | 1.00  | 5.0     | -0.7301           | 0.4675  |        |
| Week 28        | Placebo   | 14.73              | 93                   | -0.03 | 3.262 | -12.0 | -2.00 | 0.00   | 2.00  | 9.0     | -0.0826           | 0.9343  |        |
|                | 15 mg     | 13.60              | 83                   | -0.17 | 2.825 | -7.0  | -2.00 | 0.00   | 2.00  | 7.0     | -0.5556           | 0.5800  |        |
|                | 30 mg     | 14.96              | 84                   | -0.66 | 3.266 | -10.0 | -3.00 | -1.00  | 1.00  | 8.0     | -1.8407           | 0.0692  |        |
|                | 60 mg     | 13.85              | 77                   | 0.05  | 3.663 | -15.0 | -2.00 | 0.00   | 2.00  | 9.0     | 0.1307            | 0.8964  |        |
|                | 120 mg    | 14.10              | 78                   | -0.17 | 2.865 | -8.0  | -2.00 | 0.00   | 2.00  | 7.0     | -0.5336           | 0.5951  |        |
| Follow-up      | Placebo   | 14.78              | 97                   | 0.08  | 3.277 | -12.0 | -2.00 | 0.00   | 2.00  | 9.0     | 0.2293            | 0.8191  |        |
|                | 15 mg     | 13.65              | 86                   | -0.27 | 2.828 | -7.0  | -2.00 | 0.00   | 2.00  | 7.0     | -0.8886           | 0.3767  |        |
|                | 30 mg     | 14.81              | 92                   | -0.68 | 3.208 | -10.0 | -3.00 | -1.00  | 1.00  | 8.0     | -2.0180           | 0.0465  |        |
|                | 60 mg     | 13.73              | 87                   | 0.14  | 3.691 | -15.0 | -2.00 | 0.00   | 2.00  | 11.0    | 0.3602            | 0.7196  |        |
|                | 120 mg    | 14.19              | 88                   | -0.21 | 2.977 | -8.0  | -2.00 | 0.00   | 1.50  | 7.0     | -0.6624           | 0.5095  |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (14/24)

| Item          | Treatment | Summary Statistics |                      |      |      |      |      |      |         |         |         | One-sample t-test |        |
|---------------|-----------|--------------------|----------------------|------|------|------|------|------|---------|---------|---------|-------------------|--------|
|               |           | Baseline Mean      | Change from Baseline |      |      |      |      | Max  | t-value | p-value |         |                   |        |
|               |           |                    | N                    | Mean | SD   | Min  | Q1   |      |         |         | Median  | Q3                |        |
| Na<br>(mEq/L) | Week 1    | Placebo            | 140.0                | 39   | 0.6  | 1.71 | -2   | -1.0 | 1.0     | 2.0     | 4       | 2.2463            | 0.0306 |
|               |           | 15 mg              | 140.2                | 37   | -0.2 | 1.88 | -5   | -1.0 | 0.0     | 1.0     | 3       | -0.6111           | 0.5450 |
|               |           | 30 mg              | 140.5                | 39   | -0.1 | 1.65 | -4   | -1.0 | 0.0     | 1.0     | 3       | -0.3879           | 0.7002 |
|               |           | 60 mg              | 140.6                | 37   | -0.0 | 1.59 | -4   | -1.0 | 0.0     | 1.0     | 3       | -0.1034           | 0.9182 |
|               |           | 120 mg             | 140.1                | 47   | 0.4  | 1.62 | -4   | -1.0 | 1.0     | 2.0     | 3       | 1.6182            | 0.1125 |
|               | Week 2    | Placebo            | 140.0                | 96   | 0.4  | 1.79 | -3   | -1.0 | 0.0     | 1.0     | 5       | 2.2772            | 0.0250 |
|               |           | 15 mg              | 140.0                | 89   | -0.0 | 1.97 | -5   | -1.0 | 0.0     | 1.0     | 5       | -0.2158           | 0.8297 |
|               |           | 30 mg              | 140.4                | 91   | -0.1 | 1.73 | -5   | -1.0 | 0.0     | 1.0     | 5       | -0.7284           | 0.4683 |
|               |           | 60 mg              | 140.2                | 93   | 0.2  | 1.82 | -4   | -1.0 | 0.0     | 1.0     | 6       | 1.1377            | 0.2582 |
|               |           | 120 mg             | 139.9                | 89   | 0.4  | 1.54 | -3   | -1.0 | 0.0     | 1.0     | 5       | 2.2037            | 0.0302 |
|               | Week 3    | Placebo            | 139.7                | 36   | 0.6  | 1.53 | -2   | 0.0  | 1.0     | 2.0     | 4       | 2.4997            | 0.0173 |
|               |           | 15 mg              | 140.1                | 36   | -0.1 | 1.57 | -4   | -1.0 | 0.0     | 1.0     | 3       | -0.2128           | 0.8327 |
|               |           | 30 mg              | 140.4                | 38   | -0.1 | 1.73 | -4   | -1.0 | 0.0     | 2.0     | 3       | -0.2813           | 0.7801 |
|               |           | 60 mg              | 140.4                | 32   | 0.3  | 1.27 | -3   | -1.0 | 0.0     | 1.0     | 3       | 1.1136            | 0.2740 |
|               |           | 120 mg             | 140.0                | 42   | 0.3  | 1.83 | -4   | -1.0 | 0.0     | 1.0     | 5       | 1.1803            | 0.2447 |
|               | Week 4    | Placebo            | 140.0                | 95   | 0.5  | 1.71 | -4   | -1.0 | 0.0     | 2.0     | 6       | 2.8154            | 0.0059 |
|               |           | 15 mg              | 140.0                | 89   | 0.1  | 1.81 | -5   | -1.0 | 0.0     | 1.0     | 6       | 0.6449            | 0.5206 |
|               |           | 30 mg              | 140.4                | 89   | -0.0 | 1.92 | -5   | -1.0 | 0.0     | 1.0     | 4       | -0.2204           | 0.8261 |
|               |           | 60 mg              | 140.2                | 89   | 0.3  | 1.80 | -5   | -1.0 | 0.0     | 1.0     | 7       | 1.4123            | 0.1614 |
|               |           | 120 mg             | 139.9                | 87   | 0.3  | 1.62 | -4   | -1.0 | 0.0     | 1.0     | 3       | 1.8560            | 0.0669 |
| Week 6        | Placebo   | 140.0              | 40                   | 0.3  | 1.81 | -4   | -1.0 | 0.0  | 1.5     | 4       | 0.9601  | 0.3429            |        |
|               | 15 mg     | 139.9              | 40                   | -0.2 | 2.03 | -6   | -1.5 | 0.0  | 1.0     | 3       | -0.7004 | 0.4878            |        |
|               | 30 mg     | 140.5              | 38                   | -0.1 | 1.66 | -5   | -1.0 | 0.0  | 1.0     | 3       | -0.1955 | 0.8461            |        |
|               | 60 mg     | 140.4              | 31                   | -0.0 | 1.33 | -2   | -1.0 | 0.0  | 1.0     | 3       | -0.1352 | 0.8934            |        |
|               | 120 mg    | 140.1              | 47                   | 0.3  | 1.64 | -3   | -1.0 | 0.0  | 1.0     | 5       | 1.1577  | 0.2529            |        |
| Week 8        | Placebo   | 140.0              | 94                   | 0.6  | 1.86 | -3   | -1.0 | 1.0  | 2.0     | 6       | 3.3843  | 0.0010            |        |
|               | 15 mg     | 140.1              | 86                   | 0.4  | 2.21 | -7   | -1.0 | 1.0  | 1.0     | 7       | 1.7528  | 0.0832            |        |
|               | 30 mg     | 140.4              | 89                   | 0.0  | 1.91 | -4   | -1.0 | 0.0  | 1.0     | 5       | 0.2217  | 0.8250            |        |
|               | 60 mg     | 140.3              | 86                   | 0.2  | 1.93 | -4   | -1.0 | 0.0  | 1.0     | 7       | 1.1161  | 0.2675            |        |
|               | 120 mg    | 140.0              | 84                   | 0.4  | 1.69 | -3   | -1.0 | 0.0  | 2.0     | 4       | 2.0627  | 0.0423            |        |
| Week 12       | Placebo   | 140.0              | 94                   | 0.4  | 2.00 | -3   | -1.0 | 0.5  | 1.0     | 7       | 1.9070  | 0.0596            |        |
|               | 15 mg     | 140.1              | 86                   | 0.4  | 2.18 | -6   | -1.0 | 0.0  | 2.0     | 6       | 1.6847  | 0.0957            |        |
|               | 30 mg     | 140.4              | 86                   | 0.2  | 1.97 | -5   | -1.0 | 0.0  | 2.0     | 6       | 0.8743  | 0.3844            |        |
|               | 60 mg     | 140.3              | 82                   | 0.5  | 1.78 | -3   | -1.0 | 0.5  | 2.0     | 5       | 2.4823  | 0.0151            |        |
|               | 120 mg    | 139.9              | 83                   | 0.3  | 1.81 | -3   | -1.0 | 0.0  | 2.0     | 4       | 1.2710  | 0.2073            |        |
| Week 16       | Placebo   | 140.0              | 94                   | 0.5  | 1.94 | -5   | -1.0 | 0.0  | 2.0     | 7       | 2.7064  | 0.0081            |        |
|               | 15 mg     | 140.0              | 84                   | 0.4  | 1.71 | -4   | -1.0 | 0.0  | 1.0     | 5       | 2.0456  | 0.0440            |        |
|               | 30 mg     | 140.4              | 85                   | -0.2 | 1.99 | -5   | -1.0 | 0.0  | 1.0     | 4       | -0.9281 | 0.3560            |        |
|               | 60 mg     | 140.4              | 80                   | 0.3  | 1.93 | -4   | -1.0 | 0.0  | 1.5     | 5       | 1.5651  | 0.1216            |        |
|               | 120 mg    | 139.9              | 82                   | 0.4  | 1.60 | -3   | -1.0 | 0.0  | 1.0     | 5       | 2.3436  | 0.0215            |        |
| Week 20       | Placebo   | 140.0              | 94                   | 0.6  | 2.15 | -4   | -1.0 | 0.5  | 2.0     | 7       | 2.7391  | 0.0074            |        |
|               | 15 mg     | 140.0              | 84                   | 0.3  | 1.98 | -4   | -1.0 | 1.0  | 2.0     | 5       | 1.5447  | 0.1262            |        |
|               | 30 mg     | 140.4              | 85                   | 0.1  | 2.17 | -5   | -1.0 | 0.0  | 2.0     | 7       | 0.3495  | 0.7276            |        |
|               | 60 mg     | 140.4              | 79                   | 0.3  | 1.92 | -4   | -1.0 | 0.0  | 1.0     | 6       | 1.4035  | 0.1644            |        |
|               | 120 mg    | 139.9              | 80                   | 0.7  | 1.93 | -5   | 0.0  | 1.0  | 2.0     | 5       | 3.1320  | 0.0024            |        |
| Week 24       | Placebo   | 140.0              | 93                   | 0.5  | 1.98 | -5   | -1.0 | 1.0  | 2.0     | 6       | 2.4663  | 0.0155            |        |
|               | 15 mg     | 140.0              | 84                   | 0.4  | 1.84 | -6   | -1.0 | 1.0  | 1.0     | 5       | 2.0702  | 0.0415            |        |
|               | 30 mg     | 140.4              | 84                   | 0.1  | 1.92 | -6   | -1.0 | 0.0  | 2.0     | 4       | 0.6816  | 0.4974            |        |
|               | 60 mg     | 140.4              | 78                   | 0.9  | 2.07 | -3   | 0.0  | 1.0  | 2.0     | 7       | 3.7152  | 0.0004            |        |
|               | 120 mg    | 139.9              | 79                   | 0.7  | 1.78 | -4   | -1.0 | 1.0  | 2.0     | 4       | 3.5313  | 0.0007            |        |
| Week 28       | Placebo   | 140.0              | 93                   | 0.7  | 1.74 | -4   | -1.0 | 1.0  | 2.0     | 5       | 3.6940  | 0.0004            |        |
|               | 15 mg     | 140.0              | 83                   | 0.1  | 1.72 | -4   | -1.0 | 0.0  | 1.0     | 4       | 0.5750  | 0.5669            |        |
|               | 30 mg     | 140.4              | 84                   | -0.1 | 2.06 | -6   | -2.0 | 0.0  | 1.0     | 4       | -0.4231 | 0.6733            |        |
|               | 60 mg     | 140.4              | 77                   | 0.2  | 2.29 | -4   | -1.0 | 0.0  | 2.0     | 6       | 0.7469  | 0.4574            |        |
|               | 120 mg    | 139.9              | 78                   | 0.6  | 1.95 | -4   | -1.0 | 0.0  | 2.0     | 6       | 2.9075  | 0.0048            |        |
| Follow-up     | Placebo   | 140.0              | 97                   | 0.7  | 1.76 | -4   | -1.0 | 1.0  | 2.0     | 5       | 3.8093  | 0.0002            |        |
|               | 15 mg     | 140.0              | 86                   | 0.2  | 1.72 | -4   | -1.0 | 0.0  | 1.0     | 4       | 0.8157  | 0.4169            |        |
|               | 30 mg     | 140.4              | 92                   | -0.1 | 2.07 | -6   | -2.0 | 0.0  | 1.0     | 4       | -0.5528 | 0.5818            |        |
|               | 60 mg     | 140.3              | 87                   | 0.1  | 2.28 | -5   | -1.0 | 0.0  | 1.0     | 6       | 0.5169  | 0.6065            |        |
|               | 120 mg    | 139.9              | 88                   | 0.7  | 1.91 | -4   | -0.5 | 0.0  | 2.0     | 6       | 3.4081  | 0.0010            |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (15/24)

| Item         | Treatment | Summary Statistics |                      |       |       |       |       |        |       |         | One-sample t-test |         |        |
|--------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|---------|-------------------|---------|--------|
|              |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       | t-value | p-value           |         |        |
|              |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    |         |                   | Max     |        |
| K<br>(mEq/L) | Week 1    | Placebo            | 4.15                 | 39    | 0.05  | 0.268 | -0.5  | -0.20  | 0.00  | 0.20    | 0.7               | 1.1336  | 0.2641 |
|              |           | 15 mg              | 4.25                 | 37    | -0.09 | 0.255 | -1.2  | -0.20  | -0.10 | 0.10    | 0.3               | -2.0620 | 0.0465 |
|              |           | 30 mg              | 4.21                 | 39    | 0.06  | 0.371 | -0.7  | -0.20  | 0.00  | 0.30    | 1.0               | 0.9490  | 0.3486 |
|              |           | 60 mg              | 4.22                 | 37    | -0.11 | 0.243 | -0.6  | -0.30  | -0.10 | 0.10    | 0.3               | -2.6428 | 0.0121 |
|              |           | 120 mg             | 4.24                 | 47    | -0.03 | 0.363 | -1.1  | -0.20  | 0.00  | 0.20    | 0.9               | -0.4827 | 0.6316 |
|              | Week 2    | Placebo            | 4.16                 | 96    | 0.05  | 0.272 | -0.6  | -0.10  | 0.00  | 0.20    | 0.7               | 1.9122  | 0.0589 |
|              |           | 15 mg              | 4.27                 | 89    | -0.03 | 0.277 | -0.9  | -0.20  | 0.00  | 0.20    | 0.7               | -0.8802 | 0.3811 |
|              |           | 30 mg              | 4.26                 | 91    | -0.05 | 0.306 | -0.8  | -0.20  | 0.00  | 0.10    | 0.8               | -1.4031 | 0.1640 |
|              |           | 60 mg              | 4.22                 | 93    | 0.01  | 0.298 | -0.7  | -0.10  | 0.00  | 0.20    | 1.4               | 0.1742  | 0.8621 |
|              |           | 120 mg             | 4.26                 | 89    | -0.00 | 0.331 | -1.0  | -0.20  | 0.00  | 0.20    | 0.7               | -0.0000 | 1.0000 |
|              | Week 3    | Placebo            | 4.16                 | 36    | 0.11  | 0.521 | -0.3  | -0.10  | 0.00  | 0.10    | 2.8               | 1.2799  | 0.2090 |
|              |           | 15 mg              | 4.24                 | 36    | 0.00  | 0.333 | -1.1  | -0.20  | 0.10  | 0.25    | 0.4               | 0.0000  | 1.0000 |
|              |           | 30 mg              | 4.24                 | 38    | 0.03  | 0.332 | -0.8  | -0.20  | 0.10  | 0.30    | 0.6               | 0.5861  | 0.5614 |
|              |           | 60 mg              | 4.21                 | 32    | 0.03  | 0.257 | -0.5  | -0.15  | 0.00  | 0.25    | 0.5               | 0.6195  | 0.5401 |
|              |           | 120 mg             | 4.22                 | 42    | 0.06  | 0.356 | -0.8  | -0.20  | 0.00  | 0.20    | 1.2               | 1.1702  | 0.2487 |
|              | Week 4    | Placebo            | 4.16                 | 95    | 0.03  | 0.295 | -0.7  | -0.20  | 0.00  | 0.20    | 1.0               | 0.9752  | 0.3320 |
|              |           | 15 mg              | 4.27                 | 89    | -0.03 | 0.312 | -0.7  | -0.20  | 0.00  | 0.10    | 1.3               | -0.8159 | 0.4168 |
|              |           | 30 mg              | 4.26                 | 89    | -0.04 | 0.349 | -1.0  | -0.20  | 0.00  | 0.10    | 1.1               | -1.1250 | 0.2637 |
|              |           | 60 mg              | 4.20                 | 89    | -0.00 | 0.302 | -0.7  | -0.20  | 0.00  | 0.20    | 1.2               | -0.1054 | 0.9163 |
|              |           | 120 mg             | 4.26                 | 87    | 0.02  | 0.334 | -1.0  | -0.20  | 0.00  | 0.20    | 1.2               | 0.6739  | 0.5022 |
| Week 6       | Placebo   | 4.16               | 40                   | 0.04  | 0.323 | -0.7  | -0.20 | 0.05   | 0.30  | 0.8     | 0.7825            | 0.4386  |        |
|              | 15 mg     | 4.24               | 40                   | -0.06 | 0.295 | -0.9  | -0.20 | 0.00   | 0.15  | 0.3     | -1.2860           | 0.2060  |        |
|              | 30 mg     | 4.21               | 38                   | 0.07  | 0.351 | -0.4  | -0.20 | 0.00   | 0.20  | 1.2     | 1.1545            | 0.2557  |        |
|              | 60 mg     | 4.19               | 31                   | -0.01 | 0.316 | -0.5  | -0.20 | 0.00   | 0.30  | 0.7     | -0.1708           | 0.8656  |        |
|              | 120 mg    | 4.26               | 47                   | -0.02 | 0.368 | -0.8  | -0.30 | 0.00   | 0.20  | 1.1     | -0.3967           | 0.6934  |        |
| Week 8       | Placebo   | 4.16               | 94                   | 0.03  | 0.319 | -0.9  | -0.20 | 0.00   | 0.20  | 1.0     | 0.7762            | 0.4396  |        |
|              | 15 mg     | 4.27               | 86                   | -0.12 | 0.305 | -1.0  | -0.30 | -0.10  | 0.10  | 0.5     | -3.5671           | 0.0006  |        |
|              | 30 mg     | 4.26               | 89                   | -0.04 | 0.324 | -0.9  | -0.20 | 0.00   | 0.20  | 0.8     | -1.1780           | 0.2420  |        |
|              | 60 mg     | 4.19               | 86                   | -0.05 | 0.272 | -0.6  | -0.30 | -0.10  | 0.10  | 0.6     | -1.5876           | 0.1161  |        |
|              | 120 mg    | 4.26               | 84                   | -0.03 | 0.322 | -1.2  | -0.20 | 0.00   | 0.20  | 0.7     | -0.7111           | 0.4790  |        |
| Week 12      | Placebo   | 4.16               | 94                   | -0.01 | 0.324 | -1.1  | -0.20 | 0.00   | 0.20  | 0.7     | -0.1593           | 0.8738  |        |
|              | 15 mg     | 4.27               | 86                   | -0.09 | 0.365 | -1.4  | -0.30 | -0.05  | 0.20  | 0.9     | -2.2468           | 0.0272  |        |
|              | 30 mg     | 4.25               | 86                   | -0.07 | 0.351 | -1.1  | -0.30 | 0.00   | 0.10  | 1.0     | -1.8114           | 0.0736  |        |
|              | 60 mg     | 4.20               | 82                   | -0.08 | 0.383 | -0.9  | -0.40 | -0.10  | 0.10  | 1.2     | -1.8763           | 0.0642  |        |
|              | 120 mg    | 4.26               | 83                   | -0.07 | 0.345 | -1.0  | -0.30 | -0.10  | 0.10  | 1.0     | -1.9064           | 0.0601  |        |
| Week 16      | Placebo   | 4.16               | 94                   | -0.03 | 0.306 | -1.1  | -0.20 | 0.00   | 0.10  | 1.0     | -1.0797           | 0.2831  |        |
|              | 15 mg     | 4.27               | 84                   | -0.07 | 0.295 | -1.0  | -0.25 | 0.00   | 0.10  | 0.4     | -2.1833           | 0.0318  |        |
|              | 30 mg     | 4.25               | 85                   | -0.04 | 0.323 | -1.0  | -0.20 | 0.00   | 0.10  | 1.1     | -1.2073           | 0.2307  |        |
|              | 60 mg     | 4.19               | 80                   | -0.04 | 0.336 | -0.6  | -0.30 | -0.10  | 0.20  | 0.8     | -1.1328           | 0.2607  |        |
|              | 120 mg    | 4.25               | 82                   | -0.04 | 0.299 | -0.7  | -0.20 | -0.10  | 0.10  | 0.9     | -1.3277           | 0.1880  |        |
| Week 20      | Placebo   | 4.16               | 94                   | -0.01 | 0.276 | -0.7  | -0.20 | 0.00   | 0.10  | 1.2     | -0.2240           | 0.8232  |        |
|              | 15 mg     | 4.27               | 84                   | -0.08 | 0.287 | -0.9  | -0.30 | -0.10  | 0.10  | 0.7     | -2.6573           | 0.0094  |        |
|              | 30 mg     | 4.25               | 85                   | -0.05 | 0.364 | -1.1  | -0.30 | -0.10  | 0.20  | 0.9     | -1.2823           | 0.2033  |        |
|              | 60 mg     | 4.19               | 79                   | -0.05 | 0.332 | -0.8  | -0.30 | -0.10  | 0.20  | 0.9     | -1.2893           | 0.2011  |        |
|              | 120 mg    | 4.25               | 80                   | -0.05 | 0.307 | -0.6  | -0.30 | -0.10  | 0.20  | 0.8     | -1.5281           | 0.1305  |        |
| Week 24      | Placebo   | 4.16               | 93                   | -0.04 | 0.361 | -1.4  | -0.20 | -0.10  | 0.20  | 0.9     | -1.0327           | 0.3044  |        |
|              | 15 mg     | 4.27               | 84                   | -0.08 | 0.304 | -1.2  | -0.30 | -0.10  | 0.10  | 0.5     | -2.4408           | 0.0168  |        |
|              | 30 mg     | 4.25               | 84                   | -0.05 | 0.340 | -1.1  | -0.20 | -0.10  | 0.20  | 0.8     | -1.3788           | 0.1717  |        |
|              | 60 mg     | 4.20               | 78                   | -0.06 | 0.296 | -0.7  | -0.20 | -0.10  | 0.20  | 0.6     | -1.8722           | 0.0650  |        |
|              | 120 mg    | 4.26               | 79                   | -0.08 | 0.340 | -1.1  | -0.30 | -0.10  | 0.10  | 0.8     | -2.0520           | 0.0435  |        |
| Week 28      | Placebo   | 4.16               | 93                   | -0.06 | 0.347 | -1.2  | -0.30 | 0.00   | 0.10  | 0.9     | -1.7649           | 0.0809  |        |
|              | 15 mg     | 4.27               | 83                   | -0.08 | 0.297 | -0.8  | -0.30 | 0.00   | 0.10  | 0.7     | -2.4009           | 0.0186  |        |
|              | 30 mg     | 4.25               | 84                   | -0.11 | 0.308 | -1.0  | -0.30 | -0.10  | 0.10  | 0.7     | -3.2635           | 0.0016  |        |
|              | 60 mg     | 4.19               | 77                   | 0.00  | 0.330 | -0.9  | -0.20 | 0.00   | 0.20  | 1.2     | 0.0691            | 0.9451  |        |
|              | 120 mg    | 4.27               | 78                   | -0.03 | 0.339 | -0.8  | -0.20 | -0.10  | 0.20  | 0.9     | -0.8692           | 0.3875  |        |
| Follow-up    | Placebo   | 4.16               | 97                   | -0.06 | 0.340 | -1.2  | -0.30 | 0.00   | 0.10  | 0.9     | -1.7624           | 0.0812  |        |
|              | 15 mg     | 4.27               | 86                   | -0.08 | 0.290 | -0.8  | -0.30 | 0.00   | 0.10  | 0.7     | -2.6025           | 0.0109  |        |
|              | 30 mg     | 4.25               | 92                   | -0.11 | 0.324 | -1.0  | -0.30 | -0.10  | 0.10  | 0.7     | -3.1870           | 0.0020  |        |
|              | 60 mg     | 4.21               | 87                   | -0.02 | 0.350 | -1.0  | -0.30 | 0.00   | 0.20  | 1.2     | -0.5827           | 0.5616  |        |
|              | 120 mg    | 4.27               | 88                   | -0.01 | 0.335 | -0.8  | -0.20 | 0.00   | 0.20  | 0.9     | -0.3505           | 0.7268  |        |



表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (16/24)

| Item          | Treatment | Summary Statistics |                      |      |      |      |      |        |     |     |         | One-sample t-test |        |
|---------------|-----------|--------------------|----------------------|------|------|------|------|--------|-----|-----|---------|-------------------|--------|
|               |           | Baseline Mean      | Change from Baseline |      |      |      |      |        |     |     | t-value | p-value           |        |
|               |           |                    | N                    | Mean | SD   | Min  | Q1   | Median | Q3  | Max |         |                   |        |
| Cl<br>(mEq/L) | Week 1    | Placebo            | 101.8                | 39   | 0.3  | 1.59 | -2   | -1.0   | 0.0 | 1.0 | 4       | 1.1088            | 0.2745 |
|               |           | 15 mg              | 100.9                | 37   | 0.6  | 2.68 | -4   | -1.0   | 0.0 | 2.0 | 11      | 1.4726            | 0.1495 |
|               |           | 30 mg              | 101.9                | 39   | -0.1 | 2.12 | -5   | -1.0   | 0.0 | 1.0 | 4       | -0.3014           | 0.7647 |
|               |           | 60 mg              | 101.8                | 37   | 0.5  | 1.66 | -2   | -1.0   | 0.0 | 1.0 | 5       | 1.7823            | 0.0831 |
|               |           | 120 mg             | 101.8                | 47   | 0.7  | 1.58 | -2   | 0.0    | 0.0 | 1.0 | 5       | 3.1311            | 0.0030 |
|               | Week 2    | Placebo            | 101.1                | 96   | 0.4  | 1.79 | -5   | -1.0   | 0.0 | 1.0 | 5       | 2.1647            | 0.0329 |
|               |           | 15 mg              | 101.1                | 89   | 0.1  | 2.16 | -4   | -1.0   | 0.0 | 1.0 | 8       | 0.6376            | 0.5254 |
|               |           | 30 mg              | 101.8                | 91   | -0.1 | 2.18 | -6   | -2.0   | 0.0 | 1.0 | 6       | -0.6255           | 0.5332 |
|               |           | 60 mg              | 101.3                | 93   | 0.6  | 2.03 | -3   | -1.0   | 1.0 | 2.0 | 6       | 2.7601            | 0.0070 |
|               |           | 120 mg             | 101.3                | 89   | 0.8  | 1.94 | -5   | 0.0    | 1.0 | 2.0 | 5       | 3.7219            | 0.0003 |
|               | Week 3    | Placebo            | 101.7                | 36   | 0.1  | 2.09 | -4   | -1.0   | 0.0 | 1.5 | 5       | 0.3995            | 0.6920 |
|               |           | 15 mg              | 101.0                | 36   | -0.0 | 2.27 | -3   | -2.0   | 0.0 | 1.5 | 7       | -0.0733           | 0.9420 |
|               |           | 30 mg              | 101.9                | 38   | 0.1  | 2.34 | -7   | -1.0   | 0.0 | 2.0 | 5       | 0.2079            | 0.8365 |
|               |           | 60 mg              | 101.5                | 32   | 0.8  | 1.54 | -2   | 0.0    | 1.0 | 1.5 | 5       | 2.8707            | 0.0073 |
|               |           | 120 mg             | 101.7                | 42   | 0.9  | 2.18 | -4   | -1.0   | 1.0 | 3.0 | 5       | 2.6219            | 0.0122 |
|               | Week 4    | Placebo            | 101.1                | 95   | 0.4  | 2.07 | -7   | -1.0   | 1.0 | 2.0 | 7       | 1.6872            | 0.0949 |
|               |           | 15 mg              | 101.1                | 89   | 0.3  | 1.99 | -5   | -1.0   | 0.0 | 1.0 | 9       | 1.4417            | 0.1529 |
|               |           | 30 mg              | 101.8                | 89   | 0.2  | 2.09 | -4   | -1.0   | 0.0 | 1.0 | 6       | 1.0656            | 0.2895 |
|               |           | 60 mg              | 101.2                | 89   | 0.6  | 2.08 | -5   | -1.0   | 1.0 | 2.0 | 7       | 2.8537            | 0.0054 |
|               |           | 120 mg             | 101.3                | 87   | 0.6  | 2.02 | -4   | -1.0   | 1.0 | 2.0 | 6       | 2.9220            | 0.0044 |
| Week 6        | Placebo   | 101.6              | 40                   | -0.3 | 1.86 | -5   | -1.0 | -0.5   | 1.0 | 3   | -0.8483 | 0.4015            |        |
|               | 15 mg     | 100.9              | 40                   | 0.4  | 2.26 | -4   | -1.0 | 0.5    | 2.0 | 7   | 1.1876  | 0.2422            |        |
|               | 30 mg     | 102.1              | 38                   | -0.1 | 1.96 | -4   | -1.0 | 0.0    | 1.0 | 6   | -0.1657 | 0.8693            |        |
|               | 60 mg     | 101.4              | 31                   | 0.8  | 1.73 | -2   | 0.0  | 1.0    | 2.0 | 5   | 2.6932  | 0.0115            |        |
|               | 120 mg    | 101.5              | 47                   | 1.0  | 1.98 | -3   | -1.0 | 1.0    | 3.0 | 6   | 3.3828  | 0.0015            |        |
| Week 8        | Placebo   | 101.1              | 94                   | 0.4  | 2.07 | -5   | -1.0 | 0.0    | 2.0 | 6   | 2.0465  | 0.0435            |        |
|               | 15 mg     | 101.1              | 86                   | 0.6  | 2.06 | -4   | 0.0  | 1.0    | 2.0 | 6   | 2.7226  | 0.0079            |        |
|               | 30 mg     | 101.8              | 89                   | 0.4  | 1.97 | -5   | 0.0  | 0.0    | 2.0 | 4   | 1.7194  | 0.0891            |        |
|               | 60 mg     | 101.3              | 86                   | 0.3  | 2.25 | -6   | -1.0 | 0.5    | 2.0 | 5   | 1.1516  | 0.2527            |        |
|               | 120 mg    | 101.3              | 84                   | 0.6  | 2.04 | -5   | -1.0 | 1.0    | 2.0 | 6   | 2.6788  | 0.0089            |        |
| Week 12       | Placebo   | 101.1              | 94                   | 0.4  | 2.12 | -4   | -1.0 | 0.5    | 2.0 | 5   | 1.7552  | 0.0825            |        |
|               | 15 mg     | 101.1              | 86                   | 0.7  | 2.24 | -5   | -1.0 | 0.0    | 2.0 | 10  | 2.6925  | 0.0085            |        |
|               | 30 mg     | 101.8              | 86                   | 0.4  | 1.97 | -5   | -1.0 | 0.0    | 2.0 | 5   | 1.9191  | 0.0583            |        |
|               | 60 mg     | 101.3              | 82                   | 0.8  | 2.23 | -7   | -1.0 | 1.0    | 2.0 | 6   | 3.0767  | 0.0029            |        |
|               | 120 mg    | 101.3              | 83                   | 0.8  | 2.21 | -5   | 0.0  | 1.0    | 2.0 | 6   | 3.3343  | 0.0013            |        |
| Week 16       | Placebo   | 101.1              | 94                   | 0.4  | 1.92 | -3   | -1.0 | 0.0    | 2.0 | 7   | 1.9888  | 0.0497            |        |
|               | 15 mg     | 101.1              | 84                   | 0.6  | 2.22 | -3   | -1.0 | 1.0    | 1.5 | 12  | 2.6533  | 0.0096            |        |
|               | 30 mg     | 101.8              | 85                   | 0.0  | 2.03 | -7   | -1.0 | 0.0    | 1.0 | 4   | 0.2132  | 0.8317            |        |
|               | 60 mg     | 101.3              | 80                   | 0.6  | 2.04 | -5   | -1.0 | 1.0    | 2.0 | 7   | 2.7403  | 0.0076            |        |
|               | 120 mg    | 101.3              | 82                   | 0.6  | 2.34 | -5   | -1.0 | 0.0    | 2.0 | 5   | 2.3597  | 0.0207            |        |
| Week 20       | Placebo   | 101.1              | 94                   | 0.4  | 2.59 | -8   | -1.0 | 0.0    | 2.0 | 5   | 1.4309  | 0.1558            |        |
|               | 15 mg     | 101.1              | 84                   | 0.7  | 2.45 | -5   | -1.0 | 1.0    | 2.0 | 11  | 2.4928  | 0.0147            |        |
|               | 30 mg     | 101.8              | 85                   | 0.5  | 2.31 | -7   | -1.0 | 1.0    | 2.0 | 6   | 2.0637  | 0.0421            |        |
|               | 60 mg     | 101.3              | 79                   | 0.8  | 2.15 | -4   | -1.0 | 1.0    | 2.0 | 5   | 3.1950  | 0.0020            |        |
|               | 120 mg    | 101.2              | 80                   | 0.8  | 2.22 | -6   | 0.0  | 0.5    | 2.0 | 6   | 3.0755  | 0.0029            |        |
| Week 24       | Placebo   | 101.1              | 93                   | 0.5  | 2.23 | -5   | -1.0 | 1.0    | 2.0 | 5   | 2.3209  | 0.0225            |        |
|               | 15 mg     | 101.1              | 84                   | 0.7  | 2.32 | -4   | -1.0 | 0.0    | 2.0 | 10  | 2.5917  | 0.0113            |        |
|               | 30 mg     | 101.7              | 84                   | 0.4  | 2.12 | -5   | -1.0 | 0.5    | 2.0 | 5   | 1.8539  | 0.0673            |        |
|               | 60 mg     | 101.3              | 78                   | 1.2  | 2.47 | -4   | -1.0 | 1.0    | 3.0 | 7   | 4.3139  | <0.0001           |        |
|               | 120 mg    | 101.2              | 79                   | 0.8  | 2.16 | -5   | -1.0 | 1.0    | 2.0 | 6   | 3.1774  | 0.0021            |        |
| Week 28       | Placebo   | 101.1              | 93                   | 0.5  | 2.09 | -4   | -1.0 | 0.0    | 2.0 | 6   | 2.4272  | 0.0172            |        |
|               | 15 mg     | 101.2              | 83                   | 0.4  | 2.25 | -5   | -1.0 | 0.0    | 2.0 | 8   | 1.5118  | 0.1344            |        |
|               | 30 mg     | 101.7              | 84                   | -0.1 | 2.37 | -6   | -2.0 | 0.0    | 1.0 | 6   | -0.4601 | 0.6467            |        |
|               | 60 mg     | 101.3              | 77                   | 0.4  | 2.19 | -4   | -1.0 | 1.0    | 2.0 | 6   | 1.6131  | 0.1109            |        |
|               | 120 mg    | 101.2              | 78                   | 0.2  | 2.31 | -5   | -1.0 | 0.0    | 2.0 | 7   | 0.7844  | 0.4352            |        |
| Follow-up     | Placebo   | 101.1              | 97                   | 0.6  | 2.16 | -4   | -1.0 | 0.0    | 2.0 | 7   | 2.5856  | 0.0112            |        |
|               | 15 mg     | 101.1              | 86                   | 0.5  | 2.30 | -5   | -1.0 | 0.5    | 2.0 | 8   | 1.8251  | 0.0715            |        |
|               | 30 mg     | 101.8              | 92                   | -0.1 | 2.30 | -6   | -1.0 | 0.0    | 1.0 | 6   | -0.3634 | 0.7171            |        |
|               | 60 mg     | 101.2              | 87                   | 0.4  | 2.15 | -4   | -1.0 | 1.0    | 2.0 | 6   | 1.7916  | 0.0767            |        |
|               | 120 mg    | 101.3              | 88                   | 0.3  | 2.22 | -5   | -1.0 | 0.0    | 2.0 | 7   | 1.0543  | 0.2947            |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (17/24)

| Item          | Treatment | Summary Statistics |                      |       |       |       |       |       |         |         |         | One-sample t-test |        |
|---------------|-----------|--------------------|----------------------|-------|-------|-------|-------|-------|---------|---------|---------|-------------------|--------|
|               |           | Baseline Mean      | Change from Baseline |       |       |       |       | Max   | t-value | p-value |         |                   |        |
|               |           |                    | N                    | Mean  | SD    | Min   | Q1    |       |         |         | Median  | Q3                |        |
| Ca<br>(mg/dL) | Week 1    | Placebo            | 9.65                 | 39    | 0.03  | 0.346 | -0.7  | -0.20 | 0.00    | 0.20    | 1.1     | 0.4625            | 0.6463 |
|               |           | 15 mg              | 9.68                 | 37    | -0.10 | 0.315 | -0.8  | -0.30 | -0.10   | 0.10    | 0.5     | -1.9839           | 0.0549 |
|               |           | 30 mg              | 9.61                 | 39    | 0.07  | 0.308 | -0.5  | -0.20 | 0.00    | 0.30    | 0.7     | 1.4041            | 0.1684 |
|               |           | 60 mg              | 9.66                 | 37    | -0.13 | 0.253 | -0.8  | -0.20 | -0.10   | 0.00    | 0.4     | -3.1238           | 0.0035 |
|               |           | 120 mg             | 9.63                 | 47    | -0.05 | 0.327 | -0.7  | -0.30 | 0.00    | 0.10    | 0.7     | -0.9815           | 0.3315 |
|               | Week 2    | Placebo            | 9.73                 | 96    | -0.03 | 0.342 | -0.7  | -0.30 | -0.10   | 0.20    | 1.2     | -0.8642           | 0.3896 |
|               |           | 15 mg              | 9.71                 | 89    | -0.05 | 0.346 | -1.4  | -0.20 | 0.00    | 0.20    | 0.8     | -1.4381           | 0.1540 |
|               |           | 30 mg              | 9.65                 | 91    | -0.03 | 0.350 | -0.7  | -0.30 | -0.10   | 0.20    | 0.7     | -0.8979           | 0.3716 |
|               |           | 60 mg              | 9.65                 | 93    | -0.10 | 0.353 | -1.3  | -0.30 | -0.10   | 0.10    | 1.0     | -2.6760           | 0.0088 |
|               |           | 120 mg             | 9.67                 | 89    | -0.09 | 0.312 | -0.8  | -0.30 | -0.10   | 0.10    | 0.8     | -2.6515           | 0.0095 |
|               | Week 3    | Placebo            | 9.67                 | 36    | 0.08  | 0.416 | -0.6  | -0.20 | 0.05    | 0.30    | 1.3     | 1.1213            | 0.2698 |
|               |           | 15 mg              | 9.65                 | 36    | -0.10 | 0.361 | -0.8  | -0.35 | 0.00    | 0.20    | 0.6     | -1.6144           | 0.1154 |
|               |           | 30 mg              | 9.60                 | 38    | 0.01  | 0.333 | -0.6  | -0.20 | 0.00    | 0.20    | 0.8     | 0.1950            | 0.8464 |
|               |           | 60 mg              | 9.69                 | 32    | -0.05 | 0.254 | -0.6  | -0.20 | -0.10   | 0.05    | 0.6     | -1.1832           | 0.2457 |
|               |           | 120 mg             | 9.63                 | 42    | -0.05 | 0.313 | -0.7  | -0.30 | -0.05   | 0.20    | 0.6     | -1.0368           | 0.3059 |
|               | Week 4    | Placebo            | 9.73                 | 95    | 0.01  | 0.328 | -0.7  | -0.20 | 0.00    | 0.20    | 0.9     | 0.3129            | 0.7551 |
|               |           | 15 mg              | 9.71                 | 89    | -0.05 | 0.331 | -0.9  | -0.20 | 0.00    | 0.10    | 0.8     | -1.3770           | 0.1720 |
|               |           | 30 mg              | 9.66                 | 89    | -0.01 | 0.317 | -0.9  | -0.20 | 0.00    | 0.20    | 1.1     | -0.3343           | 0.7390 |
|               |           | 60 mg              | 9.66                 | 89    | -0.04 | 0.308 | -0.9  | -0.20 | -0.10   | 0.20    | 0.9     | -1.2743           | 0.2059 |
|               |           | 120 mg             | 9.68                 | 87    | -0.06 | 0.333 | -0.8  | -0.30 | -0.10   | 0.20    | 0.8     | -1.7721           | 0.0799 |
| Week 6        | Placebo   | 9.67               | 40                   | -0.03 | 0.376 | -0.8  | -0.30 | 0.00  | 0.20    | 1.1     | -0.4205 | 0.6765            |        |
|               | 15 mg     | 9.65               | 40                   | -0.12 | 0.295 | -0.7  | -0.30 | -0.10 | 0.10    | 0.5     | -2.5765 | 0.0139            |        |
|               | 30 mg     | 9.61               | 38                   | -0.01 | 0.334 | -0.7  | -0.20 | -0.05 | 0.20    | 0.7     | -0.0972 | 0.9231            |        |
|               | 60 mg     | 9.71               | 31                   | -0.11 | 0.300 | -0.9  | -0.30 | -0.10 | 0.10    | 0.5     | -2.0974 | 0.0445            |        |
|               | 120 mg    | 9.64               | 47                   | -0.06 | 0.305 | -0.6  | -0.30 | -0.10 | 0.20    | 0.5     | -1.3850 | 0.1727            |        |
| Week 8        | Placebo   | 9.73               | 94                   | -0.02 | 0.357 | -1.0  | -0.20 | 0.00  | 0.20    | 0.9     | -0.5490 | 0.5843            |        |
|               | 15 mg     | 9.72               | 86                   | -0.03 | 0.374 | -0.8  | -0.30 | 0.00  | 0.20    | 1.0     | -0.8655 | 0.3892            |        |
|               | 30 mg     | 9.66               | 89                   | -0.01 | 0.337 | -0.8  | -0.30 | 0.00  | 0.20    | 1.4     | -0.4090 | 0.6836            |        |
|               | 60 mg     | 9.66               | 86                   | -0.03 | 0.364 | -1.2  | -0.20 | 0.00  | 0.20    | 0.7     | -0.7401 | 0.4613            |        |
|               | 120 mg    | 9.67               | 84                   | -0.08 | 0.365 | -1.0  | -0.35 | -0.10 | 0.20    | 0.8     | -2.0640 | 0.0421            |        |
| Week 12       | Placebo   | 9.73               | 94                   | -0.01 | 0.334 | -0.9  | -0.20 | 0.00  | 0.20    | 1.0     | -0.4015 | 0.6890            |        |
|               | 15 mg     | 9.72               | 86                   | -0.07 | 0.363 | -1.2  | -0.30 | -0.10 | 0.20    | 0.8     | -1.8139 | 0.0732            |        |
|               | 30 mg     | 9.67               | 86                   | -0.00 | 0.336 | -0.7  | -0.20 | 0.00  | 0.20    | 0.9     | -0.0321 | 0.9745            |        |
|               | 60 mg     | 9.67               | 82                   | -0.06 | 0.360 | -1.0  | -0.30 | 0.00  | 0.20    | 1.0     | -1.4429 | 0.1529            |        |
|               | 120 mg    | 9.67               | 83                   | -0.09 | 0.336 | -0.9  | -0.30 | -0.10 | 0.10    | 0.6     | -2.4171 | 0.0179            |        |
| Week 16       | Placebo   | 9.73               | 94                   | -0.01 | 0.380 | -0.9  | -0.30 | 0.00  | 0.20    | 1.2     | -0.2442 | 0.8076            |        |
|               | 15 mg     | 9.71               | 84                   | -0.05 | 0.327 | -0.8  | -0.30 | 0.00  | 0.20    | 0.8     | -1.3684 | 0.1749            |        |
|               | 30 mg     | 9.66               | 85                   | -0.02 | 0.326 | -0.9  | -0.20 | 0.00  | 0.20    | 0.9     | -0.4992 | 0.6190            |        |
|               | 60 mg     | 9.67               | 80                   | 0.01  | 0.340 | -1.0  | -0.20 | 0.10  | 0.20    | 0.7     | 0.1975  | 0.8440            |        |
|               | 120 mg    | 9.67               | 82                   | -0.06 | 0.361 | -0.8  | -0.30 | -0.10 | 0.20    | 0.8     | -1.5003 | 0.1374            |        |
| Week 20       | Placebo   | 9.73               | 94                   | -0.01 | 0.335 | -0.7  | -0.30 | -0.10 | 0.20    | 1.2     | -0.4314 | 0.6672            |        |
|               | 15 mg     | 9.71               | 84                   | -0.07 | 0.339 | -1.0  | -0.30 | 0.00  | 0.10    | 0.8     | -1.9624 | 0.0531            |        |
|               | 30 mg     | 9.66               | 85                   | -0.02 | 0.391 | -0.8  | -0.30 | -0.10 | 0.20    | 1.0     | -0.3887 | 0.6984            |        |
|               | 60 mg     | 9.67               | 79                   | -0.03 | 0.354 | -1.0  | -0.30 | 0.00  | 0.20    | 1.0     | -0.7631 | 0.4477            |        |
|               | 120 mg    | 9.66               | 80                   | -0.06 | 0.361 | -0.9  | -0.35 | -0.10 | 0.20    | 0.7     | -1.5165 | 0.1334            |        |
| Week 24       | Placebo   | 9.74               | 93                   | -0.06 | 0.354 | -0.9  | -0.30 | -0.10 | 0.10    | 1.1     | -1.5219 | 0.1315            |        |
|               | 15 mg     | 9.71               | 84                   | -0.03 | 0.331 | -0.8  | -0.20 | -0.10 | 0.15    | 1.0     | -0.9561 | 0.3418            |        |
|               | 30 mg     | 9.66               | 84                   | -0.02 | 0.287 | -0.8  | -0.20 | 0.00  | 0.20    | 0.6     | -0.5708 | 0.5697            |        |
|               | 60 mg     | 9.68               | 78                   | -0.04 | 0.361 | -1.0  | -0.30 | 0.00  | 0.20    | 1.0     | -0.8771 | 0.3831            |        |
|               | 120 mg    | 9.67               | 79                   | -0.08 | 0.399 | -0.9  | -0.40 | -0.10 | 0.20    | 1.1     | -1.7486 | 0.0843            |        |
| Week 28       | Placebo   | 9.74               | 93                   | 0.01  | 0.380 | -0.7  | -0.20 | 0.00  | 0.20    | 1.8     | 0.1364  | 0.8918            |        |
|               | 15 mg     | 9.71               | 83                   | -0.01 | 0.361 | -1.0  | -0.20 | 0.00  | 0.20    | 1.1     | -0.3037 | 0.7621            |        |
|               | 30 mg     | 9.66               | 84                   | -0.02 | 0.347 | -0.9  | -0.30 | 0.00  | 0.20    | 0.7     | -0.5027 | 0.6165            |        |
|               | 60 mg     | 9.68               | 77                   | -0.04 | 0.351 | -0.9  | -0.30 | 0.00  | 0.20    | 0.8     | -0.9740 | 0.3331            |        |
|               | 120 mg    | 9.67               | 78                   | 0.01  | 0.346 | -0.9  | -0.20 | 0.00  | 0.20    | 0.9     | 0.2290  | 0.8195            |        |
| Follow-up     | Placebo   | 9.74               | 97                   | 0.00  | 0.373 | -0.7  | -0.20 | 0.00  | 0.20    | 1.8     | 0.1088  | 0.9136            |        |
|               | 15 mg     | 9.71               | 86                   | -0.01 | 0.357 | -1.0  | -0.20 | 0.00  | 0.20    | 1.1     | -0.2416 | 0.8096            |        |
|               | 30 mg     | 9.64               | 92                   | -0.01 | 0.341 | -0.9  | -0.30 | 0.00  | 0.20    | 0.7     | -0.3980 | 0.6915            |        |
|               | 60 mg     | 9.67               | 87                   | -0.04 | 0.347 | -0.9  | -0.30 | 0.00  | 0.20    | 0.8     | -1.1730 | 0.2440            |        |
|               | 120 mg    | 9.67               | 88                   | 0.01  | 0.343 | -0.9  | -0.20 | 0.00  | 0.20    | 0.9     | 0.1556  | 0.8767            |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (18/24)

| Item         | Treatment | Summary Statistics |                      |       |       |       |       |        |       |         | One-sample t-test |         |        |
|--------------|-----------|--------------------|----------------------|-------|-------|-------|-------|--------|-------|---------|-------------------|---------|--------|
|              |           | Baseline Mean      | Change from Baseline |       |       |       |       |        |       | t-value | p-value           |         |        |
|              |           |                    | N                    | Mean  | SD    | Min   | Q1    | Median | Q3    |         |                   | Max     |        |
| P<br>(mg/dL) | Week 1    | Placebo            | 3.62                 | 39    | -0.03 | 0.311 | -0.6  | -0.30  | -0.10 | 0.20    | 0.7               | -0.5661 | 0.5747 |
|              | 15 mg     | 3.69               | 37                   | -0.00 | 0.420 | -0.8  | -0.20 | -0.10  | 0.30  | 1.1     | -0.0391           | 0.9690  |        |
|              | 30 mg     | 3.55               | 39                   | 0.08  | 0.356 | -0.7  | -0.10 | 0.10   | 0.30  | 0.8     | 1.4391            | 0.1583  |        |
|              | 60 mg     | 3.55               | 37                   | 0.02  | 0.404 | -0.7  | -0.20 | 0.10   | 0.20  | 1.4     | 0.3259            | 0.7464  |        |
|              | 120 mg    | 3.73               | 47                   | 0.01  | 0.324 | -0.7  | -0.20 | 0.00   | 0.20  | 0.8     | 0.1799            | 0.8580  |        |
|              | Week 2    | Placebo            | 3.56                 | 96    | 0.00  | 0.416 | -1.8  | -0.25  | 0.00  | 0.30    | 1.1               | 0.0490  | 0.9610 |
|              | 15 mg     | 3.59               | 89                   | -0.04 | 0.485 | -1.9  | -0.20 | 0.00   | 0.20  | 0.9     | -0.7211           | 0.4728  |        |
|              | 30 mg     | 3.50               | 91                   | 0.10  | 0.454 | -0.8  | -0.20 | 0.10   | 0.30  | 1.6     | 2.1919            | 0.0310  |        |
|              | 60 mg     | 3.52               | 93                   | 0.05  | 0.425 | -1.4  | -0.20 | 0.10   | 0.30  | 1.2     | 1.1713            | 0.2445  |        |
|              | 120 mg    | 3.67               | 89                   | -0.01 | 0.385 | -0.8  | -0.30 | 0.00   | 0.20  | 1.2     | -0.2478           | 0.8049  |        |
|              | Week 3    | Placebo            | 3.59                 | 36    | 0.08  | 0.458 | -0.8  | -0.20  | 0.00  | 0.25    | 1.6               | 0.9835  | 0.3321 |
|              | 15 mg     | 3.68               | 36                   | 0.02  | 0.400 | -1.2  | -0.20 | 0.00   | 0.25  | 0.9     | 0.2502            | 0.8039  |        |
|              | 30 mg     | 3.53               | 38                   | 0.12  | 0.374 | -0.8  | -0.10 | 0.10   | 0.40  | 0.8     | 1.9515            | 0.0586  |        |
|              | 60 mg     | 3.53               | 32                   | 0.03  | 0.457 | -0.8  | -0.20 | 0.00   | 0.25  | 1.6     | 0.3870            | 0.7014  |        |
|              | 120 mg    | 3.74               | 42                   | -0.07 | 0.298 | -0.6  | -0.30 | 0.00   | 0.10  | 0.4     | -1.6051           | 0.1161  |        |
|              | Week 4    | Placebo            | 3.55                 | 95    | -0.03 | 0.422 | -2.0  | -0.30  | 0.00  | 0.30    | 0.9               | -0.7539 | 0.4528 |
|              | 15 mg     | 3.59               | 89                   | 0.02  | 0.442 | -1.1  | -0.30 | 0.00   | 0.30  | 1.2     | 0.3598            | 0.7199  |        |
|              | 30 mg     | 3.49               | 89                   | 0.07  | 0.455 | -1.2  | -0.20 | 0.10   | 0.30  | 1.3     | 1.4432            | 0.1525  |        |
|              | 60 mg     | 3.53               | 89                   | 0.00  | 0.410 | -1.1  | -0.20 | 0.00   | 0.30  | 0.9     | 0.0517            | 0.9589  |        |
|              | 120 mg    | 3.68               | 87                   | -0.02 | 0.370 | -0.8  | -0.30 | -0.10  | 0.20  | 0.9     | -0.5507           | 0.5833  |        |
| Week 6       | Placebo   | 3.61               | 40                   | -0.01 | 0.405 | -0.9  | -0.20 | -0.10  | 0.20  | 1.3     | -0.1172           | 0.9073  |        |
| 15 mg        | 3.70      | 40                 | -0.11                | 0.363 | -0.9  | -0.35 | -0.15 | 0.10   | 0.6   | -1.9592 | 0.0573            |         |        |
| 30 mg        | 3.54      | 38                 | 0.15                 | 0.475 | -0.8  | -0.20 | 0.10  | 0.40   | 1.1   | 1.9140  | 0.0634            |         |        |
| 60 mg        | 3.47      | 31                 | 0.12                 | 0.424 | -0.9  | -0.20 | 0.10  | 0.30   | 1.6   | 1.5266  | 0.1373            |         |        |
| 120 mg       | 3.73      | 47                 | 0.00                 | 0.368 | -1.0  | -0.20 | 0.00  | 0.30   | 0.7   | 0.0396  | 0.9686            |         |        |
| Week 8       | Placebo   | 3.54               | 94                   | -0.02 | 0.451 | -2.0  | -0.20 | 0.00   | 0.20  | 1.0     | -0.4113           | 0.6818  |        |
| 15 mg        | 3.60      | 86                 | -0.02                | 0.504 | -1.3  | -0.40 | 0.00  | 0.30   | 1.4   | -0.4496 | 0.6541            |         |        |
| 30 mg        | 3.49      | 89                 | 0.12                 | 0.462 | -1.3  | -0.20 | 0.10  | 0.40   | 1.5   | 2.4311  | 0.0171            |         |        |
| 60 mg        | 3.50      | 86                 | 0.01                 | 0.456 | -1.4  | -0.20 | 0.00  | 0.30   | 1.2   | 0.1181  | 0.9062            |         |        |
| 120 mg       | 3.65      | 84                 | -0.01                | 0.386 | -0.8  | -0.20 | 0.00  | 0.20   | 1.4   | -0.2545 | 0.7998            |         |        |
| Week 12      | Placebo   | 3.54               | 94                   | -0.09 | 0.422 | -1.5  | -0.30 | 0.00   | 0.10  | 1.1     | -2.1736           | 0.0323  |        |
| 15 mg        | 3.60      | 86                 | -0.07                | 0.412 | -1.2  | -0.30 | -0.10 | 0.20   | 0.8   | -1.5983 | 0.1137            |         |        |
| 30 mg        | 3.48      | 86                 | 0.10                 | 0.467 | -0.9  | -0.20 | 0.00  | 0.40   | 1.3   | 1.9620  | 0.0530            |         |        |
| 60 mg        | 3.49      | 82                 | 0.02                 | 0.432 | -0.9  | -0.30 | 0.00  | 0.30   | 1.4   | 0.3576  | 0.7215            |         |        |
| 120 mg       | 3.67      | 83                 | -0.04                | 0.347 | -0.7  | -0.30 | 0.00  | 0.20   | 0.8   | -1.1697 | 0.2455            |         |        |
| Week 16      | Placebo   | 3.54               | 94                   | -0.01 | 0.475 | -2.1  | -0.30 | 0.00   | 0.30  | 1.5     | -0.2604           | 0.7951  |        |
| 15 mg        | 3.60      | 84                 | 0.01                 | 0.420 | -1.5  | -0.20 | 0.00  | 0.20   | 1.0   | 0.2597  | 0.7957            |         |        |
| 30 mg        | 3.48      | 85                 | 0.11                 | 0.473 | -1.1  | -0.20 | 0.10  | 0.40   | 1.3   | 2.2024  | 0.0304            |         |        |
| 60 mg        | 3.48      | 80                 | 0.02                 | 0.450 | -1.7  | -0.20 | 0.05  | 0.30   | 0.8   | 0.3729  | 0.7103            |         |        |
| 120 mg       | 3.67      | 82                 | -0.03                | 0.384 | -0.9  | -0.30 | 0.00  | 0.30   | 0.9   | -0.6614 | 0.5102            |         |        |
| Week 20      | Placebo   | 3.54               | 94                   | -0.06 | 0.467 | -2.0  | -0.30 | -0.10  | 0.20  | 1.2     | -1.3253           | 0.1883  |        |
| 15 mg        | 3.60      | 84                 | -0.08                | 0.403 | -1.1  | -0.30 | 0.00  | 0.10   | 1.1   | -1.8940 | 0.0617            |         |        |
| 30 mg        | 3.48      | 85                 | 0.08                 | 0.480 | -1.5  | -0.20 | 0.10  | 0.40   | 1.2   | 1.4901  | 0.1399            |         |        |
| 60 mg        | 3.47      | 79                 | 0.07                 | 0.458 | -1.2  | -0.20 | 0.10  | 0.40   | 1.2   | 1.4251  | 0.1581            |         |        |
| 120 mg       | 3.66      | 80                 | -0.02                | 0.407 | -1.1  | -0.30 | 0.00  | 0.20   | 1.1   | -0.4395 | 0.6615            |         |        |
| Week 24      | Placebo   | 3.54               | 93                   | -0.06 | 0.499 | -2.4  | -0.20 | 0.00   | 0.20  | 1.2     | -1.1440           | 0.2556  |        |
| 15 mg        | 3.60      | 84                 | -0.02                | 0.425 | -1.1  | -0.25 | 0.00  | 0.25   | 1.3   | -0.3855 | 0.7009            |         |        |
| 30 mg        | 3.47      | 84                 | 0.08                 | 0.463 | -1.4  | -0.20 | 0.10  | 0.35   | 1.6   | 1.5567  | 0.1233            |         |        |
| 60 mg        | 3.47      | 78                 | 0.04                 | 0.405 | -1.2  | -0.30 | 0.10  | 0.30   | 0.9   | 0.8668  | 0.3888            |         |        |
| 120 mg       | 3.65      | 79                 | 0.02                 | 0.465 | -1.1  | -0.30 | 0.00  | 0.30   | 1.3   | 0.3386  | 0.7358            |         |        |
| Week 28      | Placebo   | 3.54               | 93                   | -0.11 | 0.448 | -2.1  | -0.40 | -0.10  | 0.10  | 1.1     | -2.3611           | 0.0203  |        |
| 15 mg        | 3.60      | 83                 | -0.08                | 0.464 | -1.2  | -0.40 | -0.10 | 0.20   | 1.2   | -1.5852 | 0.1168            |         |        |
| 30 mg        | 3.47      | 84                 | -0.01                | 0.415 | -1.0  | -0.30 | 0.00  | 0.30   | 1.2   | -0.1579 | 0.8750            |         |        |
| 60 mg        | 3.47      | 77                 | 0.12                 | 0.465 | -1.1  | -0.20 | 0.20  | 0.40   | 1.9   | 2.3038  | 0.0240            |         |        |
| 120 mg       | 3.66      | 78                 | -0.03                | 0.478 | -1.1  | -0.30 | 0.00  | 0.30   | 1.0   | -0.5447 | 0.5875            |         |        |
| Follow-up    | Placebo   | 3.55               | 97                   | -0.11 | 0.440 | -2.1  | -0.40 | -0.10  | 0.10  | 1.1     | -2.4928           | 0.0144  |        |
| 15 mg        | 3.59      | 86                 | -0.07                | 0.461 | -1.2  | -0.40 | -0.10 | 0.20   | 1.2   | -1.4735 | 0.1443            |         |        |
| 30 mg        | 3.49      | 92                 | -0.01                | 0.403 | -1.0  | -0.30 | 0.00  | 0.30   | 1.2   | -0.1813 | 0.8565            |         |        |
| 60 mg        | 3.54      | 87                 | 0.11                 | 0.486 | -1.1  | -0.20 | 0.20  | 0.40   | 1.9   | 2.0978  | 0.0389            |         |        |
| 120 mg       | 3.68      | 88                 | -0.03                | 0.482 | -1.1  | -0.35 | 0.00  | 0.30   | 1.0   | -0.4869 | 0.6276            |         |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (19/24)

| Item                 | Treatment | Summary Statistics |                      |       |       |        |        |       |         |         | One-sample t-test |         |         |        |
|----------------------|-----------|--------------------|----------------------|-------|-------|--------|--------|-------|---------|---------|-------------------|---------|---------|--------|
|                      |           | Baseline Mean      | Change from Baseline |       |       |        |        | Max   | t-value | p-value |                   |         |         |        |
|                      |           |                    | N                    | Mean  | SD    | Min    | Q1     |       |         |         | Median            | Q3      |         |        |
| Vitamin A<br>(ng/mL) | Week 1    | Placebo            | 556.4                | 39    | -3.7  | 104.48 | -259   | -51.0 | -15.0   | 39.0    | 365               | -0.2238 | 0.8241  |        |
|                      |           | 15 mg              | 587.0                | 37    | -20.8 | 83.72  | -269   | -66.0 | -20.0   | 34.0    | 113               | -1.5141 | 0.1387  |        |
|                      |           | 30 mg              | 575.9                | 39    | 7.3   | 84.27  | -135   | -41.0 | 2.0     | 55.0    | 232               | 0.5396  | 0.5926  |        |
|                      |           | 60 mg              | 557.0                | 37    | -17.7 | 79.78  | -229   | -63.0 | -3.0    | 25.0    | 131               | -1.3518 | 0.1849  |        |
|                      |           | 120 mg             | 555.2                | 47    | -14.0 | 71.90  | -147   | -67.0 | -10.0   | 28.0    | 142               | -1.3349 | 0.1885  |        |
|                      |           | Week 2             | Placebo              | 572.7 | 97    | -7.9   | 80.02  | -256  | -60.0   | -17.0   | 46.0              | 208     | -0.9707 | 0.3342 |
|                      |           |                    | 15 mg                | 593.5 | 89    | -21.5  | 112.10 | -377  | -67.0   | -29.0   | 38.0              | 299     | -1.8070 | 0.0742 |
|                      |           |                    | 30 mg                | 606.8 | 91    | -24.4  | 110.27 | -455  | -80.0   | -17.0   | 32.0              | 319     | -2.1123 | 0.0374 |
|                      |           |                    | 60 mg                | 563.8 | 93    | -29.4  | 100.07 | -409  | -75.0   | -15.0   | 18.0              | 301     | -2.8310 | 0.0057 |
|                      |           |                    | 120 mg               | 557.6 | 89    | -24.8  | 78.86  | -253  | -70.0   | -8.0    | 20.0              | 157     | -2.9613 | 0.0039 |
|                      |           | Week 3             | Placebo              | 569.1 | 36    | -18.1  | 84.54  | -160  | -63.5   | -23.5   | 26.0              | 216     | -1.2834 | 0.2078 |
|                      |           |                    | 15 mg                | 589.8 | 36    | -20.6  | 76.33  | -284  | -53.5   | -20.0   | 34.0              | 167     | -1.6158 | 0.1151 |
|                      |           |                    | 30 mg                | 585.3 | 38    | -5.8   | 81.76  | -150  | -53.0   | -15.5   | 37.0              | 203     | -0.4385 | 0.6636 |
|                      |           |                    | 60 mg                | 561.4 | 32    | -30.1  | 84.49  | -254  | -91.0   | -30.0   | 24.0              | 218     | -2.0170 | 0.0524 |
|                      |           |                    | 120 mg               | 562.1 | 42    | -18.2  | 101.94 | -304  | -58.0   | -32.5   | 15.0              | 277     | -1.1579 | 0.2536 |
|                      |           | Week 4             | Placebo              | 574.8 | 96    | -2.9   | 90.26  | -235  | -55.5   | -8.0    | 37.0              | 306     | -0.3132 | 0.7548 |
|                      |           |                    | 15 mg                | 593.5 | 89    | -9.6   | 122.21 | -447  | -63.0   | -17.0   | 74.0              | 328     | -0.7424 | 0.4598 |
|                      |           |                    | 30 mg                | 612.8 | 89    | -24.7  | 108.75 | -334  | -72.0   | -38.0   | 40.0              | 293     | -2.1464 | 0.0346 |
|                      |           |                    | 60 mg                | 565.1 | 89    | -24.9  | 90.33  | -403  | -75.0   | -16.0   | 23.0              | 269     | -2.6003 | 0.0109 |
|                      |           |                    | 120 mg               | 555.7 | 87    | -5.3   | 98.83  | -195  | -67.0   | -20.0   | 49.0              | 283     | -0.4990 | 0.6190 |
|                      |           | Week 6             | Placebo              | 554.6 | 40    | 15.7   | 119.74 | -150  | -50.5   | -0.5    | 62.5              | 529     | 0.8293  | 0.4120 |
|                      |           |                    | 15 mg                | 591.8 | 40    | -34.2  | 89.01  | -351  | -58.0   | -22.0   | 5.5               | 181     | -2.4283 | 0.0199 |
|                      |           |                    | 30 mg                | 583.9 | 38    | -8.5   | 103.27 | -222  | -68.0   | -10.5   | 55.0              | 210     | -0.5090 | 0.6138 |
|                      |           |                    | 60 mg                | 547.4 | 31    | -37.4  | 109.07 | -409  | -94.0   | -45.0   | 27.0              | 231     | -1.9085 | 0.0659 |
|                      |           |                    | 120 mg               | 543.6 | 47    | -14.5  | 82.45  | -170  | -61.0   | -20.0   | 30.0              | 263     | -1.2048 | 0.2344 |
|                      |           | Week 8             | Placebo              | 573.6 | 95    | -6.4   | 93.92  | -297  | -61.0   | -13.0   | 58.0              | 241     | -0.6675 | 0.5061 |
|                      |           |                    | 15 mg                | 593.0 | 86    | -7.7   | 115.89 | -405  | -71.0   | -5.0    | 65.0              | 243     | -0.6178 | 0.5383 |
|                      |           |                    | 30 mg                | 612.8 | 89    | -14.3  | 114.40 | -444  | -63.0   | -16.0   | 45.0              | 316     | -1.1777 | 0.2421 |
|                      |           |                    | 60 mg                | 557.8 | 86    | -24.4  | 117.62 | -431  | -93.0   | -20.5   | 52.0              | 249     | -1.9243 | 0.0577 |
|                      |           |                    | 120 mg               | 559.5 | 84    | -22.9  | 102.79 | -325  | -62.0   | -21.0   | 27.5              | 374     | -2.0402 | 0.0445 |
|                      |           | Week 12            | Placebo              | 573.6 | 95    | -11.5  | 86.64  | -277  | -69.0   | -13.0   | 39.0              | 236     | -1.2932 | 0.1991 |
|                      |           |                    | 15 mg                | 593.0 | 86    | -19.6  | 112.26 | -357  | -78.0   | -12.5   | 38.0              | 240     | -1.6166 | 0.1097 |
|                      |           |                    | 30 mg                | 617.4 | 86    | -24.8  | 114.63 | -415  | -78.0   | -16.5   | 28.0              | 362     | -2.0084 | 0.0478 |
|                      |           |                    | 60 mg                | 566.2 | 82    | -32.9  | 119.93 | -403  | -93.0   | -21.0   | 27.0              | 284     | -2.4805 | 0.0152 |
|                      |           |                    | 120 mg               | 551.2 | 83    | -15.6  | 101.93 | -305  | -71.0   | -21.0   | 32.0              | 295     | -1.3902 | 0.1682 |
|                      |           | Week 16            | Placebo              | 573.6 | 95    | -11.2  | 114.07 | -340  | -74.0   | -4.0    | 51.0              | 353     | -0.9552 | 0.3419 |
|                      |           |                    | 15 mg                | 590.5 | 84    | -26.7  | 107.32 | -335  | -82.5   | -17.0   | 47.0              | 213     | -2.2773 | 0.0253 |
|                      |           |                    | 30 mg                | 616.6 | 85    | -29.2  | 131.54 | -571  | -76.0   | -15.0   | 34.0              | 321     | -2.0458 | 0.0439 |
|                      |           |                    | 60 mg                | 556.8 | 80    | -28.4  | 118.60 | -385  | -77.0   | -19.0   | 29.5              | 592     | -2.1379 | 0.0356 |
|                      |           |                    | 120 mg               | 554.3 | 82    | -16.1  | 92.88  | -296  | -70.0   | -25.0   | 29.0              | 226     | -1.5729 | 0.1196 |
|                      |           | Week 20            | Placebo              | 573.6 | 95    | -9.2   | 101.89 | -278  | -70.0   | -10.0   | 42.0              | 312     | -0.8781 | 0.3821 |
|                      |           |                    | 15 mg                | 590.5 | 84    | -32.3  | 96.53  | -289  | -92.5   | -31.0   | 24.0              | 207     | -3.0665 | 0.0029 |
|                      |           |                    | 30 mg                | 616.6 | 85    | -17.5  | 126.46 | -501  | -85.0   | -12.0   | 45.0              | 349     | -1.2771 | 0.2051 |
|                      |           |                    | 60 mg                | 557.6 | 79    | -24.4  | 100.22 | -408  | -64.0   | -36.0   | 38.0              | 166     | -2.1599 | 0.0338 |
|                      |           |                    | 120 mg               | 554.1 | 80    | -4.7   | 103.78 | -210  | -69.0   | -2.0    | 49.0              | 384     | -0.4029 | 0.6881 |
|                      |           | Week 24            | Placebo              | 576.5 | 94    | -28.7  | 98.72  | -294  | -94.0   | -19.0   | 37.0              | 238     | -2.8168 | 0.0059 |
|                      |           |                    | 15 mg                | 590.5 | 84    | -17.9  | 110.47 | -297  | -78.5   | -5.0    | 48.0              | 222     | -1.4844 | 0.1415 |
|                      |           |                    | 30 mg                | 617.4 | 84    | -10.0  | 113.42 | -422  | -61.0   | -14.0   | 48.0              | 303     | -0.8109 | 0.4197 |
|                      |           | 60 mg              | 559.8                | 78    | -31.4 | 101.72 | -432   | -80.0 | -29.0   | 29.0    | 300               | -2.7293 | 0.0079  |        |
|                      |           | 120 mg             | 555.3                | 79    | -30.0 | 92.36  | -288   | -76.0 | -26.0   | 33.0    | 178               | -2.8894 | 0.0050  |        |
|                      | Week 28   | Placebo            | 576.5                | 94    | -12.7 | 103.00 | -303   | -68.0 | -6.0    | 44.0    | 221               | -1.1946 | 0.2353  |        |
|                      |           | 15 mg              | 591.3                | 83    | -29.1 | 108.32 | -350   | -82.0 | -24.0   | 38.0    | 231               | -2.4513 | 0.0164  |        |
|                      |           | 30 mg              | 617.4                | 84    | -30.1 | 127.72 | -527   | -85.5 | -18.0   | 31.5    | 326               | -2.1613 | 0.0336  |        |
|                      |           | 60 mg              | 557.8                | 77    | -16.4 | 120.51 | -409   | -70.0 | -16.0   | 47.0    | 328               | -1.1962 | 0.2353  |        |
|                      |           | 120 mg             | 556.2                | 78    | -22.6 | 101.32 | -366   | -84.0 | -19.5   | 43.0    | 345               | -1.9668 | 0.0528  |        |
|                      | Follow-up | Placebo            | 573.0                | 98    | -14.4 | 103.01 | -303   | -68.0 | -9.5    | 44.0    | 221               | -1.3846 | 0.1693  |        |
|                      |           | 15 mg              | 595.0                | 86    | -24.8 | 110.03 | -350   | -78.0 | -22.5   | 41.0    | 231               | -2.0923 | 0.0394  |        |
|                      |           | 30 mg              | 602.9                | 92    | -28.0 | 123.43 | -527   | -82.5 | -15.5   | 31.5    | 326               | -2.1767 | 0.0321  |        |
|                      |           | 60 mg              | 566.0                | 87    | -19.8 | 117.20 | -409   | -70.0 | -17.0   | 47.0    | 328               | -1.5761 | 0.1187  |        |
|                      |           | 120 mg             | 558.4                | 88    | -20.2 | 101.77 | -366   | -75.5 | -18.5   | 49.5    | 345               | -1.8634 | 0.0658  |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (20/24)

| Item                 | Treatment | Summary Statistics |                      |      |       |       |       |        |      |         | One-sample t-test |         |        |
|----------------------|-----------|--------------------|----------------------|------|-------|-------|-------|--------|------|---------|-------------------|---------|--------|
|                      |           | Baseline Mean      | Change from Baseline |      |       |       |       |        |      | t-value | p-value           |         |        |
|                      |           |                    | N                    | Mean | SD    | Min   | Q1    | Median | Q3   |         |                   | Max     |        |
| Vitamin D<br>(pg/mL) | Week 1    | Placebo            | 55.1                 | 39   | -6.1  | 19.08 | -59   | -15.0  | -4.0 | 6.0     | 37                | -1.9803 | 0.0549 |
|                      |           | 15 mg              | 52.6                 | 37   | -0.7  | 22.98 | -37   | -20.0  | -2.0 | 13.0    | 71                | -0.1788 | 0.8591 |
|                      |           | 30 mg              | 51.6                 | 39   | -1.3  | 16.17 | -33   | -12.0  | -3.0 | 9.0     | 43                | -0.4951 | 0.6234 |
|                      |           | 60 mg              | 54.7                 | 37   | -0.1  | 16.82 | -40   | -9.0   | -1.0 | 9.0     | 40                | -0.0196 | 0.9845 |
|                      |           | 120 mg             | 52.1                 | 47   | -0.6  | 18.02 | -36   | -15.0  | 3.0  | 13.0    | 35                | -0.2186 | 0.8279 |
|                      | Week 2    | Placebo            | 52.6                 | 97   | -3.6  | 16.96 | -56   | -14.0  | -4.0 | 8.0     | 49                | -2.0840 | 0.0398 |
|                      |           | 15 mg              | 51.4                 | 89   | 0.6   | 20.45 | -78   | -9.0   | 0.0  | 8.0     | 55                | 0.2643  | 0.7922 |
|                      |           | 30 mg              | 51.3                 | 91   | 1.3   | 19.10 | -42   | -12.0  | -1.0 | 16.0    | 68                | 0.6642  | 0.5083 |
|                      |           | 60 mg              | 53.5                 | 93   | -2.4  | 19.48 | -50   | -14.0  | -5.0 | 9.0     | 61                | -1.2032 | 0.2320 |
|                      |           | 120 mg             | 51.3                 | 89   | -0.8  | 17.35 | -38   | -14.0  | -2.0 | 12.0    | 45                | -0.4459 | 0.6568 |
|                      | Week 3    | Placebo            | 55.3                 | 36   | -4.2  | 21.28 | -61   | -11.5  | -5.0 | 3.5     | 44                | -1.1905 | 0.2419 |
|                      |           | 15 mg              | 50.9                 | 36   | 0.7   | 14.59 | -50   | -9.0   | 0.0  | 13.0    | 24                | 0.2970  | 0.7683 |
|                      |           | 30 mg              | 51.2                 | 38   | 0.8   | 15.06 | -26   | -10.0  | -3.0 | 11.0    | 29                | 0.3446  | 0.7324 |
|                      |           | 60 mg              | 52.9                 | 32   | 1.0   | 16.62 | -33   | -8.0   | 1.5  | 11.0    | 38                | 0.3510  | 0.7280 |
|                      |           | 120 mg             | 52.3                 | 42   | -2.5  | 15.93 | -37   | -10.0  | -2.5 | 6.0     | 44                | -1.0365 | 0.3060 |
|                      | Week 4    | Placebo            | 52.9                 | 96   | -3.3  | 15.77 | -54   | -13.5  | -5.5 | 8.0     | 35                | -2.0387 | 0.0443 |
|                      |           | 15 mg              | 51.4                 | 89   | -3.2  | 17.26 | -56   | -13.0  | -1.0 | 6.0     | 39                | -1.7261 | 0.0878 |
|                      |           | 30 mg              | 51.7                 | 89   | 0.3   | 19.31 | -51   | -11.0  | -2.0 | 11.0    | 87                | 0.1482  | 0.8825 |
|                      |           | 60 mg              | 53.1                 | 89   | -4.6  | 15.73 | -58   | -14.0  | -4.0 | 5.0     | 29                | -2.7428 | 0.0074 |
|                      |           | 120 mg             | 51.6                 | 87   | -0.7  | 16.13 | -60   | -12.0  | 0.0  | 8.0     | 36                | -0.4122 | 0.6812 |
|                      | Week 6    | Placebo            | 55.1                 | 40   | -3.1  | 14.82 | -46   | -8.0   | -1.5 | 6.5     | 24                | -1.3021 | 0.2005 |
|                      |           | 15 mg              | 51.8                 | 40   | 1.6   | 23.26 | -61   | -9.0   | -1.0 | 14.5    | 59                | 0.4283  | 0.6708 |
|                      |           | 30 mg              | 54.7                 | 38   | -0.2  | 17.53 | -45   | -11.0  | 0.5  | 13.0    | 30                | -0.0740 | 0.9414 |
|                      |           | 60 mg              | 54.0                 | 31   | -0.8  | 16.78 | -39   | -13.0  | 2.0  | 9.0     | 27                | -0.2675 | 0.7909 |
|                      |           | 120 mg             | 49.8                 | 47   | 2.5   | 19.41 | -40   | -6.0   | 3.0  | 17.0    | 44                | 0.8718  | 0.3878 |
|                      | Week 8    | Placebo            | 53.0                 | 95   | -4.6  | 16.66 | -55   | -14.0  | -4.0 | 5.0     | 35                | -2.6975 | 0.0083 |
|                      |           | 15 mg              | 51.5                 | 86   | -2.3  | 17.86 | -48   | -13.0  | -2.0 | 6.0     | 60                | -1.2076 | 0.2305 |
|                      |           | 30 mg              | 51.7                 | 89   | 0.7   | 18.39 | -31   | -12.0  | -1.0 | 10.0    | 75                | 0.3343  | 0.7389 |
|                      |           | 60 mg              | 52.9                 | 86   | -5.6  | 15.30 | -43   | -14.0  | -5.5 | 4.0     | 35                | -3.3960 | 0.0010 |
|                      |           | 120 mg             | 50.8                 | 84   | -0.0  | 18.16 | -43   | -11.0  | 1.0  | 10.0    | 57                | -0.0060 | 0.9952 |
|                      | Week 12   | Placebo            | 53.0                 | 95   | -1.9  | 16.78 | -39   | -13.0  | -2.0 | 9.0     | 42                | -1.1006 | 0.2739 |
|                      |           | 15 mg              | 51.5                 | 86   | -3.1  | 17.47 | -74   | -12.0  | -4.0 | 6.0     | 36                | -1.6608 | 0.1004 |
|                      |           | 30 mg              | 51.1                 | 86   | -1.5  | 18.41 | -55   | -13.0  | -1.0 | 9.0     | 51                | -0.7379 | 0.4626 |
|                      |           | 60 mg              | 52.6                 | 82   | -4.3  | 17.07 | -51   | -17.0  | -3.0 | 7.0     | 34                | -2.2776 | 0.0254 |
|                      |           | 120 mg             | 50.7                 | 83   | -0.4  | 18.19 | -35   | -12.0  | -1.0 | 9.0     | 46                | -0.1810 | 0.8568 |
|                      | Week 16   | Placebo            | 53.0                 | 95   | -1.5  | 16.35 | -41   | -13.0  | 0.0  | 11.0    | 34                | -0.9162 | 0.3619 |
|                      |           | 15 mg              | 51.8                 | 84   | -2.1  | 19.05 | -71   | -13.5  | -2.5 | 12.0    | 44                | -1.0196 | 0.3109 |
|                      |           | 30 mg              | 51.6                 | 85   | -1.1  | 21.01 | -48   | -13.0  | -5.0 | 8.0     | 65                | -0.4749 | 0.6361 |
|                      |           | 60 mg              | 53.6                 | 80   | -4.5  | 15.01 | -49   | -15.0  | -3.0 | 4.5     | 28                | -2.6735 | 0.0091 |
|                      |           | 120 mg             | 51.4                 | 82   | -6.3  | 18.06 | -49   | -17.0  | -6.0 | 7.0     | 39                | -3.1488 | 0.0023 |
|                      | Week 20   | Placebo            | 53.0                 | 95   | 0.1   | 15.03 | -35   | -10.0  | -1.0 | 11.0    | 38                | 0.0956  | 0.9241 |
|                      |           | 15 mg              | 51.8                 | 84   | -2.5  | 19.19 | -43   | -13.5  | -2.0 | 6.0     | 47                | -1.1712 | 0.2449 |
|                      |           | 30 mg              | 51.6                 | 85   | -1.1  | 21.20 | -48   | -15.0  | -3.0 | 11.0    | 54                | -0.4655 | 0.6428 |
|                      |           | 60 mg              | 53.1                 | 79   | -4.9  | 17.93 | -56   | -14.0  | -3.0 | 5.0     | 37                | -2.4158 | 0.0180 |
|                      |           | 120 mg             | 51.9                 | 80   | -3.2  | 18.91 | -42   | -17.0  | -5.0 | 11.0    | 61                | -1.5019 | 0.1371 |
|                      | Week 24   | Placebo            | 53.2                 | 94   | 0.6   | 17.23 | -71   | -9.0   | 1.0  | 11.0    | 48                | 0.3112  | 0.7563 |
|                      |           | 15 mg              | 51.8                 | 84   | -3.4  | 18.90 | -73   | -13.0  | -3.0 | 10.0    | 42                | -1.6568 | 0.1013 |
|                      |           | 30 mg              | 51.7                 | 84   | 2.0   | 20.04 | -57   | -11.0  | 2.0  | 16.5    | 46                | 0.8984  | 0.3716 |
| 60 mg                |           | 53.1               | 78                   | -4.4 | 17.83 | -53   | -15.0 | -3.0   | 8.0  | 36      | -2.1596           | 0.0339  |        |
| 120 mg               |           | 52.2               | 79                   | -0.8 | 18.80 | -41   | -14.0 | -3.0   | 10.0 | 52      | -0.4009           | 0.6896  |        |
| Week 28              | Placebo   | 53.2               | 94                   | -0.4 | 17.57 | -53   | -13.0 | -1.0   | 12.0 | 52      | -0.2172           | 0.8285  |        |
|                      | 15 mg     | 51.9               | 83                   | -2.6 | 19.48 | -85   | -10.0 | -2.0   | 6.0  | 44      | -1.1947           | 0.2356  |        |
|                      | 30 mg     | 51.7               | 84                   | 1.3  | 16.39 | -32   | -10.5 | 1.0    | 12.5 | 53      | 0.7389            | 0.4621  |        |
|                      | 60 mg     | 52.7               | 77                   | -5.8 | 17.29 | -59   | -17.0 | -5.0   | 6.0  | 42      | -2.9197           | 0.0046  |        |
|                      | 120 mg    | 52.3               | 78                   | -0.8 | 20.85 | -53   | -16.0 | -1.0   | 15.0 | 57      | -0.3530           | 0.7250  |        |
| Follow-up            | Placebo   | 52.5               | 98                   | -0.2 | 17.35 | -53   | -11.0 | -1.0   | 12.0 | 52      | -0.1339           | 0.8938  |        |
|                      | 15 mg     | 51.7               | 86                   | -2.8 | 21.15 | -85   | -11.0 | -1.5   | 6.0  | 49      | -1.2186           | 0.2264  |        |
|                      | 30 mg     | 51.5               | 92                   | 0.7  | 16.01 | -32   | -11.5 | 0.5    | 12.0 | 53      | 0.4232            | 0.6731  |        |
|                      | 60 mg     | 52.4               | 87                   | -4.3 | 18.95 | -59   | -17.0 | -5.0   | 6.0  | 71      | -2.1272           | 0.0363  |        |
|                      | 120 mg    | 51.3               | 88                   | -1.4 | 20.37 | -53   | -16.0 | -1.0   | 13.0 | 57      | -0.6436           | 0.5215  |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (21/24)

| Item                 | Treatment | Summary Statistics |                      |        |        |        |        |        |        |        |         | One-sample t-test |         |
|----------------------|-----------|--------------------|----------------------|--------|--------|--------|--------|--------|--------|--------|---------|-------------------|---------|
|                      |           | Baseline Mean      | Change from Baseline |        |        |        |        |        |        |        | t-value | p-value           |         |
|                      |           |                    | N                    | Mean   | SD     | Min    | Q1     | Median | Q3     | Max    |         |                   |         |
| Vitamin E<br>(mg/dL) | Week 1    | Placebo            | 1.601                | 39     | 0.033  | 0.2499 | -0.41  | -0.150 | -0.020 | 0.200  | 0.76    | 0.8265            | 0.4137  |
|                      |           | 15 mg              | 1.758                | 37     | -0.178 | 0.4281 | -1.80  | -0.330 | -0.190 | 0.060  | 0.81    | -2.5230           | 0.0162  |
|                      |           | 30 mg              | 1.586                | 39     | -0.107 | 0.3492 | -0.73  | -0.320 | -0.120 | 0.040  | 0.69    | -1.9125           | 0.0634  |
|                      |           | 60 mg              | 1.505                | 37     | -0.122 | 0.2533 | -0.60  | -0.260 | -0.100 | -0.020 | 0.74    | -2.9333           | 0.0058  |
|                      |           | 120 mg             | 1.569                | 47     | -0.178 | 0.2323 | -0.59  | -0.310 | -0.210 | -0.020 | 0.49    | -5.2501           | <0.0001 |
|                      | Week 2    | Placebo            | 1.544                | 97     | -0.007 | 0.2741 | -0.53  | -0.160 | -0.020 | 0.110  | 1.53    | -0.2445           | 0.8074  |
|                      |           | 15 mg              | 1.581                | 89     | -0.173 | 0.3248 | -1.78  | -0.280 | -0.120 | 0.030  | 0.38    | -5.0162           | <0.0001 |
|                      |           | 30 mg              | 1.606                | 91     | -0.209 | 0.2988 | -1.03  | -0.370 | -0.170 | -0.040 | 0.73    | -6.6826           | <0.0001 |
|                      |           | 60 mg              | 1.560                | 93     | -0.193 | 0.3286 | -1.32  | -0.350 | -0.190 | -0.040 | 0.75    | -5.6715           | <0.0001 |
|                      |           | 120 mg             | 1.592                | 89     | -0.234 | 0.2632 | -0.82  | -0.390 | -0.250 | -0.070 | 0.59    | -8.3926           | <0.0001 |
|                      | Week 3    | Placebo            | 1.617                | 36     | 0.060  | 0.2480 | -0.39  | -0.125 | 0.035  | 0.160  | 0.91    | 1.4517            | 0.1555  |
|                      |           | 15 mg              | 1.740                | 36     | -0.231 | 0.4471 | -1.42  | -0.450 | -0.170 | 0.015  | 0.87    | -3.0942           | 0.0039  |
|                      |           | 30 mg              | 1.592                | 38     | -0.182 | 0.3850 | -0.96  | -0.520 | -0.200 | 0.080  | 0.84    | -2.9113           | 0.0061  |
|                      |           | 60 mg              | 1.503                | 32     | -0.208 | 0.2883 | -0.88  | -0.320 | -0.205 | -0.095 | 0.62    | -4.0719           | 0.0003  |
|                      |           | 120 mg             | 1.602                | 42     | -0.268 | 0.3149 | -1.03  | -0.430 | -0.255 | -0.130 | 0.62    | -5.5222           | <0.0001 |
|                      | Week 4    | Placebo            | 1.551                | 96     | 0.020  | 0.2824 | -0.92  | -0.155 | 0.010  | 0.165  | 0.76    | 0.7084            | 0.4804  |
|                      |           | 15 mg              | 1.581                | 89     | -0.172 | 0.3129 | -1.06  | -0.340 | -0.130 | 0.040  | 0.62    | -5.1805           | <0.0001 |
|                      |           | 30 mg              | 1.594                | 89     | -0.223 | 0.2772 | -0.97  | -0.380 | -0.200 | -0.050 | 0.42    | -7.5835           | <0.0001 |
|                      |           | 60 mg              | 1.537                | 89     | -0.183 | 0.2951 | -1.01  | -0.360 | -0.150 | 0.010  | 0.66    | -5.8618           | <0.0001 |
|                      |           | 120 mg             | 1.595                | 87     | -0.227 | 0.2937 | -0.86  | -0.410 | -0.220 | -0.060 | 0.65    | -7.1953           | <0.0001 |
| Week 6               | Placebo   | 1.590              | 40                   | 0.012  | 0.2667 | -0.51  | -0.155 | -0.020 | 0.130  | 0.72   | 0.2905  | 0.7729            |         |
|                      | 15 mg     | 1.670              | 40                   | -0.174 | 0.3002 | -1.07  | -0.290 | -0.120 | -0.020 | 0.63   | -3.6707 | 0.0007            |         |
|                      | 30 mg     | 1.543              | 38                   | -0.244 | 0.3468 | -0.90  | -0.400 | -0.230 | -0.070 | 0.58   | -4.3363 | 0.0001            |         |
|                      | 60 mg     | 1.519              | 31                   | -0.244 | 0.3041 | -0.88  | -0.490 | -0.140 | 0.000  | 0.33   | -4.4655 | 0.0001            |         |
|                      | 120 mg    | 1.553              | 47                   | -0.263 | 0.2966 | -0.87  | -0.530 | -0.270 | -0.020 | 0.40   | -6.0887 | <0.0001           |         |
| Week 8               | Placebo   | 1.557              | 95                   | -0.011 | 0.2410 | -0.56  | -0.180 | -0.040 | 0.120  | 0.66   | -0.4469 | 0.6560            |         |
|                      | 15 mg     | 1.576              | 86                   | -0.171 | 0.3881 | -1.51  | -0.370 | -0.150 | 0.040  | 1.09   | -4.0816 | 0.0001            |         |
|                      | 30 mg     | 1.594              | 89                   | -0.209 | 0.3830 | -1.11  | -0.440 | -0.200 | -0.010 | 1.33   | -5.1481 | <0.0001           |         |
|                      | 60 mg     | 1.512              | 86                   | -0.206 | 0.3067 | -0.99  | -0.380 | -0.140 | -0.030 | 0.68   | -6.2190 | <0.0001           |         |
|                      | 120 mg    | 1.596              | 84                   | -0.258 | 0.2818 | -0.90  | -0.430 | -0.255 | -0.085 | 0.75   | -8.3983 | <0.0001           |         |
| Week 12              | Placebo   | 1.557              | 95                   | -0.025 | 0.3086 | -1.18  | -0.210 | -0.020 | 0.120  | 1.01   | -0.7780 | 0.4385            |         |
|                      | 15 mg     | 1.576              | 86                   | -0.195 | 0.3724 | -1.39  | -0.340 | -0.180 | 0.000  | 1.58   | -4.8623 | <0.0001           |         |
|                      | 30 mg     | 1.594              | 86                   | -0.219 | 0.3126 | -0.95  | -0.390 | -0.190 | -0.040 | 1.05   | -6.5053 | <0.0001           |         |
|                      | 60 mg     | 1.486              | 82                   | -0.263 | 0.2745 | -1.00  | -0.430 | -0.235 | -0.050 | 0.17   | -8.6905 | <0.0001           |         |
|                      | 120 mg    | 1.591              | 83                   | -0.253 | 0.3287 | -1.26  | -0.490 | -0.240 | -0.050 | 0.90   | -7.0224 | <0.0001           |         |
| Week 16              | Placebo   | 1.557              | 95                   | -0.065 | 0.2624 | -0.98  | -0.230 | -0.070 | 0.130  | 0.58   | -2.4012 | 0.0183            |         |
|                      | 15 mg     | 1.580              | 84                   | -0.180 | 0.3679 | -1.97  | -0.330 | -0.150 | 0.040  | 0.85   | -4.4777 | <0.0001           |         |
|                      | 30 mg     | 1.576              | 85                   | -0.210 | 0.3833 | -1.38  | -0.350 | -0.190 | -0.040 | 1.59   | -5.0452 | <0.0001           |         |
|                      | 60 mg     | 1.470              | 80                   | -0.220 | 0.2878 | -1.00  | -0.430 | -0.155 | -0.030 | 0.42   | -6.8420 | <0.0001           |         |
|                      | 120 mg    | 1.602              | 82                   | -0.268 | 0.3232 | -1.02  | -0.470 | -0.230 | -0.090 | 1.27   | -7.4959 | <0.0001           |         |
| Week 20              | Placebo   | 1.557              | 95                   | -0.053 | 0.3214 | -1.37  | -0.250 | -0.040 | 0.120  | 0.83   | -1.6152 | 0.1096            |         |
|                      | 15 mg     | 1.580              | 84                   | -0.208 | 0.3572 | -1.56  | -0.360 | -0.185 | 0.000  | 0.71   | -5.3430 | <0.0001           |         |
|                      | 30 mg     | 1.576              | 85                   | -0.194 | 0.3826 | -1.10  | -0.360 | -0.210 | 0.000  | 1.62   | -4.6715 | <0.0001           |         |
|                      | 60 mg     | 1.468              | 79                   | -0.252 | 0.3255 | -1.20  | -0.470 | -0.200 | -0.070 | 0.74   | -6.8686 | <0.0001           |         |
|                      | 120 mg    | 1.601              | 80                   | -0.255 | 0.2883 | -0.98  | -0.455 | -0.260 | -0.085 | 0.64   | -7.9140 | <0.0001           |         |
| Week 24              | Placebo   | 1.564              | 94                   | -0.125 | 0.2894 | -1.22  | -0.260 | -0.110 | 0.030  | 0.75   | -4.1952 | <0.0001           |         |
|                      | 15 mg     | 1.580              | 84                   | -0.163 | 0.3618 | -1.62  | -0.325 | -0.135 | 0.025  | 0.75   | -4.1370 | <0.0001           |         |
|                      | 30 mg     | 1.578              | 84                   | -0.223 | 0.3072 | -1.03  | -0.390 | -0.210 | -0.065 | 0.99   | -6.6386 | <0.0001           |         |
|                      | 60 mg     | 1.459              | 78                   | -0.238 | 0.3059 | -1.20  | -0.430 | -0.200 | -0.050 | 0.40   | -6.8777 | <0.0001           |         |
|                      | 120 mg    | 1.600              | 79                   | -0.286 | 0.3281 | -1.22  | -0.510 | -0.300 | -0.090 | 1.01   | -7.7368 | <0.0001           |         |
| Week 28              | Placebo   | 1.564              | 94                   | -0.102 | 0.2869 | -0.81  | -0.280 | -0.100 | 0.080  | 1.07   | -3.4617 | 0.0008            |         |
|                      | 15 mg     | 1.581              | 83                   | -0.116 | 0.2839 | -0.74  | -0.300 | -0.120 | 0.080  | 0.47   | -3.7314 | 0.0003            |         |
|                      | 30 mg     | 1.578              | 84                   | -0.131 | 0.3734 | -1.06  | -0.380 | -0.155 | 0.080  | 0.81   | -3.2255 | 0.0018            |         |
|                      | 60 mg     | 1.461              | 77                   | -0.062 | 0.3310 | -0.94  | -0.270 | 0.010  | 0.110  | 0.98   | -1.6560 | 0.1018            |         |
|                      | 120 mg    | 1.599              | 78                   | -0.140 | 0.3249 | -0.93  | -0.370 | -0.120 | 0.110  | 0.80   | -3.8128 | 0.0003            |         |
| Follow-up            | Placebo   | 1.545              | 98                   | -0.089 | 0.3035 | -0.81  | -0.280 | -0.100 | 0.080  | 1.07   | -2.8955 | 0.0047            |         |
|                      | 15 mg     | 1.587              | 86                   | -0.117 | 0.2838 | -0.74  | -0.300 | -0.115 | 0.080  | 0.47   | -3.8225 | 0.0003            |         |
|                      | 30 mg     | 1.603              | 92                   | -0.142 | 0.3708 | -1.06  | -0.375 | -0.160 | 0.060  | 0.81   | -3.6809 | 0.0004            |         |
|                      | 60 mg     | 1.508              | 87                   | -0.075 | 0.3380 | -0.94  | -0.270 | -0.030 | 0.120  | 0.98   | -2.0681 | 0.0416            |         |
|                      | 120 mg    | 1.591              | 88                   | -0.124 | 0.3305 | -0.93  | -0.365 | -0.110 | 0.130  | 0.80   | -3.5290 | 0.0007            |         |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (22/24)

| Item               | Treatment | Summary Statistics |                      |       |        |        |       |        |       |         | One-sample t-test |         |        |
|--------------------|-----------|--------------------|----------------------|-------|--------|--------|-------|--------|-------|---------|-------------------|---------|--------|
|                    |           | Baseline Mean      | Change from Baseline |       |        |        |       |        |       | t-value | p-value           |         |        |
|                    |           |                    | N                    | Mean  | SD     | Min    | Q1    | Median | Q3    |         |                   | Max     |        |
| β-Carotene (µg/dL) | Week 1    | Placebo            | 27.88                | 39    | -1.77  | 6.258  | -18.0 | -4.30  | -0.60 | 2.20    | 14.4              | -1.7708 | 0.0846 |
|                    |           | 15 mg              | 28.02                | 37    | -2.44  | 8.395  | -33.3 | -4.40  | -1.40 | 0.30    | 20.7              | -1.7682 | 0.0855 |
|                    |           | 30 mg              | 30.63                | 39    | -3.89  | 8.758  | -46.0 | -5.80  | -1.80 | 0.00    | 11.3              | -2.7773 | 0.0085 |
|                    |           | 60 mg              | 23.57                | 37    | -0.62  | 6.786  | -18.0 | -4.50  | -1.00 | 1.50    | 20.3              | -0.5524 | 0.5841 |
|                    |           | 120 mg             | 20.99                | 47    | -1.85  | 4.174  | -12.8 | -2.80  | -0.70 | 0.90    | 5.9               | -3.0370 | 0.0039 |
|                    | Week 2    | Placebo            | 23.50                | 97    | 0.50   | 7.995  | -16.2 | -2.80  | -0.40 | 2.40    | 50.4              | 0.6210  | 0.5361 |
|                    |           | 15 mg              | 24.82                | 89    | -1.02  | 11.225 | -40.9 | -4.10  | -0.30 | 3.00    | 66.5              | -0.8536 | 0.3956 |
|                    |           | 30 mg              | 28.33                | 91    | -2.22  | 13.424 | -77.2 | -3.80  | -0.90 | 2.20    | 32.6              | -1.5742 | 0.1189 |
|                    |           | 60 mg              | 22.77                | 93    | -0.47  | 9.148  | -40.7 | -3.40  | -0.80 | 1.60    | 50.8              | -0.4965 | 0.6207 |
|                    |           | 120 mg             | 22.48                | 89    | -2.03  | 7.182  | -28.0 | -4.90  | -1.30 | 1.30    | 13.5              | -2.6698 | 0.0090 |
|                    | Week 3    | Placebo            | 28.99                | 36    | -1.92  | 8.572  | -22.3 | -5.00  | -0.45 | 3.30    | 12.5              | -1.3474 | 0.1865 |
|                    |           | 15 mg              | 27.76                | 36    | -1.83  | 9.491  | -35.3 | -2.95  | 0.15  | 2.60    | 17.1              | -1.1572 | 0.2550 |
|                    |           | 30 mg              | 31.28                | 38    | -4.25  | 11.485 | -43.0 | -6.50  | -1.50 | 1.50    | 13.6              | -2.2825 | 0.0283 |
|                    |           | 60 mg              | 24.81                | 32    | -1.31  | 9.877  | -32.0 | -3.85  | -0.75 | 1.55    | 29.8              | -0.7499 | 0.4590 |
|                    |           | 120 mg             | 22.02                | 42    | -1.89  | 7.344  | -29.7 | -5.20  | -0.30 | 1.90    | 12.6              | -1.6703 | 0.1025 |
|                    | Week 4    | Placebo            | 23.74                | 96    | 2.14   | 11.187 | -53.1 | -2.20  | 1.05  | 5.75    | 41.7              | 1.8748  | 0.0639 |
|                    |           | 15 mg              | 24.82                | 89    | -0.46  | 8.342  | -41.0 | -3.20  | 0.10  | 3.00    | 33.3              | -0.5185 | 0.6054 |
|                    |           | 30 mg              | 28.51                | 89    | -2.24  | 15.235 | -88.9 | -3.40  | -1.00 | 3.40    | 26.1              | -1.3866 | 0.1691 |
|                    |           | 60 mg              | 23.07                | 89    | 0.20   | 8.499  | -23.2 | -3.00  | -0.30 | 3.40    | 32.4              | 0.2170  | 0.8287 |
|                    |           | 120 mg             | 22.61                | 87    | -2.08  | 10.204 | -35.9 | -3.80  | -0.70 | 2.00    | 26.5              | -1.9018 | 0.0605 |
| Week 6             | Placebo   | 27.76              | 40                   | 0.58  | 9.297  | -21.5  | -1.50 | 0.00   | 5.00  | 30.0    | 0.3912            | 0.6978  |        |
|                    | 15 mg     | 24.12              | 40                   | -0.21 | 13.398 | -53.6  | -3.25 | -0.60  | 3.70  | 42.8    | -0.1003           | 0.9206  |        |
|                    | 30 mg     | 31.44              | 38                   | -5.24 | 12.518 | -49.5  | -6.90 | -1.60  | 1.60  | 14.5    | -2.5828           | 0.0139  |        |
|                    | 60 mg     | 23.33              | 31                   | -1.46 | 13.570 | -40.3  | -3.50 | -0.30  | 1.90  | 33.4    | -0.6009           | 0.5524  |        |
|                    | 120 mg    | 22.01              | 47                   | -2.22 | 9.767  | -32.1  | -8.40 | -0.90  | 1.90  | 21.3    | -1.5562           | 0.1265  |        |
| Week 8             | Placebo   | 23.91              | 95                   | 2.33  | 11.026 | -17.8  | -2.20 | 0.80   | 5.80  | 53.1    | 2.0638            | 0.0418  |        |
|                    | 15 mg     | 24.87              | 86                   | 0.87  | 13.146 | -57.8  | -3.50 | 0.35   | 3.90  | 51.6    | 0.6144            | 0.5406  |        |
|                    | 30 mg     | 28.51              | 89                   | -0.11 | 19.059 | -74.6  | -4.60 | -0.40  | 4.50  | 86.3    | -0.0528           | 0.9580  |        |
|                    | 60 mg     | 22.56              | 86                   | 1.66  | 13.273 | -32.1  | -3.30 | -0.10  | 5.10  | 70.7    | 1.1602            | 0.2492  |        |
|                    | 120 mg    | 22.81              | 84                   | -2.28 | 11.121 | -34.1  | -6.50 | -1.55  | 3.10  | 36.8    | -1.8807           | 0.0635  |        |
| Week 12            | Placebo   | 23.91              | 95                   | 1.54  | 12.299 | -36.7  | -2.00 | 1.10   | 6.10  | 45.8    | 1.2171            | 0.2266  |        |
|                    | 15 mg     | 24.87              | 86                   | 0.59  | 15.154 | -65.6  | -3.50 | 0.60   | 5.20  | 47.5    | 0.3636            | 0.7171  |        |
|                    | 30 mg     | 29.06              | 86                   | -1.10 | 17.056 | -69.3  | -4.70 | -0.40  | 3.70  | 81.1    | -0.5981           | 0.5514  |        |
|                    | 60 mg     | 22.90              | 82                   | 1.03  | 14.561 | -45.9  | -2.90 | 0.50   | 3.90  | 71.5    | 0.6386            | 0.5249  |        |
|                    | 120 mg    | 23.27              | 83                   | -3.61 | 10.713 | -38.7  | -6.80 | -2.50  | 2.00  | 27.0    | -3.0676           | 0.0029  |        |
| Week 16            | Placebo   | 23.91              | 95                   | 0.74  | 16.035 | -66.7  | -3.00 | 0.30   | 5.30  | 75.5    | 0.4498            | 0.6539  |        |
|                    | 15 mg     | 25.00              | 84                   | 1.32  | 13.650 | -67.2  | -2.65 | 0.05   | 5.60  | 32.2    | 0.8849            | 0.3788  |        |
|                    | 30 mg     | 29.34              | 85                   | -1.16 | 18.600 | -75.1  | -6.40 | -1.40  | 5.50  | 59.7    | -0.5744           | 0.5672  |        |
|                    | 60 mg     | 22.94              | 80                   | 2.07  | 16.639 | -64.6  | -3.55 | 0.10   | 8.30  | 66.9    | 1.1121            | 0.2695  |        |
|                    | 120 mg    | 23.19              | 82                   | -2.68 | 10.969 | -41.6  | -5.00 | -1.40  | 2.70  | 24.5    | -2.2088           | 0.0300  |        |
| Week 20            | Placebo   | 23.91              | 95                   | 1.32  | 18.295 | -49.2  | -4.30 | 0.00   | 6.50  | 109.4   | 0.7038            | 0.4833  |        |
|                    | 15 mg     | 25.00              | 84                   | 2.44  | 17.837 | -87.9  | -3.45 | 2.00   | 7.05  | 52.5    | 1.2540            | 0.2134  |        |
|                    | 30 mg     | 29.34              | 85                   | -2.55 | 17.636 | -65.2  | -5.10 | -0.40  | 4.40  | 60.0    | -1.3322           | 0.1864  |        |
|                    | 60 mg     | 23.19              | 79                   | 3.35  | 18.233 | -58.5  | -3.30 | 0.60   | 5.10  | 92.1    | 1.6315            | 0.1068  |        |
|                    | 120 mg    | 22.26              | 80                   | -1.43 | 10.334 | -28.4  | -7.05 | -0.90  | 4.10  | 27.5    | -1.2410           | 0.2183  |        |
| Week 24            | Placebo   | 23.95              | 94                   | -0.33 | 16.867 | -68.0  | -4.60 | 0.70   | 6.20  | 61.6    | -0.1871           | 0.8520  |        |
|                    | 15 mg     | 25.00              | 84                   | 3.13  | 19.547 | -89.4  | -3.15 | 0.80   | 6.55  | 87.5    | 1.4652            | 0.1466  |        |
|                    | 30 mg     | 29.42              | 84                   | -2.48 | 18.267 | -89.9  | -5.45 | -0.75  | 2.70  | 52.4    | -1.2465           | 0.2161  |        |
|                    | 60 mg     | 23.23              | 78                   | 3.19  | 21.179 | -48.1  | -5.20 | 0.40   | 4.10  | 114.9   | 1.3290            | 0.1878  |        |
|                    | 120 mg    | 22.38              | 79                   | -0.55 | 12.976 | -33.0  | -7.40 | -1.10  | 5.60  | 42.0    | -0.3798           | 0.7051  |        |
| Week 28            | Placebo   | 23.95              | 94                   | -1.02 | 15.270 | -69.9  | -5.80 | 0.00   | 3.90  | 52.2    | -0.6505           | 0.5170  |        |
|                    | 15 mg     | 25.01              | 83                   | 5.20  | 21.133 | -60.5  | -1.90 | 1.60   | 7.50  | 104.1   | 2.2396            | 0.0278  |        |
|                    | 30 mg     | 29.42              | 84                   | -0.77 | 16.197 | -72.7  | -4.05 | 1.00   | 5.75  | 43.6    | -0.4372           | 0.6631  |        |
|                    | 60 mg     | 23.48              | 77                   | 5.99  | 21.059 | -43.6  | -2.20 | 1.90   | 9.10  | 95.9    | 2.4964            | 0.0147  |        |
|                    | 120 mg    | 22.57              | 78                   | 1.10  | 13.729 | -27.7  | -4.60 | 0.40   | 5.90  | 64.9    | 0.7051            | 0.4829  |        |
| Follow-up          | Placebo   | 23.45              | 98                   | -0.92 | 14.975 | -69.9  | -5.40 | 0.00   | 3.90  | 52.2    | -0.6078           | 0.5448  |        |
|                    | 15 mg     | 24.97              | 86                   | 5.10  | 20.860 | -60.5  | -1.90 | 1.65   | 7.70  | 104.1   | 2.2683            | 0.0258  |        |
|                    | 30 mg     | 28.08              | 92                   | -0.40 | 15.808 | -72.7  | -4.05 | 1.00   | 6.00  | 43.6    | -0.2440           | 0.8078  |        |
|                    | 60 mg     | 22.67              | 87                   | 5.86  | 19.977 | -43.6  | -2.20 | 2.60   | 9.10  | 95.9    | 2.7359            | 0.0076  |        |
|                    | 120 mg    | 22.65              | 88                   | 0.74  | 13.623 | -27.7  | -4.75 | 0.25   | 5.40  | 64.9    | 0.5063            | 0.6139  |        |

表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (23/24)

| Item                 | Treatment | Summary Statistics |                      |      |      |      |      |        |     |         | One-sample t-test |         |        |
|----------------------|-----------|--------------------|----------------------|------|------|------|------|--------|-----|---------|-------------------|---------|--------|
|                      |           | Baseline Mean      | Change from Baseline |      |      |      |      |        |     | t-value | p-value           |         |        |
|                      |           |                    | N                    | Mean | SD   | Min  | Q1   | Median | Q3  |         |                   | Max     |        |
| m-AST<br>(IU/L/37°C) | Week 1    | Placebo            | 2.8                  | 39   | -0.2 | 1.81 | -4   | -2.0   | 0.0 | 1.0     | 5                 | -0.7948 | 0.4316 |
|                      |           | 15 mg              | 2.4                  | 37   | -0.1 | 1.38 | -5   | -1.0   | 0.0 | 1.0     | 3                 | -0.5967 | 0.5545 |
|                      |           | 30 mg              | 2.4                  | 39   | 0.1  | 1.41 | -5   | -1.0   | 0.0 | 1.0     | 3                 | 0.2266  | 0.8219 |
|                      |           | 60 mg              | 2.9                  | 37   | -0.5 | 1.52 | -6   | -1.0   | 0.0 | 0.0     | 3                 | -1.8385 | 0.0742 |
|                      |           | 120 mg             | 2.7                  | 47   | 0.2  | 2.08 | -4   | -1.0   | 0.0 | 1.0     | 6                 | 0.5616  | 0.5771 |
|                      | Week 2    | Placebo            | 3.0                  | 96   | 0.2  | 1.88 | -6   | -1.0   | 0.0 | 1.0     | 7                 | 0.8157  | 0.4167 |
|                      |           | 15 mg              | 3.1                  | 89   | 0.2  | 1.90 | -5   | -1.0   | 0.0 | 1.0     | 8                 | 1.1746  | 0.2433 |
|                      |           | 30 mg              | 2.5                  | 91   | 0.2  | 1.59 | -5   | -1.0   | 0.0 | 1.0     | 5                 | 1.3887  | 0.1683 |
|                      |           | 60 mg              | 3.0                  | 93   | 0.3  | 1.92 | -6   | -1.0   | 0.0 | 2.0     | 5                 | 1.5113  | 0.1341 |
|                      |           | 120 mg             | 2.8                  | 89   | 0.0  | 1.52 | -7   | -1.0   | 0.0 | 1.0     | 5                 | 0.0698  | 0.9445 |
|                      | Week 3    | Placebo            | 2.8                  | 36   | 0.7  | 3.27 | -5   | -0.5   | 0.0 | 1.0     | 16                | 1.3270  | 0.1931 |
|                      |           | 15 mg              | 2.5                  | 36   | 0.0  | 2.10 | -7   | -1.0   | 0.0 | 1.0     | 7                 | 0.0000  | 1.0000 |
|                      |           | 30 mg              | 2.5                  | 38   | 0.4  | 2.07 | -3   | -1.0   | 0.0 | 1.0     | 10                | 1.1735  | 0.2481 |
|                      |           | 60 mg              | 2.6                  | 32   | -0.2 | 1.82 | -6   | -1.0   | 0.0 | 0.0     | 7                 | -0.4868 | 0.6299 |
|                      |           | 120 mg             | 2.7                  | 42   | 0.4  | 2.00 | -3   | 0.0    | 0.0 | 1.0     | 9                 | 1.3875  | 0.1728 |
|                      | Week 4    | Placebo            | 3.0                  | 95   | 0.3  | 2.06 | -5   | -1.0   | 0.0 | 1.0     | 11                | 1.1968  | 0.2344 |
|                      |           | 15 mg              | 3.1                  | 89   | 0.1  | 1.75 | -5   | -1.0   | 0.0 | 1.0     | 8                 | 0.6064  | 0.5458 |
|                      |           | 30 mg              | 2.6                  | 89   | 0.1  | 1.46 | -3   | -1.0   | 0.0 | 1.0     | 3                 | 0.6525  | 0.5158 |
|                      |           | 60 mg              | 3.1                  | 89   | 0.1  | 1.95 | -8   | -1.0   | 0.0 | 1.0     | 9                 | 0.2718  | 0.7864 |
|                      |           | 120 mg             | 2.8                  | 87   | 0.1  | 1.60 | -4   | -1.0   | 0.0 | 1.0     | 7                 | 0.3346  | 0.7387 |
| Week 6               | Placebo   | 3.1                | 40                   | -0.0 | 2.04 | -5   | -1.0 | 0.0    | 1.0 | 5       | -0.0773           | 0.9387  |        |
|                      | 15 mg     | 2.6                | 40                   | 0.3  | 1.80 | -3   | -1.0 | 0.0    | 1.5 | 6       | 0.9677            | 0.3391  |        |
|                      | 30 mg     | 2.6                | 38                   | 0.3  | 1.43 | -3   | -1.0 | 0.0    | 1.0 | 3       | 1.1366            | 0.2630  |        |
|                      | 60 mg     | 2.9                | 31                   | 0.1  | 1.76 | -6   | -1.0 | 0.0    | 1.0 | 3       | 0.3065            | 0.7613  |        |
|                      | 120 mg    | 2.7                | 47                   | 0.0  | 1.38 | -4   | -1.0 | 0.0    | 1.0 | 4       | 0.1061            | 0.9160  |        |
| Week 8               | Placebo   | 3.0                | 94                   | 0.1  | 2.36 | -6   | -1.0 | 0.0    | 1.0 | 10      | 0.4375            | 0.6628  |        |
|                      | 15 mg     | 3.0                | 86                   | 0.4  | 2.18 | -4   | -1.0 | 0.0    | 2.0 | 8       | 1.7338            | 0.0866  |        |
|                      | 30 mg     | 2.6                | 89                   | 0.3  | 1.75 | -4   | -1.0 | 0.0    | 1.0 | 8       | 1.8753            | 0.0641  |        |
|                      | 60 mg     | 3.0                | 86                   | 0.1  | 1.80 | -6   | -1.0 | 0.0    | 1.0 | 7       | 0.6587            | 0.5118  |        |
|                      | 120 mg    | 2.8                | 84                   | -0.1 | 1.73 | -7   | -1.0 | 0.0    | 1.0 | 4       | -0.3775           | 0.7067  |        |
| Week 12              | Placebo   | 3.0                | 94                   | 0.2  | 2.29 | -6   | -1.0 | 0.0    | 1.0 | 10      | 0.9472            | 0.3460  |        |
|                      | 15 mg     | 3.0                | 86                   | 0.7  | 5.66 | -4   | -1.0 | 0.0    | 1.0 | 50      | 1.2190            | 0.2262  |        |
|                      | 30 mg     | 2.6                | 86                   | 0.2  | 1.73 | -5   | -1.0 | 0.0    | 1.0 | 5       | 1.1833            | 0.2400  |        |
|                      | 60 mg     | 3.0                | 82                   | 0.1  | 2.06 | -8   | -1.0 | 0.0    | 1.0 | 5       | 0.5353            | 0.5939  |        |
|                      | 120 mg    | 2.8                | 83                   | 0.2  | 2.12 | -6   | -1.0 | 0.0    | 1.0 | 6       | 0.8790            | 0.3820  |        |
| Week 16              | Placebo   | 3.0                | 94                   | 0.5  | 2.33 | -5   | -1.0 | 0.0    | 2.0 | 11      | 2.2122            | 0.0294  |        |
|                      | 15 mg     | 3.0                | 84                   | 0.2  | 1.82 | -4   | -1.0 | 0.0    | 1.0 | 7       | 0.9592            | 0.3403  |        |
|                      | 30 mg     | 2.6                | 85                   | 0.2  | 1.57 | -5   | -1.0 | 0.0    | 1.0 | 5       | 1.1055            | 0.2721  |        |
|                      | 60 mg     | 3.0                | 80                   | 0.1  | 2.00 | -7   | -1.0 | 0.0    | 1.0 | 3       | 0.5584            | 0.5782  |        |
|                      | 120 mg    | 2.7                | 82                   | 0.2  | 1.93 | -6   | -1.0 | 0.0    | 1.0 | 7       | 1.1455            | 0.2554  |        |
| Week 20              | Placebo   | 3.0                | 94                   | 0.6  | 2.06 | -4   | -1.0 | 0.0    | 2.0 | 7       | 2.9566            | 0.0039  |        |
|                      | 15 mg     | 3.0                | 84                   | 0.5  | 1.87 | -4   | 0.0  | 0.0    | 1.0 | 9       | 2.4558            | 0.0161  |        |
|                      | 30 mg     | 2.6                | 85                   | 0.5  | 1.65 | -4   | 0.0  | 0.0    | 1.0 | 8       | 2.8232            | 0.0059  |        |
|                      | 60 mg     | 3.0                | 79                   | 0.3  | 1.90 | -7   | 0.0  | 0.0    | 1.0 | 4       | 1.2439            | 0.2173  |        |
|                      | 120 mg    | 2.8                | 80                   | 0.5  | 2.32 | -8   | 0.0  | 0.0    | 2.0 | 9       | 2.0754            | 0.0412  |        |
| Week 24              | Placebo   | 3.0                | 93                   | 0.7  | 2.21 | -5   | 0.0  | 0.0    | 2.0 | 10      | 2.9528            | 0.0040  |        |
|                      | 15 mg     | 3.0                | 84                   | 0.4  | 1.88 | -4   | 0.0  | 0.0    | 1.0 | 9       | 2.0276            | 0.0458  |        |
|                      | 30 mg     | 2.6                | 84                   | 0.5  | 1.62 | -3   | 0.0  | 0.5    | 1.5 | 5       | 3.0912            | 0.0027  |        |
|                      | 60 mg     | 2.9                | 78                   | 0.3  | 2.11 | -8   | -1.0 | 0.0    | 2.0 | 6       | 1.0732            | 0.2866  |        |
|                      | 120 mg    | 2.8                | 79                   | 0.2  | 2.26 | -5   | -1.0 | 0.0    | 1.0 | 13      | 0.6960            | 0.4885  |        |
| Week 28              | Placebo   | 3.0                | 93                   | 0.8  | 2.15 | -3   | -1.0 | 1.0    | 2.0 | 11      | 3.6205            | 0.0005  |        |
|                      | 15 mg     | 3.0                | 83                   | 0.5  | 1.78 | -4   | -1.0 | 1.0    | 1.0 | 4       | 2.3487            | 0.0212  |        |
|                      | 30 mg     | 2.6                | 84                   | 0.5  | 1.61 | -5   | -1.0 | 1.0    | 1.5 | 4       | 2.5772            | 0.0117  |        |
|                      | 60 mg     | 2.9                | 77                   | 0.2  | 1.91 | -7   | -1.0 | 0.0    | 1.0 | 3       | 0.8937            | 0.3743  |        |
|                      | 120 mg    | 2.8                | 78                   | 0.5  | 2.04 | -6   | 0.0  | 1.0    | 2.0 | 6       | 2.1057            | 0.0385  |        |
| Follow-up            | Placebo   | 3.0                | 97                   | 0.9  | 2.15 | -3   | -1.0 | 1.0    | 2.0 | 11      | 3.9187            | 0.0002  |        |
|                      | 15 mg     | 3.1                | 86                   | 0.3  | 1.95 | -7   | -1.0 | 0.5    | 1.0 | 4       | 1.4901            | 0.1399  |        |
|                      | 30 mg     | 2.5                | 92                   | 0.5  | 1.62 | -5   | -0.5 | 1.0    | 1.0 | 5       | 2.9593            | 0.0039  |        |
|                      | 60 mg     | 3.0                | 87                   | 0.3  | 1.86 | -7   | -1.0 | 1.0    | 1.0 | 3       | 1.2662            | 0.2089  |        |
|                      | 120 mg    | 2.8                | 88                   | 0.6  | 2.30 | -6   | 0.0  | 1.0    | 2.0 | 8       | 2.6423            | 0.0098  |        |



表 2.7.4.7-付録-14 血液生化学検査の評価時点別の変化量 (CCT-001 試験) (24/24)

| Item                    | Treatment | Summary Statistics |                      |        |        |         |         |        |        |         | One-sample t-test |         |         |        |
|-------------------------|-----------|--------------------|----------------------|--------|--------|---------|---------|--------|--------|---------|-------------------|---------|---------|--------|
|                         |           | Baseline Mean      | Change from Baseline |        |        |         |         |        | Max    | t-value | p-value           |         |         |        |
|                         |           |                    | N                    | Mean   | SD     | Min     | Q1      | Median |        |         |                   | Q3      |         |        |
| CK18 (Segment)<br>(U/L) | Week 1    | Placebo            | 171.09               | 39     | 3.44   | 65.565  | -220.5  | -21.60 | 0.00   | 26.40   | 271.5             | 0.3280  | 0.7447  |        |
|                         |           | 15 mg              | 154.66               | 37     | -7.59  | 74.767  | -197.8  | -47.90 | 0.00   | 25.50   | 239.2             | -0.6172 | 0.5410  |        |
|                         |           | 30 mg              | 145.66               | 39     | -14.95 | 44.137  | -104.6  | -42.70 | -3.00  | 0.00    | 79.5              | -2.1151 | 0.0410  |        |
|                         |           | 60 mg              | 168.86               | 37     | -4.26  | 77.187  | -284.5  | -25.90 | 0.00   | 25.00   | 156.2             | -0.3359 | 0.7389  |        |
|                         |           | 120 mg             | 151.42               | 47     | -12.77 | 80.217  | -416.8  | -33.50 | 0.00   | 27.90   | 144.0             | -1.0918 | 0.2806  |        |
|                         |           | Week 2             | Placebo              | 175.58 | 97     | 3.12    | 112.045 | -459.5 | -32.90 | 0.00    | 35.80             | 595.2   | 0.2741  | 0.7846 |
|                         |           |                    | 15 mg                | 159.05 | 89     | 5.58    | 100.988 | -322.8 | -32.90 | 0.00    | 43.60             | 610.8   | 0.5212  | 0.6035 |
|                         |           |                    | 30 mg                | 137.74 | 91     | -3.58   | 50.035  | -136.8 | -26.80 | 0.00    | 7.70              | 122.3   | -0.6830 | 0.4964 |
|                         |           |                    | 60 mg                | 153.08 | 93     | -12.05  | 82.286  | -547.3 | -28.00 | 0.00    | 8.50              | 261.1   | -1.4128 | 0.1611 |
|                         |           |                    | 120 mg               | 127.27 | 89     | 6.94    | 76.904  | -279.0 | -24.40 | 0.00    | 24.00             | 311.5   | 0.8518  | 0.3966 |
|                         |           | Week 3             | Placebo              | 177.14 | 36     | -4.84   | 101.465 | -454.8 | -16.80 | 0.00    | 40.65             | 215.9   | -0.2863 | 0.7763 |
|                         |           |                    | 15 mg                | 153.55 | 36     | -8.93   | 65.956  | -168.9 | -49.25 | 0.00    | 25.10             | 185.8   | -0.8124 | 0.4220 |
|                         |           |                    | 30 mg                | 155.28 | 38     | -29.29  | 53.733  | -159.6 | -51.20 | -12.00  | 0.00              | 151.5   | -3.3605 | 0.0018 |
|                         |           |                    | 60 mg                | 136.79 | 32     | -14.93  | 58.012  | -229.0 | -39.15 | 0.00    | 1.90              | 101.3   | -1.4557 | 0.1555 |
|                         |           |                    | 120 mg               | 151.98 | 42     | 8.54    | 43.404  | -92.7  | -9.50  | 0.00    | 23.20             | 179.0   | 1.2748  | 0.2095 |
|                         |           | Week 4             | Placebo              | 175.84 | 96     | -3.09   | 117.940 | -724.6 | -19.40 | 0.00    | 35.30             | 290.5   | -0.2568 | 0.7979 |
|                         |           |                    | 15 mg                | 159.05 | 89     | 7.16    | 109.490 | -346.0 | -42.70 | 0.00    | 44.00             | 495.7   | 0.6165  | 0.5392 |
|                         |           |                    | 30 mg                | 134.78 | 89     | -17.01  | 64.761  | -176.8 | -42.00 | 0.00    | 0.00              | 184.7   | -2.4786 | 0.0151 |
|                         |           |                    | 60 mg                | 165.41 | 89     | -9.97   | 107.995 | -690.5 | -39.40 | 0.00    | 29.60             | 301.8   | -0.8705 | 0.3864 |
|                         |           |                    | 120 mg               | 127.08 | 87     | 4.58    | 50.144  | -125.0 | -27.60 | 0.00    | 30.50             | 136.2   | 0.8518  | 0.3967 |
|                         |           | Week 6             | Placebo              | 185.73 | 40     | 5.49    | 79.398  | -223.2 | -29.65 | 0.00    | 32.50             | 172.5   | 0.4375  | 0.6642 |
|                         |           |                    | 15 mg                | 151.75 | 40     | -19.77  | 83.724  | -212.4 | -49.25 | -0.30   | 1.75              | 276.3   | -1.4933 | 0.1434 |
|                         |           |                    | 30 mg                | 147.53 | 38     | -4.09   | 86.355  | -177.6 | -72.70 | 0.00    | 12.20             | 240.1   | -0.2921 | 0.7718 |
|                         |           |                    | 60 mg                | 137.48 | 31     | -5.34   | 80.756  | -269.4 | -33.50 | 0.00    | 38.80             | 138.1   | -0.3683 | 0.7152 |
|                         |           |                    | 120 mg               | 146.09 | 47     | -14.17  | 101.107 | -545.4 | -37.40 | 0.00    | 33.00             | 179.6   | -0.9607 | 0.3417 |
|                         |           | Week 8             | Placebo              | 176.83 | 95     | -5.69   | 115.052 | -598.2 | -23.20 | 0.00    | 38.40             | 350.9   | -0.4820 | 0.6309 |
|                         |           |                    | 15 mg                | 159.70 | 86     | -3.31   | 114.832 | -501.0 | -45.20 | 0.00    | 41.00             | 552.8   | -0.2673 | 0.7899 |
|                         |           |                    | 30 mg                | 134.78 | 89     | -7.07   | 98.638  | -272.5 | -45.00 | 0.00    | 6.80              | 484.6   | -0.6765 | 0.5005 |
|                         |           |                    | 60 mg                | 155.59 | 86     | -5.64   | 118.792 | -691.8 | -45.50 | 0.00    | 11.60             | 302.3   | -0.4401 | 0.6610 |
|                         |           |                    | 120 mg               | 124.72 | 84     | 1.12    | 75.830  | -445.8 | -25.35 | 0.00    | 25.90             | 203.5   | 0.1353  | 0.8927 |
|                         |           | Week 12            | Placebo              | 176.83 | 95     | 6.90    | 150.708 | -871.3 | 0.00   | 15.60   | 56.40             | 318.3   | 0.4462  | 0.6565 |
|                         |           |                    | 15 mg                | 159.70 | 86     | 10.12   | 106.142 | -363.9 | -43.50 | 0.00    | 40.20             | 495.7   | 0.8843  | 0.3790 |
|                         |           |                    | 30 mg                | 135.16 | 86     | -6.50   | 76.434  | -241.0 | -44.70 | 0.00    | 46.40             | 200.4   | -0.7888 | 0.4324 |
|                         |           |                    | 60 mg                | 156.78 | 82     | -16.86  | 105.606 | -711.4 | -43.30 | 0.00    | 20.30             | 195.5   | -1.4457 | 0.1521 |
|                         |           |                    | 120 mg               | 126.48 | 83     | 3.25    | 133.216 | -834.7 | -40.00 | 0.00    | 37.50             | 657.0   | 0.2221  | 0.8248 |
|                         |           | Week 16            | Placebo              | 176.83 | 95     | 17.66   | 137.080 | -647.6 | -10.40 | 1.70    | 72.00             | 573.1   | 1.2557  | 0.2123 |
|                         |           |                    | 15 mg                | 156.38 | 84     | -3.96   | 99.166  | -242.5 | -54.70 | 0.00    | 47.95             | 355.3   | -0.3664 | 0.7150 |
|                         |           |                    | 30 mg                | 136.31 | 85     | 2.15    | 70.305  | -245.1 | -33.40 | 0.00    | 46.70             | 201.4   | 0.2825  | 0.7783 |
|                         |           |                    | 60 mg                | 157.09 | 80     | -6.83   | 121.820 | -625.9 | -51.30 | 0.00    | 43.15             | 510.1   | -0.5017 | 0.6172 |
|                         |           |                    | 120 mg               | 125.63 | 82     | -1.34   | 142.353 | -830.8 | -56.30 | 0.00    | 35.90             | 524.2   | -0.0854 | 0.9321 |
|                         |           | Week 20            | Placebo              | 176.83 | 95     | 35.34   | 143.397 | -530.6 | -7.10  | 25.90   | 67.30             | 740.3   | 2.4020  | 0.0183 |
|                         |           |                    | 15 mg                | 156.38 | 84     | 6.33    | 107.195 | -247.6 | -46.35 | 0.00    | 69.65             | 335.0   | 0.5408  | 0.5901 |
|                         |           |                    | 30 mg                | 136.31 | 85     | 6.91    | 87.957  | -302.2 | -19.90 | 8.00    | 56.40             | 207.3   | 0.7246  | 0.4707 |
|                         |           |                    | 60 mg                | 158.60 | 79     | -1.82   | 126.260 | -581.5 | -49.30 | 0.00    | 52.70             | 486.9   | -0.1278 | 0.8987 |
|                         |           |                    | 120 mg               | 127.83 | 80     | 3.09    | 140.543 | -851.4 | -44.20 | 0.00    | 40.80             | 569.7   | 0.1966  | 0.8447 |
|                         |           | Week 24            | Placebo              | 177.51 | 94     | 14.51   | 167.694 | -801.8 | -37.50 | 5.80    | 80.90             | 587.8   | 0.8391  | 0.4036 |
|                         |           |                    | 15 mg                | 156.38 | 84     | 7.64    | 113.899 | -254.4 | -59.80 | 0.00    | 65.90             | 340.4   | 0.6146  | 0.5405 |
|                         |           |                    | 30 mg                | 136.43 | 84     | -0.12   | 94.968  | -267.0 | -47.20 | 0.00    | 51.80             | 398.1   | -0.0114 | 0.9910 |
|                         |           | 60 mg              | 148.61               | 78     | -8.84  | 112.632 | -346.2  | -49.40 | 0.00   | 46.60   | 266.2             | -0.6928 | 0.4905  |        |
|                         |           | 120 mg             | 125.03               | 79     | -15.09 | 128.775 | -919.2  | -50.60 | 0.00   | 37.50   | 283.4             | -1.0413 | 0.3010  |        |
|                         | Week 28   | Placebo            | 177.51               | 94     | 6.94   | 122.610 | -386.3  | -47.60 | 0.00   | 56.00   | 388.2             | 0.5486  | 0.5846  |        |
|                         |           | 15 mg              | 157.81               | 83     | 6.19   | 128.292 | -619.4  | -40.00 | 0.00   | 74.90   | 303.8             | 0.4394  | 0.6615  |        |
|                         |           | 30 mg              | 136.43               | 84     | 9.61   | 100.584 | -269.4  | -26.20 | 0.00   | 55.30   | 402.0             | 0.8756  | 0.3838  |        |
|                         |           | 60 mg              | 149.31               | 77     | 7.62   | 128.212 | -265.2  | -60.30 | 0.00   | 45.80   | 539.2             | 0.5218  | 0.6033  |        |
|                         |           | 120 mg             | 126.63               | 79     | 11.09  | 145.267 | -909.7  | -28.20 | 0.00   | 45.70   | 463.0             | 0.6783  | 0.4996  |        |
|                         | Follow-up | Placebo            | 174.94               | 98     | 7.34   | 120.741 | -386.3  | -47.60 | 0.00   | 56.00   | 388.2             | 0.6015  | 0.5489  |        |
|                         |           | 15 mg              | 161.18               | 86     | 8.93   | 129.166 | -619.4  | -37.90 | 0.00   | 74.10   | 303.8             | 0.6409  | 0.5233  |        |
|                         |           | 30 mg              | 137.35               | 92     | 8.56   | 98.860  | -269.4  | -27.85 | 0.00   | 52.10   | 402.0             | 0.8304  | 0.4085  |        |
|                         |           | 60 mg              | 154.16               | 87     | 10.34  | 125.024 | -265.2  | -60.30 | 0.00   | 46.50   | 539.2             | 0.7717  | 0.4424  |        |
|                         |           | 120 mg             | 126.19               | 88     | 13.76  | 140.132 | -909.7  | -23.85 | 3.55   | 48.25   | 463.0             | 0.9215  | 0.3594  |        |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (1/24)

|               |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|---------------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|               |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|               |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Total Protein | Week 1  | Lower   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | -     | 1     |    |
|               |         | Normal  | 1      | 34    | 2     | 37     | -      | 34    | 3     | 37     | 1      | 36    | -     | 37     | -      | 34    | 1     | 35     | -      | 45    | -     | 45 |
|               |         | Upper   | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | -     | -     | -      | -      | -     | 1     | 1  |
|               |         | Total   | 1      | 35    | 3     | 39     | -      | 34    | 3     | 37     | 1      | 38    | -     | 39     | -      | 36    | 1     | 37     | -      | 46    | 1     | 47 |
|               | Week 2  | Lower   | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | 2      | 2     | -     | 4      | -      | 5     | -     | 5  |
|               |         | Normal  | 2      | 84    | 7     | 93     | -      | 81    | 4     | 85     | -      | 87    | 1     | 88     | -      | 86    | 1     | 87     | -      | 80    | 3     | 83 |
|               |         | Upper   | -      | -     | 1     | 1      | -      | 1     | 2     | 3      | -      | 1     | 1     | 2      | -      | 2     | 2     | -      | -      | -     | 1     | 1  |
|               |         | Total   | 2      | 86    | 8     | 96     | -      | 83    | 6     | 89     | -      | 89    | 2     | 91     | 2      | 88    | 3     | 93     | -      | 85    | 4     | 89 |
|               | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 2     | -     | 2  |
|               |         | Normal  | -      | 33    | 2     | 35     | -      | 34    | 2     | 36     | 1      | 35    | -     | 36     | -      | 30    | -     | 30     | -      | 38    | 1     | 39 |
|               |         | Upper   | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1  |
|               |         | Total   | -      | 33    | 3     | 36     | -      | 34    | 2     | 36     | 1      | 37    | -     | 38     | -      | 32    | -     | 32     | -      | 41    | 1     | 42 |
|               | Week 4  | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | 1      | 1     | -     | 2      | -      | 2     | -     | 2  |
|               |         | Normal  | 1      | 83    | 5     | 89     | -      | 80    | 3     | 83     | -      | 84    | 1     | 85     | -      | 79    | 4     | 83     | -      | 80    | 2     | 82 |
|               |         | Upper   | -      | 2     | 3     | 5      | -      | 3     | 3     | 6      | -      | 2     | 1     | 3      | -      | 4     | -     | 4      | -      | 1     | 2     | 3  |
|               |         | Total   | 2      | 85    | 8     | 95     | -      | 83    | 6     | 89     | -      | 87    | 2     | 89     | 1      | 84    | 4     | 89     | -      | 83    | 4     | 87 |
|               | Week 6  | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 2     | -     | 2  |
|               |         | Normal  | -      | 35    | 2     | 37     | -      | 37    | 1     | 38     | -      | 37    | -     | 37     | -      | 30    | -     | 30     | -      | 43    | -     | 43 |
|               |         | Upper   | -      | 1     | 1     | 2      | -      | 1     | 1     | 2      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | 2     | 2  |
|               |         | Total   | 1      | 36    | 3     | 40     | -      | 38    | 2     | 40     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 45    | 2     | 47 |
|               | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | 1      | 1     | -     | 2      | -      | 1     | -     | 1  |
|               |         | Normal  | 2      | 82    | 7     | 91     | -      | 79    | 4     | 83     | -      | 85    | 2     | 87     | -      | 79    | 3     | 82     | -      | 78    | 2     | 80 |
|               |         | Upper   | -      | 2     | 1     | 3      | -      | 1     | 2     | 3      | -      | 1     | -     | 1      | -      | 2     | -     | 2      | -      | 1     | 2     | 3  |
|               |         | Total   | 2      | 84    | 8     | 94     | -      | 80    | 6     | 86     | -      | 87    | 2     | 89     | 1      | 82    | 3     | 86     | -      | 80    | 4     | 84 |
|               | Week 12 | Lower   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | 1      | -     | -     | 1      | -      | 3     | -     | 3  |
|               |         | Normal  | 2      | 80    | 4     | 86     | -      | 80    | 2     | 82     | -      | 83    | 2     | 85     | -      | 78    | 3     | 81     | -      | 72    | 2     | 74 |
|               |         | Upper   | -      | 3     | 4     | 7      | -      | -     | 4     | 4      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 4     | 2     | 6  |
|               |         | Total   | 2      | 84    | 8     | 94     | -      | 80    | 6     | 86     | -      | 84    | 2     | 86     | 1      | 78    | 3     | 82     | -      | 79    | 4     | 83 |
|               | Week 16 | Lower   | 2      | 1     | -     | 3      | -      | -     | -     | -      | -      | 3     | -     | 3      | -      | 2     | -     | 2      | -      | 2     | -     | 2  |
|               |         | Normal  | -      | 82    | 5     | 87     | -      | 76    | 3     | 79     | -      | 79    | 2     | 81     | -      | 73    | 3     | 76     | -      | 72    | 2     | 74 |
|               |         | Upper   | -      | 1     | 3     | 4      | -      | 2     | 3     | 5      | -      | 1     | -     | 1      | -      | 2     | -     | 2      | -      | 4     | 2     | 6  |
|               |         | Total   | 2      | 84    | 8     | 94     | -      | 78    | 6     | 84     | -      | 83    | 2     | 85     | -      | 77    | 3     | 80     | -      | 78    | 4     | 82 |
|               | Week 20 | Lower   | 1      | 2     | -     | 3      | -      | -     | -     | -      | -      | 3     | -     | 3      | -      | -     | -     | -      | -      | 2     | -     | 2  |
|               |         | Normal  | 1      | 82    | 5     | 88     | -      | 76    | 4     | 80     | -      | 79    | 2     | 81     | -      | 75    | 3     | 78     | -      | 71    | 2     | 73 |
|               |         | Upper   | -      | -     | 3     | 3      | -      | 2     | 2     | 4      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 3     | 2     | 5  |
|               |         | Total   | 2      | 84    | 8     | 94     | -      | 78    | 6     | 84     | -      | 83    | 2     | 85     | -      | 76    | 3     | 79     | -      | 76    | 4     | 80 |
|               | Week 24 | Lower   | 1      | 1     | -     | 2      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1  |
|               |         | Normal  | 1      | 80    | 6     | 87     | -      | 77    | 3     | 80     | -      | 81    | 1     | 82     | -      | 73    | 3     | 76     | -      | 70    | 3     | 73 |
|               |         | Upper   | -      | 2     | 2     | 4      | -      | 1     | 3     | 4      | -      | -     | 1     | 1      | -      | 1     | -     | 1      | -      | 4     | 1     | 5  |
|               |         | Total   | 2      | 83    | 8     | 93     | -      | 78    | 6     | 84     | -      | 82    | 2     | 84     | -      | 75    | 3     | 78     | -      | 75    | 4     | 79 |
| Week 28       | Lower   | 1       | 3      | -     | 4     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | -      | -     | -     | -      | 2      | -     | 2     |    |
|               | Normal  | 1       | 77     | 6     | 84    | -      | 73     | 3     | 76    | -      | 79     | 1     | 80    | -      | 74     | 2     | 76    | -      | 70     | 2     | 72    |    |
|               | Upper   | -       | 3      | 2     | 5     | -      | 4      | 3     | 7     | -      | 1      | 1     | 2     | -      | 1      | 1     | -     | 2      | 2      | 4     |       |    |
|               | Total   | 2       | 83     | 8     | 93    | -      | 77     | 6     | 83    | -      | 82     | 2     | 84    | -      | 74     | 3     | 77    | -      | 74     | 4     | 78    |    |
| Follow-up     | Lower   | 1       | 3      | -     | 4     | -      | -      | -     | -     | 1      | 2      | -     | 3     | 1      | -      | -     | 1     | -      | 2      | -     | 2     |    |
|               | Normal  | 1       | 81     | 6     | 88    | -      | 76     | 3     | 79    | -      | 86     | 1     | 87    | -      | 82     | 3     | 85    | -      | 80     | 2     | 82    |    |
|               | Upper   | -       | 3      | 2     | 5     | -      | 4      | 3     | 7     | -      | 1      | 1     | 2     | -      | -      | 1     | 1     | -      | 2      | 2     | 4     |    |
|               | Total   | 2       | 87     | 8     | 97    | -      | 80     | 6     | 86    | 1      | 89     | 2     | 92    | 1      | 82     | 4     | 87    | -      | 84     | 4     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (2/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Albumin   | Week 1 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 37    | 1     | 38     | -      | 37    | -     | 37     | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 47    | -     | 47 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 1      | 37    | 1     | 39     | -      | 37    | -     | 37     | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 47    | -     | 47 |
|           | Week 2 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | 2      | 1      | -     | 3     | 2      | -      | -     | -     | 2  |
|           |        | Normal  | 1      | 93    | 2     | 96     | -      | 89    | -     | 89     | -      | 91    | -     | 91     | -      | 90    | -     | 90     | -      | 87    | -     | 87 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 1      | 93    | 2     | 96     | -      | 89    | -     | 89     | -      | 91    | -     | 91     | 2      | 91    | -     | 93     | 2      | 87    | -     | 89 |
|           | Week 3 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | -  |
|           |        | Normal  | -      | 34    | 1     | 35     | -      | 36    | -     | 36     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 42    | -     | 42 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 1      | 34    | 1     | 36     | -      | 36    | -     | 36     | -      | 38    | -     | 38     | -      | 32    | -     | 32     | -      | 42    | -     | 42 |
|           | Week 4 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | 1      | 1      | -     | 2     | 2      | -      | -     | -     | 2  |
|           |        | Normal  | -      | 92    | 2     | 94     | -      | 89    | -     | 89     | -      | 89    | -     | 89     | -      | 85    | -     | 85     | -      | 84    | -     | 84 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | -     | 1     |    |
|           |        | Total   | 1      | 92    | 2     | 95     | -      | 89    | -     | 89     | -      | 89    | -     | 89     | 1      | 88    | -     | 89     | 2      | 85    | -     | 87 |
|           | Week 6 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Normal  | -      | 38    | 1     | 39     | -      | 40    | -     | 40     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 47    | -     | 47 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 1      | 38    | 1     | 40     | -      | 40    | -     | 40     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 47    | -     | 47 |
|           | Week 8 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     | 2      | -      | -     | -     | 2  |
|           |        | Normal  | -      | 91    | 2     | 93     | -      | 86    | -     | 86     | -      | 89    | -     | 89     | -      | 83    | -     | 83     | -      | 82    | -     | 82 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | -      | -     | -     |    |
|           |        | Total   | 1      | 91    | 2     | 94     | -      | 86    | -     | 86     | -      | 89    | -     | 89     | 1      | 85    | -     | 86     | 2      | 82    | -     | 84 |
| Week 12   | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | -      | 1     | 2     | -      | -      | -     | 2     |    |
|           | Normal | -       | 91     | 2     | 93    | -      | 86     | -     | 86    | -      | 86     | -     | 86    | -      | 81     | -     | 81    | -      | 81     | -     | 81    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 1       | 91     | 2     | 94    | -      | 86     | -     | 86    | -      | 86     | -     | 86    | 1      | 81     | -     | 82    | 2      | 81     | -     | 83    |    |
| Week 16   | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | 2     | -      | -      | -     | 2     |    |
|           | Normal | -       | 91     | 2     | 93    | -      | 84     | -     | 84    | -      | 85     | -     | 85    | -      | 79     | -     | 79    | -      | 80     | -     | 80    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | 1     | -     | -      | -      | -     |       |    |
|           | Total  | 1       | 91     | 2     | 94    | -      | 84     | -     | 84    | -      | 85     | -     | 85    | -      | 80     | -     | 80    | 2      | 80     | -     | 82    |    |
| Week 20   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | 1     | 1     | -      | -      | -     | 1     |    |
|           | Normal | 1       | 91     | 2     | 94    | -      | 84     | -     | 84    | -      | 85     | -     | 85    | -      | 77     | -     | 77    | 1      | 78     | -     | 79    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | 1     | -     | -      | -      | -     |       |    |
|           | Total  | 1       | 91     | 2     | 94    | -      | 84     | -     | 84    | -      | 85     | -     | 85    | -      | 79     | -     | 79    | 2      | 78     | -     | 80    |    |
| Week 24   | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | 1     | 1     | -      | -      | -     | 1     |    |
|           | Normal | -       | 90     | 2     | 92    | -      | 84     | -     | 84    | -      | 84     | -     | 84    | -      | 77     | -     | 77    | 1      | 77     | -     | 78    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Total  | 1       | 90     | 2     | 93    | -      | 84     | -     | 84    | -      | 84     | -     | 84    | -      | 78     | -     | 78    | 2      | 77     | -     | 79    |    |
| Week 28   | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | -      | -     | 1     |    |
|           | Normal | -       | 90     | 2     | 92    | -      | 83     | -     | 83    | -      | 84     | -     | 84    | -      | 77     | -     | 77    | 1      | 76     | -     | 77    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Total  | 1       | 90     | 2     | 93    | -      | 83     | -     | 83    | -      | 84     | -     | 84    | -      | 77     | -     | 77    | 2      | 76     | -     | 78    |    |
| Follow-up | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | -      | 1     | 1     | -      | -      | -     | 1     |    |
|           | Normal | -       | 94     | 2     | 96    | -      | 86     | -     | 86    | -      | 92     | -     | 92    | -      | 86     | -     | 86    | 1      | 86     | -     | 87    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Total  | 1       | 94     | 2     | 97    | -      | 86     | -     | 86    | -      | 92     | -     | 92    | 1      | 86     | -     | 87    | 2      | 86     | -     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (3/24)

|           |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| AST(GOT)  | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 36    | 1     | 37     | -      | 36    | 1     | 37     | -      | 36    | 2     | 38     | -      | 33    | 1     | 34     | -      | 40    | 1     | 41 |
|           |         | Upper   | -      | -     | 2     | 2      | -      | -     | -     | -      | -      | -     | 1     | 1      | -      | 1     | 2     | 3      | -      | 4     | 2     | 6  |
|           |         | Total   | -      | 36    | 3     | 39     | -      | 36    | 1     | 37     | -      | 36    | 3     | 39     | -      | 34    | 3     | 37     | -      | 44    | 3     | 47 |
|           | Week 2  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 81    | 4     | 85     | -      | 72    | 3     | 75     | -      | 82    | 4     | 86     | -      | 81    | 5     | 86     | -      | 81    | 2     | 83 |
|           |         | Upper   | -      | -     | 11    | 11     | -      | 6     | 8     | 14     | -      | 2     | 3     | 5      | -      | 2     | 5     | 7      | -      | 3     | 3     | 6  |
|           |         | Total   | -      | 81    | 15    | 96     | -      | 78    | 11    | 89     | -      | 84    | 7     | 91     | -      | 83    | 10    | 93     | -      | 84    | 5     | 89 |
|           | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 33    | -     | 33     | -      | 35    | -     | 35     | -      | 33    | 4     | 37     | -      | 31    | 1     | 32     | -      | 38    | 1     | 39 |
|           |         | Upper   | -      | -     | 4     | 4      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | 2     | 3  |
|           |         | Total   | -      | 33    | 4     | 37     | -      | 36    | -     | 36     | -      | 34    | 4     | 38     | -      | 31    | 1     | 32     | -      | 39    | 3     | 42 |
|           | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 80    | 5     | 85     | -      | 74    | 4     | 78     | -      | 82    | 3     | 85     | -      | 75    | 6     | 81     | -      | 79    | 1     | 80 |
|           |         | Upper   | -      | 1     | 9     | 10     | -      | 4     | 7     | 11     | -      | -     | 4     | 4      | -      | 3     | 5     | 8      | -      | 3     | 4     | 7  |
|           |         | Total   | -      | 81    | 14    | 95     | -      | 78    | 11    | 89     | -      | 82    | 7     | 89     | -      | 78    | 11    | 89     | -      | 82    | 5     | 87 |
|           | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 35    | -     | 35     | -      | 37    | 2     | 39     | -      | 33    | 2     | 35     | -      | 29    | -     | 29     | -      | 39    | 3     | 42 |
|           |         | Upper   | -      | -     | 5     | 5      | -      | 1     | -     | 1      | -      | 1     | 2     | 3      | -      | -     | 2     | 2      | -      | 4     | 1     | 5  |
|           |         | Total   | -      | 35    | 5     | 40     | -      | 38    | 2     | 40     | -      | 34    | 4     | 38     | -      | 29    | 2     | 31     | -      | 43    | 4     | 47 |
|           | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 78    | 6     | 84     | -      | 71    | 3     | 74     | -      | 79    | 5     | 84     | -      | 75    | 4     | 79     | -      | 77    | 3     | 80 |
|           |         | Upper   | -      | 2     | 8     | 10     | -      | 5     | 7     | 12     | -      | 3     | 2     | 5      | -      | 1     | 6     | 7      | -      | 2     | 2     | 4  |
|           |         | Total   | -      | 80    | 14    | 94     | -      | 76    | 10    | 86     | -      | 82    | 7     | 89     | -      | 76    | 10    | 86     | -      | 79    | 5     | 84 |
|           | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 77    | 5     | 82     | -      | 69    | 7     | 76     | -      | 79    | 5     | 84     | -      | 73    | 5     | 78     | -      | 75    | 3     | 78 |
|           |         | Upper   | -      | 3     | 9     | 12     | -      | 7     | 3     | 10     | -      | -     | 2     | 2      | -      | 4     | 4     | -      | -      | 3     | 2     | 5  |
|           |         | Total   | -      | 80    | 14    | 94     | -      | 76    | 10    | 86     | -      | 79    | 7     | 86     | -      | 73    | 9     | 82     | -      | 78    | 5     | 83 |
|           | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 77    | 8     | 85     | -      | 70    | 5     | 75     | -      | 75    | 4     | 79     | -      | 70    | 5     | 75     | -      | 74    | 2     | 76 |
|           |         | Upper   | -      | 3     | 6     | 9      | -      | 4     | 5     | 9      | -      | 3     | 3     | 6      | -      | 2     | 3     | 5      | -      | 3     | 3     | 6  |
|           |         | Total   | -      | 80    | 14    | 94     | -      | 74    | 10    | 84     | -      | 78    | 7     | 85     | -      | 72    | 8     | 80     | -      | 77    | 5     | 82 |
|           | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 76    | 3     | 79     | -      | 67    | 7     | 74     | -      | 76    | 4     | 80     | -      | 71    | 4     | 75     | -      | 72    | 4     | 76 |
|           |         | Upper   | -      | 4     | 11    | 15     | -      | 7     | 3     | 10     | -      | 2     | 3     | 5      | -      | -     | 4     | 4      | -      | 3     | 1     | 4  |
|           |         | Total   | -      | 80    | 14    | 94     | -      | 74    | 10    | 84     | -      | 78    | 7     | 85     | -      | 71    | 8     | 79     | -      | 75    | 5     | 80 |
| Week 24   | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 74     | 6     | 80    | -      | 68     | 5     | 73    | -      | 74     | 4     | 78    | -      | 71     | 6     | 77    | -      | 72     | 4     | 76    |    |
|           | Upper   | -       | 5      | 8     | 13    | -      | 6      | 5     | 11    | -      | 3      | 3     | 6     | -      | -      | 1     | 1     | -      | 3      | -     | 3     |    |
|           | Total   | -       | 79     | 14    | 93    | -      | 74     | 10    | 84    | -      | 77     | 7     | 84    | -      | 71     | 7     | 78    | -      | 75     | 4     | 79    |    |
| Week 28   | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 76     | 4     | 80    | -      | 68     | 6     | 74    | -      | 74     | 4     | 78    | -      | 68     | 5     | 73    | -      | 70     | 3     | 73    |    |
|           | Upper   | -       | 3      | 10    | 13    | -      | 5      | 4     | 9     | -      | 3      | 3     | 6     | -      | 2      | 2     | 4     | -      | 4      | 2     | 6     |    |
|           | Total   | -       | 79     | 14    | 93    | -      | 73     | 10    | 83    | -      | 77     | 7     | 84    | -      | 70     | 7     | 77    | -      | 74     | 5     | 79    |    |
| Follow-up | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 79     | 4     | 83    | -      | 70     | 7     | 77    | -      | 82     | 4     | 86    | -      | 76     | 6     | 82    | -      | 78     | 3     | 81    |    |
|           | Upper   | -       | 3      | 11    | 14    | -      | 5      | 4     | 9     | -      | 3      | 3     | 6     | -      | 2      | 3     | 5     | -      | 5      | 2     | 7     |    |
|           | Total   | -       | 82     | 15    | 97    | -      | 75     | 11    | 86    | -      | 85     | 7     | 92    | -      | 78     | 9     | 87    | -      | 83     | 5     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (4/24)

|          |           | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|----------|-----------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|          |           | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|          |           | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| ALT(GPT) | Week 1    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 32    | 3     | 35     | -      | 31    | 2     | 33     | -      | 36    | -     | 36     | -      | 28    | 2     | 30     | -      | 35    | 1     | 36 |
|          |           | Upper   | -      | -     | 4     | 4      | -      | 1     | 3     | 4      | -      | 1     | 2     | 3      | -      | 1     | 6     | 7      | -      | 3     | 8     | 11 |
|          |           | Total   | -      | 32    | 7     | 39     | -      | 32    | 5     | 37     | -      | 37    | 2     | 39     | -      | 29    | 8     | 37     | -      | 38    | 9     | 47 |
|          | Week 2    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 71    | 6     | 77     | -      | 66    | 2     | 68     | -      | 78    | -     | 78     | -      | 57    | 8     | 65     | -      | 74    | 2     | 76 |
|          |           | Upper   | -      | 4     | 15    | 19     | -      | 5     | 16    | 21     | -      | 5     | 8     | 13     | -      | 5     | 23    | 28     | -      | 3     | 10    | 13 |
|          |           | Total   | -      | 75    | 21    | 96     | -      | 71    | 18    | 89     | -      | 83    | 8     | 91     | -      | 62    | 31    | 93     | -      | 77    | 12    | 89 |
|          | Week 3    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 29    | 3     | 32     | -      | 29    | 2     | 31     | -      | 32    | 1     | 33     | -      | 23    | 4     | 27     | -      | 30    | 4     | 34 |
|          |           | Upper   | -      | 1     | 3     | 4      | -      | 1     | 4     | 5      | -      | 3     | 2     | 5      | -      | 3     | 2     | 5      | -      | 6     | 2     | 8  |
|          |           | Total   | -      | 30    | 6     | 36     | -      | 30    | 6     | 36     | -      | 35    | 3     | 38     | -      | 26    | 6     | 32     | -      | 36    | 6     | 42 |
|          | Week 4    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 68    | 2     | 70     | -      | 64    | 5     | 69     | -      | 77    | 1     | 78     | -      | 52    | 8     | 60     | -      | 72    | 2     | 74 |
|          |           | Upper   | -      | 6     | 19    | 25     | -      | 7     | 13    | 20     | -      | 4     | 7     | 11     | -      | 6     | 23    | 29     | -      | 3     | 10    | 13 |
|          |           | Total   | -      | 74    | 21    | 95     | -      | 71    | 18    | 89     | -      | 81    | 8     | 89     | -      | 58    | 31    | 89     | -      | 75    | 12    | 87 |
|          | Week 6    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 29    | 3     | 32     | -      | 29    | 1     | 30     | -      | 33    | 1     | 34     | -      | 25    | 4     | 29     | -      | 37    | 2     | 39 |
|          |           | Upper   | -      | 2     | 6     | 8      | -      | 3     | 7     | 10     | -      | 2     | 2     | 4      | -      | 1     | 2     | 3      | -      | 2     | 6     | 8  |
|          |           | Total   | -      | 31    | 9     | 40     | -      | 32    | 8     | 40     | -      | 35    | 3     | 38     | -      | 26    | 6     | 32     | -      | 39    | 8     | 47 |
|          | Week 8    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 67    | 7     | 74     | -      | 64    | 3     | 67     | -      | 76    | 2     | 78     | -      | 51    | 12    | 63     | -      | 71    | 6     | 77 |
|          |           | Upper   | -      | 6     | 14    | 20     | -      | 5     | 14    | 19     | -      | 5     | 6     | 11     | -      | 6     | 17    | 23     | -      | 2     | 5     | 7  |
|          |           | Total   | -      | 73    | 21    | 94     | -      | 69    | 17    | 86     | -      | 81    | 8     | 89     | -      | 57    | 29    | 86     | -      | 73    | 11    | 84 |
|          | Week 12   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 70    | 6     | 76     | -      | 64    | 6     | 70     | -      | 73    | 2     | 75     | -      | 49    | 13    | 62     | -      | 66    | 6     | 72 |
|          |           | Upper   | -      | 3     | 15    | 18     | -      | 5     | 11    | 16     | -      | 5     | 6     | 11     | -      | 6     | 14    | 20     | -      | 6     | 5     | 11 |
|          |           | Total   | -      | 73    | 21    | 94     | -      | 69    | 17    | 86     | -      | 78    | 8     | 86     | -      | 55    | 27    | 82     | -      | 72    | 11    | 83 |
|          | Week 16   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 66    | 4     | 70     | -      | 63    | 8     | 71     | -      | 70    | 4     | 74     | -      | 49    | 13    | 62     | -      | 68    | 5     | 73 |
|          |           | Upper   | -      | 7     | 17    | 24     | -      | 5     | 8     | 13     | -      | 7     | 4     | 11     | -      | 5     | 13    | 18     | -      | 4     | 5     | 9  |
|          |           | Total   | -      | 73    | 21    | 94     | -      | 68    | 16    | 84     | -      | 77    | 8     | 85     | -      | 54    | 26    | 80     | -      | 72    | 10    | 82 |
|          | Week 20   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 68    | 3     | 71     | -      | 60    | 8     | 68     | -      | 69    | 3     | 72     | -      | 48    | 15    | 63     | -      | 63    | 6     | 69 |
|          |           | Upper   | -      | 5     | 18    | 23     | -      | 8     | 8     | 16     | -      | 8     | 5     | 13     | -      | 5     | 11    | 16     | -      | 7     | 4     | 11 |
|          |           | Total   | -      | 73    | 21    | 94     | -      | 68    | 16    | 84     | -      | 77    | 8     | 85     | -      | 53    | 26    | 79     | -      | 70    | 10    | 80 |
|          | Week 24   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 64    | 4     | 68     | -      | 60    | 7     | 67     | -      | 70    | 2     | 72     | -      | 52    | 16    | 68     | -      | 65    | 7     | 72 |
|          |           | Upper   | -      | 8     | 17    | 25     | -      | 8     | 9     | 17     | -      | 6     | 6     | 12     | -      | 1     | 9     | 10     | -      | 5     | 2     | 7  |
|          |           | Total   | -      | 72    | 21    | 93     | -      | 68    | 16    | 84     | -      | 76    | 8     | 84     | -      | 53    | 25    | 78     | -      | 70    | 9     | 79 |
|          | Week 28   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 65    | 7     | 72     | -      | 62    | 8     | 70     | -      | 69    | 4     | 73     | -      | 50    | 17    | 67     | -      | 67    | 6     | 73 |
|          |           | Upper   | -      | 7     | 14    | 21     | -      | 5     | 8     | 13     | -      | 7     | 4     | 11     | -      | 2     | 8     | 10     | -      | 2     | 4     | 6  |
|          |           | Total   | -      | 72    | 21    | 93     | -      | 67    | 16    | 83     | -      | 76    | 8     | 84     | -      | 52    | 25    | 77     | -      | 69    | 10    | 79 |
|          | Follow-up | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|          |           | Normal  | -      | 69    | 7     | 76     | -      | 63    | 10    | 73     | -      | 77    | 4     | 81     | -      | 57    | 19    | 76     | -      | 74    | 7     | 81 |
|          |           | Upper   | -      | 7     | 14    | 21     | -      | 5     | 8     | 13     | -      | 7     | 4     | 11     | -      | 2     | 9     | 11     | -      | 2     | 5     | 7  |
|          |           | Total   | -      | 76    | 21    | 97     | -      | 68    | 18    | 86     | -      | 84    | 8     | 92     | -      | 59    | 28    | 87     | -      | 76    | 12    | 88 |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (5/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| γ-GTP     | Week 1 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 22    | 1     | 23     | -      | 24    | 2     | 26     | -      | 28    | -     | 28     | -      | 28    | 1     | 29     | -      | 31    | 2     | 33 |
|           |        | Upper   | -      | 2     | 14    | 16     | -      | 2     | 9     | 11     | -      | 1     | 10    | 11     | -      | -     | 8     | 8      | -      | 1     | 13    | 14 |
|           |        | Total   | -      | 24    | 15    | 39     | -      | 26    | 11    | 37     | -      | 29    | 10    | 39     | -      | 28    | 9     | 37     | -      | 32    | 15    | 47 |
|           | Week 2 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 59    | 6     | 65     | -      | 56    | 4     | 60     | -      | 60    | 4     | 64     | -      | 60    | 6     | 66     | -      | 57    | 1     | 58 |
|           |        | Upper   | -      | 3     | 28    | 31     | -      | 3     | 26    | 29     | -      | 4     | 23    | 27     | -      | 3     | 24    | 27     | -      | 2     | 29    | 31 |
|           |        | Total   | -      | 62    | 34    | 96     | -      | 59    | 30    | 89     | -      | 64    | 27    | 91     | -      | 63    | 30    | 93     | -      | 59    | 30    | 89 |
|           | Week 3 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 21    | 3     | 24     | -      | 23    | 2     | 25     | -      | 27    | 2     | 29     | -      | 25    | 2     | 27     | -      | 28    | 1     | 29 |
|           |        | Upper   | -      | 1     | 11    | 12     | -      | 3     | 8     | 11     | -      | 2     | 7     | 9      | -      | -     | 5     | 5      | -      | -     | 13    | 13 |
|           |        | Total   | -      | 22    | 14    | 36     | -      | 26    | 10    | 36     | -      | 29    | 9     | 38     | -      | 25    | 7     | 32     | -      | 28    | 14    | 42 |
|           | Week 4 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 60    | 6     | 66     | -      | 57    | 5     | 62     | -      | 60    | 6     | 66     | -      | 58    | 5     | 63     | -      | 56    | 3     | 59 |
|           |        | Upper   | -      | 3     | 26    | 29     | -      | 2     | 25    | 27     | -      | 2     | 21    | 23     | -      | 3     | 23    | 26     | -      | 1     | 27    | 28 |
|           |        | Total   | -      | 63    | 32    | 95     | -      | 59    | 30    | 89     | -      | 62    | 27    | 89     | -      | 61    | 28    | 89     | -      | 57    | 30    | 87 |
|           | Week 6 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 23    | 5     | 28     | -      | 26    | 2     | 28     | -      | 27    | 2     | 29     | -      | 23    | 2     | 25     | -      | 28    | 2     | 30 |
|           |        | Upper   | -      | 1     | 11    | 12     | -      | 1     | 11    | 12     | -      | 1     | 8     | 9      | -      | -     | 6     | 6      | -      | 2     | 15    | 17 |
|           |        | Total   | -      | 24    | 16    | 40     | -      | 27    | 13    | 40     | -      | 28    | 10    | 38     | -      | 23    | 8     | 31     | -      | 30    | 17    | 47 |
|           | Week 8 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 61    | 5     | 66     | -      | 53    | 5     | 58     | -      | 57    | 6     | 63     | -      | 56    | 5     | 61     | -      | 54    | 3     | 57 |
|           |        | Upper   | -      | 1     | 27    | 28     | -      | 4     | 24    | 28     | -      | 5     | 21    | 26     | -      | 4     | 21    | 25     | -      | 2     | 25    | 27 |
|           |        | Total   | -      | 62    | 32    | 94     | -      | 57    | 29    | 86     | -      | 62    | 27    | 89     | -      | 60    | 26    | 86     | -      | 56    | 28    | 84 |
| Week 12   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 59     | 4     | 63    | -      | 54     | 5     | 59    | -      | 56     | 6     | 62    | -      | 55     | 8     | 63    | -      | 53     | 6     | 59    |    |
|           | Upper  | -       | 3      | 28    | 31    | -      | 3      | 24    | 27    | -      | 4      | 20    | 24    | -      | 4      | 15    | 19    | -      | 2      | 22    | 24    |    |
|           | Total  | -       | 62     | 32    | 94    | -      | 57     | 29    | 86    | -      | 60     | 26    | 86    | -      | 59     | 23    | 82    | -      | 55     | 28    | 83    |    |
| Week 16   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 58     | 6     | 64    | -      | 54     | 6     | 60    | -      | 55     | 8     | 63    | -      | 55     | 7     | 62    | -      | 51     | 4     | 55    |    |
|           | Upper  | -       | 4      | 26    | 30    | -      | 2      | 22    | 24    | -      | 4      | 18    | 22    | -      | 4      | 14    | 18    | -      | 4      | 23    | 27    |    |
|           | Total  | -       | 62     | 32    | 94    | -      | 56     | 28    | 84    | -      | 59     | 26    | 85    | -      | 59     | 21    | 80    | -      | 55     | 27    | 82    |    |
| Week 20   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 59     | 6     | 65    | -      | 50     | 3     | 53    | -      | 55     | 10    | 65    | -      | 56     | 6     | 62    | -      | 53     | 4     | 57    |    |
|           | Upper  | -       | 3      | 26    | 29    | -      | 6      | 25    | 31    | -      | 4      | 16    | 20    | -      | 2      | 15    | 17    | -      | 1      | 22    | 23    |    |
|           | Total  | -       | 62     | 32    | 94    | -      | 56     | 28    | 84    | -      | 59     | 26    | 85    | -      | 58     | 21    | 79    | -      | 54     | 26    | 80    |    |
| Week 24   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 58     | 5     | 63    | -      | 52     | 4     | 56    | -      | 54     | 7     | 61    | -      | 57     | 7     | 64    | -      | 52     | 5     | 57    |    |
|           | Upper  | -       | 3      | 27    | 30    | -      | 4      | 24    | 28    | -      | 5      | 18    | 23    | -      | 1      | 13    | 14    | -      | 2      | 20    | 22    |    |
|           | Total  | -       | 61     | 32    | 93    | -      | 56     | 28    | 84    | -      | 59     | 25    | 84    | -      | 58     | 20    | 78    | -      | 54     | 25    | 79    |    |
| Week 28   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 58     | 4     | 62    | -      | 52     | 6     | 58    | -      | 54     | 9     | 63    | -      | 56     | 9     | 65    | -      | 51     | 7     | 58    |    |
|           | Upper  | -       | 3      | 28    | 31    | -      | 3      | 22    | 25    | -      | 5      | 16    | 21    | -      | 1      | 11    | 12    | -      | 2      | 19    | 21    |    |
|           | Total  | -       | 61     | 32    | 93    | -      | 55     | 28    | 83    | -      | 59     | 25    | 84    | -      | 57     | 20    | 77    | -      | 53     | 26    | 79    |    |
| Follow-up | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 60     | 4     | 64    | -      | 53     | 6     | 59    | -      | 59     | 10    | 69    | -      | 60     | 11    | 71    | -      | 55     | 8     | 63    |    |
|           | Upper  | -       | 3      | 30    | 33    | -      | 3      | 24    | 27    | -      | 6      | 17    | 23    | -      | 1      | 15    | 16    | -      | 3      | 22    | 25    |    |
|           | Total  | -       | 63     | 34    | 97    | -      | 56     | 30    | 86    | -      | 65     | 27    | 92    | -      | 61     | 26    | 87    | -      | 58     | 30    | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (6/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| ALP       | Week 1 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     |    |
|           |        | Normal  | -      | 32    | 2     | 34     | -      | 33    | 2     | 35     | -      | 37    | -     | 37     | -      | 34    | 1     | 35     | -      | 41    | 1     | 42 |
|           |        | Upper   | -      | 2     | 2     | 4      | -      | 1     | 1     | 2      | -      | -     | 2     | 2      | -      | 1     | 1     | 2      | -      | -     | 4     | 4  |
|           |        | Total   | 1      | 34    | 4     | 39     | -      | 34    | 3     | 37     | -      | 37    | 2     | 39     | -      | 35    | 2     | 37     | 1      | 41    | 5     | 47 |
|           | Week 2 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     |    |
|           |        | Normal  | -      | 81    | 4     | 85     | -      | 75    | 4     | 79     | -      | 83    | 2     | 85     | -      | 81    | -     | 81     | -      | 76    | 5     | 81 |
|           |        | Upper   | -      | 3     | 7     | 10     | -      | 4     | 6     | 10     | -      | -     | 6     | 6      | -      | 3     | 9     | 12     | -      | 1     | 6     | 7  |
|           |        | Total   | 1      | 84    | 11    | 96     | -      | 79    | 10    | 89     | -      | 83    | 8     | 91     | -      | 84    | 9     | 93     | 1      | 77    | 11    | 89 |
|           | Week 3 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     |    |
|           |        | Normal  | -      | 28    | 4     | 32     | -      | 31    | 3     | 34     | -      | 36    | -     | 36     | -      | 30    | 2     | 32     | -      | 36    | 1     | 37 |
|           |        | Upper   | -      | 3     | 1     | 4      | -      | 2     | -     | 2      | -      | 1     | 1     | 2      | -      | -     | -     | -      | -      | -     | 4     | 4  |
|           |        | Total   | 1      | 31    | 5     | 37     | -      | 33    | 3     | 36     | -      | 37    | 1     | 38     | -      | 30    | 2     | 32     | 1      | 36    | 5     | 42 |
| Week 4    | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | -      | 1     |       |    |
|           | Normal | -       | 81     | 5     | 86    | -      | 76     | 2     | 78    | -      | 80     | 1     | 81    | -      | 79     | 2     | 81    | -      | 74     | 5     | 79    |    |
|           | Upper  | -       | 2      | 6     | 8     | -      | 3      | 8     | 11    | -      | 2      | 6     | 8     | -      | 3      | 5     | 8     | -      | 1      | 6     | 7     |    |
|           | Total  | 1       | 83     | 11    | 95    | -      | 79     | 10    | 89    | -      | 82     | 7     | 89    | -      | 82     | 7     | 89    | 1      | 75     | 11    | 87    |    |
| Week 6    | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | -      | 2     |       |    |
|           | Normal | -       | 34     | 1     | 35    | -      | 35     | 3     | 38    | -      | 34     | -     | 34    | -      | 30     | -     | 30    | -      | 39     | 1     | 40    |    |
|           | Upper  | -       | 1      | 3     | 4     | -      | 1      | 1     | 2     | -      | 1      | 3     | 4     | -      | 2      | -     | 2     | -      | 1      | 4     | 5     |    |
|           | Total  | 1       | 35     | 4     | 40    | -      | 36     | 4     | 40    | -      | 35     | 3     | 38    | -      | 32     | -     | 32    | 1      | 41     | 5     | 47    |    |
| Week 8    | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | -      | 1     |       |    |
|           | Normal | -       | 77     | 2     | 79    | -      | 74     | 3     | 77    | -      | 79     | 1     | 80    | -      | 73     | 2     | 75    | -      | 70     | 4     | 74    |    |
|           | Upper  | -       | 4      | 9     | 13    | -      | 2      | 7     | 9     | -      | 3      | 6     | 9     | -      | 6      | 5     | 11    | -      | 2      | 7     | 9     |    |
|           | Total  | 1       | 82     | 11    | 94    | -      | 76     | 10    | 86    | -      | 82     | 7     | 89    | -      | 79     | 7     | 86    | 1      | 72     | 11    | 84    |    |
| Week 12   | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 79     | 5     | 84    | -      | 75     | 4     | 79    | -      | 80     | 2     | 82    | -      | 68     | 2     | 70    | 1      | 67     | 3     | 71    |    |
|           | Upper  | -       | 3      | 6     | 9     | -      | 1      | 6     | 7     | -      | -      | 4     | 4     | -      | 7      | 5     | 12    | -      | 4      | 8     | 12    |    |
|           | Total  | 1       | 82     | 11    | 94    | -      | 76     | 10    | 86    | -      | 80     | 6     | 86    | -      | 75     | 7     | 82    | 1      | 71     | 11    | 83    |    |
| Week 16   | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | -      | 1     |       |    |
|           | Normal | -       | 79     | 3     | 82    | -      | 72     | 6     | 78    | -      | 77     | 2     | 79    | -      | 67     | 1     | 68    | -      | 65     | 3     | 68    |    |
|           | Upper  | -       | 3      | 8     | 11    | -      | 2      | 4     | 6     | -      | 2      | 4     | 6     | -      | 8      | 4     | 12    | -      | 6      | 7     | 13    |    |
|           | Total  | 1       | 82     | 11    | 94    | -      | 74     | 10    | 84    | -      | 79     | 6     | 85    | -      | 75     | 5     | 80    | 1      | 71     | 10    | 82    |    |
| Week 20   | Lower  | 1       | -      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | -      | 2     |       |    |
|           | Normal | -       | 80     | 7     | 87    | -      | 69     | 5     | 74    | -      | 75     | 2     | 77    | -      | 68     | 2     | 70    | -      | 66     | 3     | 69    |    |
|           | Upper  | -       | 2      | 4     | 6     | -      | 5      | 5     | 10    | -      | 4      | 4     | 8     | -      | 6      | 3     | 9     | -      | 2      | 7     | 9     |    |
|           | Total  | 1       | 82     | 11    | 94    | -      | 74     | 10    | 84    | -      | 79     | 6     | 85    | -      | 74     | 5     | 79    | 1      | 69     | 10    | 80    |    |
| Week 24   | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | -      | 1     |       |    |
|           | Normal | -       | 76     | 3     | 79    | -      | 68     | 8     | 76    | -      | 75     | 2     | 77    | -      | 68     | 3     | 71    | -      | 65     | 3     | 68    |    |
|           | Upper  | -       | 4      | 8     | 12    | -      | 6      | 2     | 8     | -      | 4      | 3     | 7     | -      | 6      | 1     | 7     | -      | 3      | 7     | 10    |    |
|           | Total  | 1       | 81     | 11    | 93    | -      | 74     | 10    | 84    | -      | 79     | 5     | 84    | -      | 74     | 4     | 78    | 1      | 68     | 10    | 79    |    |
| Week 28   | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 76     | 2     | 78    | -      | 71     | 5     | 76    | -      | 78     | 2     | 80    | -      | 68     | 2     | 70    | 1      | 65     | 5     | 71    |    |
|           | Upper  | -       | 4      | 9     | 13    | -      | 2      | 5     | 7     | -      | 1      | 3     | 4     | -      | 5      | 2     | 7     | -      | 2      | 5     | 7     |    |
|           | Total  | 1       | 81     | 11    | 93    | -      | 73     | 10    | 83    | -      | 79     | 5     | 84    | -      | 73     | 4     | 77    | 1      | 67     | 10    | 78    |    |
| Follow-up | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 80     | 2     | 82    | -      | 75     | 5     | 80    | -      | 84     | 2     | 86    | -      | 74     | 3     | 77    | 1      | 73     | 6     | 80    |    |
|           | Upper  | -       | 4      | 9     | 13    | -      | 2      | 4     | 6     | -      | 1      | 5     | 6     | -      | 6      | 4     | 10    | -      | 3      | 5     | 8     |    |
|           | Total  | 1       | 85     | 11    | 97    | -      | 77     | 9     | 86    | -      | 85     | 7     | 92    | -      | 80     | 7     | 87    | 1      | 76     | 11    | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (7/24)

|     |           | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----|-----------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|     |           | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|     |           | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| LDH | Week 1    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|     |           | Normal  | -      | 30    | 2     | 32     | -      | 30    | 1     | 31     | -      | 34    | 1     | 35     | -      | 31    | 1     | 32     | -      | 38    | 4     | 42 |
|     |           | Upper   | -      | 2     | 5     | 7      | -      | 2     | 3     | 5      | -      | 2     | 2     | 4      | -      | 1     | 2     | 3      | -      | 3     | 2     | 5  |
|     |           | Total   | -      | 32    | 7     | 39     | -      | 33    | 4     | 37     | -      | 36    | 3     | 39     | -      | 34    | 3     | 37     | -      | 41    | 6     | 47 |
|     | Week 2    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     | -  |
|     |           | Normal  | -      | 75    | 6     | 81     | -      | 79    | 3     | 82     | -      | 83    | 3     | 86     | -      | 80    | 1     | 81     | -      | 80    | 3     | 83 |
|     |           | Upper   | -      | 5     | 10    | 15     | -      | -     | 7     | 7      | -      | 2     | 3     | 5      | -      | 4     | 7     | 11     | -      | 2     | 4     | 6  |
|     |           | Total   | -      | 80    | 16    | 96     | -      | 79    | 10    | 89     | -      | 85    | 6     | 91     | 1      | 84    | 8     | 93     | -      | 82    | 7     | 89 |
|     | Week 3    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -  |
|     |           | Normal  | -      | 28    | 1     | 29     | -      | 32    | -     | 32     | -      | 35    | -     | 35     | -      | 29    | -     | 29     | -      | 35    | 3     | 38 |
|     |           | Upper   | -      | 2     | 5     | 7      | -      | 1     | 3     | 4      | -      | 1     | 2     | 3      | -      | 2     | -     | 2      | -      | 1     | 3     | 4  |
|     |           | Total   | -      | 30    | 6     | 36     | -      | 33    | 3     | 36     | -      | 36    | 2     | 38     | -      | 32    | -     | 32     | -      | 36    | 6     | 42 |
|     | Week 4    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | 2     | 3     | -      | -      | -     | -     | -  |
|     |           | Normal  | -      | 77    | 6     | 83     | -      | 76    | 3     | 79     | -      | 81    | 3     | 84     | -      | 75    | 5     | 80     | -      | 75    | 2     | 77 |
|     |           | Upper   | -      | 3     | 9     | 12     | -      | 3     | 7     | 10     | -      | 2     | 3     | 5      | -      | 3     | 3     | 6      | -      | 5     | 5     | 10 |
|     |           | Total   | -      | 80    | 15    | 95     | -      | 79    | 10    | 89     | -      | 83    | 6     | 89     | 1      | 80    | 8     | 89     | -      | 80    | 7     | 87 |
|     | Week 6    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -  |
|     |           | Normal  | -      | 31    | 2     | 33     | -      | 33    | 1     | 34     | -      | 36    | -     | 36     | -      | 28    | -     | 28     | -      | 39    | 1     | 40 |
|     |           | Upper   | -      | 1     | 6     | 7      | -      | 3     | 3     | 6      | -      | 1     | 1     | 2      | -      | 2     | -     | 2      | -      | 2     | 5     | 7  |
|     |           | Total   | -      | 32    | 8     | 40     | -      | 36    | 4     | 40     | -      | 37    | 1     | 38     | -      | 31    | -     | 31     | -      | 41    | 6     | 47 |
|     | Week 8    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | 1     | -     | 2      | -      | -     | -     | -  |
|     |           | Normal  | -      | 75    | 5     | 80     | -      | 73    | 4     | 77     | -      | 79    | 4     | 83     | -      | 74    | 3     | 77     | -      | 75    | 1     | 76 |
|     |           | Upper   | -      | 4     | 10    | 14     | -      | 3     | 6     | 9      | -      | 4     | 2     | 6      | -      | 3     | 4     | 7      | -      | 2     | 6     | 8  |
|     |           | Total   | -      | 79    | 15    | 94     | -      | 76    | 10    | 86     | -      | 83    | 6     | 89     | 1      | 78    | 7     | 86     | -      | 77    | 7     | 84 |
|     | Week 12   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|     |           | Normal  | -      | 77    | 3     | 80     | -      | 72    | 2     | 74     | -      | 73    | 3     | 76     | 1      | 74    | 2     | 77     | -      | 72    | 2     | 74 |
|     |           | Upper   | -      | 2     | 12    | 14     | -      | 4     | 8     | 12     | -      | 6     | 3     | 9      | -      | 1     | 4     | 5      | -      | 4     | 5     | 9  |
|     |           | Total   | -      | 79    | 15    | 94     | -      | 76    | 10    | 86     | -      | 79    | 6     | 85     | 1      | 75    | 6     | 82     | -      | 76    | 7     | 83 |
|     | Week 16   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | 2     | -     | 3      | -      | -     | -     | -  |
|     |           | Normal  | -      | 74    | 8     | 82     | -      | 71    | 3     | 74     | -      | 75    | 2     | 77     | -      | 69    | 1     | 70     | -      | 71    | -     | 71 |
|     |           | Upper   | -      | 5     | 7     | 12     | -      | 3     | 7     | 10     | -      | 4     | 4     | 8      | -      | 3     | 4     | 7      | -      | 5     | 6     | 11 |
|     |           | Total   | -      | 79    | 15    | 94     | -      | 74    | 10    | 84     | -      | 79    | 6     | 85     | 1      | 74    | 5     | 80     | -      | 76    | 6     | 82 |
|     | Week 20   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | -  |
|     |           | Normal  | -      | 76    | 6     | 82     | -      | 69    | 1     | 70     | -      | 75    | 2     | 77     | -      | 72    | -     | 72     | -      | 70    | 3     | 73 |
|     |           | Upper   | -      | 3     | 9     | 12     | -      | 5     | 9     | 14     | -      | 4     | 4     | 8      | -      | 1     | 5     | 6      | -      | 4     | 3     | 7  |
|     |           | Total   | -      | 79    | 15    | 94     | -      | 74    | 10    | 84     | -      | 79    | 6     | 85     | 1      | 73    | 5     | 79     | -      | 74    | 6     | 80 |
|     | Week 24   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | 1     | -     | 2      | -      | 1     | -     | 1  |
|     |           | Normal  | -      | 75    | 7     | 82     | -      | 68    | 3     | 71     | -      | 76    | 2     | 78     | -      | 70    | -     | 70     | -      | 70    | 1     | 71 |
|     |           | Upper   | -      | 3     | 8     | 11     | -      | 6     | 7     | 13     | -      | 2     | 4     | 6      | -      | 1     | 5     | 6      | -      | 2     | 5     | 7  |
|     |           | Total   | -      | 78    | 15    | 93     | -      | 74    | 10    | 84     | -      | 78    | 6     | 84     | 1      | 72    | 5     | 78     | -      | 73    | 6     | 79 |
|     | Week 28   | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | 1     | -     | 2      | -      | -     | -     | -  |
|     |           | Normal  | -      | 76    | 6     | 82     | -      | 70    | 4     | 74     | -      | 75    | 3     | 78     | -      | 68    | -     | 68     | -      | 68    | 1     | 69 |
|     |           | Upper   | -      | 2     | 9     | 11     | -      | 3     | 6     | 9      | -      | 3     | 3     | 6      | -      | 2     | 5     | 7      | -      | 4     | 5     | 9  |
|     |           | Total   | -      | 78    | 15    | 93     | -      | 73    | 10    | 83     | -      | 78    | 6     | 84     | 1      | 71    | 5     | 77     | -      | 72    | 6     | 78 |
|     | Follow-up | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | 1      | 1     | -     | 2      | -      | -     | -     | -  |
|     |           | Normal  | -      | 79    | 6     | 85     | -      | 72    | 4     | 76     | -      | 82    | 4     | 86     | -      | 76    | 2     | 78     | -      | 77    | 2     | 79 |
|     |           | Upper   | -      | 2     | 10    | 12     | -      | 3     | 6     | 9      | -      | 3     | 3     | 6      | -      | 2     | 5     | 7      | -      | 4     | 5     | 9  |
|     |           | Total   | -      | 81    | 16    | 97     | -      | 76    | 10    | 86     | -      | 85    | 7     | 92     | 1      | 79    | 7     | 87     | -      | 81    | 7     | 88 |



表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (8/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| CPK       | Week 1 | Lower   | 2      | -     | -     | 2      | 1      | 1     | -     | 2      | 2      | 1     | -     | 3      | 1      | 1     | -     | 2      | -      | 2     | -     | 2  |
|           |        | Normal  | -      | 31    | 4     | 35     | -      | 26    | 1     | 27     | 2      | 29    | 1     | 32     | -      | 26    | 3     | 29     | 1      | 38    | 1     | 40 |
|           |        | Upper   | -      | -     | 2     | 2      | -      | 1     | 7     | 8      | -      | 1     | 3     | 4      | -      | 4     | 2     | 6      | -      | 2     | 3     | 5  |
|           |        | Total   | 2      | 31    | 6     | 39     | 1      | 28    | 8     | 37     | 4      | 31    | 4     | 39     | 1      | 31    | 5     | 37     | 1      | 42    | 4     | 47 |
|           | Week 2 | Lower   | 6      | -     | -     | 6      | 1      | -     | -     | 1      | 2      | 3     | -     | 5      | 1      | 6     | -     | 7      | 1      | 2     | -     | 3  |
|           |        | Normal  | 6      | 68    | 4     | 78     | 3      | 61    | 8     | 72     | 1      | 70    | 3     | 74     | 1      | 68    | 8     | 77     | 1      | 77    | 1     | 79 |
|           |        | Upper   | -      | 5     | 7     | 12     | -      | 7     | 9     | 16     | -      | 6     | 6     | 12     | -      | 5     | 4     | 9      | -      | 2     | 5     | 7  |
|           |        | Total   | 12     | 73    | 11    | 96     | 4      | 68    | 17    | 89     | 3      | 79    | 9     | 91     | 2      | 79    | 12    | 93     | 2      | 81    | 6     | 89 |
|           | Week 3 | Lower   | 1      | -     | -     | 1      | 1      | 1     | -     | 2      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -  |
|           |        | Normal  | 1      | 26    | 3     | 30     | -      | 25    | 3     | 28     | 4      | 29    | -     | 33     | 1      | 26    | -     | 27     | 1      | 38    | 2     | 41 |
|           |        | Upper   | -      | 4     | 2     | 6      | -      | 1     | 5     | 6      | -      | 2     | 3     | 5      | -      | 2     | 2     | 4      | -      | 1     | -     | 1  |
|           |        | Total   | 2      | 30    | 5     | 37     | 1      | 27    | 8     | 36     | 4      | 31    | 3     | 38     | 1      | 29    | 2     | 32     | 1      | 39    | 2     | 42 |
|           | Week 4 | Lower   | 9      | -     | -     | 9      | 1      | 1     | -     | 2      | 1      | -     | -     | 1      | 1      | 5     | -     | 6      | -      | 2     | -     | 2  |
|           |        | Normal  | 2      | 71    | 3     | 76     | 3      | 65    | 8     | 76     | 2      | 72    | 4     | 78     | 1      | 68    | 4     | 73     | 2      | 75    | 1     | 78 |
|           |        | Upper   | -      | 2     | 8     | 10     | -      | 2     | 9     | 11     | -      | 5     | 5     | 10     | -      | 3     | 7     | 10     | -      | 2     | 5     | 7  |
|           |        | Total   | 11     | 73    | 11    | 95     | 4      | 68    | 17    | 89     | 3      | 77    | 9     | 89     | 2      | 76    | 11    | 89     | 2      | 79    | 6     | 87 |
|           | Week 6 | Lower   | 2      | -     | -     | 2      | 1      | 1     | -     | 2      | 1      | 1     | -     | 2      | 1      | -     | -     | 1      | -      | 1     | -     | 1  |
|           |        | Normal  | -      | 32    | 3     | 35     | -      | 29    | 2     | 31     | 2      | 30    | 1     | 33     | 1      | 25    | 1     | 27     | 1      | 39    | 1     | 41 |
|           |        | Upper   | -      | 1     | 2     | 3      | -      | -     | 7     | 7      | -      | 1     | 2     | 3      | -      | 1     | 2     | 3      | -      | 3     | 2     | 5  |
|           |        | Total   | 2      | 33    | 5     | 40     | 1      | 30    | 9     | 40     | 3      | 32    | 3     | 38     | 2      | 26    | 3     | 31     | 1      | 43    | 3     | 47 |
|           | Week 8 | Lower   | 7      | 1     | -     | 8      | 3      | 2     | -     | 5      | 2      | 3     | -     | 5      | -      | 1     | -     | 1      | 1      | -     | -     | 1  |
|           |        | Normal  | 3      | 69    | 5     | 77     | 1      | 59    | 8     | 68     | 1      | 69    | 4     | 74     | 2      | 68    | 8     | 78     | 1      | 71    | 1     | 73 |
|           |        | Upper   | -      | 3     | 6     | 9      | -      | 4     | 9     | 13     | -      | 5     | 5     | 10     | -      | 4     | 3     | 7      | -      | 5     | 5     | 10 |
|           |        | Total   | 10     | 73    | 11    | 94     | 4      | 65    | 17    | 86     | 3      | 77    | 9     | 89     | 2      | 73    | 11    | 86     | 2      | 76    | 6     | 84 |
| Week 12   | Lower  | 6       | -      | -     | 6     | 2      | 2      | -     | 4     | 1      | 1      | -     | 2     | -      | 1      | -     | 1     | -      | 1      | -     | 1     |    |
|           | Normal | 4       | 70     | 3     | 77    | 2      | 60     | 8     | 70    | 2      | 69     | 3     | 74    | 2      | 67     | 6     | 75    | 2      | 71     | -     | 73    |    |
|           | Upper  | -       | 3      | 8     | 11    | -      | 3      | 9     | 12    | -      | 4      | 5     | 9     | -      | 1      | 5     | 6     | -      | 3      | 6     | 9     |    |
|           | Total  | 10      | 73     | 11    | 94    | 4      | 65     | 17    | 86    | 3      | 74     | 8     | 85    | 2      | 69     | 11    | 82    | 2      | 75     | 6     | 83    |    |
| Week 16   | Lower  | 3       | -      | -     | 3     | 2      | 1      | -     | 3     | 1      | 1      | -     | 2     | 1      | 1      | -     | 2     | -      | 4      | -     | 4     |    |
|           | Normal | 7       | 67     | 4     | 78    | 2      | 57     | 8     | 67    | 2      | 66     | 2     | 70    | 1      | 65     | 3     | 69    | 2      | 66     | -     | 68    |    |
|           | Upper  | -       | 6      | 7     | 13    | -      | 5      | 9     | 14    | -      | 7      | 6     | 13    | -      | 2      | 7     | 9     | -      | 4      | 6     | 10    |    |
|           | Total  | 10      | 73     | 11    | 94    | 4      | 63     | 17    | 84    | 3      | 74     | 8     | 85    | 2      | 68     | 10    | 80    | 2      | 74     | 6     | 82    |    |
| Week 20   | Lower  | 6       | -      | -     | 6     | 2      | 1      | -     | 3     | -      | 1      | -     | 1     | 1      | -      | -     | 1     | 1      | 3      | -     | 4     |    |
|           | Normal | 4       | 68     | 4     | 76    | 2      | 53     | 6     | 61    | 3      | 66     | 3     | 72    | 1      | 63     | 4     | 68    | 1      | 64     | 2     | 67    |    |
|           | Upper  | -       | 5      | 7     | 12    | -      | 9      | 11    | 20    | -      | 7      | 5     | 12    | -      | 4      | 6     | 10    | -      | 5      | 4     | 9     |    |
|           | Total  | 10      | 73     | 11    | 94    | 4      | 63     | 17    | 84    | 3      | 74     | 8     | 85    | 2      | 67     | 10    | 79    | 2      | 72     | 6     | 80    |    |
| Week 24   | Lower  | 4       | -      | -     | 4     | 2      | 1      | -     | 3     | 1      | 2      | -     | 3     | -      | 1      | -     | 1     | -      | 3      | -     | 3     |    |
|           | Normal | 6       | 68     | 5     | 79    | 2      | 56     | 8     | 66    | 2      | 66     | -     | 68    | 2      | 62     | 5     | 69    | 2      | 62     | -     | 64    |    |
|           | Upper  | -       | 4      | 6     | 10    | -      | 6      | 9     | 15    | -      | 5      | 8     | 13    | -      | 3      | 5     | 8     | -      | 6      | 6     | 12    |    |
|           | Total  | 10      | 72     | 11    | 93    | 4      | 63     | 17    | 84    | 3      | 73     | 8     | 84    | 2      | 66     | 10    | 78    | 2      | 71     | 6     | 79    |    |
| Week 28   | Lower  | 6       | -      | -     | 6     | 2      | 2      | -     | 4     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | 4      | -     | 4     |    |
|           | Normal | 4       | 68     | 6     | 78    | 2      | 55     | 11    | 68    | 3      | 69     | 2     | 74    | 2      | 61     | 6     | 69    | 2      | 64     | -     | 66    |    |
|           | Upper  | -       | 4      | 5     | 9     | -      | 5      | 6     | 11    | -      | 2      | 6     | 8     | -      | 3      | 4     | 7     | -      | 2      | 6     | 8     |    |
|           | Total  | 10      | 72     | 11    | 93    | 4      | 62     | 17    | 83    | 3      | 73     | 8     | 84    | 2      | 65     | 10    | 77    | 2      | 70     | 6     | 78    |    |
| Follow-up | Lower  | 7       | -      | -     | 7     | 2      | 3      | -     | 5     | -      | 2      | -     | 2     | -      | 3      | -     | 3     | -      | 5      | -     | 5     |    |
|           | Normal | 5       | 70     | 6     | 81    | 2      | 57     | 11    | 70    | 4      | 74     | 3     | 81    | 2      | 68     | 7     | 77    | 2      | 72     | -     | 74    |    |
|           | Upper  | -       | 4      | 5     | 9     | -      | 5      | 6     | 11    | -      | 2      | 7     | 9     | -      | 3      | 4     | 7     | -      | 3      | 6     | 9     |    |
|           | Total  | 12      | 74     | 11    | 97    | 4      | 65     | 17    | 86    | 4      | 78     | 10    | 92    | 2      | 74     | 11    | 87    | 2      | 80     | 6     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (9/24)

|           |           | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|-----------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |           | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |           | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Uric Acid | Week 1    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 34    | 1     | 35     | -      | 28    | 1     | 29     | -      | 32    | 3     | 35     | -      | 29    | 1     | 30     | -      | 38    | 2     | 40 |
|           |           | Upper   | -      | -     | 4     | 4      | -      | -     | 8     | 8      | -      | -     | 4     | 4      | -      | -     | 7     | 7      | -      | 1     | 6     | 7  |
|           |           | Total   | -      | 34    | 5     | 39     | -      | 28    | 9     | 37     | -      | 32    | 7     | 39     | -      | 29    | 8     | 37     | -      | 39    | 8     | 47 |
|           | Week 2    | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 71    | 7     | 78     | -      | 72    | 2     | 74     | -      | 77    | 5     | 82     | -      | 72    | 3     | 75     | -      | 73    | 5     | 78 |
|           |           | Upper   | -      | 8     | 9     | 17     | -      | 4     | 11    | 15     | -      | 1     | 8     | 9      | -      | 7     | 11    | 18     | -      | 6     | 5     | 11 |
|           |           | Total   | 1      | 79    | 16    | 96     | -      | 76    | 13    | 89     | -      | 78    | 13    | 91     | -      | 79    | 14    | 93     | -      | 79    | 10    | 89 |
|           | Week 3    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 30    | -     | 30     | -      | 27    | 3     | 30     | -      | 30    | 2     | 32     | -      | 24    | 1     | 25     | -      | 32    | 3     | 35 |
|           |           | Upper   | -      | 2     | 4     | 6      | -      | -     | 6     | 6      | -      | 1     | 5     | 6      | -      | -     | 7     | 7      | -      | 2     | 5     | 7  |
|           |           | Total   | -      | 32    | 4     | 36     | -      | 27    | 9     | 36     | -      | 31    | 7     | 38     | -      | 24    | 8     | 32     | -      | 34    | 8     | 42 |
|           | Week 4    | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 73    | 7     | 80     | -      | 71    | 4     | 75     | -      | 73    | 4     | 77     | -      | 71    | 4     | 75     | -      | 73    | 4     | 77 |
|           |           | Upper   | -      | 5     | 9     | 14     | -      | 5     | 9     | 14     | -      | 3     | 9     | 12     | -      | 4     | 10    | 14     | -      | 4     | 6     | 10 |
|           |           | Total   | 1      | 78    | 16    | 95     | -      | 76    | 13    | 89     | -      | 76    | 13    | 89     | -      | 75    | 14    | 89     | -      | 77    | 10    | 87 |
|           | Week 6    | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 33    | 4     | 37     | -      | 32    | 3     | 35     | -      | 30    | 3     | 33     | -      | 24    | 2     | 26     | -      | 37    | 5     | 42 |
|           |           | Upper   | -      | -     | 3     | 3      | -      | -     | 5     | 5      | -      | 1     | 4     | 5      | -      | -     | 5     | 5      | -      | 2     | 3     | 5  |
|           |           | Total   | -      | 33    | 7     | 40     | -      | 32    | 8     | 40     | -      | 31    | 7     | 38     | -      | 24    | 7     | 31     | -      | 39    | 8     | 47 |
|           | Week 8    | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 73    | 8     | 81     | -      | 71    | 2     | 73     | -      | 73    | 6     | 79     | -      | 72    | 4     | 76     | -      | 71    | 3     | 74 |
|           |           | Upper   | -      | 4     | 8     | 12     | -      | 3     | 10    | 13     | -      | 3     | 7     | 10     | -      | 1     | 9     | 10     | -      | 3     | 7     | 10 |
|           |           | Total   | 1      | 77    | 16    | 94     | -      | 74    | 12    | 86     | -      | 76    | 13    | 89     | -      | 73    | 13    | 86     | -      | 74    | 10    | 84 |
|           | Week 12   | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 73    | 6     | 79     | -      | 72    | 2     | 74     | -      | 70    | 7     | 77     | -      | 67    | 6     | 73     | -      | 70    | 4     | 74 |
|           |           | Upper   | -      | 4     | 10    | 14     | -      | 2     | 10    | 12     | -      | 3     | 6     | 9      | -      | 2     | 7     | 9      | -      | 3     | 6     | 9  |
|           |           | Total   | 1      | 77    | 16    | 94     | -      | 74    | 12    | 86     | -      | 73    | 13    | 86     | -      | 69    | 13    | 82     | -      | 73    | 10    | 83 |
|           | Week 16   | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 72    | 10    | 82     | -      | 70    | 5     | 75     | -      | 68    | 5     | 73     | -      | 63    | 6     | 69     | -      | 70    | 4     | 74 |
|           |           | Upper   | -      | 5     | 6     | 11     | -      | 3     | 6     | 9      | -      | 4     | 8     | 12     | -      | 4     | 7     | 11     | -      | 2     | 6     | 8  |
|           |           | Total   | 1      | 77    | 16    | 94     | -      | 73    | 11    | 84     | -      | 72    | 13    | 85     | -      | 67    | 13    | 80     | -      | 72    | 10    | 82 |
|           | Week 20   | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 74    | 10    | 84     | -      | 68    | 6     | 74     | -      | 68    | 4     | 72     | -      | 64    | 6     | 70     | -      | 67    | 2     | 69 |
|           |           | Upper   | -      | 3     | 6     | 9      | -      | 5     | 5     | 10     | -      | 4     | 9     | 13     | -      | 2     | 7     | 9      | -      | 3     | 8     | 11 |
|           |           | Total   | 1      | 77    | 16    | 94     | -      | 73    | 11    | 84     | -      | 72    | 13    | 85     | -      | 66    | 13    | 79     | -      | 70    | 10    | 80 |
|           | Week 24   | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 73    | 6     | 79     | -      | 67    | 3     | 70     | -      | 69    | 4     | 73     | -      | 62    | 6     | 68     | -      | 65    | 6     | 71 |
|           |           | Upper   | -      | 3     | 10    | 13     | -      | 6     | 8     | 14     | -      | 1     | 9     | 10     | -      | 3     | 7     | 10     | -      | 4     | 4     | 8  |
|           |           | Total   | 1      | 76    | 16    | 93     | -      | 73    | 11    | 84     | -      | 70    | 13    | 83     | -      | 65    | 13    | 78     | -      | 69    | 10    | 79 |
|           | Week 28   | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 71    | 8     | 79     | -      | 67    | 7     | 74     | -      | 69    | 4     | 73     | -      | 61    | 6     | 67     | -      | 65    | 5     | 70 |
|           |           | Upper   | -      | 5     | 8     | 13     | -      | 5     | 4     | 9      | -      | 2     | 9     | 11     | -      | 3     | 7     | 10     | -      | 3     | 5     | 8  |
|           |           | Total   | 1      | 76    | 16    | 93     | -      | 72    | 11    | 83     | -      | 71    | 13    | 84     | -      | 64    | 13    | 77     | -      | 68    | 10    | 78 |
|           | Follow-up | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |           | Normal  | -      | 75    | 8     | 83     | -      | 68    | 7     | 75     | -      | 77    | 4     | 81     | -      | 69    | 6     | 75     | -      | 75    | 5     | 80 |
|           |           | Upper   | -      | 5     | 8     | 13     | -      | 5     | 6     | 11     | -      | 2     | 9     | 11     | -      | 5     | 7     | 12     | -      | 3     | 5     | 8  |
|           |           | Total   | 1      | 80    | 16    | 97     | -      | 73    | 13    | 86     | -      | 79    | 13    | 92     | -      | 74    | 13    | 87     | -      | 78    | 10    | 88 |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (10/24)

|                 |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|                 |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|                 |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Total Bilirubin | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 38    | -     | 38     | -      | 37    | -     | 37     | -      | 38    | 1     | 39     | -      | 37    | -     | 37     | -      | 46    | 1     | 47 |
|                 |         | Upper   | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|                 |         | Total   | -      | 38    | 1     | 39     | -      | 37    | -     | 37     | -      | 38    | 1     | 39     | -      | 37    | -     | 37     | -      | 46    | 1     | 47 |
|                 | Week 2  | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 89    | 1     | 90     | -      | 85    | 1     | 86     | -      | 87    | 2     | 89     | -      | 90    | -     | 90     | -      | 86    | 1     | 87 |
|                 |         | Upper   | -      | 3     | 3     | 6      | -      | -     | 2     | 2      | -      | 2     | -     | 2      | -      | 2     | 1     | 3      | -      | -     | 2     | 2  |
|                 |         | Total   | -      | 92    | 4     | 96     | -      | 86    | 3     | 89     | -      | 89    | 2     | 91     | -      | 92    | 1     | 93     | -      | 86    | 3     | 89 |
|                 | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 35    | -     | 35     | -      | 35    | 1     | 36     | -      | 37    | 1     | 38     | -      | 32    | -     | 32     | -      | 41    | 1     | 42 |
|                 |         | Upper   | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|                 |         | Total   | -      | 35    | 1     | 36     | -      | 35    | 1     | 36     | -      | 37    | 1     | 38     | -      | 32    | -     | 32     | -      | 41    | 1     | 42 |
|                 | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 88    | 1     | 89     | -      | 85    | 2     | 87     | -      | 84    | 2     | 86     | -      | 87    | -     | 87     | -      | 85    | 2     | 87 |
|                 |         | Upper   | -      | 3     | 3     | 6      | -      | 1     | 1     | 2      | -      | 3     | -     | 3      | -      | 1     | 1     | 2      | -      | -     | -     | -  |
|                 |         | Total   | -      | 91    | 4     | 95     | -      | 86    | 3     | 89     | -      | 87    | 2     | 89     | -      | 88    | 1     | 89     | -      | 85    | 2     | 87 |
|                 | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 38    | -     | 38     | -      | 39    | 1     | 40     | -      | 36    | 1     | 37     | -      | 31    | -     | 31     | -      | 46    | 1     | 47 |
|                 |         | Upper   | -      | 1     | 1     | 2      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -  |
|                 |         | Total   | -      | 39    | 1     | 40     | -      | 39    | 1     | 40     | -      | 37    | 1     | 38     | -      | 31    | -     | 31     | -      | 46    | 1     | 47 |
|                 | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 88    | -     | 88     | -      | 83    | 1     | 84     | -      | 85    | 2     | 87     | -      | 83    | -     | 83     | -      | 82    | 1     | 83 |
|                 |         | Upper   | -      | 2     | 4     | 6      | -      | -     | 2     | 2      | -      | 2     | -     | 2      | -      | 2     | 1     | 3      | -      | -     | 1     | 1  |
|                 |         | Total   | -      | 90    | 4     | 94     | -      | 83    | 3     | 86     | -      | 87    | 2     | 89     | -      | 85    | 1     | 86     | -      | 82    | 2     | 84 |
|                 | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 86    | -     | 86     | -      | 83    | 3     | 86     | -      | 82    | 2     | 84     | -      | 79    | 1     | 80     | -      | 80    | 2     | 82 |
|                 |         | Upper   | -      | 4     | 4     | 8      | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | 2     | -     | 2      | -      | 1     | -     | 1  |
|                 |         | Total   | -      | 90    | 4     | 94     | -      | 83    | 3     | 86     | -      | 84    | 2     | 86     | -      | 81    | 1     | 82     | -      | 81    | 2     | 83 |
|                 | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 87    | -     | 87     | -      | 80    | 2     | 82     | -      | 79    | 2     | 81     | -      | 76    | -     | 76     | -      | 79    | 1     | 80 |
|                 |         | Upper   | -      | 3     | 4     | 7      | -      | 1     | 1     | 2      | -      | 4     | -     | 4      | -      | 3     | 1     | 4      | -      | 1     | 1     | 2  |
|                 |         | Total   | -      | 90    | 4     | 94     | -      | 81    | 3     | 84     | -      | 83    | 2     | 85     | -      | 79    | 1     | 80     | -      | 80    | 2     | 82 |
|                 | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 86    | 1     | 87     | -      | 80    | 3     | 83     | -      | 78    | 1     | 79     | -      | 75    | -     | 75     | -      | 76    | 2     | 78 |
|                 |         | Upper   | -      | 4     | 3     | 7      | -      | 1     | -     | 1      | -      | 4     | 1     | 5      | -      | 3     | 1     | 4      | -      | 2     | -     | 2  |
|                 |         | Total   | -      | 90    | 4     | 94     | -      | 81    | 3     | 84     | -      | 83    | 2     | 85     | -      | 78    | 1     | 79     | -      | 78    | 2     | 80 |
|                 | Week 24 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                 |         | Normal  | -      | 85    | 1     | 86     | -      | 78    | 3     | 81     | -      | 76    | 2     | 78     | -      | 73    | -     | 73     | -      | 76    | 1     | 77 |
|                 |         | Upper   | -      | 4     | 3     | 7      | -      | 3     | -     | 3      | -      | 6     | -     | 6      | -      | 4     | 1     | 5      | -      | 1     | 1     | 2  |
|                 |         | Total   | -      | 89    | 4     | 93     | -      | 81    | 3     | 84     | -      | 82    | 2     | 84     | -      | 77    | 1     | 78     | -      | 77    | 2     | 79 |
| Week 28         | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                 | Normal  | -       | 87     | 1     | 88    | -      | 78     | 2     | 80    | -      | 76     | 1     | 77    | -      | 74     | -     | 74    | -      | 74     | -     | 74    |    |
|                 | Upper   | -       | 2      | 3     | 5     | -      | 2      | 1     | 3     | -      | 6      | 1     | 7     | -      | 2      | 1     | 3     | -      | 2      | 2     | 4     |    |
|                 | Total   | -       | 89     | 4     | 93    | -      | 80     | 3     | 83    | -      | 82     | 2     | 84    | -      | 76     | 1     | 77    | -      | 76     | 2     | 78    |    |
| Follow-up       | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                 | Normal  | -       | 91     | 1     | 92    | -      | 81     | 2     | 83    | -      | 84     | 1     | 85    | -      | 84     | -     | 84    | -      | 83     | -     | 83    |    |
|                 | Upper   | -       | 2      | 3     | 5     | -      | 2      | 1     | 3     | -      | 6      | 1     | 7     | -      | 2      | 1     | 3     | -      | 2      | 3     | 5     |    |
|                 | Total   | -       | 93     | 4     | 97    | -      | 83     | 3     | 86    | -      | 90     | 2     | 92    | -      | 86     | 1     | 87    | -      | 85     | 3     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (11/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |    |   |   |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|----|---|---|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |    |   |   |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |    |   |   |
| ChE       | Week 1 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     | 2     | -  | -  | - |   |
|           |        | Normal  | -      | 29    | 2     | 31     | -      | 28    | 1     | 29     | -      | 29    | 1     | 30     | -      | 28    | 3     | 31     | -      | 39    | 1     | 40 | -  | - | - |
|           |        | Upper   | -      | 3     | 4     | 7      | -      | 1     | 7     | 8      | -      | 1     | 7     | 8      | -      | -     | 4     | 4      | -      | -     | -     | 7  | 7  | - | - |
|           |        | Total   | 1      | 32    | 6     | 39     | -      | 29    | 8     | 37     | -      | 31    | 8     | 39     | 1      | 29    | 7     | 37     | -      | 39    | 8     | 47 | -  | - | - |
|           | Week 2 | Lower   | 1      | 1     | -     | 2      | -      | -     | -     | -      | 1      | -     | -     | 1      | 1      | 1     | -     | 2      | -      | -     | -     | -  | -  | - | - |
|           |        | Normal  | -      | 72    | 7     | 79     | -      | 67    | 7     | 74     | -      | 71    | 5     | 76     | -      | 69    | 7     | 76     | -      | 71    | 6     | 77 | -  | - | - |
|           |        | Upper   | -      | 2     | 13    | 15     | -      | 3     | 12    | 15     | -      | -     | 14    | 14     | -      | 3     | 12    | 15     | -      | -     | -     | 12 | 12 | - | - |
|           |        | Total   | 1      | 75    | 20    | 96     | -      | 70    | 19    | 89     | 1      | 71    | 19    | 91     | 1      | 73    | 19    | 93     | -      | 71    | 18    | 89 | -  | - | - |
|           | Week 3 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     | -  | -  | - | - |
|           |        | Normal  | -      | 29    | 2     | 31     | -      | 29    | 3     | 32     | -      | 30    | 4     | 34     | -      | 26    | 2     | 28     | -      | 34    | 3     | 37 | -  | - | - |
|           |        | Upper   | -      | -     | 4     | 4      | -      | -     | 4     | 4      | -      | -     | 4     | 4      | -      | -     | 3     | 3      | -      | 1     | 4     | 5  | -  | - | - |
|           |        | Total   | 1      | 29    | 6     | 36     | -      | 29    | 7     | 36     | -      | 30    | 8     | 38     | 1      | 26    | 5     | 32     | -      | 35    | 7     | 42 | -  | - | - |
|           | Week 4 | Lower   | 1      | 1     | -     | 2      | -      | -     | -     | -      | 1      | -     | -     | 1      | 1      | 1     | -     | 2      | -      | -     | -     | -  | -  | - | - |
|           |        | Normal  | -      | 70    | 7     | 77     | -      | 68    | 9     | 77     | -      | 68    | 7     | 75     | -      | 68    | 5     | 73     | -      | 69    | 5     | 74 | -  | - | - |
|           |        | Upper   | -      | 3     | 13    | 16     | -      | 2     | 10    | 12     | -      | 1     | 12    | 13     | -      | 1     | 13    | 14     | -      | -     | 13    | 13 | -  | - | - |
|           |        | Total   | 1      | 74    | 20    | 95     | -      | 70    | 19    | 89     | 1      | 69    | 19    | 89     | 1      | 70    | 18    | 89     | -      | 69    | 18    | 87 | -  | - | - |
|           | Week 6 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     | -  | -  | - | - |
|           |        | Normal  | -      | 30    | -     | 30     | -      | 31    | 5     | 36     | -      | 29    | 5     | 34     | -      | 25    | 4     | 29     | -      | 41    | 5     | 46 | -  | - | - |
|           |        | Upper   | -      | 2     | 7     | 9      | -      | -     | 4     | 4      | -      | -     | 4     | 4      | -      | -     | 1     | 1      | -      | -     | 1     | 1  | -  | - | - |
|           |        | Total   | 1      | 32    | 7     | 40     | -      | 31    | 9     | 40     | -      | 29    | 9     | 38     | 1      | 25    | 5     | 31     | -      | 41    | 6     | 47 | -  | - | - |
|           | Week 8 | Lower   | 1      | 1     | -     | 2      | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -  | -  | - | - |
|           |        | Normal  | -      | 71    | 6     | 77     | -      | 65    | 5     | 70     | -      | 67    | 7     | 74     | 1      | 67    | 7     | 75     | -      | 67    | 5     | 72 | -  | - | - |
|           |        | Upper   | -      | 1     | 14    | 15     | -      | 3     | 13    | 16     | -      | 2     | 12    | 14     | -      | 2     | 9     | 11     | -      | 1     | 11    | 12 | -  | - | - |
|           |        | Total   | 1      | 73    | 20    | 94     | -      | 68    | 18    | 86     | 1      | 69    | 19    | 89     | 1      | 69    | 16    | 86     | -      | 68    | 16    | 84 | -  | - | - |
| Week 12   | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | 1      | 1      | -     | 2     | 1      | -      | -     | 1     | -      | -      | -     | -     | -  | -  | - |   |
|           | Normal | -       | 69     | 11    | 80    | -      | 67     | 8     | 75    | -      | 65     | 7     | 72    | -      | 64     | 8     | 72    | -      | 64     | 6     | 70    | -  | -  | - |   |
|           | Upper  | -       | 3      | 9     | 12    | -      | 1      | 10    | 11    | -      | 1      | 11    | 12    | -      | 2      | 7     | 9     | -      | 3      | 10    | 13    | -  | -  | - |   |
|           | Total  | 1       | 73     | 20    | 94    | -      | 68     | 18    | 86    | 1      | 67     | 18    | 86    | 1      | 66     | 15    | 82    | -      | 67     | 16    | 83    | -  | -  | - |   |
| Week 16   | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | 1      | 2      | -     | 3     | 1      | -      | -     | 1     | -      | -      | -     | -     | -  | -  | - |   |
|           | Normal | -       | 69     | 13    | 82    | -      | 65     | 8     | 73    | -      | 62     | 4     | 66    | -      | 63     | 9     | 72    | -      | 64     | 5     | 69    | -  | -  | - |   |
|           | Upper  | -       | 3      | 7     | 10    | -      | 1      | 10    | 11    | -      | 2      | 14    | 16    | -      | 2      | 5     | 7     | -      | 1      | 12    | 13    | -  | -  | - |   |
|           | Total  | 1       | 73     | 20    | 94    | -      | 66     | 18    | 84    | 1      | 66     | 18    | 85    | 1      | 65     | 14    | 80    | -      | 65     | 17    | 82    | -  | -  | - |   |
| Week 20   | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | 1      | 1      | -     | 2     | 1      | -      | -     | 1     | -      | -      | -     | -     | -  | -  | - |   |
|           | Normal | -       | 68     | 9     | 77    | -      | 66     | 5     | 71    | -      | 65     | 8     | 73    | -      | 62     | 10    | 72    | -      | 63     | 8     | 71    | -  | -  | - |   |
|           | Upper  | -       | 4      | 11    | 15    | -      | -      | 13    | 13    | -      | -      | 10    | 10    | -      | 2      | 4     | 6     | -      | 2      | 7     | 9     | -  | -  | - |   |
|           | Total  | 1       | 73     | 20    | 94    | -      | 66     | 18    | 84    | 1      | 66     | 18    | 85    | 1      | 64     | 14    | 79    | -      | 65     | 15    | 80    | -  | -  | - |   |
| Week 24   | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | 1      | 2      | -     | 3     | 1      | -      | -     | 1     | -      | -      | -     | -     | -  | -  | - |   |
|           | Normal | -       | 70     | 10    | 80    | -      | 64     | 5     | 69    | -      | 63     | 6     | 69    | -      | 62     | 11    | 73    | -      | 61     | 6     | 67    | -  | -  | - |   |
|           | Upper  | -       | 1      | 10    | 11    | -      | 2      | 13    | 15    | -      | 1      | 11    | 12    | -      | 1      | 3     | 4     | -      | 2      | 9     | 11    | -  | -  | - |   |
|           | Total  | 1       | 72     | 20    | 93    | -      | 66     | 18    | 84    | 1      | 66     | 17    | 84    | 1      | 63     | 14    | 78    | -      | 64     | 15    | 79    | -  | -  | - |   |
| Week 28   | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | 1      | -      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     | -     | -  | -  | - |   |
|           | Normal | -       | 68     | 9     | 77    | -      | 63     | 8     | 71    | -      | 65     | 7     | 72    | -      | 61     | 8     | 69    | -      | 61     | 7     | 68    | -  | -  | - |   |
|           | Upper  | -       | 3      | 11    | 14    | -      | 2      | 10    | 12    | -      | 1      | 10    | 11    | -      | 1      | 6     | 7     | -      | 2      | 8     | 10    | -  | -  | - |   |
|           | Total  | 1       | 72     | 20    | 93    | -      | 65     | 18    | 83    | 1      | 66     | 17    | 84    | 1      | 62     | 14    | 77    | -      | 63     | 15    | 78    | -  | -  | - |   |
| Follow-up | Lower  | 1       | 2      | -     | 3     | -      | -      | -     | -     | 1      | -      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     | -     | -  | -  | - |   |
|           | Normal | -       | 71     | 9     | 80    | -      | 65     | 9     | 74    | -      | 71     | 8     | 79    | -      | 67     | 11    | 78    | -      | 68     | 8     | 76    | -  | -  | - |   |
|           | Upper  | -       | 3      | 11    | 14    | -      | 2      | 10    | 12    | -      | 1      | 11    | 12    | -      | 1      | 7     | 8     | -      | 2      | 10    | 12    | -  | -  | - |   |
|           | Total  | 1       | 76     | 20    | 97    | -      | 67     | 19    | 86    | 1      | 72     | 19    | 92    | 1      | 68     | 18    | 87    | -      | 70     | 18    | 88    | -  | -  | - |   |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (12/24)

|            |           | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|------------|-----------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|            |           | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|            |           | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Creatinine | Week 1    | Lower   | 1      | 1     | -     | 2      | 5      | -     | -     | 5      | 1      | -     | -     | 1      | 3      | -     | -     | 3      | 2      | -     | -     | 2  |
|            |           | Normal  | 1      | 31    | 1     | 33     | -      | 24    | 1     | 25     | 3      | 28    | -     | 31     | 1      | 30    | -     | 31     | 1      | 39    | -     | 40 |
|            |           | Upper   | -      | -     | 4     | 4      | -      | 1     | 6     | 7      | -      | 3     | 4     | 7      | -      | -     | 3     | 3      | -      | -     | 5     | 5  |
|            |           | Total   | 2      | 32    | 5     | 39     | 5      | 25    | 7     | 37     | 4      | 31    | 4     | 39     | 4      | 30    | 3     | 37     | 3      | 39    | 5     | 47 |
|            | Week 2    | Lower   | 1      | 4     | -     | 5      | 6      | 2     | -     | 8      | 5      | 2     | -     | 7      | 7      | 1     | -     | 8      | 4      | 3     | -     | 7  |
|            |           | Normal  | 2      | 77    | 4     | 83     | 1      | 69    | -     | 70     | 3      | 64    | 1     | 68     | 2      | 73    | -     | 75     | 3      | 69    | 1     | 73 |
|            |           | Upper   | -      | 1     | 7     | 8      | -      | 3     | 8     | 11     | -      | 4     | 12    | 16     | -      | 1     | 9     | 10     | -      | 1     | 8     | 9  |
|            |           | Total   | 3      | 82    | 11    | 96     | 7      | 74    | 8     | 89     | 8      | 70    | 13    | 91     | 9      | 75    | 9     | 93     | 7      | 73    | 9     | 89 |
|            | Week 3    | Lower   | 1      | 1     | -     | 2      | 5      | 1     | -     | 6      | 1      | -     | -     | 1      | 2      | -     | -     | 2      | 1      | 2     | -     | 3  |
|            |           | Normal  | 1      | 26    | 1     | 28     | -      | 25    | -     | 25     | 3      | 29    | 1     | 33     | 1      | 25    | 1     | 27     | 1      | 33    | -     | 34 |
|            |           | Upper   | -      | 2     | 4     | 6      | -      | -     | -     | 5      | -      | 1     | 3     | 4      | -      | 1     | 2     | 3      | -      | -     | -     | 5  |
|            |           | Total   | 2      | 29    | 5     | 36     | 5      | 26    | 5     | 36     | 4      | 30    | 4     | 38     | 3      | 26    | 3     | 32     | 2      | 35    | 5     | 42 |
|            | Week 4    | Lower   | 1      | 4     | -     | 5      | 7      | 3     | -     | 10     | 6      | 4     | -     | 10     | 7      | 2     | -     | 9      | 5      | 1     | -     | 6  |
|            |           | Normal  | 2      | 77    | 3     | 82     | -      | 69    | 1     | 70     | 2      | 63    | 3     | 68     | 2      | 68    | 1     | 71     | 2      | 70    | 3     | 75 |
|            |           | Upper   | -      | 1     | 7     | 8      | -      | 2     | 7     | 9      | -      | 1     | 10    | 11     | -      | 3     | 6     | 9      | -      | -     | 6     | 6  |
|            |           | Total   | 3      | 82    | 10    | 95     | 7      | 74    | 8     | 89     | 8      | 68    | 13    | 89     | 9      | 73    | 7     | 89     | 7      | 71    | 9     | 87 |
|            | Week 6    | Lower   | 2      | 1     | -     | 3      | 4      | -     | -     | 4      | 2      | -     | -     | 2      | 3      | 1     | -     | 4      | 3      | 2     | -     | 5  |
|            |           | Normal  | 1      | 31    | 2     | 34     | 1      | 29    | 2     | 32     | 3      | 29    | 1     | 33     | -      | 25    | -     | 25     | 1      | 35    | 2     | 38 |
|            |           | Upper   | -      | 1     | 2     | 3      | -      | 1     | 3     | 4      | -      | -     | 3     | 3      | -      | -     | 3     | 3      | -      | -     | 4     | 4  |
|            |           | Total   | 3      | 33    | 4     | 40     | 5      | 30    | 5     | 40     | 5      | 29    | 4     | 38     | 3      | 26    | 3     | 32     | 4      | 37    | 6     | 47 |
|            | Week 8    | Lower   | 2      | 4     | -     | 6      | 6      | 1     | -     | 7      | 5      | 6     | -     | 11     | 5      | -     | -     | 5      | 4      | 2     | -     | 6  |
|            |           | Normal  | 1      | 75    | 3     | 79     | 1      | 68    | 1     | 70     | 3      | 60    | 2     | 65     | 3      | 71    | 1     | 75     | 2      | 67    | 1     | 70 |
|            |           | Upper   | -      | 2     | 7     | 9      | -      | 2     | 7     | 9      | -      | 2     | 11    | 13     | -      | 1     | 5     | 6      | -      | -     | 8     | 8  |
|            |           | Total   | 3      | 81    | 10    | 94     | 7      | 71    | 8     | 86     | 8      | 68    | 13    | 89     | 8      | 72    | 6     | 86     | 6      | 69    | 9     | 84 |
|            | Week 12   | Lower   | 1      | 3     | -     | 4      | 7      | 2     | -     | 9      | 3      | 4     | -     | 7      | 5      | 2     | -     | 7      | 3      | 3     | -     | 6  |
|            |           | Normal  | 2      | 77    | 2     | 81     | -      | 66    | 1     | 67     | 4      | 61    | 1     | 66     | 2      | 66    | 1     | 69     | 3      | 63    | 1     | 67 |
|            |           | Upper   | -      | 1     | 8     | 9      | -      | 3     | 7     | 10     | -      | 1     | 12    | 13     | -      | 1     | 5     | 6      | -      | 2     | 8     | 10 |
|            |           | Total   | 3      | 81    | 10    | 94     | 7      | 71    | 8     | 86     | 7      | 66    | 13    | 86     | 7      | 69    | 6     | 82     | 6      | 68    | 9     | 83 |
|            | Week 16   | Lower   | 1      | 3     | -     | 4      | 5      | 1     | -     | 6      | 2      | 6     | -     | 8      | 5      | 2     | -     | 7      | 3      | 4     | -     | 7  |
|            |           | Normal  | 2      | 75    | 2     | 79     | 2      | 65    | 1     | 68     | 5      | 56    | 1     | 62     | 1      | 63    | 2     | 66     | 3      | 63    | 2     | 68 |
|            |           | Upper   | -      | 3     | 8     | 11     | -      | 3     | 7     | 10     | -      | 4     | 11    | 15     | -      | 4     | 3     | 7      | -      | -     | 7     | 7  |
|            |           | Total   | 3      | 81    | 10    | 94     | 7      | 69    | 8     | 84     | 7      | 66    | 12    | 85     | 6      | 69    | 5     | 80     | 6      | 67    | 9     | 82 |
|            | Week 20   | Lower   | 2      | 4     | -     | 6      | 4      | 4     | -     | 8      | 5      | 7     | -     | 12     | 5      | 2     | -     | 7      | -      | 3     | -     | 3  |
|            |           | Normal  | 1      | 76    | 4     | 81     | 3      | 62    | -     | 65     | 2      | 54    | 1     | 57     | 1      | 64    | 2     | 67     | 5      | 63    | 3     | 71 |
|            |           | Upper   | -      | 1     | 6     | 7      | -      | 3     | 8     | 11     | -      | 5     | 11    | 16     | -      | 2     | 3     | 5      | -      | -     | 6     | 6  |
|            |           | Total   | 3      | 81    | 10    | 94     | 7      | 69    | 8     | 84     | 7      | 66    | 12    | 85     | 6      | 68    | 5     | 79     | 5      | 66    | 9     | 80 |
|            | Week 24   | Lower   | 1      | 7     | -     | 8      | 4      | 5     | -     | 9      | 4      | 4     | -     | 8      | 5      | 1     | -     | 6      | 1      | 3     | -     | 4  |
|            |           | Normal  | 1      | 71    | 5     | 77     | 3      | 64    | 3     | 70     | 2      | 60    | 1     | 63     | 1      | 63    | 3     | 67     | 4      | 61    | 1     | 66 |
|            |           | Upper   | -      | 3     | 5     | 8      | -      | -     | 5     | 5      | -      | 2     | 11    | 13     | -      | 3     | 2     | 5      | -      | 1     | 8     | 9  |
|            |           | Total   | 2      | 81    | 10    | 93     | 7      | 69    | 8     | 84     | 6      | 66    | 12    | 84     | 6      | 67    | 5     | 78     | 5      | 65    | 9     | 79 |
|            | Week 28   | Lower   | 1      | 4     | -     | 5      | 6      | 1     | -     | 7      | 3      | 2     | -     | 5      | 4      | 1     | -     | 5      | 2      | 3     | -     | 5  |
|            |           | Normal  | 1      | 76    | 3     | 80     | 1      | 66    | 1     | 68     | 3      | 63    | 1     | 67     | 2      | 63    | -     | 65     | 3      | 61    | 1     | 65 |
|            |           | Upper   | -      | 1     | 7     | 8      | -      | 1     | 7     | 8      | -      | 1     | 11    | 12     | -      | 2     | 5     | 7      | -      | -     | 8     | 8  |
|            |           | Total   | 2      | 81    | 10    | 93     | 7      | 68    | 8     | 83     | 6      | 66    | 12    | 84     | 6      | 66    | 5     | 77     | 5      | 64    | 9     | 78 |
|            | Follow-up | Lower   | 1      | 4     | -     | 5      | 6      | 1     | -     | 7      | 4      | 2     | -     | 6      | 5      | 2     | -     | 7      | 4      | 3     | -     | 7  |
|            |           | Normal  | 2      | 77    | 4     | 83     | 1      | 68    | 1     | 70     | 4      | 67    | 1     | 72     | 4      | 67    | -     | 71     | 3      | 69    | 1     | 73 |
|            |           | Upper   | -      | 2     | 7     | 9      | -      | 2     | 7     | 9      | -      | 2     | 12    | 14     | -      | 2     | 7     | 9      | -      | -     | 8     | 8  |
|            |           | Total   | 3      | 83    | 11    | 97     | 7      | 71    | 8     | 86     | 8      | 71    | 13    | 92     | 9      | 71    | 7     | 87     | 7      | 72    | 9     | 88 |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (13/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| BUN       | Week 1 | Lower   | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1  |
|           |        | Normal  | -      | 36    | 1     | 37     | -      | 34    | 1     | 35     | -      | 35    | 1     | 36     | -      | 34    | 1     | 35     | -      | 42    | 2     | 44 |
|           |        | Upper   | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | 1     | 1     | 2      | -      | -     | 2     | 2      | -      | 2     | -     | 2  |
|           |        | Total   | -      | 37    | 2     | 39     | 1      | 34    | 2     | 37     | 1      | 36    | 2     | 39     | -      | 34    | 3     | 37     | -      | 45    | 2     | 47 |
|           | Week 2 | Lower   | -      | 3     | -     | 3      | 1      | -     | -     | 1      | 1      | -     | -     | 1      | -      | 3     | -     | 3      | -      | -     | -     | -  |
|           |        | Normal  | -      | 88    | 4     | 92     | -      | 84    | 1     | 85     | 1      | 83    | 2     | 86     | 1      | 83    | 1     | 85     | 1      | 84    | 2     | 87 |
|           |        | Upper   | -      | 1     | -     | 1      | -      | 2     | 1     | 3      | -      | 3     | 1     | 4      | -      | 3     | 2     | 5      | -      | 1     | 1     | 2  |
|           |        | Total   | -      | 92    | 4     | 96     | 1      | 86    | 2     | 89     | 2      | 86    | 3     | 91     | 1      | 89    | 3     | 93     | 1      | 85    | 3     | 89 |
|           | Week 3 | Lower   | -      | 1     | -     | 1      | 1      | 1     | -     | 2      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -  |
|           |        | Normal  | -      | 33    | 2     | 35     | -      | 31    | -     | 31     | 1      | 35    | 1     | 37     | -      | 28    | 1     | 29     | -      | 39    | 2     | 41 |
|           |        | Upper   | -      | -     | -     | -      | -      | 1     | 2     | 3      | -      | -     | 1     | 1      | -      | 2     | 2     | 2      | -      | 1     | -     | 1  |
|           |        | Total   | -      | 34    | 2     | 36     | 1      | 33    | 2     | 36     | 1      | 35    | 2     | 38     | -      | 29    | 3     | 32     | -      | 40    | 2     | 42 |
|           | Week 4 | Lower   | -      | -     | -     | -      | 1      | 1     | -     | 2      | 1      | -     | -     | 1      | 1      | 2     | -     | 3      | -      | 2     | -     | 2  |
|           |        | Normal  | -      | 91    | 2     | 93     | -      | 84    | -     | 84     | 1      | 84    | 1     | 86     | -      | 83    | 1     | 84     | 1      | 78    | 1     | 80 |
|           |        | Upper   | -      | -     | 2     | 2      | -      | 1     | 2     | 3      | -      | -     | 2     | 2      | -      | -     | 2     | 2      | -      | 3     | 2     | 5  |
|           |        | Total   | -      | 91    | 4     | 95     | 1      | 86    | 2     | 89     | 2      | 84    | 3     | 89     | 1      | 85    | 3     | 89     | 1      | 83    | 3     | 87 |
|           | Week 6 | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |        | Normal  | -      | 37    | 2     | 39     | -      | 37    | -     | 37     | 1      | 34    | -     | 35     | -      | 29    | 1     | 30     | -      | 43    | 1     | 44 |
|           |        | Upper   | -      | 1     | -     | 1      | -      | -     | 2     | 2      | -      | -     | 2     | 2      | -      | -     | 1     | 1      | -      | 1     | 1     | 2  |
|           |        | Total   | -      | 38    | 2     | 40     | -      | 38    | 2     | 40     | 1      | 35    | 2     | 38     | -      | 29    | 2     | 31     | -      | 45    | 2     | 47 |
|           | Week 8 | Lower   | -      | -     | -     | -      | 1      | 1     | -     | 2      | 1      | -     | -     | 1      | -      | 2     | -     | 2      | -      | -     | -     | -  |
|           |        | Normal  | -      | 89    | 1     | 90     | -      | 82    | 1     | 83     | 1      | 82    | 1     | 84     | 1      | 80    | 1     | 82     | 1      | 78    | 3     | 82 |
|           |        | Upper   | -      | 1     | 3     | 4      | -      | -     | 1     | 1      | -      | 2     | 2     | 4      | -      | 1     | 1     | 2      | -      | 2     | -     | 2  |
|           |        | Total   | -      | 90    | 4     | 94     | 1      | 83    | 2     | 86     | 2      | 84    | 3     | 89     | 1      | 83    | 2     | 86     | 1      | 80    | 3     | 84 |
| Week 12   | Lower  | -       | 2      | -     | 2     | -      | 2      | -     | 2     | 1      | 2      | -     | 3     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Normal | -       | 85     | 1     | 86    | 1      | 80     | -     | 81    | 1      | 79     | 2     | 82    | 1      | 79     | 1     | 81    | 1      | 76     | 3     | 80    |    |
|           | Upper  | -       | 3      | 3     | 6     | -      | 1      | 2     | 3     | -      | -      | 1     | 1     | -      | -      | 1     | 1     | -      | 3      | -     | 3     |    |
|           | Total  | -       | 90     | 4     | 94    | 1      | 83     | 2     | 86    | 2      | 81     | 3     | 86    | 1      | 79     | 2     | 82    | 1      | 79     | 3     | 83    |    |
| Week 16   | Lower  | -       | 1      | -     | 1     | -      | 2      | -     | 2     | 1      | 1      | -     | 2     | -      | 1      | -     | 1     | -      | 1      | -     | 1     |    |
|           | Normal | -       | 89     | 4     | 93    | 1      | 78     | -     | 79    | 1      | 77     | 1     | 79    | 1      | 76     | 1     | 78    | 1      | 75     | 2     | 78    |    |
|           | Upper  | -       | -      | -     | -     | -      | 1      | 2     | 3     | -      | 2      | 2     | 4     | -      | -      | 1     | 1     | -      | 3      | -     | 3     |    |
|           | Total  | -       | 90     | 4     | 94    | 1      | 81     | 2     | 84    | 2      | 80     | 3     | 85    | 1      | 77     | 2     | 80    | 1      | 79     | 2     | 82    |    |
| Week 20   | Lower  | -       | -      | -     | -     | 1      | 1      | -     | 2     | 1      | 2      | -     | 3     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           | Normal | -       | 89     | 3     | 92    | -      | 80     | 1     | 81    | 1      | 78     | 2     | 81    | 1      | 73     | 1     | 75    | 1      | 75     | 1     | 77    |    |
|           | Upper  | -       | 1      | 1     | 2     | -      | -      | 1     | 1     | -      | -      | 1     | 1     | -      | 2      | 1     | 3     | -      | 2      | 1     | 3     |    |
|           | Total  | -       | 90     | 4     | 94    | 1      | 81     | 2     | 84    | 2      | 80     | 3     | 85    | 1      | 76     | 2     | 79    | 1      | 77     | 2     | 80    |    |
| Week 24   | Lower  | -       | 3      | -     | 3     | 1      | 2      | -     | 3     | 2      | 1      | -     | 3     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           | Normal | -       | 84     | 4     | 88    | -      | 77     | -     | 77    | -      | 76     | 2     | 78    | 1      | 71     | 1     | 73    | 1      | 75     | -     | 76    |    |
|           | Upper  | -       | 2      | -     | 2     | -      | 2      | 2     | 4     | -      | 2      | 1     | 3     | -      | 3      | 1     | 4     | -      | 1      | 2     | 3     |    |
|           | Total  | -       | 89     | 4     | 93    | 1      | 81     | 2     | 84    | 2      | 79     | 3     | 84    | 1      | 75     | 2     | 78    | 1      | 76     | 2     | 79    |    |
| Week 28   | Lower  | -       | 1      | -     | 1     | 1      | 2      | -     | 3     | 1      | -      | -     | 1     | -      | -      | -     | 1     | -      | -      | -     | 1     |    |
|           | Normal | -       | 86     | 3     | 89    | -      | 78     | -     | 78    | 1      | 78     | 1     | 80    | 1      | 72     | 1     | 74    | -      | 74     | -     | 74    |    |
|           | Upper  | -       | 2      | 1     | 3     | -      | -      | 2     | 2     | -      | 1      | 2     | 3     | -      | 2      | 1     | 3     | -      | 1      | 2     | 3     |    |
|           | Total  | -       | 89     | 4     | 93    | 1      | 80     | 2     | 83    | 2      | 79     | 3     | 84    | 1      | 74     | 2     | 77    | 1      | 75     | 2     | 78    |    |
| Follow-up | Lower  | -       | 1      | -     | 1     | 1      | 3      | -     | 4     | 1      | -      | -     | 1     | -      | 1      | -     | 1     | 1      | -      | -     | 1     |    |
|           | Normal | -       | 89     | 3     | 92    | -      | 80     | -     | 80    | 1      | 86     | 1     | 88    | 1      | 80     | 1     | 82    | -      | 82     | 1     | 83    |    |
|           | Upper  | -       | 3      | 1     | 4     | -      | -      | 2     | 2     | -      | 1      | 2     | 3     | -      | 2      | 2     | 4     | -      | 2      | 2     | 4     |    |
|           | Total  | -       | 93     | 4     | 97    | 1      | 83     | 2     | 86    | 2      | 87     | 3     | 92    | 1      | 83     | 3     | 87    | 1      | 84     | 3     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (14/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Na        | Week 1 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | 2      | 37    | -     | 39     | -      | 36    | -     | 36     | 1      | 37    | -     | 38     | -      | 35    | -     | 35     | 1      | 45    | -     | 46 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 2      | 37    | -     | 39     | -      | 37    | -     | 37     | 1      | 38    | -     | 39     | 1      | 36    | -     | 37     | 1      | 46    | -     | 47 |
|           | Week 2 | Lower   | 1      | -     | -     | 1      | -      | 3     | -     | 3      | 2      | 1     | -     | 3      | 2      | 1     | -     | 3      | -      | -     | -     |    |
|           |        | Normal  | 4      | 91    | -     | 95     | 1      | 85    | -     | 86     | 1      | 87    | -     | 88     | 3      | 87    | -     | 90     | 1      | 88    | -     | 89 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 5      | 91    | -     | 96     | 1      | 88    | -     | 89     | 3      | 88    | -     | 91     | 5      | 88    | -     | 93     | 1      | 88    | -     | 89 |
|           | Week 3 | Lower   | 1      | -     | -     | 1      | -      | 1     | -     | 1      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | 1      | 34    | -     | 35     | -      | 35    | -     | 35     | -      | 37    | -     | 37     | 1      | 31    | -     | 32     | 1      | 41    | -     | 42 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 2      | 34    | -     | 36     | -      | 36    | -     | 36     | 1      | 37    | -     | 38     | 1      | 31    | -     | 32     | 1      | 41    | -     | 42 |
|           | Week 4 | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | 2      | -     | -     | 2      | -      | 2     | -     | 2  |
|           |        | Normal  | 5      | 90    | -     | 95     | 1      | 87    | -     | 88     | 3      | 85    | -     | 88     | 2      | 85    | -     | 87     | 1      | 84    | -     | 85 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 5      | 90    | -     | 95     | 1      | 88    | -     | 89     | 3      | 86    | -     | 89     | 4      | 85    | -     | 89     | 1      | 86    | -     | 87 |
|           | Week 6 | Lower   | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | 2      | 37    | -     | 39     | -      | 39    | -     | 39     | 1      | 36    | -     | 37     | 1      | 30    | -     | 31     | 1      | 46    | -     | 47 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 2      | 38    | -     | 40     | -      | 40    | -     | 40     | 1      | 37    | -     | 38     | 1      | 30    | -     | 31     | 1      | 46    | -     | 47 |
| Week 8    | Lower  | 1       | 1      | -     | 2     | -      | 2      | -     | 2     | 2      | -      | -     | 2     | 1      | 1      | -     | 2     | -      | 1      | -     | 1     |    |
|           | Normal | 4       | 88     | -     | 92    | 1      | 83     | -     | 84    | 1      | 86     | -     | 87    | 3      | 81     | -     | 84    | 1      | 82     | -     | 83    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 5       | 89     | -     | 94    | 1      | 85     | -     | 86    | 3      | 86     | -     | 89    | 4      | 82     | -     | 86    | 1      | 83     | -     | 84    |    |
| Week 12   | Lower  | 1       | 2      | -     | 3     | -      | 1      | -     | 1     | 2      | 1      | -     | 3     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Normal | 4       | 86     | -     | 90    | 1      | 84     | -     | 85    | 1      | 82     | -     | 83    | 4      | 78     | -     | 82    | 1      | 81     | -     | 82    |    |
|           | Upper  | -       | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 5       | 89     | -     | 94    | 1      | 85     | -     | 86    | 3      | 83     | -     | 86    | 4      | 78     | -     | 82    | 1      | 82     | -     | 83    |    |
| Week 16   | Lower  | 1       | 2      | -     | 3     | -      | 1      | -     | 1     | 1      | 2      | -     | 3     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Normal | 4       | 87     | -     | 91    | 1      | 82     | -     | 83    | 2      | 80     | -     | 82    | 3      | 77     | -     | 80    | 1      | 80     | -     | 81    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 5       | 89     | -     | 94    | 1      | 83     | -     | 84    | 3      | 82     | -     | 85    | 3      | 77     | -     | 80    | 1      | 81     | -     | 82    |    |
| Week 20   | Lower  | 1       | 1      | -     | 2     | -      | 3      | -     | 3     | -      | 1      | -     | 1     | 1      | -      | -     | 1     | -      | 2      | -     | 2     |    |
|           | Normal | 4       | 88     | -     | 92    | 1      | 80     | -     | 81    | 3      | 81     | -     | 84    | 2      | 76     | -     | 78    | 1      | 77     | -     | 78    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 5       | 89     | -     | 94    | 1      | 83     | -     | 84    | 3      | 82     | -     | 85    | 3      | 76     | -     | 79    | 1      | 79     | -     | 80    |    |
| Week 24   | Lower  | 1       | 1      | -     | 2     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Normal | 4       | 87     | -     | 91    | 1      | 83     | -     | 84    | 3      | 80     | -     | 83    | 3      | 75     | -     | 78    | 1      | 78     | -     | 79    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 5       | 88     | -     | 93    | 1      | 83     | -     | 84    | 3      | 81     | -     | 84    | 3      | 75     | -     | 78    | 1      | 78     | -     | 79    |    |
| Week 28   | Lower  | -       | 2      | -     | 2     | -      | 2      | -     | 2     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           | Normal | 5       | 86     | -     | 91    | 1      | 80     | -     | 81    | 3      | 79     | -     | 82    | 3      | 73     | -     | 76    | 1      | 77     | -     | 78    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 5       | 88     | -     | 93    | 1      | 82     | -     | 83    | 3      | 81     | -     | 84    | 3      | 74     | -     | 77    | 1      | 77     | -     | 78    |    |
| Follow-up | Lower  | -       | 2      | -     | 2     | -      | 2      | -     | 2     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           | Normal | 5       | 90     | -     | 95    | 1      | 83     | -     | 84    | 3      | 87     | -     | 90    | 4      | 82     | -     | 86    | 1      | 87     | -     | 88    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 5       | 92     | -     | 97    | 1      | 85     | -     | 86    | 3      | 89     | -     | 92    | 4      | 83     | -     | 87    | 1      | 87     | -     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (15/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| K         | Week 1 | Lower   | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | -      | -      | 1     | -     |    |
|           |        | Normal  | 1      | 38    | -     | 39     | -      | 34    | 2     | 36     | 1      | 37    | -     | 38     | -      | 36    | -     | 36     | -      | 45    | 1     | 46 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |        | Total   | 1      | 38    | -     | 39     | 1      | 34    | 2     | 37     | 1      | 38    | -     | 39     | -      | 37    | -     | 37     | -      | 46    | 1     | 47 |
|           | Week 2 | Lower   | -      | 1     | -     | 1      | 2      | -     | -     | 2      | -      | -     | -     | -      | 1      | -     | 1     | 1      | 1      | -     | 2     |    |
|           |        | Normal  | 2      | 90    | 1     | 93     | -      | 83    | 3     | 86     | -      | 89    | -     | 89     | -      | 90    | -     | 90     | -      | 83    | 2     | 85 |
|           |        | Upper   | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | 2     | -     | 2      | -      | 2     | -     | 2      | -      | 2     | -     | 2  |
|           |        | Total   | 2      | 93    | 1     | 96     | 2      | 84    | 3     | 89     | -      | 91    | -     | 91     | -      | 93    | -     | 93     | 1      | 86    | 2     | 89 |
|           | Week 3 | Lower   | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | 1      | 34    | -     | 35     | -      | 33    | 2     | 35     | -      | 37    | -     | 37     | -      | 31    | -     | 31     | -      | 40    | 1     | 41 |
|           |        | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1  |
|           |        | Total   | 1      | 35    | -     | 36     | 1      | 33    | 2     | 36     | -      | 38    | -     | 38     | -      | 32    | -     | 32     | -      | 41    | 1     | 42 |
|           | Week 4 | Lower   | 1      | -     | -     | 1      | 2      | -     | -     | 2      | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     |    |
|           |        | Normal  | 1      | 91    | 1     | 93     | -      | 82    | 3     | 85     | -      | 87    | -     | 87     | -      | 86    | -     | 86     | -      | 81    | 2     | 83 |
|           |        | Upper   | -      | 1     | -     | 1      | -      | 2     | -     | 2      | -      | 2     | -     | 2      | -      | 3     | -     | 3      | -      | 3     | -     | 3  |
|           |        | Total   | 2      | 92    | 1     | 95     | 2      | 84    | 3     | 89     | -      | 89    | -     | 89     | -      | 89    | -     | 89     | 1      | 84    | 2     | 87 |
|           | Week 6 | Lower   | -      | 1     | -     | 1      | 1      | 1     | -     | 2      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | 1      | 37    | 1     | 39     | -      | 36    | 2     | 38     | -      | 36    | -     | 36     | -      | 31    | -     | 31     | -      | 44    | 1     | 45 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | -     | -     | -      | 2      | -     | 2     |    |
|           |        | Total   | 1      | 38    | 1     | 40     | 1      | 37    | 2     | 40     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 46    | 1     | 47 |
| Week 8    | Lower  | -       | -      | -     | -     | 2      | 1      | -     | 3     | -      | -      | -     | -     | 3      | -      | 3     | 1     | -      | -      | 1     |       |    |
|           | Normal | 2       | 89     | 1     | 92    | -      | 80     | 3     | 83    | -      | 88     | -     | 88    | -      | 83     | -     | 83    | -      | 80     | 2     | 82    |    |
|           | Upper  | -       | 2      | -     | 2     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | 1      | -      | 1     |       |    |
|           | Total  | 2       | 91     | 1     | 94    | 2      | 81     | 3     | 86    | -      | 89     | -     | 89    | -      | 86     | -     | 86    | 1      | 81     | 2     | 84    |    |
| Week 12   | Lower  | 1       | 1      | -     | 2     | 2      | 2      | -     | 4     | -      | -      | -     | -     | 4      | -      | 4     | 1     | -      | -      | 1     |       |    |
|           | Normal | 1       | 88     | 1     | 90    | -      | 76     | 3     | 79    | -      | 84     | -     | 84    | -      | 76     | -     | 76    | -      | 79     | 2     | 81    |    |
|           | Upper  | -       | 2      | -     | 2     | -      | 3      | -     | 3     | -      | 2      | -     | 2     | -      | 2      | -     | 2     | -      | 1      | -     | 1     |    |
|           | Total  | 2       | 91     | 1     | 94    | 2      | 81     | 3     | 86    | -      | 86     | -     | 86    | -      | 82     | -     | 82    | 1      | 80     | 2     | 83    |    |
| Week 16   | Lower  | 1       | 1      | -     | 2     | 1      | 2      | -     | 3     | -      | 1      | -     | 1     | -      | 2      | -     | 2     | 1      | -      | -     | 1     |    |
|           | Normal | 1       | 88     | 1     | 90    | 1      | 75     | 3     | 79    | -      | 82     | -     | 82    | -      | 78     | -     | 78    | -      | 78     | 1     | 79    |    |
|           | Upper  | -       | 2      | -     | 2     | -      | 2      | -     | 2     | -      | 2      | -     | 2     | -      | -      | -     | -     | 1      | 1      | 2     |       |    |
|           | Total  | 2       | 91     | 1     | 94    | 2      | 79     | 3     | 84    | -      | 85     | -     | 85    | -      | 80     | -     | 80    | 1      | 79     | 2     | 82    |    |
| Week 20   | Lower  | 1       | 1      | -     | 2     | 1      | -      | -     | 1     | -      | 1      | -     | 1     | -      | 2      | -     | 2     | -      | 1      | -     | 1     |    |
|           | Normal | 1       | 89     | 1     | 91    | 1      | 77     | 2     | 80    | -      | 83     | -     | 83    | -      | 76     | -     | 76    | 1      | 74     | 2     | 77    |    |
|           | Upper  | -       | 1      | -     | 1     | -      | 2      | 1     | 3     | -      | 1      | -     | 1     | -      | 1      | -     | 1     | -      | 2      | -     | 2     |    |
|           | Total  | 2       | 91     | 1     | 94    | 2      | 79     | 3     | 84    | -      | 85     | -     | 85    | -      | 79     | -     | 79    | 1      | 77     | 2     | 80    |    |
| Week 24   | Lower  | 1       | 3      | -     | 4     | 1      | -      | -     | 1     | -      | -      | -     | -     | 1      | -      | 1     | 1     | -      | -      | 1     |       |    |
|           | Normal | 1       | 86     | 1     | 88    | 1      | 78     | 3     | 82    | -      | 82     | -     | 82    | -      | 76     | -     | 76    | -      | 75     | 2     | 77    |    |
|           | Upper  | -       | 1      | -     | 1     | -      | 1      | -     | 1     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | 1      | -     | 1     |    |
|           | Total  | 2       | 90     | 1     | 93    | 2      | 79     | 3     | 84    | -      | 84     | -     | 84    | -      | 78     | -     | 78    | 1      | 76     | 2     | 79    |    |
| Week 28   | Lower  | -       | 2      | -     | 2     | 1      | 1      | -     | 2     | -      | -      | -     | -     | 2      | -      | 2     | 1     | -      | -      | 1     |       |    |
|           | Normal | 2       | 87     | 1     | 90    | 1      | 76     | 2     | 79    | -      | 84     | -     | 84    | -      | 73     | -     | 73    | -      | 74     | 2     | 76    |    |
|           | Upper  | -       | 1      | -     | 1     | -      | 1      | 1     | 2     | -      | -      | -     | -     | 2      | -      | 2     | -     | 1      | -      | 1     |       |    |
|           | Total  | 2       | 90     | 1     | 93    | 2      | 78     | 3     | 83    | -      | 84     | -     | 84    | -      | 77     | -     | 77    | 1      | 75     | 2     | 78    |    |
| Follow-up | Lower  | -       | 2      | -     | 2     | 1      | 1      | -     | 2     | -      | -      | -     | -     | 3      | -      | 3     | 1     | -      | -      | 1     |       |    |
|           | Normal | 2       | 91     | 1     | 94    | 1      | 79     | 2     | 82    | 1      | 91     | -     | 92    | -      | 82     | -     | 82    | -      | 83     | 2     | 85    |    |
|           | Upper  | -       | 1      | -     | 1     | -      | 1      | 1     | 2     | -      | -      | -     | -     | 2      | -      | 2     | -     | 2      | -      | 2     |       |    |
|           | Total  | 2       | 94     | 1     | 97    | 2      | 81     | 3     | 86    | 1      | 91     | -     | 92    | -      | 87     | -     | 87    | 1      | 85     | 2     | 88    |    |



表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (16/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| CI        | Week 1 | Lower   | 1      | -     | -     | 1      | 1      | -     | -     | 1      | 1      | -     | -     | 2      | -      | 3     | -     | 3      | -      | -     | -     | -  |
|           |        | Normal  | 1      | 37    | -     | 38     | 2      | 34    | -     | 36     | -      | 37    | -     | 37     | 2      | 32    | -     | 34     | 3      | 44    | -     | 47 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 2      | 37    | -     | 39     | 3      | 34    | -     | 37     | 1      | 38    | -     | 39     | 2      | 35    | -     | 37     | 3      | 44    | -     | 47 |
|           | Week 2 | Lower   | 1      | 1     | -     | 2      | 4      | 3     | -     | 7      | -      | 3     | -     | 3      | 2      | 3     | -     | 5      | 1      | 1     | -     | 2  |
|           |        | Normal  | 5      | 89    | -     | 94     | 3      | 79    | -     | 82     | 2      | 86    | -     | 88     | 6      | 80    | -     | 86     | 5      | 82    | -     | 87 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 6      | 90    | -     | 96     | 7      | 82    | -     | 89     | 2      | 89    | -     | 91     | 8      | 85    | -     | 93     | 6      | 83    | -     | 89 |
|           | Week 3 | Lower   | 1      | 1     | -     | 2      | 1      | -     | -     | 1      | 1      | -     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -  |
|           |        | Normal  | 1      | 32    | -     | 33     | 2      | 33    | -     | 35     | -      | 37    | -     | 37     | 1      | 30    | -     | 31     | 3      | 39    | -     | 42 |
|           |        | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 2      | 34    | -     | 36     | 3      | 33    | -     | 36     | 1      | 37    | -     | 38     | 1      | 31    | -     | 32     | 3      | 39    | -     | 42 |
|           | Week 4 | Lower   | 4      | 4     | -     | 8      | 2      | 2     | -     | 4      | 1      | 1     | -     | 2      | 1      | 3     | -     | 4      | -      | -     | -     | -  |
|           |        | Normal  | 2      | 85    | -     | 87     | 5      | 80    | -     | 85     | 1      | 86    | -     | 87     | 7      | 78    | -     | 85     | 6      | 81    | -     | 87 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 6      | 89    | -     | 95     | 7      | 82    | -     | 89     | 2      | 87    | -     | 89     | 8      | 81    | -     | 89     | 6      | 81    | -     | 87 |
|           | Week 6 | Lower   | 1      | 2     | -     | 3      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | 1      | -     | -     | 1  |
|           |        | Normal  | 1      | 36    | -     | 37     | 4      | 35    | -     | 39     | 1      | 36    | -     | 37     | 2      | 28    | -     | 30     | 2      | 44    | -     | 46 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 2      | 38    | -     | 40     | 4      | 36    | -     | 40     | 1      | 37    | -     | 38     | 2      | 29    | -     | 31     | 3      | 44    | -     | 47 |
|           | Week 8 | Lower   | 2      | 4     | -     | 6      | 1      | 2     | -     | 3      | -      | 1     | -     | 1      | -      | 4     | -     | 4      | 1      | -     | -     | 1  |
|           |        | Normal  | 4      | 83    | -     | 87     | 6      | 77    | -     | 83     | 2      | 86    | -     | 88     | 7      | 74    | -     | 81     | 5      | 78    | -     | 83 |
|           |        | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -  |
|           |        | Total   | 6      | 88    | -     | 94     | 7      | 79    | -     | 86     | 2      | 87    | -     | 89     | 7      | 79    | -     | 86     | 6      | 78    | -     | 84 |
| Week 12   | Lower  | 3       | 6      | -     | 9     | 2      | 2      | -     | 4     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           | Normal | 3       | 82     | -     | 85    | 5      | 77     | -     | 82    | 2      | 84     | -     | 86    | 7      | 73     | -     | 80    | 6      | 77     | -     | 83    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 6       | 88     | -     | 94    | 7      | 79     | -     | 86    | 2      | 84     | -     | 86    | 7      | 75     | -     | 82    | 6      | 77     | -     | 83    |    |
| Week 16   | Lower  | 3       | 3      | -     | 6     | 2      | 1      | -     | 3     | 1      | 2      | -     | 3     | -      | -      | -     | -     | -      | 2      | -     | 2     |    |
|           | Normal | 3       | 85     | -     | 88    | 5      | 76     | -     | 81    | 1      | 81     | -     | 82    | 7      | 73     | -     | 80    | 6      | 74     | -     | 80    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 6       | 88     | -     | 94    | 7      | 77     | -     | 84    | 2      | 83     | -     | 85    | 7      | 73     | -     | 80    | 6      | 76     | -     | 82    |    |
| Week 20   | Lower  | 2       | 5      | -     | 7     | 1      | 1      | -     | 2     | 1      | -      | -     | 1     | -      | 1      | -     | 1     | -      | 2      | -     | 2     |    |
|           | Normal | 4       | 81     | -     | 85    | 6      | 75     | -     | 81    | 1      | 82     | -     | 83    | 7      | 71     | -     | 78    | 6      | 72     | -     | 78    |    |
|           | Upper  | -       | 2      | -     | 2     | -      | 1      | -     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 6       | 88     | -     | 94    | 7      | 77     | -     | 84    | 2      | 83     | -     | 85    | 7      | 72     | -     | 79    | 6      | 74     | -     | 80    |    |
| Week 24   | Lower  | 1       | 5      | -     | 6     | 1      | 1      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | 2      | 1      | -     | 3     |    |
|           | Normal | 5       | 82     | -     | 87    | 6      | 76     | -     | 82    | 2      | 82     | -     | 84    | 7      | 71     | -     | 78    | 4      | 72     | -     | 76    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 6       | 87     | -     | 93    | 7      | 77     | -     | 84    | 2      | 82     | -     | 84    | 7      | 71     | -     | 78    | 6      | 73     | -     | 79    |    |
| Week 28   | Lower  | 2       | 4      | -     | 6     | 1      | 2      | -     | 3     | 1      | 2      | -     | 3     | 1      | 2      | -     | 3     | 1      | 3      | -     | 4     |    |
|           | Normal | 4       | 81     | -     | 85    | 6      | 74     | -     | 80    | 1      | 80     | -     | 81    | 6      | 68     | -     | 74    | 5      | 69     | -     | 74    |    |
|           | Upper  | -       | 2      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 6       | 87     | -     | 93    | 7      | 76     | -     | 83    | 2      | 82     | -     | 84    | 7      | 70     | -     | 77    | 6      | 72     | -     | 78    |    |
| Follow-up | Lower  | 2       | 4      | -     | 6     | 1      | 2      | -     | 3     | 1      | 2      | -     | 3     | 1      | 3      | -     | 4     | 1      | 3      | -     | 4     |    |
|           | Normal | 4       | 84     | -     | 88    | 6      | 77     | -     | 83    | 1      | 88     | -     | 89    | 7      | 76     | -     | 83    | 5      | 79     | -     | 84    |    |
|           | Upper  | -       | 3      | -     | 3     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | 6       | 91     | -     | 97    | 7      | 79     | -     | 86    | 2      | 90     | -     | 92    | 8      | 79     | -     | 87    | 6      | 82     | -     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (17/24)

|           |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Ca        | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 37    | 1     | 38     | -      | 36    | 1     | 37     | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 46    | 1     | 47 |
|           |         | Upper   | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | -      | 37    | 2     | 39     | -      | 36    | 1     | 37     | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 46    | 1     | 47 |
|           | Week 2  | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 88    | 6     | 94     | -      | 85    | 2     | 87     | -      | 88    | 1     | 89     | -      | 91    | -     | 91     | -      | 87    | 2     | 89 |
|           |         | Upper   | -      | -     | 2     | 2      | -      | 1     | -     | 1      | -      | 1     | 1     | 2      | -      | 2     | -     | 2      | -      | -     | -     | -  |
|           |         | Total   | -      | 88    | 8     | 96     | -      | 87    | 2     | 89     | -      | 89    | 2     | 91     | -      | 93    | -     | 93     | -      | 87    | 2     | 89 |
|           | Week 3  | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 33    | 1     | 34     | -      | 34    | -     | 34     | -      | 37    | -     | 37     | -      | 32    | -     | 32     | -      | 41    | 1     | 42 |
|           |         | Upper   | -      | 1     | 1     | 2      | -      | -     | 1     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | -      | 34    | 2     | 36     | -      | 35    | 1     | 36     | -      | 38    | -     | 38     | -      | 32    | -     | 32     | -      | 41    | 1     | 42 |
|           | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 84    | 5     | 89     | -      | 87    | 2     | 89     | -      | 86    | 2     | 88     | -      | 88    | -     | 88     | -      | 84    | 2     | 86 |
|           |         | Upper   | -      | 3     | 3     | 6      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 1     | -     | 1  |
|           |         | Total   | -      | 87    | 8     | 95     | -      | 87    | 2     | 89     | -      | 87    | 2     | 89     | -      | 89    | -     | 89     | -      | 85    | 2     | 87 |
|           | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 39    | -     | 39     | -      | 39    | -     | 39     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 46    | 1     | 47 |
|           |         | Upper   | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | -      | 39    | 1     | 40     | -      | 39    | 1     | 40     | -      | 38    | -     | 38     | -      | 31    | -     | 31     | -      | 46    | 1     | 47 |
|           | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |         | Normal  | -      | 84    | 7     | 91     | -      | 83    | 1     | 84     | -      | 85    | 2     | 87     | -      | 84    | -     | 84     | -      | 80    | 2     | 82 |
|           |         | Upper   | -      | 2     | 1     | 3      | -      | 1     | 1     | 2      | -      | 2     | -     | 2      | -      | 2     | -     | 2      | -      | 1     | -     | 1  |
|           |         | Total   | -      | 86    | 8     | 94     | -      | 84    | 2     | 86     | -      | 87    | 2     | 89     | -      | 86    | -     | 86     | -      | 82    | 2     | 84 |
|           | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |         | Normal  | -      | 86    | 5     | 91     | -      | 83    | 2     | 85     | -      | 84    | 1     | 85     | -      | 82    | -     | 82     | -      | 80    | 2     | 82 |
|           |         | Upper   | -      | -     | 3     | 3      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | -      | 86    | 8     | 94     | -      | 84    | 2     | 86     | -      | 84    | 2     | 86     | -      | 82    | -     | 82     | -      | 81    | 2     | 83 |
|           | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |         | Normal  | -      | 84    | 5     | 89     | -      | 81    | 1     | 82     | -      | 83    | 1     | 84     | -      | 77    | -     | 77     | -      | 77    | 2     | 79 |
|           |         | Upper   | -      | 2     | 3     | 5      | -      | 1     | 1     | 2      | -      | 1     | -     | 1      | -      | 3     | -     | 3      | -      | 2     | -     | 2  |
|           |         | Total   | -      | 86    | 8     | 94     | -      | 82    | 2     | 84     | -      | 84    | 1     | 85     | -      | 80    | -     | 80     | -      | 80    | 2     | 82 |
|           | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Normal  | -      | 85    | 6     | 91     | -      | 82    | 2     | 84     | -      | 81    | 1     | 82     | -      | 79    | -     | 79     | -      | 77    | 2     | 79 |
|           |         | Upper   | -      | 1     | 2     | 3      | -      | -     | -     | -      | -      | 3     | -     | 3      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |         | Total   | -      | 86    | 8     | 94     | -      | 82    | 2     | 84     | -      | 84    | 1     | 85     | -      | 79    | -     | 79     | -      | 78    | 2     | 80 |
|           | Week 24 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Normal  | -      | 84    | 5     | 89     | -      | 81    | 2     | 83     | -      | 82    | -     | 82     | -      | 76    | -     | 76     | -      | 75    | 2     | 77 |
|           |         | Upper   | -      | 1     | 3     | 4      | -      | 1     | -     | 1      | -      | 1     | 1     | 2      | -      | 2     | -     | 2      | -      | 2     | -     | 2  |
|           |         | Total   | -      | 85    | 8     | 93     | -      | 82    | 2     | 84     | -      | 83    | 1     | 84     | -      | 78    | -     | 78     | -      | 77    | 2     | 79 |
| Week 28   | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Normal  | -       | 82     | 4     | 86    | -      | 78     | 1     | 79    | -      | 80     | 1     | 81    | -      | 77     | -     | 77    | -      | 74     | 2     | 76    |    |
|           | Upper   | -       | 3      | 4     | 7     | -      | 3      | 1     | 4     | -      | 3      | -     | 3     | -      | -      | -     | -     | -      | 2      | -     | 2     |    |
|           | Total   | -       | 85     | 8     | 93    | -      | 81     | 2     | 83    | -      | 83     | 1     | 84    | -      | 77     | -     | 77    | -      | 76     | 2     | 78    |    |
| Follow-up | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Normal  | -       | 86     | 4     | 90    | -      | 81     | 1     | 82    | -      | 87     | 1     | 88    | -      | 87     | -     | 87    | -      | 84     | 2     | 86    |    |
|           | Upper   | -       | 3      | 4     | 7     | -      | 3      | 1     | 4     | -      | 3      | 1     | 4     | -      | -      | -     | -     | -      | 2      | -     | 2     |    |
|           | Total   | -       | 89     | 8     | 97    | -      | 84     | 2     | 86    | -      | 90     | 2     | 92    | -      | 87     | -     | 87    | -      | 86     | 2     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (18/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| P         | Week 1 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 37    | 1     | 38     | -      | 35    | 1     | 36     | -      | 38    | -     | 38     | -      | 35    | -     | 35     | -      | 43    | 1     | 44 |
|           |        | Upper   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | -     | 2     | 2      | -      | 2     | 1     | 3  |
|           |        | Total   | 1      | 37    | 1     | 39     | -      | 36    | 1     | 37     | -      | 38    | 1     | 39     | -      | 35    | 2     | 37     | -      | 45    | 2     | 47 |
|           | Week 2 | Lower   | 1      | -     | -     | 1      | -      | 1     | -     | 1      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           |        | Normal  | 2      | 89    | 2     | 93     | -      | 85    | 2     | 87     | 1      | 84    | -     | 85     | -      | 86    | 2     | 88     | -      | 84    | 2     | 86 |
|           |        | Upper   | -      | 1     | 1     | 2      | -      | 1     | -     | 1      | -      | 5     | 1     | 6      | -      | 3     | 1     | 4      | -      | 3     | -     | 3  |
|           |        | Total   | 3      | 90    | 3     | 96     | -      | 87    | 2     | 89     | 1      | 89    | 1     | 91     | -      | 90    | 3     | 93     | -      | 87    | 2     | 89 |
|           | Week 3 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 33    | -     | 33     | -      | 34    | 1     | 35     | -      | 36    | -     | 36     | -      | 30    | -     | 30     | -      | 40    | 2     | 42 |
|           |        | Upper   | -      | 1     | 1     | 2      | -      | 1     | -     | 1      | -      | 1     | 1     | 2      | -      | 1     | 1     | 2      | -      | -     | -     | -  |
|           |        | Total   | 1      | 34    | 1     | 36     | -      | 35    | 1     | 36     | -      | 37    | 1     | 38     | -      | 31    | 1     | 32     | -      | 40    | 2     | 42 |
|           | Week 4 | Lower   | 1      | 1     | -     | 2      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | 2      | 87    | 1     | 90     | -      | 83    | 2     | 85     | 1      | 85    | -     | 86     | -      | 84    | 2     | 86     | -      | 84    | 1     | 85 |
|           |        | Upper   | -      | 2     | 1     | 3      | -      | 2     | -     | 2      | -      | 1     | 1     | 2      | -      | 1     | 2     | 3      | -      | 1     | 1     | 2  |
|           |        | Total   | 3      | 90    | 2     | 95     | -      | 87    | 2     | 89     | 1      | 87    | 1     | 89     | -      | 85    | 4     | 89     | -      | 85    | 2     | 87 |
|           | Week 6 | Lower   | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 37    | -     | 37     | -      | 39    | 1     | 40     | -      | 34    | -     | 34     | -      | 30    | -     | 30     | -      | 44    | 2     | 46 |
|           |        | Upper   | -      | 1     | 1     | 2      | -      | -     | -     | -      | -      | 2     | 1     | 3      | -      | 1     | -     | 1      | -      | 1     | -     | 1  |
|           |        | Total   | 1      | 38    | 1     | 40     | -      | 39    | 1     | 40     | -      | 37    | 1     | 38     | -      | 31    | -     | 31     | -      | 45    | 2     | 47 |
|           | Week 8 | Lower   | 1      | 2     | -     | 3      | -      | 3     | -     | 3      | -      | -     | -     | -      | 1      | -     | 1     | -      | 1      | -     | 1     |    |
|           |        | Normal  | 2      | 87    | 1     | 90     | -      | 81    | 2     | 83     | 1      | 86    | 1     | 88     | -      | 82    | 2     | 84     | -      | 79    | 2     | 81 |
|           |        | Upper   | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 1     | -     | 1      | -      | 2     | -     | 2  |
|           |        | Total   | 3      | 89    | 2     | 94     | -      | 84    | 2     | 86     | 1      | 87    | 1     | 89     | -      | 84    | 2     | 86     | -      | 82    | 2     | 84 |
| Week 12   | Lower  | 1       | 1      | -     | 2     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | 2       | 86     | 1     | 89    | -      | 81     | 2     | 83    | 1      | 81     | -     | 82    | -      | 76     | 1     | 77    | -      | 80     | 1     | 81    |    |
|           | Upper  | -       | 2      | 1     | 3     | -      | 1      | -     | 1     | -      | 2      | 1     | 3     | -      | 4      | 1     | 5     | -      | 1      | 1     | 2     |    |
|           | Total  | 3       | 89     | 2     | 94    | -      | 84     | 2     | 86    | 1      | 84     | 1     | 86    | -      | 80     | 2     | 82    | -      | 81     | 2     | 83    |    |
| Week 16   | Lower  | -       | 1      | -     | 1     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | 1      | -     | 1     | -      | 1      | -     | 1     |    |
|           | Normal | 3       | 86     | 1     | 90    | -      | 79     | 2     | 81    | 1      | 80     | -     | 81    | -      | 78     | 1     | 79    | -      | 78     | 2     | 80    |    |
|           | Upper  | -       | 2      | 1     | 3     | -      | 1      | -     | 1     | -      | 2      | 1     | 3     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Total  | 3       | 89     | 2     | 94    | -      | 82     | 2     | 84    | 1      | 83     | 1     | 85    | -      | 79     | 1     | 80    | -      | 80     | 2     | 82    |    |
| Week 20   | Lower  | 1       | 2      | -     | 3     | -      | 2      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | 2       | 86     | 1     | 89    | -      | 80     | 2     | 82    | 1      | 79     | 1     | 81    | -      | 74     | 1     | 75    | -      | 75     | 1     | 76    |    |
|           | Upper  | -       | 1      | 1     | 2     | -      | -      | -     | -     | -      | 4      | -     | 4     | -      | 4      | -     | 4     | -      | 3      | 1     | 4     |    |
|           | Total  | 3       | 89     | 2     | 94    | -      | 82     | 2     | 84    | 1      | 83     | 1     | 85    | -      | 78     | 1     | 79    | -      | 78     | 2     | 80    |    |
| Week 24   | Lower  | 1       | 1      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Normal | 2       | 84     | 1     | 87    | -      | 81     | 2     | 83    | 1      | 81     | 1     | 83    | -      | 77     | 1     | 78    | -      | 74     | 2     | 76    |    |
|           | Upper  | -       | 3      | 1     | 4     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | 2      | -     | 2     |    |
|           | Total  | 3       | 88     | 2     | 93    | -      | 82     | 2     | 84    | 1      | 82     | 1     | 84    | -      | 77     | 1     | 78    | -      | 77     | 2     | 79    |    |
| Week 28   | Lower  | 1       | 1      | -     | 2     | -      | 3      | -     | 3     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | 2       | 87     | 2     | 91    | -      | 76     | 2     | 78    | 1      | 81     | -     | 82    | -      | 74     | 1     | 75    | -      | 76     | 2     | 78    |    |
|           | Upper  | -       | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | 1     | 2     | -      | 2      | -     | 2     | -      | -      | -     | -     |    |
|           | Total  | 3       | 88     | 2     | 93    | -      | 81     | 2     | 83    | 1      | 82     | 1     | 84    | -      | 76     | 1     | 77    | -      | 76     | 2     | 78    |    |
| Follow-up | Lower  | 1       | 1      | -     | 2     | -      | 3      | -     | 3     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | 2       | 90     | 3     | 95    | -      | 79     | 2     | 81    | 1      | 89     | -     | 90    | -      | 79     | 3     | 82    | -      | 84     | 2     | 86    |    |
|           | Upper  | -       | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | 1     | 2     | -      | 4      | 1     | 5     | -      | 2      | -     | 2     |    |
|           | Total  | 3       | 91     | 3     | 97    | -      | 84     | 2     | 86    | 1      | 90     | 1     | 92    | -      | 83     | 4     | 87    | -      | 86     | 2     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (19/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Vitamin A | Week 1 | Lower   | 6      | 3     | -     | 9      | 2      | 4     | -     | 6      | 6      | 2     | -     | 8      | 6      | 1     | -     | 7      | 6      | 6     | -     | 12 |
|           |        | Normal  | 5      | 22    | 1     | 28     | -      | 29    | 1     | 30     | 3      | 27    | -     | 30     | 2      | 27    | 1     | 30     | 1      | 34    | -     | 35 |
|           |        | Upper   | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 11     | 27    | 1     | 39     | 2      | 34    | 1     | 37     | 9      | 29    | 1     | 39     | 8      | 28    | 1     | 37     | 7      | 40    | -     | 47 |
|           | Week 2 | Lower   | 15     | 8     | -     | 23     | 8      | 10    | -     | 18     | 12     | 6     | -     | 18     | 23     | 4     | -     | 27     | 13     | 11    | -     | 24 |
|           |        | Normal  | 7      | 62    | 1     | 70     | 2      | 68    | -     | 70     | 3      | 65    | 3     | 71     | 3      | 58    | 3     | 64     | 3      | 61    | -     | 64 |
|           |        | Upper   | -      | 2     | 2     | 4      | -      | -     | 1     | 1      | -      | 1     | 1     | 2      | -      | -     | 2     | 2      | -      | 1     | -     | 1  |
|           |        | Total   | 22     | 72    | 3     | 97     | 10     | 78    | 1     | 89     | 15     | 72    | 4     | 91     | 26     | 62    | 5     | 93     | 16     | 73    | -     | 89 |
|           | Week 3 | Lower   | 5      | 6     | -     | 11     | 2      | 4     | -     | 6      | 6      | 1     | -     | 7      | 6      | 1     | -     | 7      | 4      | 5     | -     | 9  |
|           |        | Normal  | 3      | 21    | -     | 24     | -      | 29    | -     | 29     | 2      | 28    | -     | 30     | 1      | 23    | 1     | 25     | 2      | 31    | -     | 33 |
|           |        | Upper   | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 8      | 27    | 1     | 36     | 2      | 33    | 1     | 36     | 8      | 29    | 1     | 38     | 7      | 24    | 1     | 32     | 6      | 36    | -     | 42 |
|           | Week 4 | Lower   | 12     | 5     | -     | 17     | 6      | 6     | -     | 12     | 9      | 4     | -     | 13     | 19     | 6     | -     | 25     | 11     | 12    | -     | 23 |
|           |        | Normal  | 10     | 64    | 1     | 75     | 4      | 72    | -     | 76     | 4      | 68    | 2     | 74     | 4      | 55    | 3     | 62     | 5      | 58    | -     | 63 |
|           |        | Upper   | -      | 2     | 2     | 4      | -      | -     | 1     | 1      | -      | -     | 2     | 2      | -      | -     | 2     | 2      | -      | 1     | -     | 1  |
|           |        | Total   | 22     | 71    | 3     | 96     | 10     | 78    | 1     | 89     | 13     | 72    | 4     | 89     | 23     | 61    | 5     | 89     | 16     | 71    | -     | 87 |
|           | Week 6 | Lower   | 5      | 2     | -     | 7      | 3      | 4     | -     | 7      | 6      | 2     | -     | 8      | 3      | 4     | -     | 7      | 6      | 7     | -     | 13 |
|           |        | Normal  | 6      | 25    | -     | 31     | -      | 32    | -     | 32     | 2      | 27    | 1     | 30     | 3      | 20    | 1     | 24     | 2      | 32    | -     | 34 |
|           |        | Upper   | -      | 1     | 1     | 2      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | 11     | 28    | 1     | 40     | 3      | 36    | 1     | 40     | 8      | 29    | 1     | 38     | 6      | 24    | 1     | 31     | 8      | 39    | -     | 47 |
|           | Week 8 | Lower   | 13     | 6     | -     | 19     | 7      | 6     | -     | 13     | 8      | 3     | -     | 11     | 18     | 10    | -     | 28     | 10     | 12    | -     | 22 |
|           |        | Normal  | 9      | 63    | 1     | 73     | 2      | 69    | -     | 71     | 5      | 67    | 3     | 75     | 5      | 48    | 3     | 56     | 5      | 55    | -     | 60 |
|           |        | Upper   | -      | 1     | 2     | 3      | -      | 1     | 1     | 2      | -      | 2     | 1     | 3      | -      | -     | 2     | 2      | -      | 2     | -     | 2  |
|           |        | Total   | 22     | 70    | 3     | 95     | 9      | 76    | 1     | 86     | 13     | 72    | 4     | 89     | 23     | 58    | 5     | 86     | 15     | 69    | -     | 84 |
| Week 12   | Lower  | 13      | 9      | -     | 22    | 7      | 9      | -     | 16    | 8      | 4      | -     | 12    | 15     | 9      | -     | 24    | 12     | 10     | -     | 22    |    |
|           | Normal | 9       | 60     | 1     | 70    | 2      | 66     | -     | 68    | 4      | 65     | 2     | 71    | 6      | 47     | 3     | 56    | 3      | 57     | -     | 60    |    |
|           | Upper  | -       | 1      | 2     | 3     | -      | 1      | 1     | 2     | -      | 1      | 2     | 3     | -      | -      | 2     | 2     | -      | 1      | -     | 1     |    |
|           | Total  | 22      | 70     | 3     | 95    | 9      | 76     | 1     | 86    | 12     | 70     | 4     | 86    | 21     | 56     | 5     | 82    | 15     | 68     | -     | 83    |    |
| Week 16   | Lower  | 13      | 11     | -     | 24    | 6      | 9      | -     | 15    | 8      | 4      | -     | 12    | 15     | 5      | -     | 20    | 10     | 11     | -     | 21    |    |
|           | Normal | 9       | 58     | 1     | 68    | 3      | 65     | -     | 68    | 4      | 65     | 3     | 72    | 6      | 49     | 3     | 58    | 5      | 55     | -     | 60    |    |
|           | Upper  | -       | 1      | 2     | 3     | -      | -      | 1     | 1     | -      | -      | 1     | 1     | -      | 1      | 1     | 2     | -      | 1      | -     | 1     |    |
|           | Total  | 22      | 70     | 3     | 95    | 9      | 74     | 1     | 84    | 12     | 69     | 4     | 85    | 21     | 55     | 4     | 80    | 15     | 67     | -     | 82    |    |
| Week 20   | Lower  | 12      | 9      | -     | 21    | 5      | 7      | -     | 12    | 10     | 6      | -     | 16    | 16     | 7      | -     | 23    | 10     | 11     | -     | 21    |    |
|           | Normal | 10      | 61     | 2     | 73    | 4      | 67     | -     | 71    | 2      | 62     | 2     | 66    | 5      | 47     | 3     | 55    | 5      | 54     | -     | 59    |    |
|           | Upper  | -       | -      | 1     | 1     | -      | -      | 1     | 1     | -      | 1      | 2     | 3     | -      | -      | 1     | 1     | -      | -      | -     | -     |    |
|           | Total  | 22      | 70     | 3     | 95    | 9      | 74     | 1     | 84    | 12     | 69     | 4     | 85    | 21     | 54     | 4     | 79    | 15     | 65     | -     | 80    |    |
| Week 24   | Lower  | 11      | 13     | -     | 24    | 7      | 10     | -     | 17    | 9      | 2      | -     | 11    | 18     | 9      | -     | 27    | 10     | 12     | -     | 22    |    |
|           | Normal | 10      | 56     | 1     | 67    | 2      | 64     | -     | 66    | 3      | 66     | 1     | 70    | 2      | 44     | 2     | 48    | 5      | 52     | -     | 57    |    |
|           | Upper  | -       | 1      | 2     | 3     | -      | -      | 1     | 1     | -      | -      | 3     | 3     | -      | 1      | 2     | 3     | -      | -      | -     | -     |    |
|           | Total  | 21      | 70     | 3     | 94    | 9      | 74     | 1     | 84    | 12     | 68     | 4     | 84    | 20     | 54     | 4     | 78    | 15     | 64     | -     | 79    |    |
| Week 28   | Lower  | 10      | 9      | -     | 19    | 6      | 9      | -     | 15    | 9      | 5      | -     | 14    | 12     | 8      | -     | 20    | 9      | 12     | -     | 21    |    |
|           | Normal | 11      | 61     | 1     | 73    | 3      | 64     | 1     | 68    | 3      | 63     | 3     | 69    | 8      | 45     | 2     | 55    | 6      | 51     | -     | 57    |    |
|           | Upper  | -       | -      | 2     | 2     | -      | -      | -     | -     | -      | -      | 1     | 1     | -      | -      | 2     | 2     | -      | -      | -     | -     |    |
|           | Total  | 21      | 70     | 3     | 94    | 9      | 73     | 1     | 83    | 12     | 68     | 4     | 84    | 20     | 53     | 4     | 77    | 15     | 63     | -     | 78    |    |
| Follow-up | Lower  | 11      | 10     | -     | 21    | 6      | 9      | -     | 15    | 13     | 6      | -     | 19    | 14     | 10     | -     | 24    | 10     | 13     | -     | 23    |    |
|           | Normal | 11      | 63     | 1     | 75    | 3      | 67     | 1     | 71    | 3      | 66     | 3     | 72    | 8      | 50     | 2     | 60    | 6      | 59     | -     | 65    |    |
|           | Upper  | -       | -      | 2     | 2     | -      | -      | -     | -     | -      | -      | 1     | 1     | -      | -      | 3     | 3     | -      | -      | -     | -     |    |
|           | Total  | 22      | 73     | 3     | 98    | 9      | 76     | 1     | 86    | 16     | 72     | 4     | 92    | 22     | 60     | 5     | 87    | 16     | 72     | -     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (20/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Vitamin D | Week 1 | Lower   | -      | 1     | -     | 1      | -      | 2     | -     | 2      | 1      | -     | 1     | -      | 2      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 26    | 5     | 31     | -      | 16    | 5     | 21     | 1      | 20    | 4     | 25     | 1      | 19    | -     | 20     | 1      | 23    | 9     | 33 |
|           |        | Upper   | -      | 2     | 5     | 7      | -      | 8     | 6     | 14     | -      | 4     | 8     | 12     | -      | 6     | 11    | 17     | -      | 8     | 6     | 14 |
|           |        | Total   | -      | 29    | 10    | 39     | -      | 26    | 11    | 37     | 2      | 25    | 12    | 39     | 1      | 25    | 11    | 37     | 1      | 31    | 15    | 47 |
|           | Week 2 | Lower   | -      | 2     | -     | 2      | -      | -     | -     | -      | 1      | 2     | -     | 3      | 1      | 4     | -     | 5      | -      | 2     | -     | 2  |
|           |        | Normal  | 2      | 60    | 11    | 73     | -      | 47    | 12    | 59     | 2      | 46    | 15    | 63     | 1      | 44    | 19    | 64     | 1      | 49    | 17    | 67 |
|           |        | Upper   | -      | 6     | 16    | 22     | -      | 17    | 13    | 30     | -      | 15    | 10    | 25     | -      | 9     | 15    | 24     | -      | 9     | 11    | 20 |
|           |        | Total   | 2      | 68    | 27    | 97     | -      | 64    | 25    | 89     | 3      | 63    | 25    | 91     | 2      | 57    | 34    | 93     | 1      | 60    | 28    | 89 |
|           | Week 3 | Lower   | -      | 2     | -     | 2      | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     | -      | 1      | -     | -     | 1  |
|           |        | Normal  | -      | 20    | 7     | 27     | -      | 21    | 3     | 24     | 1      | 19    | 5     | 25     | 1      | 18    | 2     | 21     | -      | 24    | 7     | 31 |
|           |        | Upper   | -      | 5     | 2     | 7      | -      | 6     | 6     | 12     | -      | 5     | 7     | 12     | -      | 4     | 7     | 11     | -      | 3     | 7     | 10 |
|           |        | Total   | -      | 27    | 9     | 36     | -      | 27    | 9     | 36     | 2      | 24    | 12    | 38     | 1      | 22    | 9     | 32     | 1      | 27    | 14    | 42 |
|           | Week 4 | Lower   | 1      | 1     | -     | 2      | -      | 1     | -     | 1      | 1      | 1     | -     | 2      | 2      | 3     | -     | 5      | -      | -     | -     | -  |
|           |        | Normal  | 1      | 56    | 11    | 68     | -      | 57    | 13    | 70     | 2      | 48    | 11    | 61     | -      | 52    | 11    | 63     | 1      | 47    | 15    | 63 |
|           |        | Upper   | -      | 10    | 16    | 26     | -      | 6     | 12    | 18     | -      | 12    | 14    | 26     | -      | 2     | 19    | 21     | -      | 11    | 13    | 24 |
|           |        | Total   | 2      | 67    | 27    | 96     | -      | 64    | 25    | 89     | 3      | 61    | 25    | 89     | 2      | 57    | 30    | 89     | 1      | 58    | 28    | 87 |
|           | Week 6 | Lower   | -      | 1     | -     | 1      | -      | -     | 1     | 1      | 1      | -     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Normal  | -      | 24    | 4     | 28     | -      | 22    | 4     | 26     | 1      | 19    | 6     | 26     | 1      | 17    | 3     | 21     | 1      | 28    | 8     | 37 |
|           |        | Upper   | -      | 4     | 7     | 11     | -      | 8     | 5     | 13     | -      | 4     | 7     | 11     | -      | 4     | 6     | 10     | -      | 5     | 5     | 10 |
|           |        | Total   | -      | 29    | 11    | 40     | -      | 30    | 10    | 40     | 2      | 23    | 13    | 38     | 1      | 21    | 9     | 31     | 1      | 33    | 13    | 47 |
| Week 8    | Lower  | -       | 2      | -     | 2     | -      | 2      | -     | 2     | 2      | -      | -     | 2     | 2      | 2      | -     | 4     | -      | 2      | -     | 2     |    |
|           | Normal | 2       | 58     | 13    | 73    | -      | 51     | 10    | 61    | 1      | 48     | 10    | 59    | -      | 48     | 15    | 63    | 1      | 45     | 12    | 58    |    |
|           | Upper  | -       | 6      | 14    | 20    | -      | 9      | 14    | 23    | -      | 13     | 15    | 28    | -      | 5      | 14    | 19    | -      | 11     | 13    | 24    |    |
|           | Total  | 2       | 66     | 27    | 95    | -      | 62     | 24    | 86    | 3      | 61     | 25    | 89    | 2      | 55     | 29    | 86    | 1      | 58     | 25    | 84    |    |
| Week 12   | Lower  | -       | 3      | -     | 3     | -      | 2      | -     | 2     | 1      | -      | -     | 1     | 1      | 1      | -     | 2     | -      | 2      | -     | 2     |    |
|           | Normal | 2       | 53     | 12    | 67    | -      | 51     | 12    | 63    | 2      | 48     | 10    | 60    | 1      | 46     | 12    | 59    | 1      | 44     | 14    | 59    |    |
|           | Upper  | -       | 10     | 15    | 25    | -      | 9      | 12    | 21    | -      | 12     | 13    | 25    | -      | 6      | 15    | 21    | -      | 11     | 11    | 22    |    |
|           | Total  | 2       | 66     | 27    | 95    | -      | 62     | 24    | 86    | 3      | 60     | 23    | 86    | 2      | 53     | 27    | 82    | 1      | 57     | 25    | 83    |    |
| Week 16   | Lower  | 1       | 4      | -     | 5     | -      | 3      | -     | 3     | -      | 1      | -     | 1     | -      | 3      | -     | 3     | -      | 3      | -     | 3     |    |
|           | Normal | 1       | 49     | 9     | 59    | -      | 44     | 13    | 57    | 2      | 48     | 13    | 63    | 1      | 45     | 12    | 58    | 1      | 44     | 15    | 60    |    |
|           | Upper  | -       | 13     | 18    | 31    | -      | 13     | 11    | 24    | -      | 11     | 10    | 21    | -      | 4      | 15    | 19    | -      | 8      | 11    | 19    |    |
|           | Total  | 2       | 66     | 27    | 95    | -      | 60     | 24    | 84    | 2      | 60     | 23    | 85    | 1      | 52     | 27    | 80    | 1      | 55     | 26    | 82    |    |
| Week 20   | Lower  | -       | 1      | -     | 1     | -      | 3      | -     | 3     | 1      | 1      | -     | 2     | 1      | 1      | -     | 2     | -      | 3      | -     | 3     |    |
|           | Normal | 2       | 54     | 9     | 65    | -      | 49     | 12    | 61    | 1      | 47     | 13    | 61    | -      | 45     | 14    | 59    | 1      | 40     | 16    | 57    |    |
|           | Upper  | -       | 11     | 18    | 29    | -      | 8      | 12    | 20    | -      | 12     | 10    | 22    | -      | 6      | 12    | 18    | -      | 10     | 10    | 20    |    |
|           | Total  | 2       | 66     | 27    | 95    | -      | 60     | 24    | 84    | 2      | 60     | 23    | 85    | 1      | 52     | 26    | 79    | 1      | 53     | 26    | 80    |    |
| Week 24   | Lower  | -       | 1      | -     | 1     | -      | -      | -     | -     | -      | 1      | -     | 1     | 1      | 2      | -     | 3     | -      | 2      | -     | 2     |    |
|           | Normal | 2       | 54     | 8     | 64    | -      | 52     | 17    | 69    | 2      | 43     | 6     | 51    | -      | 45     | 9     | 54    | 1      | 41     | 15    | 57    |    |
|           | Upper  | -       | 10     | 19    | 29    | -      | 8      | 7     | 15    | -      | 15     | 17    | 32    | -      | 4      | 17    | 21    | -      | 9      | 11    | 20    |    |
|           | Total  | 2       | 65     | 27    | 94    | -      | 60     | 24    | 84    | 2      | 59     | 23    | 84    | 1      | 51     | 26    | 78    | 1      | 52     | 26    | 79    |    |
| Week 28   | Lower  | -       | 1      | -     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     | 1      | 3      | -     | 4     | -      | 1      | -     | 1     |    |
|           | Normal | 2       | 50     | 12    | 64    | -      | 50     | 11    | 61    | 2      | 46     | 9     | 57    | -      | 45     | 14    | 59    | 1      | 40     | 14    | 55    |    |
|           | Upper  | -       | 14     | 15    | 29    | -      | 8      | 13    | 21    | -      | 13     | 14    | 27    | -      | 3      | 11    | 14    | -      | 10     | 12    | 22    |    |
|           | Total  | 2       | 65     | 27    | 94    | -      | 59     | 24    | 83    | 2      | 59     | 23    | 84    | 1      | 51     | 25    | 77    | 1      | 51     | 26    | 78    |    |
| Follow-up | Lower  | -       | 1      | -     | 1     | -      | 1      | 1     | 2     | 1      | -      | -     | 1     | 2      | 3      | -     | 5     | -      | 1      | -     | 1     |    |
|           | Normal | 2       | 53     | 12    | 67    | -      | 51     | 11    | 62    | 2      | 50     | 10    | 62    | -      | 47     | 16    | 63    | 1      | 48     | 16    | 65    |    |
|           | Upper  | -       | 15     | 15    | 30    | -      | 9      | 13    | 22    | -      | 13     | 16    | 29    | -      | 6      | 13    | 19    | -      | 10     | 12    | 22    |    |
|           | Total  | 2       | 69     | 27    | 98    | -      | 61     | 25    | 86    | 3      | 63     | 26    | 92    | 2      | 56     | 29    | 87    | 1      | 59     | 28    | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (21/24)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Vitamin E | Week 1 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 13    | 1     | 14     | -      | 9     | 8     | 17     | -      | 11    | 10    | 21     | -      | 14    | 4     | 18     | -      | 18    | 9     | 27 |
|           |        | Upper   | -      | 3     | 22    | 25     | -      | -     | 20    | 20     | -      | 3     | 15    | 18     | -      | -     | 17    | 17     | -      | 1     | 19    | 20 |
|           |        | Total   | -      | 16    | 23    | 39     | -      | 9     | 28    | 37     | -      | 14    | 25    | 39     | -      | 16    | 21    | 37     | -      | 19    | 28    | 47 |
|           | Week 2 | Lower   | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     | -      | 2      | -     | 2     | -      | 1      | -     | 1     |    |
|           |        | Normal  | -      | 36    | 7     | 43     | -      | 39    | 15    | 54     | -      | 27    | 25    | 52     | -      | 33    | 18    | 51     | 1      | 29    | 22    | 52 |
|           |        | Upper   | -      | 8     | 46    | 54     | -      | 1     | 34    | 35     | -      | 3     | 35    | 38     | -      | 5     | 35    | 40     | -      | 2     | 34    | 36 |
|           |        | Total   | -      | 44    | 53    | 97     | -      | 40    | 49    | 89     | 1      | 30    | 60    | 91     | -      | 40    | 53    | 93     | 1      | 32    | 56    | 89 |
|           | Week 3 | Lower   | -      | -     | -     | -      | -      | -     | 1     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 13    | 1     | 14     | -      | 9     | 9     | 18     | -      | 12    | 10    | 22     | -      | 12    | 10    | 22     | -      | 14    | 15    | 29 |
|           |        | Upper   | -      | 2     | 20    | 22     | -      | -     | 17    | 17     | -      | 2     | 14    | 16     | -      | 2     | 8     | 10     | -      | 1     | 12    | 13 |
|           |        | Total   | -      | 15    | 21    | 36     | -      | 9     | 27    | 36     | -      | 14    | 24    | 38     | -      | 14    | 18    | 32     | -      | 15    | 27    | 42 |
|           | Week 4 | Lower   | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 30    | 8     | 38     | -      | 33    | 17    | 50     | -      | 29    | 26    | 55     | -      | 34    | 20    | 54     | 1      | 28    | 23    | 52 |
|           |        | Upper   | -      | 12    | 46    | 58     | -      | 7     | 32    | 39     | -      | 1     | 32    | 33     | -      | 4     | 30    | 34     | -      | 3     | 32    | 35 |
|           |        | Total   | -      | 42    | 54    | 96     | -      | 40    | 49    | 89     | 1      | 30    | 58    | 89     | -      | 39    | 50    | 89     | 1      | 31    | 55    | 87 |
|           | Week 6 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | 1     | -      | 1      | -     | 1     | -      | 2      | -     | 2     |    |
|           |        | Normal  | -      | 14    | 2     | 16     | -      | 14    | 4     | 18     | -      | 11    | 13    | 24     | -      | 12    | 8     | 20     | -      | 14    | 14    | 28 |
|           |        | Upper   | -      | 3     | 21    | 24     | -      | 1     | 21    | 22     | -      | 5     | 8     | 13     | -      | 1     | 9     | 10     | -      | 2     | 15    | 17 |
|           |        | Total   | -      | 17    | 23    | 40     | -      | 15    | 25    | 40     | -      | 16    | 22    | 38     | -      | 14    | 17    | 31     | -      | 18    | 29    | 47 |
|           | Week 8 | Lower   | -      | -     | -     | -      | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |        | Normal  | -      | 34    | 9     | 43     | -      | 31    | 16    | 47     | -      | 25    | 31    | 56     | -      | 36    | 24    | 60     | 1      | 27    | 27    | 55 |
|           |        | Upper   | -      | 7     | 45    | 52     | -      | 7     | 32    | 39     | -      | 5     | 27    | 32     | -      | 3     | 23    | 26     | -      | 1     | 27    | 28 |
|           |        | Total   | -      | 41    | 54    | 95     | -      | 38    | 48    | 86     | 1      | 30    | 58    | 89     | -      | 39    | 47    | 86     | 1      | 29    | 54    | 84 |
| Week 12   | Lower  | -       | -      | -     | -     | -      | -      | -     | 1     | 1      | -      | 2     | -     | 1      | -      | 1     | -     | 1      | -      | 1     |       |    |
|           | Normal | -       | 35     | 13    | 48    | -      | 35     | 16    | 51    | -      | 27     | 28    | 55    | -      | 37     | 22    | 59    | -      | 26     | 26    | 52    |    |
|           | Upper  | -       | 6      | 41    | 47    | -      | 3      | 32    | 35    | -      | 1      | 28    | 29    | -      | 1      | 21    | 22    | 1      | 2      | 27    | 30    |    |
|           | Total  | -       | 41     | 54    | 95    | -      | 38     | 48    | 86    | 1      | 29     | 56    | 86    | -      | 39     | 43    | 82    | 1      | 29     | 53    | 83    |    |
| Week 16   | Lower  | -       | 1      | -     | 1     | -      | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     | -     | -      | 1      | 1     | 2     |    |
|           | Normal | -       | 33     | 12    | 45    | -      | 34     | 16    | 50    | -      | 25     | 26    | 51    | -      | 37     | 19    | 56    | -      | 24     | 26    | 50    |    |
|           | Upper  | -       | 7      | 42    | 49    | -      | 3      | 31    | 34    | -      | 4      | 29    | 33    | -      | 2      | 22    | 24    | 1      | 3      | 26    | 30    |    |
|           | Total  | -       | 41     | 54    | 95    | -      | 37     | 47    | 84    | 1      | 29     | 55    | 85    | -      | 39     | 41    | 80    | 1      | 28     | 53    | 82    |    |
| Week 20   | Lower  | -       | -      | -     | -     | -      | -      | -     | 1     | 1      | -      | 2     | -     | -      | 1      | 1     | -     | 2      | -      | 2     |       |    |
|           | Normal | -       | 32     | 11    | 43    | -      | 35     | 20    | 55    | -      | 25     | 29    | 54    | -      | 36     | 21    | 57    | 1      | 21     | 27    | 49    |    |
|           | Upper  | -       | 9      | 43    | 52    | -      | 2      | 27    | 29    | -      | 3      | 26    | 29    | -      | 3      | 18    | 21    | -      | 5      | 24    | 29    |    |
|           | Total  | -       | 41     | 54    | 95    | -      | 37     | 47    | 84    | 1      | 29     | 55    | 85    | -      | 39     | 40    | 79    | 1      | 28     | 51    | 80    |    |
| Week 24   | Lower  | -       | -      | -     | -     | -      | -      | -     | 1     | 1      | -      | 2     | -     | 1      | -      | 1     | -     | 2      | -      | 2     |       |    |
|           | Normal | -       | 33     | 16    | 49    | -      | 33     | 14    | 47    | -      | 25     | 25    | 50    | -      | 36     | 24    | 60    | -      | 24     | 28    | 52    |    |
|           | Upper  | -       | 7      | 38    | 45    | -      | 4      | 33    | 37    | -      | 2      | 30    | 32    | -      | 2      | 15    | 17    | 1      | 2      | 22    | 25    |    |
|           | Total  | -       | 40     | 54    | 94    | -      | 37     | 47    | 84    | 1      | 28     | 55    | 84    | -      | 39     | 39    | 78    | 1      | 28     | 50    | 79    |    |
| Week 28   | Lower  | -       | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Normal | -       | 35     | 17    | 52    | -      | 29     | 14    | 43    | 1      | 19     | 22    | 42    | -      | 25     | 17    | 42    | -      | 19     | 18    | 37    |    |
|           | Upper  | -       | 5      | 37    | 42    | -      | 7      | 32    | 39    | -      | 9      | 33    | 42    | -      | 13     | 22    | 35    | 1      | 8      | 31    | 40    |    |
|           | Total  | -       | 40     | 54    | 94    | -      | 37     | 46    | 83    | 1      | 28     | 55    | 84    | -      | 38     | 39    | 77    | 1      | 28     | 49    | 78    |    |
| Follow-up | Lower  | -       | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Normal | -       | 38     | 17    | 55    | -      | 30     | 15    | 45    | 1      | 21     | 24    | 46    | -      | 26     | 18    | 44    | -      | 22     | 20    | 42    |    |
|           | Upper  | -       | 6      | 37    | 43    | -      | 7      | 33    | 40    | -      | 9      | 37    | 46    | -      | 13     | 30    | 43    | 1      | 9      | 35    | 45    |    |
|           | Total  | -       | 44     | 54    | 98    | -      | 38     | 48    | 86    | 1      | 30     | 61    | 92    | -      | 39     | 48    | 87    | 1      | 32     | 55    | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (22/24)

|            |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|------------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|            |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|            |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| β-Carotene | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 38    | 1     | 39     | -      | 35    | -     | 35     | -      | 35    | -     | 35     | -      | 35    | -     | 35     | -      | 45    | 1     | 46 |
|            |         | Upper   | -      | -     | -     | -      | -      | -     | 2     | 2      | -      | -     | 4     | 4      | -      | -     | 2     | 2      | -      | -     | 1     | 1  |
|            |         | Total   | -      | 38    | 1     | 39     | -      | 35    | 2     | 37     | -      | 35    | 4     | 39     | -      | 35    | 2     | 37     | -      | 45    | 2     | 47 |
|            | Week 2  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 93    | 1     | 94     | -      | 86    | 1     | 87     | -      | 84    | 2     | 86     | -      | 89    | -     | 89     | -      | 85    | 1     | 86 |
|            |         | Upper   | -      | -     | 3     | 3      | -      | 1     | 1     | 2      | -      | -     | 5     | 5      | -      | -     | 4     | 4      | -      | 1     | 2     | 3  |
|            |         | Total   | -      | 93    | 4     | 97     | -      | 87    | 2     | 89     | -      | 84    | 7     | 91     | -      | 89    | 4     | 93     | -      | 86    | 3     | 89 |
|            | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 35    | 1     | 36     | -      | 34    | -     | 34     | -      | 34    | 1     | 35     | -      | 30    | -     | 30     | -      | 40    | 2     | 42 |
|            |         | Upper   | -      | -     | -     | -      | -      | -     | 2     | 2      | -      | -     | 3     | 3      | -      | -     | 2     | 2      | -      | -     | -     | -  |
|            |         | Total   | -      | 35    | 1     | 36     | -      | 34    | 2     | 36     | -      | 34    | 4     | 38     | -      | 30    | 2     | 32     | -      | 40    | 2     | 42 |
|            | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 92    | 2     | 94     | -      | 87    | -     | 87     | -      | 82    | 3     | 85     | -      | 85    | -     | 85     | -      | 82    | 1     | 83 |
|            |         | Upper   | -      | -     | 2     | 2      | -      | -     | 2     | 2      | -      | -     | 4     | 4      | -      | -     | 4     | 4      | -      | 2     | 2     | 4  |
|            |         | Total   | -      | 92    | 4     | 96     | -      | 87    | 2     | 89     | -      | 82    | 7     | 89     | -      | 85    | 4     | 89     | -      | 84    | 3     | 87 |
|            | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 39    | -     | 39     | -      | 38    | -     | 38     | -      | 34    | 1     | 35     | -      | 29    | 1     | 30     | -      | 45    | 1     | 46 |
|            |         | Upper   | -      | -     | 1     | 1      | -      | 1     | 1     | 2      | -      | -     | 3     | 3      | -      | -     | 1     | 1      | -      | -     | 1     | 1  |
|            |         | Total   | -      | 39    | 1     | 40     | -      | 39    | 1     | 40     | -      | 34    | 4     | 38     | -      | 29    | 2     | 31     | -      | 45    | 2     | 47 |
|            | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 89    | -     | 89     | -      | 81    | 1     | 82     | -      | 80    | 3     | 83     | -      | 80    | -     | 80     | -      | 79    | 1     | 80 |
|            |         | Upper   | -      | 2     | 4     | 6      | -      | 3     | 1     | 4      | -      | 2     | 4     | 6      | -      | 2     | 4     | 6      | -      | 2     | 2     | 4  |
|            |         | Total   | -      | 91    | 4     | 95     | -      | 84    | 2     | 86     | -      | 82    | 7     | 89     | -      | 82    | 4     | 86     | -      | 81    | 3     | 84 |
|            | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 90    | -     | 90     | -      | 83    | 2     | 85     | -      | 78    | 2     | 80     | -      | 76    | 1     | 77     | -      | 80    | 1     | 81 |
|            |         | Upper   | -      | 1     | 4     | 5      | -      | 1     | -     | 1      | -      | 1     | 5     | 6      | -      | 2     | 3     | 5      | -      | -     | 2     | 2  |
|            |         | Total   | -      | 91    | 4     | 95     | -      | 84    | 2     | 86     | -      | 79    | 7     | 86     | -      | 78    | 4     | 82     | -      | 80    | 3     | 83 |
|            | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 90    | 3     | 93     | -      | 81    | 1     | 82     | -      | 76    | 2     | 78     | -      | 76    | 2     | 78     | -      | 79    | 1     | 80 |
|            |         | Upper   | -      | 1     | 1     | 2      | -      | 1     | 1     | 2      | -      | 2     | 5     | 7      | -      | -     | 2     | 2      | -      | -     | 2     | 2  |
|            |         | Total   | -      | 91    | 4     | 95     | -      | 82    | 2     | 84     | -      | 78    | 7     | 85     | -      | 76    | 4     | 80     | -      | 79    | 3     | 82 |
|            | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 89    | 2     | 91     | -      | 80    | 2     | 82     | -      | 76    | 3     | 79     | -      | 74    | 2     | 76     | -      | 77    | 1     | 78 |
|            |         | Upper   | -      | 2     | 2     | 4      | -      | 2     | -     | 2      | -      | 2     | 4     | 6      | -      | 1     | 2     | 3      | -      | -     | 2     | 2  |
|            |         | Total   | -      | 91    | 4     | 95     | -      | 82    | 2     | 84     | -      | 78    | 7     | 85     | -      | 75    | 4     | 79     | -      | 77    | 3     | 80 |
|            | Week 24 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|            |         | Normal  | -      | 87    | 3     | 90     | -      | 79    | 1     | 80     | -      | 76    | 5     | 81     | -      | 71    | 1     | 72     | -      | 76    | 1     | 77 |
|            |         | Upper   | -      | 3     | 1     | 4      | -      | 3     | 1     | 4      | -      | 1     | 2     | 3      | -      | 3     | 3     | 6      | -      | -     | 2     | 2  |
|            |         | Total   | -      | 90    | 4     | 94     | -      | 82    | 2     | 84     | -      | 77    | 7     | 84     | -      | 74    | 4     | 78     | -      | 76    | 3     | 79 |
| Week 28    | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|            | Normal  | -       | 88     | 3     | 91    | -      | 76     | 1     | 77    | -      | 74     | 2     | 76    | -      | 68     | -     | 68    | -      | 75     | 1     | 76    |    |
|            | Upper   | -       | 2      | 1     | 3     | -      | 5      | 1     | 6     | -      | 3      | 5     | 8     | -      | 5      | 4     | 9     | -      | -      | 2     | 2     |    |
|            | Total   | -       | 90     | 4     | 94    | -      | 81     | 2     | 83    | -      | 77     | 7     | 84    | -      | 73     | 4     | 77    | -      | 75     | 3     | 78    |    |
| Follow-up  | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|            | Normal  | -       | 92     | 3     | 95    | -      | 79     | 1     | 80    | -      | 82     | 2     | 84    | -      | 78     | -     | 78    | -      | 85     | 1     | 86    |    |
|            | Upper   | -       | 2      | 1     | 3     | -      | 5      | 1     | 6     | -      | 3      | 5     | 8     | -      | 5      | 4     | 9     | -      | -      | 2     | 2     |    |
|            | Total   | -       | 94     | 4     | 98    | -      | 84     | 2     | 86    | -      | 85     | 7     | 92    | -      | 83     | 4     | 87    | -      | 85     | 3     | 88    |    |

表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (23/24)

|           |         |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |
|-----------|---------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|
|           |         |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |
|           |         |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |
| m-AST     | Week 1  | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 39     | -     | 39    | -      | 36     | 1     | 37    | -      | 38     | -     | 38    | -      | 36     | 1     | 37    | -      | 45     | -     | 45    |
|           |         | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | 1     | -      | -      | -     | -     | -      | 1      | 1     | 2     |
|           |         | Total  | -       | 39     | -     | 39    | -      | 36     | 1     | 37    | -      | 38     | 1     | 39    | -      | 36     | 1     | 37    | -      | 46     | 1     | 47    |
|           | Week 2  | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 91     | 1     | 92    | -      | 83     | 3     | 86    | -      | 86     | 3     | 89    | -      | 87     | 3     | 90    | -      | 85     | 3     | 88    |
|           |         | Upper  | -       | 2      | 2     | 4     | -      | 2      | 1     | 3     | -      | 2      | -     | 2     | -      | 3      | -     | 3     | -      | -      | 1     | 1     |
|           |         | Total  | -       | 93     | 3     | 96    | -      | 85     | 4     | 89    | -      | 88     | 3     | 91    | -      | 90     | 3     | 93    | -      | 85     | 4     | 89    |
|           | Week 3  | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 34     | -     | 34    | -      | 34     | 1     | 35    | -      | 36     | -     | 36    | -      | 30     | 1     | 31    | -      | 41     | -     | 41    |
|           |         | Upper  | -       | 2      | -     | 2     | -      | 1      | -     | 1     | -      | 1      | 1     | 2     | -      | 1      | -     | 1     | -      | -      | 1     | 1     |
|           |         | Total  | -       | 36     | -     | 36    | -      | 35     | 1     | 36    | -      | 37     | 1     | 38    | -      | 31     | 1     | 32    | -      | 41     | 1     | 42    |
|           | Week 4  | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 90     | 2     | 92    | -      | 84     | 1     | 85    | -      | 86     | 1     | 87    | -      | 84     | 3     | 87    | -      | 82     | 2     | 84    |
|           |         | Upper  | -       | 2      | 1     | 3     | -      | 1      | 3     | 4     | -      | -      | 2     | 2     | -      | 2      | -     | 2     | -      | 1      | 2     | 3     |
|           |         | Total  | -       | 92     | 3     | 95    | -      | 85     | 4     | 89    | -      | 86     | 3     | 89    | -      | 86     | 3     | 89    | -      | 83     | 4     | 87    |
|           | Week 6  | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 39     | -     | 39    | -      | 38     | 1     | 39    | -      | 37     | 1     | 38    | -      | 29     | 1     | 30    | -      | 45     | -     | 45    |
|           |         | Upper  | -       | -      | 1     | 1     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | 1      | 1     | -     | -      | 1      | 1     | 2     |
|           |         | Total  | -       | 39     | 1     | 40    | -      | 39     | 1     | 40    | -      | 37     | 1     | 38    | -      | 29     | 2     | 31    | -      | 46     | 1     | 47    |
|           | Week 8  | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 88     | 2     | 90    | -      | 78     | -     | 78    | -      | 84     | 1     | 85    | -      | 82     | 1     | 83    | -      | 79     | 4     | 83    |
|           |         | Upper  | -       | 3      | 1     | 4     | -      | 5      | 3     | 8     | -      | 2      | 2     | 4     | -      | 1      | 2     | 3     | -      | 1      | -     | 1     |
|           |         | Total  | -       | 91     | 3     | 94    | -      | 83     | 3     | 86    | -      | 86     | 3     | 89    | -      | 83     | 3     | 86    | -      | 80     | 4     | 84    |
|           | Week 12 | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 90     | 2     | 92    | -      | 79     | 2     | 81    | -      | 82     | 3     | 85    | -      | 76     | 3     | 79    | -      | 76     | 3     | 79    |
|           |         | Upper  | -       | 1      | 1     | 2     | -      | 4      | 1     | 5     | -      | 1      | -     | 1     | -      | 3      | -     | 3     | -      | 3      | 1     | 4     |
|           |         | Total  | -       | 91     | 3     | 94    | -      | 83     | 3     | 86    | -      | 83     | 3     | 86    | -      | 79     | 3     | 82    | -      | 79     | 4     | 83    |
|           | Week 16 | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 87     | 3     | 90    | -      | 78     | 2     | 80    | -      | 82     | 2     | 84    | -      | 75     | 3     | 78    | -      | 77     | 3     | 80    |
|           |         | Upper  | -       | 4      | -     | 4     | -      | 3      | 1     | 4     | -      | -      | 1     | 1     | -      | 2      | -     | 2     | -      | 2      | -     | 2     |
|           |         | Total  | -       | 91     | 3     | 94    | -      | 81     | 3     | 84    | -      | 82     | 3     | 85    | -      | 77     | 3     | 80    | -      | 79     | 3     | 82    |
|           | Week 20 | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 85     | 1     | 86    | -      | 77     | 1     | 78    | -      | 81     | 1     | 82    | -      | 75     | 3     | 78    | -      | 76     | 2     | 78    |
|           |         | Upper  | -       | 6      | 2     | 8     | -      | 4      | 2     | 6     | -      | 1      | 2     | 3     | -      | 1      | -     | 1     | -      | 1      | 1     | 2     |
|           |         | Total  | -       | 91     | 3     | 94    | -      | 81     | 3     | 84    | -      | 82     | 3     | 85    | -      | 76     | 3     | 79    | -      | 77     | 3     | 80    |
|           | Week 24 | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |
|           |         | Normal | -       | 85     | 2     | 87    | -      | 77     | 1     | 78    | -      | 81     | 3     | 84    | -      | 74     | 3     | 77    | -      | 75     | 2     | 77    |
|           |         | Upper  | -       | 5      | 1     | 6     | -      | 4      | 2     | 6     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | 1      | 1     | 2     |
|           |         | Total  | -       | 90     | 3     | 93    | -      | 81     | 3     | 84    | -      | 81     | 3     | 84    | -      | 75     | 3     | 78    | -      | 76     | 3     | 79    |
| Week 28   | Lower   | -      | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      |       |       |
|           | Normal  | -      | 89      | 2      | 91    | -     | 78     | 2      | 80    | -     | 80     | 2      | 82    | -     | 73     | 3      | 76    | -     | 73     | 3      | 76    |       |
|           | Upper   | -      | 1       | 1      | 2     | -     | 2      | 1      | 3     | -     | 1      | 1      | 2     | -     | 1      | -      | 1     | -     | 2      | -      | 2     |       |
|           | Total   | -      | 90      | 3      | 93    | -     | 80     | 3      | 83    | -     | 81     | 3      | 84    | -     | 74     | 3      | 77    | -     | 75     | 3      | 78    |       |
| Follow-up | Lower   | -      | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      |       |       |
|           | Normal  | -      | 93      | 2      | 95    | -     | 80     | 3      | 83    | -     | 88     | 2      | 90    | -     | 82     | 3      | 85    | -     | 80     | 4      | 84    |       |
|           | Upper   | -      | 1       | 1      | 2     | -     | 2      | 1      | 3     | -     | 1      | 1      | 2     | -     | 2      | -      | 2     | -     | 4      | -      | 4     |       |
|           | Total   | -      | 94      | 3      | 97    | -     | 82     | 4      | 86    | -     | 89     | 3      | 92    | -     | 84     | 3      | 87    | -     | 84     | 4      | 88    |       |



表 2.7.4.7-付録-15 血液生化学検査の評価時点別正異判定クロス表 (CCT-001 試験) (24/24)

|                   |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-------------------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|                   |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|                   |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| CK18<br>(Segment) | Week 1 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                   |        | Normal  | -      | 33    | 2     | 35     | -      | 33    | 2     | 35     | -      | 36    | 1     | 37     | -      | 32    | 1     | 33     | -      | 45    | -     | 45 |
|                   |        | Upper   | -      | 1     | 3     | 4      | -      | 2     | -     | 2      | -      | -     | 2     | 2      | -      | 1     | 3     | 4      | -      | -     | 2     | 2  |
|                   |        | Total   | -      | 34    | 5     | 39     | -      | 35    | 2     | 37     | -      | 36    | 3     | 39     | -      | 33    | 4     | 37     | -      | 45    | 2     | 47 |
|                   | Week 2 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                   |        | Normal  | -      | 81    | 5     | 86     | -      | 82    | 1     | 83     | -      | 82    | 1     | 83     | -      | 85    | 4     | 89     | -      | 84    | -     | 84 |
|                   |        | Upper   | -      | 4     | 7     | 11     | -      | 1     | 5     | 6      | -      | 2     | 6     | 8      | -      | 1     | 3     | 4      | -      | 3     | 2     | 5  |
|                   |        | Total   | -      | 85    | 12    | 97     | -      | 83    | 6     | 89     | -      | 84    | 7     | 91     | -      | 86    | 7     | 93     | -      | 87    | 2     | 89 |
|                   | Week 3 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                   |        | Normal  | -      | 31    | 2     | 33     | -      | 33    | 1     | 34     | -      | 34    | 2     | 36     | -      | 29    | 2     | 31     | -      | 39    | -     | 39 |
|                   |        | Upper   | -      | -     | 3     | 3      | -      | 1     | 1     | 2      | -      | -     | 2     | 2      | -      | 1     | -     | 1      | -      | 1     | 2     | 3  |
|                   |        | Total   | -      | 31    | 5     | 36     | -      | 34    | 2     | 36     | -      | 34    | 4     | 38     | -      | 30    | 2     | 32     | -      | 40    | 2     | 42 |
| Week 4            | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                   | Normal | -       | 81     | 5     | 86    | -      | 77     | 2     | 79    | -      | 82     | 3     | 85    | -      | 77     | 5     | 82    | -      | 83     | -     | 83    |    |
|                   | Upper  | -       | 3      | 7     | 10    | -      | 6      | 4     | 10    | -      | 1      | 3     | 4     | -      | 4      | 3     | 7     | -      | 2      | 2     | 4     |    |
|                   | Total  | -       | 84     | 12    | 96    | -      | 83     | 6     | 89    | -      | 83     | 6     | 89    | -      | 81     | 8     | 89    | -      | 85     | 2     | 87    |    |
| Week 6            | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                   | Normal | -       | 34     | 2     | 36    | -      | 37     | 2     | 39    | -      | 35     | 1     | 36    | -      | 28     | 1     | 29    | -      | 44     | 1     | 45    |    |
|                   | Upper  | -       | -      | 4     | 4     | -      | 1      | -     | 1     | -      | -      | 2     | 2     | -      | 1      | 1     | 2     | -      | 1      | 1     | 2     |    |
|                   | Total  | -       | 34     | 6     | 40    | -      | 38     | 2     | 40    | -      | 35     | 3     | 38    | -      | 29     | 2     | 31    | -      | 45     | 2     | 47    |    |
| Week 8            | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                   | Normal | -       | 80     | 5     | 85    | -      | 77     | 3     | 80    | -      | 80     | 3     | 83    | -      | 73     | 6     | 79    | -      | 79     | 1     | 80    |    |
|                   | Upper  | -       | 3      | 7     | 10    | -      | 3      | 3     | 6     | -      | 3      | 3     | 6     | -      | 6      | 1     | 7     | -      | 3      | 1     | 4     |    |
|                   | Total  | -       | 83     | 12    | 95    | -      | 80     | 6     | 86    | -      | 83     | 6     | 89    | -      | 79     | 7     | 86    | -      | 82     | 2     | 84    |    |
| Week 12           | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                   | Normal | -       | 78     | 5     | 83    | -      | 78     | 3     | 81    | -      | 79     | 4     | 83    | -      | 74     | 4     | 78    | -      | 79     | 2     | 81    |    |
|                   | Upper  | -       | 5      | 7     | 12    | -      | 2      | 3     | 5     | -      | 1      | 2     | 3     | -      | 1      | 3     | 4     | -      | 2      | -     | 2     |    |
|                   | Total  | -       | 83     | 12    | 95    | -      | 80     | 6     | 86    | -      | 80     | 6     | 86    | -      | 75     | 7     | 82    | -      | 81     | 2     | 83    |    |
| Week 16           | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                   | Normal | -       | 80     | 4     | 84    | -      | 75     | 2     | 77    | -      | 75     | 3     | 78    | -      | 72     | 5     | 77    | -      | 76     | 2     | 78    |    |
|                   | Upper  | -       | 3      | 8     | 11    | -      | 4      | 3     | 7     | -      | 4      | 3     | 7     | -      | 1      | 2     | 3     | -      | 4      | -     | 4     |    |
|                   | Total  | -       | 83     | 12    | 95    | -      | 79     | 5     | 84    | -      | 79     | 6     | 85    | -      | 73     | 7     | 80    | -      | 80     | 2     | 82    |    |
| Week 20           | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                   | Normal | -       | 74     | 3     | 77    | -      | 75     | 4     | 79    | -      | 77     | 2     | 79    | -      | 69     | 5     | 74    | -      | 74     | 2     | 76    |    |
|                   | Upper  | -       | 9      | 9     | 18    | -      | 4      | 1     | 5     | -      | 2      | 4     | 6     | -      | 3      | 2     | 5     | -      | 4      | -     | 4     |    |
|                   | Total  | -       | 83     | 12    | 95    | -      | 79     | 5     | 84    | -      | 79     | 6     | 85    | -      | 72     | 7     | 79    | -      | 78     | 2     | 80    |    |
| Week 24           | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                   | Normal | -       | 73     | 7     | 80    | -      | 75     | 3     | 78    | -      | 76     | 3     | 79    | -      | 70     | 6     | 76    | -      | 76     | 2     | 78    |    |
|                   | Upper  | -       | 9      | 5     | 14    | -      | 4      | 2     | 6     | -      | 2      | 3     | 5     | -      | 2      | -     | 2     | -      | 1      | -     | 1     |    |
|                   | Total  | -       | 82     | 12    | 94    | -      | 79     | 5     | 84    | -      | 78     | 6     | 84    | -      | 72     | 6     | 78    | -      | 77     | 2     | 79    |    |
| Week 28           | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                   | Normal | -       | 72     | 5     | 77    | -      | 75     | 3     | 78    | -      | 73     | 3     | 76    | -      | 66     | 4     | 70    | -      | 72     | 1     | 73    |    |
|                   | Upper  | -       | 10     | 7     | 17    | -      | 3      | 2     | 5     | -      | 5      | 3     | 8     | -      | 5      | 2     | 7     | -      | 5      | 1     | 6     |    |
|                   | Total  | -       | 82     | 12    | 94    | -      | 78     | 5     | 83    | -      | 78     | 6     | 84    | -      | 71     | 6     | 77    | -      | 77     | 2     | 79    |    |
| Follow-up         | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|                   | Normal | -       | 76     | 5     | 81    | -      | 77     | 3     | 80    | -      | 80     | 3     | 83    | -      | 73     | 4     | 77    | -      | 80     | 1     | 81    |    |
|                   | Upper  | -       | 10     | 7     | 17    | -      | 3      | 3     | 6     | -      | 5      | 4     | 9     | -      | 7      | 3     | 10    | -      | 6      | 1     | 7     |    |
|                   | Total  | -       | 86     | 12    | 98    | -      | 80     | 6     | 86    | -      | 85     | 7     | 92    | -      | 80     | 7     | 87    | -      | 86     | 2     | 88    |    |

表 2.7.4.7-付録-16 尿検査の評価時点別の変化量 (CCT-001 試験)

| Item             | Treatment | Summary Statistics |                      |         |         |         |         |         |         |         |         | One-sample t-test |        |
|------------------|-----------|--------------------|----------------------|---------|---------|---------|---------|---------|---------|---------|---------|-------------------|--------|
|                  |           | Baseline Mean      | Change from Baseline |         |         |         |         | Q3      | Max     | t-value | p-value |                   |        |
|                  |           |                    | N                    | Mean    | SD      | Min     | Q1      |         |         |         |         | Median            |        |
| Specific Gravity | Week 1    | Placebo            | 1.0179               | 39      | -0.0005 | 0.00412 | -0.009  | -0.0030 | -0.0010 | 0.0020  | 0.010   | -0.8170           | 0.4190 |
|                  |           | 15 mg              | 1.0154               | 37      | 0.0008  | 0.00623 | -0.013  | -0.0030 | 0.0010  | 0.0060  | 0.013   | 0.7921            | 0.4335 |
|                  |           | 30 mg              | 1.0179               | 39      | 0.0006  | 0.00731 | -0.018  | -0.0050 | 0.0010  | 0.0040  | 0.016   | 0.5259            | 0.6020 |
|                  |           | 60 mg              | 1.0187               | 37      | -0.0002 | 0.00796 | -0.026  | -0.0020 | 0.0000  | 0.0040  | 0.017   | -0.1653           | 0.8697 |
|                  |           | 120 mg             | 1.0178               | 46      | 0.0001  | 0.00759 | -0.017  | -0.0050 | 0.0000  | 0.0050  | 0.018   | 0.1165            | 0.9077 |
|                  | Week 2    | Placebo            | 1.0190               | 97      | -0.0012 | 0.00622 | -0.022  | -0.0030 | -0.0010 | 0.0020  | 0.019   | -1.8281           | 0.0706 |
|                  |           | 15 mg              | 1.0168               | 89      | -0.0002 | 0.00603 | -0.015  | -0.0040 | -0.0010 | 0.0040  | 0.018   | -0.3518           | 0.7258 |
|                  |           | 30 mg              | 1.0180               | 90      | 0.0007  | 0.00675 | -0.021  | -0.0020 | 0.0020  | 0.0040  | 0.020   | 0.9679            | 0.3357 |
|                  |           | 60 mg              | 1.0183               | 92      | 0.0006  | 0.00686 | -0.016  | -0.0030 | 0.0000  | 0.0040  | 0.026   | 0.7901            | 0.4315 |
|                  |           | 120 mg             | 1.0170               | 88      | 0.0012  | 0.00729 | -0.014  | -0.0030 | 0.0010  | 0.0060  | 0.028   | 1.5945            | 0.1145 |
|                  | Week 3    | Placebo            | 1.0182               | 37      | -0.0028 | 0.00555 | -0.017  | -0.0070 | -0.0020 | 0.0000  | 0.009   | -3.0519           | 0.0043 |
|                  |           | 15 mg              | 1.0161               | 36      | 0.0009  | 0.00658 | -0.010  | -0.0040 | 0.0000  | 0.0030  | 0.017   | 0.8105            | 0.4231 |
|                  |           | 30 mg              | 1.0178               | 38      | 0.0018  | 0.00627 | -0.022  | -0.0020 | 0.0030  | 0.0050  | 0.014   | 1.8120            | 0.0781 |
|                  |           | 60 mg              | 1.0172               | 33      | 0.0019  | 0.00690 | -0.011  | -0.0010 | 0.0020  | 0.0060  | 0.017   | 1.5888            | 0.1219 |
|                  |           | 120 mg             | 1.0179               | 41      | -0.0008 | 0.00692 | -0.025  | -0.0040 | 0.0000  | 0.0030  | 0.011   | -0.7219           | 0.4746 |
|                  | Week 4    | Placebo            | 1.0189               | 96      | 0.0004  | 0.00665 | -0.014  | -0.0030 | 0.0000  | 0.0040  | 0.021   | 0.5676            | 0.5716 |
|                  |           | 15 mg              | 1.0168               | 89      | -0.0010 | 0.00666 | -0.019  | -0.0050 | -0.0010 | 0.0020  | 0.013   | -1.3692           | 0.1744 |
|                  |           | 30 mg              | 1.0180               | 88      | -0.0001 | 0.00672 | -0.016  | -0.0030 | 0.0000  | 0.0040  | 0.021   | -0.1110           | 0.9118 |
|                  |           | 60 mg              | 1.0187               | 89      | -0.0010 | 0.00772 | -0.035  | -0.0020 | 0.0000  | 0.0030  | 0.014   | -1.1953           | 0.2352 |
|                  |           | 120 mg             | 1.0169               | 86      | 0.0009  | 0.00626 | -0.013  | -0.0030 | 0.0000  | 0.0040  | 0.016   | 1.3088            | 0.1941 |
|                  | Week 6    | Placebo            | 1.0181               | 40      | -0.0012 | 0.00627 | -0.018  | -0.0045 | 0.0000  | 0.0025  | 0.011   | -1.2108           | 0.2333 |
|                  |           | 15 mg              | 1.0164               | 40      | -0.0012 | 0.00557 | -0.015  | -0.0045 | -0.0010 | 0.0020  | 0.009   | -1.3915           | 0.1720 |
|                  |           | 30 mg              | 1.0192               | 38      | -0.0010 | 0.00781 | -0.021  | -0.0060 | -0.0010 | 0.0040  | 0.014   | -0.7898           | 0.4347 |
|                  |           | 60 mg              | 1.0183               | 32      | -0.0014 | 0.00650 | -0.019  | -0.0050 | 0.0000  | 0.0030  | 0.009   | -1.2229           | 0.2306 |
|                  |           | 120 mg             | 1.0179               | 46      | 0.0002  | 0.00716 | -0.013  | -0.0040 | 0.0000  | 0.0050  | 0.020   | 0.1442            | 0.8860 |
|                  | Week 8    | Placebo            | 1.0189               | 95      | -0.0008 | 0.00622 | -0.020  | -0.0050 | 0.0000  | 0.0040  | 0.015   | -1.2857           | 0.2017 |
|                  |           | 15 mg              | 1.0168               | 86      | 0.0009  | 0.00705 | -0.018  | -0.0030 | 0.0010  | 0.0050  | 0.018   | 1.1619            | 0.2485 |
|                  |           | 30 mg              | 1.0180               | 88      | 0.0007  | 0.00743 | -0.019  | -0.0030 | 0.0000  | 0.0050  | 0.017   | 0.8748            | 0.3841 |
|                  |           | 60 mg              | 1.0186               | 86      | -0.0006 | 0.00721 | -0.025  | -0.0030 | 0.0000  | 0.0040  | 0.024   | -0.7479           | 0.4566 |
|                  |           | 120 mg             | 1.0168               | 83      | 0.0008  | 0.00741 | -0.021  | -0.0030 | 0.0020  | 0.0060  | 0.017   | 1.0215            | 0.3100 |
|                  | Week 12   | Placebo            | 1.0189               | 95      | -0.0007 | 0.00613 | -0.017  | -0.0050 | -0.0010 | 0.0030  | 0.015   | -1.0540           | 0.2946 |
|                  |           | 15 mg              | 1.0168               | 86      | -0.0002 | 0.00670 | -0.019  | -0.0040 | -0.0010 | 0.0030  | 0.016   | -0.2895           | 0.7729 |
|                  |           | 30 mg              | 1.0179               | 85      | 0.0000  | 0.00693 | -0.015  | -0.0050 | -0.0010 | 0.0050  | 0.015   | 0.0157            | 0.9875 |
|                  |           | 60 mg              | 1.0183               | 82      | -0.0001 | 0.00684 | -0.029  | -0.0030 | 0.0005  | 0.0040  | 0.014   | -0.1775           | 0.8596 |
|                  |           | 120 mg             | 1.0167               | 82      | 0.0005  | 0.00707 | -0.017  | -0.0050 | 0.0010  | 0.0050  | 0.017   | 0.6245            | 0.5341 |
|                  | Week 16   | Placebo            | 1.0189               | 95      | 0.0001  | 0.00697 | -0.021  | -0.0040 | 0.0000  | 0.0050  | 0.020   | 0.1325            | 0.8948 |
|                  |           | 15 mg              | 1.0169               | 84      | 0.0002  | 0.00618 | -0.015  | -0.0025 | 0.0000  | 0.0030  | 0.019   | 0.3354            | 0.7382 |
|                  |           | 30 mg              | 1.0180               | 84      | 0.0009  | 0.00702 | -0.016  | -0.0040 | 0.0010  | 0.0050  | 0.017   | 1.1814            | 0.2408 |
|                  |           | 60 mg              | 1.0186               | 80      | 0.0004  | 0.00599 | -0.017  | -0.0030 | 0.0000  | 0.0050  | 0.015   | 0.5413            | 0.5898 |
|                  |           | 120 mg             | 1.0168               | 81      | 0.0021  | 0.00675 | -0.014  | -0.0030 | 0.0020  | 0.0070  | 0.018   | 2.7483            | 0.0074 |
|                  | Week 20   | Placebo            | 1.0189               | 95      | -0.0007 | 0.00672 | -0.018  | -0.0050 | 0.0000  | 0.0030  | 0.016   | -0.9776           | 0.3308 |
|                  |           | 15 mg              | 1.0169               | 84      | -0.0002 | 0.00638 | -0.016  | -0.0040 | 0.0000  | 0.0040  | 0.020   | -0.2905           | 0.7722 |
|                  |           | 30 mg              | 1.0180               | 84      | 0.0007  | 0.00710 | -0.020  | -0.0030 | 0.0020  | 0.0055  | 0.016   | 0.8453            | 0.4004 |
|                  |           | 60 mg              | 1.0185               | 79      | 0.0007  | 0.00627 | -0.017  | -0.0020 | 0.0010  | 0.0030  | 0.015   | 0.9509            | 0.3446 |
|                  |           | 120 mg             | 1.0169               | 79      | 0.0006  | 0.00679 | -0.026  | -0.0040 | 0.0000  | 0.0050  | 0.015   | 0.7621            | 0.4483 |
|                  | Week 24   | Placebo            | 1.0189               | 94      | 0.0004  | 0.00723 | -0.018  | -0.0040 | 0.0000  | 0.0050  | 0.017   | 0.5133            | 0.6089 |
|                  |           | 15 mg              | 1.0169               | 84      | 0.0004  | 0.00704 | -0.018  | -0.0040 | 0.0000  | 0.0050  | 0.019   | 0.5114            | 0.6104 |
|                  |           | 30 mg              | 1.0178               | 83      | 0.0002  | 0.00669 | -0.018  | -0.0040 | 0.0000  | 0.0040  | 0.019   | 0.2953            | 0.7685 |
| 60 mg            |           | 1.0185             | 78                   | 0.0011  | 0.00622 | -0.017  | -0.0030 | 0.0000  | 0.0050  | 0.022   | 1.5470  | 0.1260            |        |
| 120 mg           |           | 1.0168             | 78                   | 0.0013  | 0.00652 | -0.015  | -0.0030 | 0.0005  | 0.0070  | 0.021   | 1.8055  | 0.0749            |        |
| Week 28          | Placebo   | 1.0189             | 94                   | 0.0010  | 0.00777 | -0.020  | -0.0050 | 0.0010  | 0.0060  | 0.019   | 1.2608  | 0.2105            |        |
|                  | 15 mg     | 1.0167             | 83                   | 0.0014  | 0.00704 | -0.015  | -0.0040 | 0.0010  | 0.0060  | 0.016   | 1.7940  | 0.0765            |        |
|                  | 30 mg     | 1.0178             | 83                   | 0.0003  | 0.00714 | -0.017  | -0.0050 | 0.0000  | 0.0050  | 0.020   | 0.3381  | 0.7361            |        |
|                  | 60 mg     | 1.0185             | 77                   | 0.0002  | 0.00646 | -0.013  | -0.0040 | 0.0000  | 0.0040  | 0.017   | 0.2292  | 0.8193            |        |
|                  | 120 mg    | 1.0168             | 77                   | 0.0014  | 0.00696 | -0.013  | -0.0030 | 0.0010  | 0.0070  | 0.017   | 1.7528  | 0.0837            |        |
| Follow-up        | Placebo   | 1.0190             | 98                   | 0.0008  | 0.00774 | -0.020  | -0.0050 | 0.0010  | 0.0060  | 0.019   | 1.0177  | 0.3113            |        |
|                  | 15 mg     | 1.0167             | 86                   | 0.0011  | 0.00710 | -0.015  | -0.0040 | 0.0010  | 0.0060  | 0.016   | 1.3815  | 0.1708            |        |
|                  | 30 mg     | 1.0178             | 91                   | 0.0002  | 0.00706 | -0.017  | -0.0040 | 0.0000  | 0.0040  | 0.020   | 0.2376  | 0.8127            |        |
|                  | 60 mg     | 1.0187             | 87                   | -0.0002 | 0.00751 | -0.036  | -0.0050 | 0.0000  | 0.0040  | 0.017   | -0.2570 | 0.7978            |        |
|                  | 120 mg    | 1.0170             | 87                   | 0.0012  | 0.00712 | -0.013  | -0.0040 | 0.0010  | 0.0070  | 0.017   | 1.6266  | 0.1075            |        |

表 2.7.4.7-付録-17 尿検査の評価時点別クロス表 (CCT-001 試験) (1/6)

|                  |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|------------------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|                  |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|                  |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Specific Gravity | Week 1 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                  |        | Normal  | -      | 37    | 2     | 39     | -      | 34    | 2     | 36     | -      | 36    | 1     | 37     | 1      | 30    | 3     | 34     | -      | 43    | 1     | 44 |
|                  |        | Upper   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | 2     | -     | 2      | -      | 3     | -     | 3      | -      | 2     | -     | 2  |
|                  |        | Total   | -      | 37    | 2     | 39     | -      | 35    | 2     | 37     | -      | 38    | 1     | 39     | 1      | 33    | 3     | 37     | -      | 45    | 1     | 46 |
|                  | Week 2 | Lower   | 1      | -     | -     | 1      | 1      | 1     | -     | 2      | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     | -     |    |
|                  |        | Normal  | -      | 87    | 5     | 92     | -      | 82    | 1     | 83     | -      | 83    | 3     | 86     | 1      | 84    | 3     | 88     | -      | 82    | -     | 82 |
|                  |        | Upper   | -      | 3     | 1     | 4      | -      | 2     | 2     | 4      | -      | 4     | -     | 4      | -      | 2     | 1     | 3      | -      | 4     | 2     | 6  |
|                  |        | Total   | 1      | 90    | 6     | 97     | 1      | 85    | 3     | 89     | -      | 87    | 3     | 90     | 2      | 86    | 4     | 92     | -      | 86    | 2     | 88 |
|                  | Week 3 | Lower   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|                  |        | Normal  | -      | 34    | 3     | 37     | -      | 33    | 2     | 35     | -      | 35    | 1     | 36     | 1      | 29    | 1     | 31     | -      | 40    | 1     | 41 |
|                  |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 2     | -     | 2      | -      | 2     | -     | 2      | -      | -     | -     |    |
|                  |        | Total   | -      | 34    | 3     | 37     | -      | 34    | 2     | 36     | -      | 37    | 1     | 38     | 1      | 31    | 1     | 33     | -      | 40    | 1     | 41 |
|                  | Week 4 | Lower   | 1      | -     | -     | 1      | 1      | -     | -     | 1      | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     |    |
|                  |        | Normal  | -      | 87    | 2     | 89     | -      | 83    | 3     | 86     | -      | 84    | 3     | 87     | 1      | 79    | 5     | 85     | -      | 82    | 2     | 84 |
|                  |        | Upper   | -      | 2     | 4     | 6      | -      | 2     | -     | 2      | -      | 1     | -     | 1      | -      | 3     | -     | 3      | -      | 2     | -     | 2  |
|                  |        | Total   | 1      | 89    | 6     | 96     | 1      | 85    | 3     | 89     | -      | 85    | 3     | 88     | 2      | 82    | 5     | 89     | -      | 84    | 2     | 86 |
|                  | Week 6 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | -     | 1      | -      | -     | -     |    |
|                  |        | Normal  | -      | 36    | 2     | 38     | -      | 37    | 3     | 40     | -      | 36    | 2     | 38     | -      | 29    | 2     | 31     | -      | 42    | 2     | 44 |
|                  |        | Upper   | -      | -     | 2     | 2      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 2     | -     | 2  |
|                  |        | Total   | -      | 36    | 4     | 40     | -      | 37    | 3     | 40     | -      | 36    | 2     | 38     | 1      | 29    | 2     | 32     | -      | 44    | 2     | 46 |
| Week 8           | Lower  | -       | -      | -     | -     | 1      | 1      | -     | 2     | -      | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     |       |    |
|                  | Normal | 1       | 84     | 4     | 89    | -      | 77     | 2     | 79    | -      | 81     | 3     | 84    | 1      | 77     | 3     | 81    | -      | 80     | -     | 80    |    |
|                  | Upper  | -       | 4      | 2     | 6     | -      | 4      | 1     | 5     | -      | 4      | -     | 4     | -      | 3      | 1     | 4     | -      | 1      | 2     | 3     |    |
|                  | Total  | 1       | 88     | 6     | 95    | 1      | 82     | 3     | 86    | -      | 85     | 3     | 88    | 2      | 80     | 4     | 86    | -      | 81     | 2     | 83    |    |
| Week 12          | Lower  | -       | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     |       |    |
|                  | Normal | 1       | 88     | 5     | 94    | -      | 80     | 2     | 82    | -      | 81     | 3     | 84    | 1      | 77     | 3     | 81    | -      | 78     | 2     | 80    |    |
|                  | Upper  | -       | -      | 1     | 1     | -      | 2      | 1     | 3     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | 2      | -     | 2     |    |
|                  | Total  | 1       | 88     | 6     | 95    | 1      | 82     | 3     | 86    | -      | 82     | 3     | 85    | 2      | 77     | 3     | 82    | -      | 80     | 2     | 82    |    |
| Week 16          | Lower  | -       | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     | -     | 1      | -      | -     | 1     | -      | -      | -     |       |    |
|                  | Normal | 1       | 82     | 4     | 87    | -      | 79     | 2     | 81    | -      | 80     | 3     | 83    | 1      | 74     | 3     | 78    | -      | 76     | 2     | 78    |    |
|                  | Upper  | -       | 6      | 2     | 8     | -      | 1      | 1     | 2     | -      | 1      | -     | 1     | -      | 1      | -     | 1     | -      | 3      | -     | 3     |    |
|                  | Total  | 1       | 88     | 6     | 95    | 1      | 80     | 3     | 84    | -      | 81     | 3     | 84    | 2      | 75     | 3     | 80    | -      | 79     | 2     | 81    |    |
| Week 20          | Lower  | 1       | -      | -     | 1     | 1      | -      | -     | 1     | -      | 1      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     |       |    |
|                  | Normal | -       | 86     | 3     | 89    | -      | 78     | 2     | 80    | -      | 78     | 3     | 81    | 1      | 71     | 2     | 74    | -      | 74     | 2     | 76    |    |
|                  | Upper  | -       | 2      | 3     | 5     | -      | 2      | 1     | 3     | -      | 2      | -     | 2     | -      | 3      | 1     | 4     | -      | 3      | -     | 3     |    |
|                  | Total  | 1       | 88     | 6     | 95    | 1      | 80     | 3     | 84    | -      | 81     | 3     | 84    | 2      | 74     | 3     | 79    | -      | 77     | 2     | 79    |    |
| Week 24          | Lower  | -       | -      | -     | -     | 1      | -      | -     | 1     | -      | 1      | -     | 1     | 1      | -      | -     | 1     | -      | -      | -     |       |    |
|                  | Normal | 1       | 79     | 4     | 84    | -      | 78     | 2     | 80    | -      | 76     | 2     | 78    | 1      | 70     | 3     | 74    | -      | 72     | 1     | 73    |    |
|                  | Upper  | -       | 8      | 2     | 10    | -      | 2      | 1     | 3     | -      | 4      | -     | 4     | -      | 3      | -     | 3     | -      | 4      | 1     | 5     |    |
|                  | Total  | 1       | 87     | 6     | 94    | 1      | 80     | 3     | 84    | -      | 81     | 2     | 83    | 2      | 73     | 3     | 78    | -      | 76     | 2     | 78    |    |
| Week 28          | Lower  | -       | -      | -     | -     | 1      | 1      | -     | 2     | -      | -      | -     | -     | 2      | -      | -     | 2     | -      | -      | -     |       |    |
|                  | Normal | 1       | 80     | 6     | 87    | -      | 73     | 1     | 74    | -      | 80     | 2     | 82    | -      | 69     | 3     | 72    | -      | 75     | 2     | 77    |    |
|                  | Upper  | -       | 7      | -     | 7     | -      | 5      | 2     | 7     | -      | 1      | -     | 1     | -      | 3      | -     | 3     | -      | -      | -     |       |    |
|                  | Total  | 1       | 87     | 6     | 94    | 1      | 79     | 3     | 83    | -      | 81     | 2     | 83    | 2      | 72     | 3     | 77    | -      | 75     | 2     | 77    |    |
| Follow-up        | Lower  | -       | -      | -     | -     | 1      | 1      | -     | 2     | -      | -      | -     | -     | 2      | -      | -     | 2     | -      | -      | -     |       |    |
|                  | Normal | 1       | 84     | 6     | 91    | -      | 76     | 1     | 77    | -      | 88     | 2     | 90    | -      | 77     | 5     | 82    | -      | 85     | 2     | 87    |    |
|                  | Upper  | -       | 7      | -     | 7     | -      | 5      | 2     | 7     | -      | 1      | -     | 1     | -      | 3      | -     | 3     | -      | -      | -     |       |    |
|                  | Total  | 1       | 91     | 6     | 98    | 1      | 82     | 3     | 86    | -      | 89     | 2     | 91    | 2      | 80     | 5     | 87    | -      | 85     | 2     | 87    |    |

表 2.7.4.7-付録-17 尿検査の評価時点別クロス表 (CCT-001 試験) (2/6)

|           |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| pH        | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 46    | -     | 46 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 46    | -     | 46 |
|           | Week 2  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 96    | -     | 96     | -      | 89    | -     | 89     | -      | 90    | -     | 90     | -      | 92    | -     | 92     | -      | 88    | -     | 88 |
|           |         | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |         | Total   | -      | 97    | -     | 97     | -      | 89    | -     | 89     | -      | 90    | -     | 90     | -      | 92    | -     | 92     | -      | 88    | -     | 88 |
|           | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 37    | -     | 37     | -      | 36    | -     | 36     | -      | 38    | -     | 38     | -      | 33    | -     | 33     | -      | 41    | -     | 41 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 37    | -     | 37     | -      | 36    | -     | 36     | -      | 38    | -     | 38     | -      | 33    | -     | 33     | -      | 41    | -     | 41 |
|           | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 95    | -     | 95     | -      | 89    | -     | 89     | -      | 88    | -     | 88     | -      | 89    | -     | 89     | -      | 86    | -     | 86 |
|           |         | Upper   | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 96    | -     | 96     | -      | 89    | -     | 89     | -      | 88    | -     | 88     | -      | 89    | -     | 89     | -      | 86    | -     | 86 |
|           | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 40    | -     | 40     | -      | 40    | -     | 40     | -      | 38    | -     | 38     | -      | 32    | -     | 32     | -      | 46    | -     | 46 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 40    | -     | 40     | -      | 40    | -     | 40     | -      | 38    | -     | 38     | -      | 32    | -     | 32     | -      | 46    | -     | 46 |
|           | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 95    | -     | 95     | -      | 86    | -     | 86     | -      | 88    | -     | 88     | -      | 86    | -     | 86     | -      | 83    | -     | 83 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 95    | -     | 95     | -      | 86    | -     | 86     | -      | 88    | -     | 88     | -      | 86    | -     | 86     | -      | 83    | -     | 83 |
|           | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 95    | -     | 95     | -      | 86    | -     | 86     | -      | 85    | -     | 85     | -      | 82    | -     | 82     | -      | 82    | -     | 82 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 95    | -     | 95     | -      | 86    | -     | 86     | -      | 85    | -     | 85     | -      | 82    | -     | 82     | -      | 82    | -     | 82 |
|           | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 95    | -     | 95     | -      | 84    | -     | 84     | -      | 84    | -     | 84     | -      | 80    | -     | 80     | -      | 81    | -     | 81 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 95    | -     | 95     | -      | 84    | -     | 84     | -      | 84    | -     | 84     | -      | 80    | -     | 80     | -      | 81    | -     | 81 |
|           | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 95    | -     | 95     | -      | 84    | -     | 84     | -      | 84    | -     | 84     | -      | 79    | -     | 79     | -      | 79    | -     | 79 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 95    | -     | 95     | -      | 84    | -     | 84     | -      | 84    | -     | 84     | -      | 79    | -     | 79     | -      | 79    | -     | 79 |
|           | Week 24 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 94    | -     | 94     | -      | 84    | -     | 84     | -      | 83    | -     | 83     | -      | 78    | -     | 78     | -      | 78    | -     | 78 |
|           |         | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Total   | -      | 94    | -     | 94     | -      | 84    | -     | 84     | -      | 83    | -     | 83     | -      | 78    | -     | 78     | -      | 78    | -     | 78 |
| Week 28   | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 94     | -     | 94    | -      | 81     | -     | 81    | -      | 83     | -     | 83    | -      | 77     | -     | 77    | -      | 77     | -     | 77    |    |
|           | Upper   | -       | -      | -     | -     | -      | 2      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Total   | -       | 94     | -     | 94    | -      | 83     | -     | 83    | -      | 83     | -     | 83    | -      | 77     | -     | 77    | -      | 77     | -     | 77    |    |
| Follow-up | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 98     | -     | 98    | -      | 84     | -     | 84    | -      | 91     | -     | 91    | -      | 87     | -     | 87    | -      | 87     | -     | 87    |    |
|           | Upper   | -       | -      | -     | -     | -      | 2      | -     | 2     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Total   | -       | 98     | -     | 98    | -      | 86     | -     | 86    | -      | 91     | -     | 91    | -      | 87     | -     | 87    | -      | 87     | -     | 87    |    |

表 2.7.4.7-付録-17 尿検査の評価時点別クロス表 (CCT-001 試験) (3/6)

|           |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Glucose   | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 24    | 4     | 28     | -      | 29    | 3     | 32     | -      | 26    | 4     | 30     | -      | 20    | 5     | 25     | -      | 31    | 6     | 37 |
|           |         | Upper   | -      | 2     | 9     | 11     | -      | 2     | 3     | 5      | -      | 5     | 4     | 9      | -      | 2     | 10    | 12     | -      | 2     | 7     | 9  |
|           |         | Total   | -      | 26    | 13    | 39     | -      | 31    | 6     | 37     | -      | 31    | 8     | 39     | -      | 22    | 15    | 37     | -      | 33    | 13    | 46 |
|           | Week 2  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 53    | 14    | 67     | -      | 59    | 9     | 68     | -      | 58    | 11    | 69     | -      | 52    | 13    | 65     | -      | 54    | 15    | 69 |
|           |         | Upper   | -      | 6     | 24    | 30     | -      | 5     | 16    | 21     | -      | 7     | 14    | 21     | -      | 5     | 22    | 27     | -      | 1     | 18    | 19 |
|           |         | Total   | -      | 59    | 38    | 97     | -      | 64    | 25    | 89     | -      | 65    | 25    | 90     | -      | 57    | 35    | 92     | -      | 55    | 33    | 88 |
|           | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 23    | 4     | 27     | -      | 26    | 2     | 28     | -      | 29    | 6     | 35     | -      | 17    | 5     | 22     | -      | 29    | 5     | 34 |
|           |         | Upper   | -      | 1     | 9     | 10     | -      | 3     | 5     | 8      | -      | 1     | 2     | 3      | -      | 3     | 8     | 11     | -      | 2     | 5     | 7  |
|           |         | Total   | -      | 24    | 13    | 37     | -      | 29    | 7     | 36     | -      | 30    | 8     | 38     | -      | 20    | 13    | 33     | -      | 31    | 10    | 41 |
|           | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 53    | 15    | 68     | -      | 62    | 9     | 71     | -      | 56    | 13    | 69     | -      | 49    | 16    | 65     | -      | 51    | 18    | 69 |
|           |         | Upper   | -      | 5     | 23    | 28     | -      | 2     | 16    | 18     | -      | 7     | 12    | 19     | -      | 5     | 19    | 24     | -      | 4     | 13    | 17 |
|           |         | Total   | -      | 58    | 38    | 96     | -      | 64    | 25    | 89     | -      | 63    | 25    | 88     | -      | 54    | 35    | 89     | -      | 55    | 31    | 86 |
|           | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 22    | 5     | 27     | -      | 29    | 3     | 32     | -      | 23    | 8     | 31     | -      | 17    | 7     | 24     | -      | 30    | 9     | 39 |
|           |         | Upper   | -      | 5     | 8     | 13     | -      | 3     | 5     | 8      | -      | 4     | 3     | 7      | -      | 1     | 7     | 8      | -      | 3     | 4     | 7  |
|           |         | Total   | -      | 27    | 13    | 40     | -      | 32    | 8     | 40     | -      | 27    | 11    | 38     | -      | 18    | 14    | 32     | -      | 33    | 13    | 46 |
|           | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 50    | 15    | 65     | -      | 58    | 7     | 65     | -      | 54    | 12    | 66     | -      | 47    | 17    | 64     | -      | 49    | 16    | 65 |
|           |         | Upper   | -      | 8     | 22    | 30     | -      | 4     | 17    | 21     | -      | 9     | 13    | 22     | -      | 6     | 16    | 22     | -      | 4     | 14    | 18 |
|           |         | Total   | -      | 58    | 37    | 95     | -      | 62    | 24    | 86     | -      | 63    | 25    | 88     | -      | 53    | 33    | 86     | -      | 53    | 30    | 83 |
|           | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 50    | 15    | 65     | -      | 59    | 13    | 72     | -      | 54    | 13    | 67     | -      | 46    | 19    | 65     | -      | 49    | 13    | 62 |
|           |         | Upper   | -      | 8     | 22    | 30     | -      | 3     | 11    | 14     | -      | 7     | 11    | 18     | -      | 6     | 11    | 17     | -      | 5     | 15    | 20 |
|           |         | Total   | -      | 58    | 37    | 95     | -      | 62    | 24    | 86     | -      | 61    | 24    | 85     | -      | 52    | 30    | 82     | -      | 54    | 28    | 82 |
|           | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 47    | 18    | 65     | -      | 56    | 11    | 67     | -      | 53    | 13    | 66     | -      | 45    | 18    | 63     | -      | 48    | 18    | 66 |
|           |         | Upper   | -      | 11    | 19    | 30     | -      | 5     | 12    | 17     | -      | 7     | 11    | 18     | -      | 6     | 11    | 17     | -      | 4     | 11    | 15 |
|           |         | Total   | -      | 58    | 37    | 95     | -      | 61    | 23    | 84     | -      | 60    | 24    | 84     | -      | 51    | 29    | 80     | -      | 52    | 29    | 81 |
|           | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 48    | 13    | 61     | -      | 58    | 11    | 69     | -      | 49    | 12    | 61     | -      | 44    | 19    | 63     | -      | 47    | 16    | 63 |
|           |         | Upper   | -      | 10    | 24    | 34     | -      | 3     | 12    | 15     | -      | 11    | 12    | 23     | -      | 7     | 9     | 16     | -      | 3     | 13    | 16 |
|           |         | Total   | -      | 58    | 37    | 95     | -      | 61    | 23    | 84     | -      | 60    | 24    | 84     | -      | 51    | 28    | 79     | -      | 50    | 29    | 79 |
|           | Week 24 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 48    | 13    | 61     | -      | 54    | 13    | 67     | -      | 52    | 17    | 69     | -      | 45    | 18    | 63     | -      | 46    | 15    | 61 |
|           |         | Upper   | -      | 9     | 24    | 33     | -      | 7     | 10    | 17     | -      | 8     | 6     | 14     | -      | 5     | 10    | 15     | -      | 3     | 14    | 17 |
|           |         | Total   | -      | 57    | 37    | 94     | -      | 61    | 23    | 84     | -      | 60    | 23    | 83     | -      | 50    | 28    | 78     | -      | 49    | 29    | 78 |
| Week 28   | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 49     | 9     | 58    | -      | 50     | 9     | 59    | -      | 50     | 11    | 61    | -      | 42     | 15    | 57    | -      | 45     | 13    | 58    |    |
|           | Upper   | -       | 8      | 28    | 36    | -      | 11     | 13    | 24    | -      | 10     | 12    | 22    | -      | 7      | 13    | 20    | -      | 4      | 15    | 19    |    |
|           | Total   | -       | 57     | 37    | 94    | -      | 61     | 22    | 83    | -      | 60     | 23    | 83    | -      | 49     | 28    | 77    | -      | 49     | 28    | 77    |    |
| Follow-up | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 52     | 9     | 61    | -      | 51     | 10    | 61    | -      | 56     | 11    | 67    | -      | 47     | 17    | 64    | -      | 48     | 16    | 64    |    |
|           | Upper   | -       | 8      | 29    | 37    | -      | 11     | 14    | 25    | -      | 10     | 14    | 24    | -      | 7      | 16    | 23    | -      | 7      | 16    | 23    |    |
|           | Total   | -       | 60     | 38    | 98    | -      | 62     | 24    | 86    | -      | 66     | 25    | 91    | -      | 54     | 33    | 87    | -      | 55     | 32    | 87    |    |

表 2.7.4.7-付録-17 尿検査の評価時点別クロス表 (CCT-001 試験) (4/6)

|           |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Protein   | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 16    | 3     | 19     | -      | 18    | 1     | 19     | -      | 18    | 3     | 21     | -      | 20    | 1     | 21     | -      | 25    | 2     | 27 |
|           |         | Upper   | -      | 6     | 14    | 20     | -      | 4     | 14    | 18     | -      | 4     | 14    | 18     | -      | 5     | 11    | 16     | -      | 5     | 14    | 19 |
|           |         | Total   | -      | 22    | 17    | 39     | -      | 22    | 15    | 37     | -      | 22    | 17    | 39     | -      | 25    | 12    | 37     | -      | 30    | 16    | 46 |
|           | Week 2  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 45    | 19    | 64     | -      | 41    | 5     | 46     | -      | 33    | 12    | 45     | -      | 43    | 4     | 47     | -      | 53    | 5     | 58 |
|           |         | Upper   | -      | 6     | 27    | 33     | -      | 13    | 30    | 43     | -      | 13    | 32    | 45     | -      | 13    | 32    | 45     | -      | 7     | 23    | 30 |
|           |         | Total   | -      | 51    | 46    | 97     | -      | 54    | 35    | 89     | -      | 46    | 44    | 90     | -      | 56    | 36    | 92     | -      | 60    | 28    | 88 |
|           | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 18    | 5     | 23     | -      | 18    | 2     | 20     | -      | 17    | 5     | 22     | -      | 22    | 2     | 24     | -      | 22    | 6     | 28 |
|           |         | Upper   | -      | 3     | 11    | 14     | -      | 4     | 12    | 16     | -      | 5     | 11    | 16     | -      | 1     | 8     | 9      | -      | 5     | 8     | 13 |
|           |         | Total   | -      | 21    | 16    | 37     | -      | 22    | 14    | 36     | -      | 22    | 16    | 38     | -      | 23    | 10    | 33     | -      | 27    | 14    | 41 |
|           | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 39    | 16    | 55     | -      | 44    | 11    | 55     | -      | 40    | 11    | 51     | -      | 47    | 9     | 56     | -      | 49    | 6     | 55 |
|           |         | Upper   | -      | 11    | 30    | 41     | -      | 10    | 24    | 34     | -      | 6     | 31    | 37     | -      | 8     | 25    | 33     | -      | 10    | 21    | 31 |
|           |         | Total   | -      | 50    | 46    | 96     | -      | 54    | 35    | 89     | -      | 46    | 42    | 88     | -      | 55    | 34    | 89     | -      | 59    | 27    | 86 |
|           | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 18    | 5     | 23     | -      | 21    | 4     | 25     | -      | 17    | 4     | 21     | -      | 21    | 1     | 22     | -      | 26    | 5     | 31 |
|           |         | Upper   | -      | 5     | 12    | 17     | -      | 5     | 10    | 15     | -      | 5     | 12    | 17     | -      | 1     | 9     | 10     | -      | 4     | 11    | 15 |
|           |         | Total   | -      | 23    | 17    | 40     | -      | 26    | 14    | 40     | -      | 22    | 16    | 38     | -      | 22    | 10    | 32     | -      | 30    | 16    | 46 |
|           | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 41    | 18    | 59     | -      | 40    | 7     | 47     | -      | 31    | 15    | 46     | -      | 48    | 8     | 56     | -      | 45    | 9     | 54 |
|           |         | Upper   | -      | 9     | 27    | 36     | -      | 13    | 26    | 39     | -      | 15    | 27    | 42     | -      | 7     | 23    | 30     | -      | 12    | 17    | 29 |
|           |         | Total   | -      | 50    | 45    | 95     | -      | 53    | 33    | 86     | -      | 46    | 42    | 88     | -      | 55    | 31    | 86     | -      | 57    | 26    | 83 |
|           | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 39    | 16    | 55     | -      | 43    | 11    | 54     | -      | 36    | 12    | 48     | -      | 43    | 9     | 52     | -      | 47    | 7     | 54 |
|           |         | Upper   | -      | 11    | 29    | 40     | -      | 10    | 22    | 32     | -      | 8     | 29    | 37     | -      | 10    | 20    | 30     | -      | 11    | 17    | 28 |
|           |         | Total   | -      | 50    | 45    | 95     | -      | 53    | 33    | 86     | -      | 44    | 41    | 85     | -      | 53    | 29    | 82     | -      | 58    | 24    | 82 |
|           | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 36    | 16    | 52     | -      | 43    | 11    | 54     | -      | 35    | 9     | 44     | -      | 41    | 8     | 49     | -      | 46    | 4     | 50 |
|           |         | Upper   | -      | 14    | 29    | 43     | -      | 8     | 21    | 29     | -      | 8     | 32    | 40     | -      | 10    | 21    | 31     | -      | 10    | 21    | 31 |
|           |         | Total   | -      | 50    | 45    | 95     | -      | 51    | 32    | 83     | -      | 43    | 41    | 84     | -      | 51    | 29    | 80     | -      | 56    | 25    | 81 |
|           | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 41    | 11    | 52     | -      | 43    | 10    | 53     | -      | 33    | 11    | 44     | -      | 46    | 9     | 55     | -      | 48    | 6     | 54 |
|           |         | Upper   | -      | 9     | 34    | 43     | -      | 9     | 22    | 31     | -      | 10    | 30    | 40     | -      | 5     | 19    | 24     | -      | 6     | 19    | 25 |
|           |         | Total   | -      | 50    | 45    | 95     | -      | 52    | 32    | 84     | -      | 43    | 41    | 84     | -      | 51    | 28    | 79     | -      | 54    | 25    | 79 |
|           | Week 24 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |         | Normal  | -      | 40    | 15    | 55     | -      | 44    | 11    | 55     | -      | 37    | 14    | 51     | -      | 45    | 9     | 54     | -      | 39    | 7     | 46 |
|           |         | Upper   | -      | 10    | 29    | 39     | -      | 8     | 21    | 29     | -      | 5     | 27    | 32     | -      | 6     | 18    | 24     | -      | 15    | 17    | 32 |
|           |         | Total   | -      | 50    | 44    | 94     | -      | 52    | 32    | 84     | -      | 42    | 41    | 83     | -      | 51    | 27    | 78     | -      | 54    | 24    | 78 |
| Week 28   | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 40     | 12    | 52    | -      | 42     | 10    | 52    | -      | 32     | 14    | 46    | -      | 43     | 11    | 54    | -      | 42     | 5     | 47    |    |
|           | Upper   | -       | 10     | 31    | 41    | -      | 10     | 21    | 31    | -      | 10     | 27    | 37    | -      | 7      | 16    | 23    | -      | 11     | 19    | 30    |    |
|           | Total   | -       | 50     | 43    | 93    | -      | 52     | 31    | 83    | -      | 42     | 41    | 83    | -      | 50     | 27    | 77    | -      | 53     | 24    | 77    |    |
| Follow-up | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal  | -       | 41     | 13    | 54    | -      | 42     | 10    | 52    | -      | 34     | 14    | 48    | -      | 45     | 12    | 57    | -      | 48     | 7     | 55    |    |
|           | Upper   | -       | 10     | 33    | 43    | -      | 10     | 24    | 34    | -      | 12     | 31    | 43    | -      | 8      | 22    | 30    | -      | 11     | 21    | 32    |    |
|           | Total   | -       | 51     | 46    | 97    | -      | 52     | 34    | 86    | -      | 46     | 45    | 91    | -      | 53     | 34    | 87    | -      | 59     | 28    | 87    |    |

表 2.7.4.7-付録-17 尿検査の評価時点別クロス表 (CCT-001 試験) (5/6)

|           |        | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|-----------|--------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|           |        | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|           |        | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Ketones   | Week 1 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 45    | -     | 45 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |        | Total   | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 39    | -     | 39     | -      | 37    | -     | 37     | -      | 46    | -     | 46 |
|           | Week 2 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 94    | 1     | 95     | -      | 89    | -     | 89     | -      | 90    | -     | 90     | -      | 92    | -     | 92     | -      | 88    | -     | 88 |
|           |        | Upper   | -      | 2     | -     | 2      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -  |
|           |        | Total   | -      | 96    | 1     | 97     | -      | 89    | -     | 89     | -      | 90    | -     | 90     | -      | 92    | -     | 92     | -      | 88    | -     | 88 |
|           | Week 3 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 37    | -     | 37     | -      | 36    | -     | 36     | -      | 37    | -     | 37     | -      | 33    | -     | 33     | -      | 41    | -     | 41 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Total   | -      | 37    | -     | 37     | -      | 36    | -     | 36     | -      | 38    | -     | 38     | -      | 33    | -     | 33     | -      | 41    | -     | 41 |
|           | Week 4 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 95    | 1     | 96     | -      | 89    | -     | 89     | -      | 87    | -     | 87     | -      | 89    | -     | 89     | -      | 85    | -     | 85 |
|           |        | Upper   | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |        | Total   | -      | 95    | 1     | 96     | -      | 89    | -     | 89     | -      | 88    | -     | 88     | -      | 89    | -     | 89     | -      | 86    | -     | 86 |
|           | Week 6 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           |        | Normal  | -      | 40    | -     | 40     | -      | 39    | -     | 39     | -      | 38    | -     | 38     | -      | 32    | -     | 32     | -      | 45    | -     | 45 |
|           |        | Upper   | -      | -     | -     | -      | -      | 1     | -     | 1      | -      | -     | -     | -      | -      | -     | -     | -      | -      | 1     | -     | 1  |
|           |        | Total   | -      | 40    | -     | 40     | -      | 40    | -     | 40     | -      | 38    | -     | 38     | -      | 32    | -     | 32     | -      | 46    | -     | 46 |
| Week 8    | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 93     | 1     | 94    | -      | 86     | -     | 86    | -      | 88     | -     | 88    | -      | 86     | -     | 86    | -      | 83     | -     | 83    |    |
|           | Upper  | -       | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|           | Total  | -       | 94     | 1     | 95    | -      | 86     | -     | 86    | -      | 88     | -     | 88    | -      | 86     | -     | 86    | -      | 83     | -     | 83    |    |
| Week 12   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 94     | 1     | 95    | -      | 85     | -     | 85    | -      | 85     | -     | 85    | -      | 80     | -     | 80    | -      | 81     | -     | 81    |    |
|           | Upper  | -       | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | -     | 1     |    |
|           | Total  | -       | 94     | 1     | 95    | -      | 86     | -     | 86    | -      | 85     | -     | 85    | -      | 82     | -     | 82    | -      | 82     | -     | 82    |    |
| Week 16   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 94     | 1     | 95    | -      | 84     | -     | 84    | -      | 84     | -     | 84    | -      | 80     | -     | 80    | -      | 80     | -     | 80    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Total  | -       | 94     | 1     | 95    | -      | 84     | -     | 84    | -      | 84     | -     | 84    | -      | 80     | -     | 80    | -      | 81     | -     | 81    |    |
| Week 20   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 94     | 1     | 95    | -      | 84     | -     | 84    | -      | 82     | -     | 82    | -      | 78     | -     | 78    | -      | 77     | -     | 77    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | 2      | -     | 2     |    |
|           | Total  | -       | 94     | 1     | 95    | -      | 84     | -     | 84    | -      | 84     | -     | 84    | -      | 79     | -     | 79    | -      | 79     | -     | 79    |    |
| Week 24   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 93     | 1     | 94    | -      | 84     | -     | 84    | -      | 82     | -     | 82    | -      | 78     | -     | 78    | -      | 78     | -     | 78    |    |
|           | Upper  | -       | -      | -     | -     | -      | -      | -     | -     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Total  | -       | 93     | 1     | 94    | -      | 84     | -     | 84    | -      | 83     | -     | 83    | -      | 78     | -     | 78    | -      | 78     | -     | 78    |    |
| Week 28   | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 93     | 1     | 94    | -      | 81     | -     | 81    | -      | 82     | -     | 82    | -      | 77     | -     | 77    | -      | 76     | -     | 76    |    |
|           | Upper  | -       | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     | 1      | -      | 1     |       |    |
|           | Total  | -       | 93     | 1     | 94    | -      | 83     | -     | 83    | -      | 83     | -     | 83    | -      | 77     | -     | 77    | -      | 77     | -     | 77    |    |
| Follow-up | Lower  | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|           | Normal | -       | 97     | 1     | 98    | -      | 84     | -     | 84    | -      | 90     | -     | 90    | -      | 87     | -     | 87    | -      | 86     | -     | 86    |    |
|           | Upper  | -       | -      | -     | -     | -      | 2      | -     | 2     | -      | 1      | -     | 1     | -      | -      | -     | -     | -      | 1      | -     | 1     |    |
|           | Total  | -       | 97     | 1     | 98    | -      | 86     | -     | 86    | -      | 91     | -     | 91    | -      | 87     | -     | 87    | -      | 87     | -     | 87    |    |

表 2.7.4.7-付録-17 尿検査の評価時点別クロス表 (CCT-001 試験) (6/6)

|              |         | Placebo |        |       |       | 15 mg  |        |       |       | 30 mg  |        |       |       | 60 mg  |        |       |       | 120 mg |        |       |       |    |
|--------------|---------|---------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|--------|--------|-------|-------|----|
|              |         | Week 0  |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       | Week 0 |        |       |       |    |
|              |         | Lower   | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total | Lower  | Normal | Upper | Total |    |
| Occult Blood | Week 1  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 33    | -     | 33     | -      | 31    | 2     | 33     | -      | 30    | 4     | 34     | -      | 32    | 1     | 33     | -      | 41    | 1     | 42 |
|              |         | Upper   | -      | 1     | 4     | 5      | -      | 2     | 1     | 3      | -      | 2     | 3     | 5      | -      | 1     | 2     | 3      | -      | 2     | 1     | 3  |
|              |         | Total   | -      | 34    | 4     | 38     | -      | 33    | 3     | 36     | -      | 32    | 7     | 39     | -      | 33    | 3     | 36     | -      | 43    | 2     | 45 |
|              | Week 2  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 82    | 5     | 87     | -      | 73    | 7     | 80     | -      | 78    | 4     | 82     | -      | 77    | 7     | 84     | -      | 77    | 1     | 78 |
|              |         | Upper   | -      | 4     | 5     | 9      | -      | 4     | 3     | 7      | -      | 3     | 4     | 7      | -      | 4     | 3     | 7      | -      | 3     | 4     | 7  |
|              |         | Total   | -      | 86    | 10    | 96     | -      | 77    | 10    | 87     | -      | 81    | 8     | 89     | -      | 81    | 10    | 91     | -      | 80    | 5     | 85 |
|              | Week 3  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 31    | 1     | 32     | -      | 30    | 3     | 33     | -      | 30    | 3     | 33     | -      | 29    | 2     | 31     | -      | 34    | -     | 34 |
|              |         | Upper   | -      | 3     | 1     | 4      | -      | 2     | -     | 2      | -      | 2     | 3     | 5      | -      | 1     | 1     | 2      | -      | 4     | 2     | 6  |
|              |         | Total   | -      | 34    | 2     | 36     | -      | 32    | 3     | 35     | -      | 32    | 6     | 38     | -      | 30    | 3     | 33     | -      | 38    | 2     | 40 |
|              | Week 4  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 82    | 3     | 85     | -      | 75    | 9     | 84     | -      | 73    | 2     | 75     | -      | 79    | 2     | 81     | -      | 77    | 2     | 79 |
|              |         | Upper   | -      | 3     | 6     | 9      | -      | 3     | 1     | 4      | -      | 7     | 6     | 13     | -      | 2     | 6     | 8      | -      | 4     | 2     | 6  |
|              |         | Total   | -      | 85    | 9     | 94     | -      | 78    | 10    | 88     | -      | 80    | 8     | 88     | -      | 81    | 8     | 89     | -      | 81    | 4     | 85 |
|              | Week 6  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 36    | 1     | 37     | -      | 29    | 5     | 34     | -      | 31    | 3     | 34     | -      | 30    | 1     | 31     | -      | 40    | 1     | 41 |
|              |         | Upper   | -      | 1     | 1     | 2      | -      | 3     | 1     | 4      | -      | 1     | 3     | 4      | -      | -     | 1     | 1      | -      | 2     | 1     | 3  |
|              |         | Total   | -      | 37    | 2     | 39     | -      | 32    | 6     | 38     | -      | 32    | 6     | 38     | -      | 30    | 2     | 32     | -      | 42    | 2     | 44 |
|              | Week 8  | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 83    | 4     | 87     | -      | 71    | 7     | 78     | -      | 78    | 4     | 82     | -      | 74    | 2     | 76     | -      | 70    | 3     | 73 |
|              |         | Upper   | -      | 2     | 4     | 6      | -      | 5     | 2     | 7      | -      | 2     | 4     | 6      | -      | 5     | 5     | 10     | -      | 8     | 1     | 9  |
|              |         | Total   | -      | 85    | 8     | 93     | -      | 76    | 9     | 85     | -      | 80    | 8     | 88     | -      | 79    | 7     | 86     | -      | 78    | 4     | 82 |
|              | Week 12 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 82    | 4     | 86     | -      | 75    | 6     | 81     | -      | 76    | 5     | 81     | -      | 71    | 1     | 72     | -      | 72    | 1     | 73 |
|              |         | Upper   | -      | 3     | 5     | 8      | -      | 2     | 3     | 5      | -      | 1     | 3     | 4      | -      | 5     | 5     | 10     | -      | 6     | 2     | 8  |
|              |         | Total   | -      | 85    | 9     | 94     | -      | 77    | 9     | 86     | -      | 77    | 8     | 85     | -      | 76    | 6     | 82     | -      | 78    | 3     | 81 |
|              | Week 16 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 81    | 4     | 85     | -      | 72    | 7     | 79     | -      | 73    | 4     | 77     | -      | 69    | 2     | 71     | -      | 69    | 1     | 70 |
|              |         | Upper   | -      | 4     | 4     | 8      | -      | 1     | 1     | 2      | -      | 3     | 4     | 7      | -      | 6     | 3     | 9      | -      | 8     | 2     | 10 |
|              |         | Total   | -      | 85    | 8     | 93     | -      | 73    | 8     | 81     | -      | 76    | 8     | 84     | -      | 75    | 5     | 80     | -      | 77    | 3     | 80 |
|              | Week 20 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 82    | 5     | 87     | -      | 71    | 8     | 79     | -      | 73    | 4     | 77     | -      | 67    | 3     | 70     | -      | 69    | 1     | 70 |
|              |         | Upper   | -      | 3     | 3     | 6      | -      | 2     | -     | 2      | -      | 3     | 4     | 7      | -      | 6     | 2     | 8      | -      | 6     | 2     | 8  |
|              |         | Total   | -      | 85    | 8     | 93     | -      | 73    | 8     | 81     | -      | 76    | 8     | 84     | -      | 73    | 5     | 78     | -      | 75    | 3     | 78 |
|              | Week 24 | Lower   | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     |    |
|              |         | Normal  | -      | 83    | 5     | 88     | -      | 70    | 6     | 76     | -      | 73    | 5     | 78     | -      | 70    | 3     | 73     | -      | 68    | 1     | 69 |
|              |         | Upper   | -      | 1     | 4     | 5      | -      | 4     | 2     | 6      | -      | 2     | 3     | 5      | -      | 2     | 2     | 4      | -      | 5     | 2     | 7  |
|              |         | Total   | -      | 84    | 9     | 93     | -      | 74    | 8     | 82     | -      | 75    | 8     | 83     | -      | 72    | 5     | 77     | -      | 73    | 3     | 76 |
| Week 28      | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|              | Normal  | -       | 78     | 4     | 82    | -      | 74     | 4     | 78    | -      | 73     | 4     | 77    | -      | 69     | 3     | 72    | -      | 66     | 1     | 67    |    |
|              | Upper   | -       | 6      | 4     | 10    | -      | 1      | 3     | 4     | -      | 2      | 4     | 6     | -      | 3      | 2     | 5     | -      | 6      | 2     | 8     |    |
|              | Total   | -       | 84     | 8     | 92    | -      | 75     | 7     | 82    | -      | 75     | 8     | 83    | -      | 72     | 5     | 77    | -      | 72     | 3     | 75    |    |
| Follow-up    | Lower   | -       | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     | -     | -      | -      | -     |       |    |
|              | Normal  | -       | 81     | 4     | 85    | -      | 76     | 4     | 80    | -      | 80     | 5     | 85    | -      | 76     | 5     | 81    | -      | 74     | 1     | 75    |    |
|              | Upper   | -       | 6      | 5     | 11    | -      | 1      | 4     | 5     | -      | 2      | 4     | 6     | -      | 3      | 3     | 6     | -      | 6      | 4     | 10    |    |
|              | Total   | -       | 87     | 9     | 96    | -      | 77     | 8     | 85    | -      | 82     | 9     | 91    | -      | 79     | 8     | 87    | -      | 80     | 5     | 85    |    |



表 2.7.4.7-付録-18 血液学的検査の評価時点別の変化量 (CCT-002 試験) (1/6)

| Hemoglobin (g/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2            | Placebo        | 65                 | -0.06 | 0.466 | -1.5 | -0.30 | -0.10  | 0.20 | 1.5 |
|                   | ATL-962 120 mg | 141                | -0.13 | 0.489 | -1.9 | -0.40 | -0.10  | 0.10 | 1.3 |
| Week 4            | Placebo        | 63                 | -0.05 | 0.582 | -1.7 | -0.40 | 0.00   | 0.30 | 1.4 |
|                   | ATL-962 120 mg | 138                | -0.17 | 0.505 | -1.4 | -0.50 | -0.15  | 0.20 | 1.0 |
| Week 8            | Placebo        | 62                 | -0.09 | 0.571 | -1.5 | -0.50 | -0.10  | 0.30 | 1.3 |
|                   | ATL-962 120 mg | 137                | -0.19 | 0.608 | -1.5 | -0.50 | -0.20  | 0.10 | 3.1 |
| Week 12           | Placebo        | 59                 | -0.15 | 0.575 | -1.4 | -0.50 | -0.20  | 0.20 | 1.4 |
|                   | ATL-962 120 mg | 137                | -0.20 | 0.689 | -1.8 | -0.50 | -0.20  | 0.10 | 4.2 |
| Week 16           | Placebo        | 59                 | -0.13 | 0.617 | -1.3 | -0.50 | -0.20  | 0.30 | 1.6 |
|                   | ATL-962 120 mg | 134                | -0.26 | 0.713 | -1.9 | -0.70 | -0.30  | 0.10 | 4.7 |
| Week 20           | Placebo        | 59                 | -0.21 | 0.714 | -3.2 | -0.60 | -0.20  | 0.20 | 1.5 |
|                   | ATL-962 120 mg | 133                | -0.19 | 0.741 | -1.5 | -0.60 | -0.20  | 0.10 | 4.9 |
| Week 24           | Placebo        | 59                 | -0.29 | 0.795 | -3.2 | -0.70 | -0.30  | 0.10 | 1.9 |
|                   | ATL-962 120 mg | 130                | -0.28 | 0.752 | -1.8 | -0.80 | -0.30  | 0.00 | 4.8 |
| Week 28           | Placebo        | 59                 | -0.13 | 0.785 | -2.0 | -0.70 | -0.20  | 0.40 | 1.8 |
|                   | ATL-962 120 mg | 127                | -0.25 | 0.781 | -1.6 | -0.70 | -0.30  | 0.10 | 5.5 |
| Week 32           | Placebo        | 58                 | 0.02  | 0.679 | -1.7 | -0.30 | 0.00   | 0.30 | 3.0 |
|                   | ATL-962 120 mg | 126                | -0.09 | 0.796 | -1.7 | -0.50 | -0.10  | 0.30 | 5.3 |
| Week 36           | Placebo        | 57                 | -0.06 | 0.501 | -1.4 | -0.30 | -0.10  | 0.30 | 1.0 |
|                   | ATL-962 120 mg | 124                | 0.00  | 0.818 | -1.5 | -0.50 | 0.00   | 0.40 | 5.0 |
| Week 40           | Placebo        | 58                 | 0.07  | 0.705 | -1.8 | -0.20 | 0.05   | 0.30 | 3.2 |
|                   | ATL-962 120 mg | 122                | 0.07  | 0.920 | -2.0 | -0.40 | 0.00   | 0.50 | 5.3 |
| Week 44           | Placebo        | 58                 | 0.19  | 0.544 | -0.8 | -0.20 | 0.20   | 0.50 | 2.1 |
|                   | ATL-962 120 mg | 119                | 0.05  | 0.793 | -1.7 | -0.40 | -0.10  | 0.50 | 3.6 |
| Week 48           | Placebo        | 57                 | 0.15  | 0.610 | -1.5 | -0.10 | 0.10   | 0.40 | 2.0 |
|                   | ATL-962 120 mg | 119                | 0.00  | 0.761 | -1.5 | -0.30 | 0.00   | 0.30 | 3.8 |
| Week 52           | Placebo        | 57                 | 0.08  | 0.612 | -1.4 | -0.20 | 0.20   | 0.40 | 1.8 |
|                   | ATL-962 120 mg | 118                | -0.00 | 0.866 | -2.2 | -0.50 | 0.00   | 0.40 | 4.1 |

| Hematocrit (%) |                | Summary Statistics |       |       |      |       |        |      |      |
|----------------|----------------|--------------------|-------|-------|------|-------|--------|------|------|
| Visit          | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max  |
| Week 2         | Placebo        | 65                 | -0.05 | 1.534 | -4.0 | -1.00 | -0.10  | 0.60 | 5.5  |
|                | ATL-962 120 mg | 141                | -0.32 | 1.490 | -5.6 | -1.10 | -0.30  | 0.50 | 3.0  |
| Week 4         | Placebo        | 63                 | -0.17 | 1.650 | -4.4 | -1.20 | -0.10  | 0.70 | 4.4  |
|                | ATL-962 120 mg | 138                | -0.35 | 1.483 | -3.9 | -1.40 | -0.25  | 0.70 | 3.1  |
| Week 8         | Placebo        | 62                 | 0.03  | 1.857 | -4.6 | -1.20 | -0.20  | 1.30 | 5.1  |
|                | ATL-962 120 mg | 137                | -0.29 | 1.914 | -5.1 | -1.40 | -0.40  | 0.70 | 9.0  |
| Week 12        | Placebo        | 59                 | -0.09 | 1.936 | -3.3 | -1.30 | -0.40  | 1.10 | 5.7  |
|                | ATL-962 120 mg | 137                | -0.14 | 2.097 | -6.4 | -1.40 | -0.30  | 0.70 | 10.9 |
| Week 16        | Placebo        | 59                 | 0.16  | 1.858 | -4.3 | -0.80 | 0.20   | 1.30 | 5.0  |
|                | ATL-962 120 mg | 134                | -0.31 | 2.091 | -5.6 | -1.60 | -0.40  | 0.90 | 12.1 |
| Week 20        | Placebo        | 59                 | -0.02 | 2.229 | -9.3 | -1.50 | 0.00   | 1.50 | 5.6  |
|                | ATL-962 120 mg | 133                | -0.04 | 2.089 | -4.1 | -1.30 | -0.10  | 1.20 | 12.3 |
| Week 24        | Placebo        | 59                 | -0.26 | 2.264 | -7.9 | -1.50 | -0.30  | 1.00 | 5.7  |
|                | ATL-962 120 mg | 130                | -0.42 | 2.136 | -6.0 | -1.60 | -0.55  | 0.70 | 12.3 |
| Week 28        | Placebo        | 59                 | -0.09 | 2.355 | -5.7 | -1.50 | -0.10  | 1.00 | 6.5  |
|                | ATL-962 120 mg | 127                | -0.29 | 2.251 | -5.2 | -1.60 | -0.40  | 0.90 | 14.2 |
| Week 32        | Placebo        | 58                 | 0.30  | 2.040 | -5.0 | -0.70 | 0.35   | 1.10 | 10.2 |
|                | ATL-962 120 mg | 126                | -0.02 | 2.059 | -5.2 | -1.20 | -0.10  | 0.90 | 12.6 |
| Week 36        | Placebo        | 58                 | 0.24  | 1.863 | -3.8 | -0.50 | 0.00   | 1.10 | 8.9  |
|                | ATL-962 120 mg | 124                | 0.19  | 2.274 | -4.2 | -1.20 | 0.20   | 1.60 | 11.6 |
| Week 40        | Placebo        | 58                 | 0.21  | 2.154 | -6.0 | -0.90 | 0.20   | 1.00 | 9.8  |
|                | ATL-962 120 mg | 122                | 0.25  | 2.641 | -6.6 | -1.30 | 0.05   | 1.70 | 13.2 |
| Week 44        | Placebo        | 58                 | 0.39  | 1.778 | -2.7 | -0.80 | 0.50   | 1.30 | 6.5  |
|                | ATL-962 120 mg | 119                | 0.05  | 2.285 | -5.8 | -1.40 | -0.10  | 1.30 | 8.6  |
| Week 48        | Placebo        | 57                 | 0.42  | 2.041 | -6.1 | -0.60 | 0.40   | 1.50 | 6.7  |
|                | ATL-962 120 mg | 119                | -0.03 | 2.149 | -4.9 | -1.10 | -0.10  | 0.90 | 9.3  |
| Week 52        | Placebo        | 57                 | 0.14  | 1.871 | -4.2 | -0.80 | 0.00   | 1.10 | 6.3  |
|                | ATL-962 120 mg | 118                | -0.01 | 2.463 | -5.9 | -1.50 | -0.15  | 1.30 | 10.0 |

表 2.7.4.7-付録-18 血液学的検査の評価時点別の変化量 (CCT-002 試験) (2/6)

| RBC (x10 <sup>4</sup> /μL) |                | Summary Statistics |       |       |      |       |        |      |     |
|----------------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit                      | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2                     | Placebo        | 65                 | -1.6  | 15.34 | -51  | -10.0 | -4.0   | 6.0  | 46  |
|                            | ATL-962 120 mg | 141                | -4.2  | 16.01 | -60  | -14.0 | -3.0   | 5.0  | 40  |
| Week 4                     | Placebo        | 63                 | -1.2  | 18.27 | -46  | -15.0 | 1.0    | 10.0 | 54  |
|                            | ATL-962 120 mg | 138                | -4.4  | 15.62 | -46  | -15.0 | -2.5   | 6.0  | 36  |
| Week 8                     | Placebo        | 62                 | 0.5   | 19.89 | -45  | -16.0 | 1.0    | 15.0 | 55  |
|                            | ATL-962 120 mg | 137                | -3.8  | 18.84 | -60  | -13.0 | -4.0   | 6.0  | 68  |
| Week 12                    | Placebo        | 59                 | -1.2  | 20.55 | -37  | -15.0 | -4.0   | 12.0 | 47  |
|                            | ATL-962 120 mg | 137                | -2.9  | 20.82 | -67  | -15.0 | -4.0   | 8.0  | 62  |
| Week 16                    | Placebo        | 59                 | 0.8   | 20.15 | -38  | -13.0 | 0.0    | 14.0 | 60  |
|                            | ATL-962 120 mg | 134                | -5.3  | 19.42 | -76  | -17.0 | -4.5   | 6.0  | 45  |
| Week 20                    | Placebo        | 59                 | -1.8  | 23.56 | -107 | -12.0 | 1.0    | 16.0 | 54  |
|                            | ATL-962 120 mg | 133                | -3.6  | 19.11 | -58  | -15.0 | -4.0   | 9.0  | 46  |
| Week 24                    | Placebo        | 59                 | -6.3  | 24.18 | -97  | -20.0 | -5.0   | 8.0  | 53  |
|                            | ATL-962 120 mg | 130                | -9.0  | 20.62 | -76  | -23.0 | -8.0   | 3.0  | 54  |
| Week 28                    | Placebo        | 59                 | -5.6  | 23.34 | -71  | -19.0 | -7.0   | 8.0  | 45  |
|                            | ATL-962 120 mg | 127                | -10.0 | 20.34 | -81  | -23.0 | -12.0  | 4.0  | 42  |
| Week 32                    | Placebo        | 58                 | -4.2  | 19.85 | -66  | -13.0 | -6.0   | 5.0  | 69  |
|                            | ATL-962 120 mg | 126                | -8.7  | 20.14 | -83  | -20.0 | -9.0   | 2.0  | 47  |
| Week 36                    | Placebo        | 58                 | -5.4  | 19.34 | -54  | -16.0 | -6.5   | 6.0  | 63  |
|                            | ATL-962 120 mg | 124                | -7.1  | 21.22 | -65  | -20.0 | -7.5   | 5.5  | 63  |
| Week 40                    | Placebo        | 58                 | -5.4  | 21.98 | -79  | -14.0 | -7.0   | 5.0  | 70  |
|                            | ATL-962 120 mg | 122                | -6.2  | 24.01 | -81  | -22.0 | -6.5   | 10.0 | 82  |
| Week 44                    | Placebo        | 58                 | -2.8  | 18.20 | -42  | -17.0 | -2.5   | 5.0  | 66  |
|                            | ATL-962 120 mg | 119                | -8.3  | 22.33 | -84  | -23.0 | -8.0   | 4.0  | 47  |
| Week 48                    | Placebo        | 57                 | -2.7  | 21.71 | -68  | -15.0 | -4.0   | 9.0  | 48  |
|                            | ATL-962 120 mg | 119                | -8.3  | 20.60 | -79  | -20.0 | -6.0   | 5.0  | 43  |
| Week 52                    | Placebo        | 57                 | -4.5  | 19.26 | -55  | -15.0 | -7.0   | 7.0  | 38  |
|                            | ATL-962 120 mg | 118                | -7.0  | 23.70 | -93  | -21.0 | -7.0   | 8.0  | 62  |

| WBC (/μL) |                | Summary Statistics |        |         |       |         |        |        |      |
|-----------|----------------|--------------------|--------|---------|-------|---------|--------|--------|------|
| Visit     | Treatment      | N                  | Mean   | SD      | Min   | Q1      | Median | Q3     | Max  |
| Week 2    | Placebo        | 65                 | 75.4   | 1166.19 | -2800 | -800.0  | 100.0  | 800.0  | 3300 |
|           | ATL-962 120 mg | 141                | 323.4  | 1031.62 | -3900 | -200.0  | 300.0  | 800.0  | 3900 |
| Week 4    | Placebo        | 63                 | -169.8 | 1315.62 | -6700 | -700.0  | -100.0 | 700.0  | 1700 |
|           | ATL-962 120 mg | 138                | 444.9  | 1280.40 | -3500 | -200.0  | 300.0  | 1100.0 | 8100 |
| Week 8    | Placebo        | 62                 | -272.6 | 1670.79 | -8300 | -800.0  | -150.0 | 600.0  | 4400 |
|           | ATL-962 120 mg | 137                | 235.8  | 1012.26 | -2500 | -400.0  | 100.0  | 700.0  | 4700 |
| Week 12   | Placebo        | 59                 | -49.2  | 1587.17 | -7800 | -800.0  | 0.0    | 600.0  | 4800 |
|           | ATL-962 120 mg | 137                | 365.7  | 1218.57 | -2900 | -300.0  | 200.0  | 800.0  | 4400 |
| Week 16   | Placebo        | 59                 | -81.4  | 1657.43 | -7800 | -800.0  | 0.0    | 700.0  | 3000 |
|           | ATL-962 120 mg | 134                | 196.3  | 928.69  | -3800 | -200.0  | 200.0  | 700.0  | 2700 |
| Week 20   | Placebo        | 59                 | 1.7    | 1615.82 | -6800 | -700.0  | -200.0 | 800.0  | 4300 |
|           | ATL-962 120 mg | 133                | 184.2  | 974.51  | -2600 | -400.0  | 100.0  | 800.0  | 3000 |
| Week 24   | Placebo        | 59                 | -155.9 | 1600.78 | -7600 | -700.0  | -300.0 | 400.0  | 4400 |
|           | ATL-962 120 mg | 130                | 300.8  | 1087.67 | -3000 | -400.0  | 200.0  | 900.0  | 3700 |
| Week 28   | Placebo        | 59                 | -305.1 | 1472.56 | -7400 | -700.0  | -100.0 | 400.0  | 2400 |
|           | ATL-962 120 mg | 127                | 187.4  | 1205.15 | -2900 | -700.0  | 100.0  | 800.0  | 6000 |
| Week 32   | Placebo        | 58                 | -405.2 | 1422.31 | -7600 | -1100.0 | -150.0 | 500.0  | 2100 |
|           | ATL-962 120 mg | 126                | 54.0   | 1083.01 | -3300 | -500.0  | 100.0  | 700.0  | 3100 |
| Week 36   | Placebo        | 58                 | -463.8 | 1457.19 | -7200 | -1000.0 | -350.0 | 400.0  | 3700 |
|           | ATL-962 120 mg | 124                | 36.3   | 1131.75 | -2900 | -600.0  | -100.0 | 500.0  | 4600 |
| Week 40   | Placebo        | 58                 | -491.4 | 1585.71 | -8800 | -1000.0 | -200.0 | 200.0  | 3200 |
|           | ATL-962 120 mg | 122                | 181.1  | 1168.05 | -2700 | -600.0  | 200.0  | 1000.0 | 4400 |
| Week 44   | Placebo        | 58                 | -189.7 | 1767.12 | -6700 | -900.0  | -150.0 | 400.0  | 7500 |
|           | ATL-962 120 mg | 119                | 37.0   | 1224.80 | -4000 | -600.0  | -100.0 | 600.0  | 6300 |
| Week 48   | Placebo        | 57                 | -491.2 | 1513.31 | -7600 | -900.0  | -200.0 | 300.0  | 1700 |
|           | ATL-962 120 mg | 119                | 77.3   | 1152.26 | -2900 | -600.0  | 0.0    | 700.0  | 5000 |
| Week 52   | Placebo        | 57                 | -719.3 | 1514.74 | -6900 | -1200.0 | -500.0 | -100.0 | 2300 |
|           | ATL-962 120 mg | 118                | 56.8   | 1146.04 | -4300 | -500.0  | 0.0    | 600.0  | 5100 |

表 2.7.4.7-付録-18 血液学的検査の評価時点別の変化量 (CCT-002 試験) (3/6)

| Neutrophils (%) |                | Summary Statistics |       |       |       |       |        |      |      |
|-----------------|----------------|--------------------|-------|-------|-------|-------|--------|------|------|
| Visit           | Treatment      | N                  | Mean  | SD    | Min   | Q1    | Median | Q3   | Max  |
| Week 2          | Placebo        | 65                 | -1.77 | 6.312 | -19.3 | -6.00 | -0.90  | 2.50 | 11.0 |
|                 | ATL-962 120 mg | 141                | -0.53 | 5.837 | -20.5 | -4.60 | 0.00   | 3.00 | 16.4 |
| Week 4          | Placebo        | 63                 | -0.97 | 5.848 | -18.5 | -4.80 | -0.40  | 3.20 | 12.6 |
|                 | ATL-962 120 mg | 138                | 0.39  | 6.159 | -15.0 | -3.50 | 0.25   | 3.70 | 31.5 |
| Week 8          | Placebo        | 62                 | -1.52 | 7.765 | -23.9 | -4.20 | -1.50  | 2.70 | 22.4 |
|                 | ATL-962 120 mg | 137                | 0.61  | 5.590 | -13.6 | -3.00 | 1.00   | 4.70 | 12.8 |
| Week 12         | Placebo        | 59                 | -0.25 | 8.251 | -21.2 | -5.20 | 0.00   | 5.00 | 19.1 |
|                 | ATL-962 120 mg | 137                | 1.19  | 7.071 | -17.0 | -3.00 | 0.70   | 5.30 | 40.6 |
| Week 16         | Placebo        | 59                 | -0.99 | 8.079 | -23.5 | -4.40 | -0.40  | 2.80 | 17.0 |
|                 | ATL-962 120 mg | 134                | 0.09  | 5.644 | -21.0 | -3.70 | 0.35   | 3.10 | 17.9 |
| Week 20         | Placebo        | 59                 | -0.67 | 7.650 | -23.8 | -4.60 | -0.50  | 4.00 | 17.3 |
|                 | ATL-962 120 mg | 133                | -0.27 | 5.930 | -17.0 | -4.00 | -0.50  | 3.80 | 17.9 |
| Week 24         | Placebo        | 59                 | -0.57 | 7.340 | -22.3 | -4.00 | -0.60  | 4.40 | 15.7 |
|                 | ATL-962 120 mg | 130                | 0.28  | 6.482 | -23.0 | -4.20 | -0.20  | 4.60 | 18.2 |
| Week 28         | Placebo        | 59                 | -1.05 | 7.530 | -25.5 | -5.00 | -0.90  | 4.30 | 12.5 |
|                 | ATL-962 120 mg | 127                | -0.08 | 5.900 | -20.0 | -3.90 | -0.70  | 3.50 | 16.0 |
| Week 32         | Placebo        | 58                 | -1.72 | 6.930 | -21.5 | -5.10 | -1.30  | 2.00 | 16.8 |
|                 | ATL-962 120 mg | 126                | 0.09  | 5.460 | -17.0 | -3.60 | -0.30  | 3.40 | 14.1 |
| Week 36         | Placebo        | 58                 | -1.30 | 6.689 | -20.3 | -4.10 | -1.10  | 3.00 | 11.3 |
|                 | ATL-962 120 mg | 124                | 0.35  | 6.314 | -22.0 | -3.60 | -0.35  | 3.85 | 25.9 |
| Week 40         | Placebo        | 58                 | -1.78 | 7.302 | -21.9 | -5.90 | -1.20  | 3.90 | 10.6 |
|                 | ATL-962 120 mg | 122                | 0.70  | 6.338 | -18.1 | -3.20 | 0.70   | 4.80 | 17.1 |
| Week 44         | Placebo        | 58                 | -0.13 | 7.978 | -20.8 | -4.10 | 0.00   | 4.60 | 20.7 |
|                 | ATL-962 120 mg | 119                | 0.19  | 7.855 | -40.3 | -3.90 | -0.30  | 4.10 | 28.7 |
| Week 48         | Placebo        | 57                 | -1.36 | 7.993 | -27.7 | -3.60 | 0.00   | 3.30 | 14.2 |
|                 | ATL-962 120 mg | 119                | 1.42  | 7.033 | -21.0 | -2.60 | 1.50   | 5.20 | 21.0 |
| Week 52         | Placebo        | 57                 | -1.90 | 7.588 | -23.0 | -5.60 | -1.50  | 2.50 | 17.1 |
|                 | ATL-962 120 mg | 118                | 1.78  | 7.048 | -21.8 | -2.70 | 1.30   | 6.80 | 23.7 |

| Lymphocytes (%) |                | Summary Statistics |       |       |       |       |        |      |      |
|-----------------|----------------|--------------------|-------|-------|-------|-------|--------|------|------|
| Visit           | Treatment      | N                  | Mean  | SD    | Min   | Q1    | Median | Q3   | Max  |
| Week 2          | Placebo        | 65                 | 1.51  | 5.934 | -12.1 | -1.90 | 1.30   | 5.00 | 21.9 |
|                 | ATL-962 120 mg | 141                | 0.24  | 5.213 | -14.8 | -2.90 | 0.20   | 3.90 | 14.6 |
| Week 4          | Placebo        | 63                 | 0.79  | 5.433 | -8.3  | -3.10 | -0.10  | 3.90 | 22.2 |
|                 | ATL-962 120 mg | 138                | -0.39 | 5.300 | -22.4 | -3.00 | -0.50  | 2.70 | 14.0 |
| Week 8          | Placebo        | 62                 | 1.47  | 6.641 | -21.2 | -1.60 | 1.35   | 4.00 | 24.4 |
|                 | ATL-962 120 mg | 137                | -0.44 | 5.207 | -13.5 | -3.90 | -0.70  | 2.90 | 14.8 |
| Week 12         | Placebo        | 59                 | 0.40  | 7.548 | -17.5 | -3.60 | -0.40  | 3.90 | 22.8 |
|                 | ATL-962 120 mg | 137                | -1.37 | 6.441 | -36.6 | -4.40 | -0.60  | 2.60 | 12.7 |
| Week 16         | Placebo        | 59                 | 1.25  | 7.136 | -14.7 | -1.90 | 0.20   | 3.30 | 26.2 |
|                 | ATL-962 120 mg | 134                | 0.24  | 5.091 | -16.0 | -3.10 | 0.25   | 3.80 | 15.0 |
| Week 20         | Placebo        | 59                 | 0.93  | 7.276 | -15.3 | -3.20 | 1.00   | 4.80 | 19.4 |
|                 | ATL-962 120 mg | 133                | 0.48  | 5.208 | -15.2 | -2.90 | 0.50   | 3.60 | 14.3 |
| Week 24         | Placebo        | 59                 | 0.71  | 6.696 | -15.5 | -4.00 | -0.30  | 4.00 | 22.6 |
|                 | ATL-962 120 mg | 130                | 0.26  | 5.810 | -14.4 | -3.80 | 0.60   | 3.80 | 21.0 |
| Week 28         | Placebo        | 59                 | 1.27  | 6.949 | -12.0 | -3.30 | 1.20   | 4.50 | 24.1 |
|                 | ATL-962 120 mg | 127                | 0.42  | 5.429 | -22.0 | -2.50 | 0.60   | 3.40 | 15.0 |
| Week 32         | Placebo        | 58                 | 2.02  | 6.561 | -15.5 | -2.30 | 1.60   | 6.30 | 23.2 |
|                 | ATL-962 120 mg | 126                | 0.20  | 5.066 | -10.8 | -2.90 | 0.20   | 3.20 | 15.0 |
| Week 36         | Placebo        | 58                 | 1.16  | 6.437 | -12.0 | -2.40 | 1.25   | 4.00 | 22.8 |
|                 | ATL-962 120 mg | 124                | -0.24 | 5.582 | -18.2 | -3.85 | -0.15  | 3.30 | 20.5 |
| Week 40         | Placebo        | 58                 | 2.33  | 6.540 | -9.1  | -2.10 | 1.30   | 4.80 | 23.3 |
|                 | ATL-962 120 mg | 122                | -0.32 | 5.690 | -14.7 | -3.60 | -0.30  | 3.40 | 14.6 |
| Week 44         | Placebo        | 58                 | 0.09  | 7.478 | -17.7 | -4.70 | 0.20   | 3.50 | 23.0 |
|                 | ATL-962 120 mg | 119                | -0.17 | 6.616 | -22.4 | -4.90 | 0.70   | 3.40 | 29.5 |
| Week 48         | Placebo        | 57                 | 0.92  | 7.393 | -13.0 | -4.30 | -0.30  | 2.60 | 24.4 |
|                 | ATL-962 120 mg | 119                | -1.29 | 6.058 | -20.0 | -4.70 | -1.00  | 2.00 | 19.0 |
| Week 52         | Placebo        | 57                 | 1.69  | 7.057 | -17.4 | -2.80 | 1.80   | 4.10 | 27.0 |
|                 | ATL-962 120 mg | 118                | -1.43 | 6.490 | -22.9 | -6.00 | -0.45  | 3.00 | 19.1 |

表 2.7.4.7-付録-18 血液学的検査の評価時点別の変化量 (CCT-002 試験) (4/6)

| Monocytes (%) |                | Summary Statistics |       |       |      |       |        |      |     |
|---------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit         | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2        | Placebo        | 65                 | 0.23  | 1.435 | -4.3 | -0.70 | 0.10   | 1.00 | 3.5 |
|               | ATL-962 120 mg | 141                | 0.03  | 1.438 | -3.7 | -0.70 | 0.00   | 1.00 | 4.9 |
| Week 4        | Placebo        | 63                 | 0.11  | 1.568 | -3.8 | -0.80 | 0.10   | 0.90 | 4.5 |
|               | ATL-962 120 mg | 138                | -0.28 | 1.444 | -4.8 | -1.20 | -0.30  | 0.50 | 3.6 |
| Week 8        | Placebo        | 62                 | 0.09  | 1.467 | -3.9 | -0.90 | 0.10   | 1.10 | 3.2 |
|               | ATL-962 120 mg | 137                | -0.26 | 1.360 | -3.5 | -1.10 | -0.50  | 0.60 | 4.7 |
| Week 12       | Placebo        | 59                 | -0.02 | 1.969 | -6.7 | -0.90 | 0.30   | 1.00 | 3.8 |
|               | ATL-962 120 mg | 137                | -0.04 | 1.562 | -4.0 | -1.10 | -0.20  | 0.60 | 5.7 |
| Week 16       | Placebo        | 59                 | -0.06 | 1.354 | -4.4 | -0.60 | 0.10   | 0.60 | 3.0 |
|               | ATL-962 120 mg | 134                | -0.27 | 1.395 | -3.3 | -1.20 | -0.30  | 0.50 | 4.5 |
| Week 20       | Placebo        | 59                 | -0.03 | 1.712 | -4.4 | -0.60 | 0.00   | 0.70 | 3.9 |
|               | ATL-962 120 mg | 133                | -0.05 | 1.436 | -4.1 | -1.00 | -0.10  | 0.90 | 4.0 |
| Week 24       | Placebo        | 59                 | -0.02 | 1.409 | -3.1 | -1.00 | 0.00   | 0.80 | 4.2 |
|               | ATL-962 120 mg | 130                | -0.34 | 1.437 | -4.5 | -1.10 | -0.40  | 0.40 | 3.8 |
| Week 28       | Placebo        | 59                 | -0.15 | 1.520 | -4.6 | -1.00 | 0.00   | 0.90 | 3.5 |
|               | ATL-962 120 mg | 127                | -0.32 | 1.346 | -4.4 | -1.10 | -0.20  | 0.40 | 4.5 |
| Week 32       | Placebo        | 58                 | -0.17 | 1.935 | -5.2 | -1.10 | -0.10  | 1.00 | 4.2 |
|               | ATL-962 120 mg | 126                | -0.33 | 1.412 | -4.2 | -1.40 | -0.30  | 0.50 | 2.9 |
| Week 36       | Placebo        | 58                 | -0.03 | 1.563 | -4.9 | -0.80 | 0.05   | 0.80 | 3.5 |
|               | ATL-962 120 mg | 124                | -0.24 | 1.581 | -6.0 | -1.00 | -0.10  | 0.60 | 6.3 |
| Week 40       | Placebo        | 58                 | -0.37 | 1.853 | -6.2 | -1.10 | -0.30  | 0.90 | 2.7 |
|               | ATL-962 120 mg | 122                | -0.42 | 1.261 | -4.1 | -1.10 | -0.40  | 0.50 | 2.3 |
| Week 44       | Placebo        | 58                 | 0.12  | 1.785 | -5.1 | -0.60 | 0.30   | 1.00 | 4.9 |
|               | ATL-962 120 mg | 119                | -0.21 | 1.582 | -4.3 | -1.10 | -0.30  | 0.70 | 6.0 |
| Week 48       | Placebo        | 57                 | 0.24  | 1.800 | -4.9 | -0.80 | 0.30   | 1.50 | 5.2 |
|               | ATL-962 120 mg | 119                | -0.20 | 1.533 | -4.9 | -1.20 | -0.30  | 0.70 | 3.6 |
| Week 52       | Placebo        | 57                 | 0.10  | 1.594 | -5.0 | -0.60 | 0.30   | 1.10 | 3.6 |
|               | ATL-962 120 mg | 118                | -0.24 | 1.480 | -3.6 | -1.30 | -0.40  | 0.80 | 5.0 |

  

| Eosinophils (%) |                | Summary Statistics |       |       |      |       |        |      |      |
|-----------------|----------------|--------------------|-------|-------|------|-------|--------|------|------|
| Visit           | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max  |
| Week 2          | Placebo        | 65                 | 0.08  | 1.126 | -3.7 | -0.40 | 0.10   | 0.80 | 2.9  |
|                 | ATL-962 120 mg | 141                | 0.28  | 1.318 | -2.7 | -0.40 | 0.10   | 0.70 | 6.8  |
| Week 4          | Placebo        | 63                 | 0.12  | 1.427 | -5.0 | -0.40 | 0.20   | 0.80 | 4.5  |
|                 | ATL-962 120 mg | 138                | 0.27  | 1.390 | -3.6 | -0.40 | 0.10   | 0.70 | 7.2  |
| Week 8          | Placebo        | 62                 | -0.04 | 1.836 | -6.9 | -0.70 | 0.05   | 0.60 | 5.5  |
|                 | ATL-962 120 mg | 137                | 0.17  | 1.581 | -3.5 | -0.60 | 0.10   | 0.50 | 11.0 |
| Week 12         | Placebo        | 59                 | -0.05 | 1.899 | -9.1 | -0.80 | 0.10   | 0.90 | 3.4  |
|                 | ATL-962 120 mg | 137                | 0.24  | 1.714 | -3.5 | -0.70 | 0.10   | 0.50 | 9.7  |
| Week 16         | Placebo        | 59                 | -0.19 | 1.708 | -7.3 | -0.70 | -0.10  | 0.50 | 4.0  |
|                 | ATL-962 120 mg | 134                | -0.00 | 1.270 | -4.3 | -0.60 | 0.10   | 0.70 | 4.0  |
| Week 20         | Placebo        | 59                 | -0.16 | 1.993 | -7.3 | -0.40 | 0.00   | 0.60 | 5.4  |
|                 | ATL-962 120 mg | 133                | -0.14 | 1.080 | -4.6 | -0.70 | -0.10  | 0.50 | 2.2  |
| Week 24         | Placebo        | 59                 | -0.07 | 1.773 | -7.1 | -0.80 | 0.00   | 0.70 | 3.3  |
|                 | ATL-962 120 mg | 130                | -0.13 | 1.178 | -4.5 | -0.60 | 0.00   | 0.50 | 3.9  |
| Week 28         | Placebo        | 59                 | 0.01  | 1.583 | -5.0 | -0.60 | -0.10  | 1.00 | 3.1  |
|                 | ATL-962 120 mg | 127                | 0.02  | 1.555 | -4.8 | -0.90 | -0.10  | 0.60 | 6.5  |
| Week 32         | Placebo        | 58                 | -0.08 | 1.576 | -6.9 | -0.70 | 0.00   | 0.60 | 2.9  |
|                 | ATL-962 120 mg | 126                | 0.10  | 1.415 | -3.8 | -0.70 | 0.00   | 0.80 | 6.4  |
| Week 36         | Placebo        | 58                 | 0.25  | 1.641 | -4.9 | -0.40 | 0.10   | 1.00 | 4.3  |
|                 | ATL-962 120 mg | 124                | 0.11  | 1.350 | -4.8 | -0.55 | 0.00   | 0.70 | 5.0  |
| Week 40         | Placebo        | 58                 | -0.09 | 1.577 | -6.9 | -0.50 | 0.10   | 0.60 | 3.5  |
|                 | ATL-962 120 mg | 122                | 0.03  | 1.334 | -3.5 | -0.70 | -0.10  | 0.40 | 4.4  |
| Week 44         | Placebo        | 58                 | -0.07 | 1.855 | -6.7 | -0.70 | -0.20  | 0.90 | 5.5  |
|                 | ATL-962 120 mg | 119                | 0.19  | 1.716 | -2.7 | -0.60 | 0.00   | 0.70 | 14.0 |
| Week 48         | Placebo        | 57                 | 0.26  | 1.523 | -5.5 | -0.40 | 0.30   | 0.80 | 4.8  |
|                 | ATL-962 120 mg | 119                | 0.04  | 1.437 | -3.8 | -0.60 | -0.10  | 0.80 | 5.3  |
| Week 52         | Placebo        | 57                 | -0.01 | 1.392 | -3.7 | -0.80 | 0.00   | 0.90 | 3.5  |
|                 | ATL-962 120 mg | 118                | -0.05 | 1.348 | -4.3 | -0.80 | -0.10  | 0.40 | 6.3  |

表 2.7.4.7-付録-18 血液学的検査の評価時点別の変化量 (CCT-002 試験) (5/6)

| Basophils (%) |                | Summary Statistics |       |       |      |       |        |      |     |
|---------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit         | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2        | Placebo        | 65                 | -0.05 | 0.479 | -1.3 | -0.30 | 0.00   | 0.20 | 1.6 |
|               | ATL-962 120 mg | 141                | -0.01 | 0.374 | -1.3 | -0.20 | 0.00   | 0.20 | 1.0 |
| Week 4        | Placebo        | 63                 | -0.04 | 0.416 | -1.0 | -0.30 | 0.00   | 0.20 | 1.1 |
|               | ATL-962 120 mg | 138                | 0.01  | 0.412 | -1.3 | -0.20 | 0.00   | 0.20 | 1.6 |
| Week 8        | Placebo        | 62                 | 0.01  | 0.514 | -1.0 | -0.20 | 0.00   | 0.20 | 2.0 |
|               | ATL-962 120 mg | 137                | -0.07 | 0.358 | -1.5 | -0.30 | 0.00   | 0.10 | 1.2 |
| Week 12       | Placebo        | 59                 | -0.07 | 0.432 | -1.0 | -0.40 | 0.00   | 0.10 | 1.0 |
|               | ATL-962 120 mg | 137                | -0.02 | 0.411 | -1.0 | -0.30 | 0.00   | 0.20 | 1.4 |
| Week 16       | Placebo        | 59                 | -0.01 | 0.456 | -1.0 | -0.20 | 0.00   | 0.20 | 1.5 |
|               | ATL-962 120 mg | 134                | -0.05 | 0.368 | -1.2 | -0.20 | -0.10  | 0.10 | 1.3 |
| Week 20       | Placebo        | 59                 | -0.07 | 0.434 | -1.3 | -0.30 | 0.00   | 0.10 | 1.2 |
|               | ATL-962 120 mg | 133                | -0.02 | 0.415 | -1.1 | -0.20 | 0.00   | 0.20 | 1.5 |
| Week 24       | Placebo        | 59                 | -0.05 | 0.491 | -1.3 | -0.30 | -0.10  | 0.20 | 1.6 |
|               | ATL-962 120 mg | 130                | -0.08 | 0.366 | -1.4 | -0.30 | 0.00   | 0.10 | 1.0 |
| Week 28       | Placebo        | 59                 | -0.08 | 0.404 | -1.0 | -0.30 | -0.10  | 0.10 | 1.4 |
|               | ATL-962 120 mg | 127                | -0.05 | 0.369 | -1.1 | -0.30 | -0.10  | 0.10 | 1.3 |
| Week 32       | Placebo        | 58                 | -0.06 | 0.441 | -1.1 | -0.30 | -0.05  | 0.20 | 1.3 |
|               | ATL-962 120 mg | 126                | -0.06 | 0.426 | -1.5 | -0.30 | -0.10  | 0.10 | 1.7 |
| Week 36       | Placebo        | 58                 | -0.07 | 0.456 | -1.3 | -0.30 | -0.10  | 0.20 | 1.0 |
|               | ATL-962 120 mg | 124                | 0.02  | 0.438 | -1.1 | -0.20 | 0.00   | 0.20 | 2.1 |
| Week 40       | Placebo        | 58                 | -0.09 | 0.385 | -1.0 | -0.40 | -0.05  | 0.10 | 1.0 |
|               | ATL-962 120 mg | 122                | 0.00  | 0.459 | -1.3 | -0.20 | 0.00   | 0.20 | 2.1 |
| Week 44       | Placebo        | 58                 | -0.00 | 0.459 | -0.8 | -0.30 | -0.05  | 0.20 | 2.0 |
|               | ATL-962 120 mg | 119                | 0.01  | 0.441 | -1.7 | -0.20 | 0.00   | 0.20 | 1.5 |
| Week 48       | Placebo        | 57                 | -0.06 | 0.464 | -1.3 | -0.30 | 0.00   | 0.20 | 1.3 |
|               | ATL-962 120 mg | 119                | 0.03  | 0.456 | -1.7 | -0.20 | 0.00   | 0.30 | 1.7 |
| Week 52       | Placebo        | 57                 | 0.12  | 0.498 | -0.9 | -0.20 | 0.10   | 0.40 | 1.6 |
|               | ATL-962 120 mg | 118                | -0.05 | 0.393 | -1.2 | -0.30 | 0.00   | 0.10 | 1.1 |

| Platelet Count (x10 <sup>4</sup> /μL) |                | Summary Statistics |       |       |       |       |        |       |      |
|---------------------------------------|----------------|--------------------|-------|-------|-------|-------|--------|-------|------|
| Visit                                 | Treatment      | N                  | Mean  | SD    | Min   | Q1    | Median | Q3    | Max  |
| Week 2                                | Placebo        | 65                 | 0.86  | 2.305 | -4.3  | -0.40 | 0.90   | 2.10  | 8.5  |
|                                       | ATL-962 120 mg | 138                | 0.77  | 2.825 | -12.6 | -0.50 | 0.65   | 2.00  | 9.9  |
| Week 4                                | Placebo        | 63                 | 0.79  | 1.890 | -4.4  | -0.50 | 0.90   | 1.80  | 7.2  |
|                                       | ATL-962 120 mg | 136                | 0.96  | 2.273 | -8.7  | -0.20 | 1.05   | 2.45  | 6.7  |
| Week 8                                | Placebo        | 61                 | 0.31  | 2.332 | -6.3  | -1.20 | 0.30   | 1.90  | 5.6  |
|                                       | ATL-962 120 mg | 133                | 0.84  | 2.549 | -7.0  | -0.80 | 0.80   | 2.80  | 8.5  |
| Week 12                               | Placebo        | 58                 | 0.34  | 2.058 | -4.3  | -1.00 | 0.70   | 1.80  | 4.3  |
|                                       | ATL-962 120 mg | 134                | 0.41  | 3.032 | -11.4 | -1.20 | 0.30   | 2.10  | 15.3 |
| Week 16                               | Placebo        | 57                 | 0.35  | 1.603 | -4.2  | -0.90 | 0.50   | 1.40  | 4.2  |
|                                       | ATL-962 120 mg | 131                | 0.30  | 2.969 | -17.0 | -1.10 | 0.50   | 1.80  | 9.5  |
| Week 20                               | Placebo        | 59                 | 0.06  | 2.329 | -5.6  | -1.60 | -0.30  | 1.70  | 5.6  |
|                                       | ATL-962 120 mg | 129                | 0.34  | 3.377 | -17.6 | -1.80 | 0.20   | 2.10  | 9.8  |
| Week 24                               | Placebo        | 59                 | -0.16 | 2.483 | -5.7  | -1.90 | -0.40  | 1.70  | 7.4  |
|                                       | ATL-962 120 mg | 124                | -0.09 | 2.985 | -18.0 | -1.75 | 0.10   | 1.55  | 6.3  |
| Week 28                               | Placebo        | 59                 | -0.95 | 2.454 | -6.7  | -2.50 | -1.20  | 0.70  | 5.5  |
|                                       | ATL-962 120 mg | 124                | -0.50 | 3.313 | -17.4 | -2.20 | -0.70  | 1.05  | 9.7  |
| Week 32                               | Placebo        | 58                 | -1.31 | 2.387 | -8.9  | -2.80 | -1.55  | 0.10  | 4.3  |
|                                       | ATL-962 120 mg | 124                | -1.01 | 3.225 | -18.6 | -2.35 | -0.85  | 0.80  | 5.5  |
| Week 36                               | Placebo        | 57                 | -1.41 | 2.298 | -6.3  | -3.00 | -1.60  | 0.10  | 5.7  |
|                                       | ATL-962 120 mg | 122                | -1.51 | 3.195 | -16.5 | -3.10 | -1.35  | 0.30  | 11.4 |
| Week 40                               | Placebo        | 57                 | -1.54 | 2.369 | -7.3  | -3.00 | -1.40  | 0.00  | 4.3  |
|                                       | ATL-962 120 mg | 120                | -1.20 | 3.355 | -17.6 | -2.90 | -1.20  | 1.10  | 7.5  |
| Week 44                               | Placebo        | 58                 | -0.88 | 2.408 | -5.8  | -2.60 | -1.10  | 0.50  | 6.2  |
|                                       | ATL-962 120 mg | 117                | -1.11 | 3.216 | -15.3 | -2.90 | -0.90  | 0.80  | 6.5  |
| Week 48                               | Placebo        | 57                 | -0.95 | 2.446 | -7.0  | -2.40 | -1.20  | 0.10  | 6.7  |
|                                       | ATL-962 120 mg | 117                | -1.20 | 3.319 | -18.0 | -2.90 | -0.90  | 0.60  | 7.9  |
| Week 52                               | Placebo        | 57                 | -1.46 | 2.318 | -7.8  | -2.80 | -1.60  | -0.10 | 4.5  |
|                                       | ATL-962 120 mg | 115                | -1.48 | 3.510 | -18.3 | -3.20 | -1.30  | 0.30  | 7.5  |

表 2.7.4.7-付録-18 血液学的検査の評価時点別の変化量 (CCT-002 試験) (6/6)

| Prothrombin Time (sec) |                | Summary Statistics |       |       |       |       |        |      |      |
|------------------------|----------------|--------------------|-------|-------|-------|-------|--------|------|------|
| Visit                  | Treatment      | N                  | Mean  | SD    | Min   | Q1    | Median | Q3   | Max  |
| Week 2                 | Placebo        | 65                 | -0.13 | 1.372 | -9.3  | -0.30 | 0.00   | 0.40 | 3.5  |
|                        | ATL-962 120 mg | 141                | 0.03  | 0.570 | -1.7  | -0.40 | 0.00   | 0.40 | 1.2  |
| Week 4                 | Placebo        | 63                 | -0.40 | 3.122 | -24.1 | -0.30 | 0.10   | 0.30 | 2.7  |
|                        | ATL-962 120 mg | 138                | 0.10  | 0.692 | -2.7  | -0.30 | 0.05   | 0.50 | 4.4  |
| Week 8                 | Placebo        | 62                 | 0.15  | 2.036 | -6.8  | -0.30 | 0.20   | 0.40 | 13.3 |
|                        | ATL-962 120 mg | 137                | 0.17  | 0.684 | -2.2  | -0.30 | 0.20   | 0.60 | 3.2  |
| Week 12                | Placebo        | 59                 | 0.19  | 1.208 | -6.7  | -0.10 | 0.30   | 0.60 | 4.0  |
|                        | ATL-962 120 mg | 137                | 0.27  | 0.727 | -1.3  | -0.20 | 0.20   | 0.70 | 4.3  |
| Week 16                | Placebo        | 59                 | 0.10  | 1.453 | -9.1  | -0.20 | 0.20   | 0.80 | 2.2  |
|                        | ATL-962 120 mg | 134                | 0.53  | 2.022 | -1.5  | -0.10 | 0.40   | 0.80 | 20.9 |
| Week 20                | Placebo        | 59                 | 0.70  | 1.842 | -2.9  | 0.10  | 0.50   | 1.00 | 13.4 |
|                        | ATL-962 120 mg | 133                | 0.47  | 0.663 | -0.9  | 0.00  | 0.50   | 0.80 | 5.1  |
| Week 24                | Placebo        | 59                 | 0.46  | 0.910 | -3.2  | -0.10 | 0.50   | 0.90 | 3.5  |
|                        | ATL-962 120 mg | 130                | 0.43  | 1.411 | -1.7  | -0.10 | 0.35   | 0.80 | 14.6 |
| Week 28                | Placebo        | 59                 | -0.02 | 2.382 | -16.9 | -0.20 | 0.30   | 0.70 | 2.8  |
|                        | ATL-962 120 mg | 127                | 0.40  | 1.145 | -1.7  | -0.20 | 0.40   | 0.80 | 10.2 |
| Week 32                | Placebo        | 58                 | 0.26  | 0.846 | -2.8  | -0.10 | 0.30   | 0.70 | 2.4  |
|                        | ATL-962 120 mg | 127                | 0.35  | 1.100 | -1.8  | -0.10 | 0.30   | 0.70 | 9.6  |
| Week 36                | Placebo        | 57                 | 0.08  | 1.334 | -7.9  | -0.20 | 0.30   | 0.70 | 1.7  |
|                        | ATL-962 120 mg | 124                | 0.29  | 0.898 | -1.7  | -0.20 | 0.25   | 0.70 | 6.4  |
| Week 40                | Placebo        | 58                 | -0.08 | 1.189 | -7.0  | -0.40 | 0.15   | 0.50 | 2.3  |
|                        | ATL-962 120 mg | 122                | 0.12  | 0.719 | -1.5  | -0.40 | 0.20   | 0.60 | 3.1  |
| Week 44                | Placebo        | 58                 | -0.24 | 1.869 | -13.1 | -0.50 | 0.10   | 0.50 | 1.3  |
|                        | ATL-962 120 mg | 119                | 0.09  | 0.693 | -1.5  | -0.20 | 0.10   | 0.30 | 4.9  |
| Week 48                | Placebo        | 58                 | -0.11 | 1.976 | -13.6 | -0.30 | 0.10   | 0.50 | 2.3  |
|                        | ATL-962 120 mg | 119                | 0.14  | 0.709 | -1.6  | -0.40 | 0.00   | 0.40 | 2.8  |
| Week 52                | Placebo        | 58                 | -0.24 | 2.129 | -15.4 | -0.20 | 0.00   | 0.30 | 1.9  |
|                        | ATL-962 120 mg | 118                | 0.21  | 0.745 | -2.0  | -0.30 | 0.10   | 0.60 | 3.3  |

表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表 (CCT-002 試験) (1/11)

|            |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|            |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|            |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Hemoglobin | Week 2  | Lower   | -      | -     | -     | -              | 10     | 2     | -     | 12  |
|            |         | Normal  | 2      | 61    | -     | 63             | 1      | 118   | 5     | 124 |
|            |         | Upper   | -      | -     | 2     | 2              | -      | 2     | 3     | 5   |
|            |         | Total   | 2      | 61    | 2     | 65             | 11     | 122   | 8     | 141 |
|            | Week 4  | Lower   | 1      | 2     | -     | 3              | 9      | 2     | -     | 11  |
|            |         | Normal  | 1      | 56    | -     | 57             | 2      | 118   | 6     | 126 |
|            |         | Upper   | -      | 1     | 2     | 3              | -      | -     | 1     | 1   |
|            |         | Total   | 2      | 59    | 2     | 63             | 11     | 120   | 7     | 138 |
|            | Week 8  | Lower   | -      | 1     | -     | 1              | 9      | 1     | -     | 10  |
|            |         | Normal  | 2      | 56    | 1     | 59             | 2      | 117   | 4     | 123 |
|            |         | Upper   | -      | 1     | 1     | 2              | -      | 1     | 3     | 4   |
|            |         | Total   | 2      | 58    | 2     | 62             | 11     | 119   | 7     | 137 |
|            | Week 12 | Lower   | 1      | -     | -     | 1              | 8      | 2     | -     | 10  |
|            |         | Normal  | 1      | 54    | 1     | 56             | 3      | 116   | 6     | 125 |
|            |         | Upper   | -      | 1     | 1     | 2              | -      | 1     | 1     | 2   |
|            |         | Total   | 2      | 55    | 2     | 59             | 11     | 119   | 7     | 137 |
|            | Week 16 | Lower   | 1      | 1     | -     | 2              | 6      | 1     | -     | 7   |
|            |         | Normal  | 1      | 52    | 1     | 54             | 4      | 116   | 4     | 124 |
|            |         | Upper   | -      | 2     | 1     | 3              | -      | -     | 3     | 3   |
|            |         | Total   | 2      | 55    | 2     | 59             | 10     | 117   | 7     | 134 |
|            | Week 20 | Lower   | 1      | 1     | -     | 2              | 7      | 1     | -     | 8   |
|            |         | Normal  | 1      | 51    | 1     | 53             | 3      | 114   | 5     | 122 |
|            |         | Upper   | -      | 3     | 1     | 4              | -      | 1     | 2     | 3   |
|            |         | Total   | 2      | 55    | 2     | 59             | 10     | 116   | 7     | 133 |
|            | Week 24 | Lower   | 1      | 2     | -     | 3              | 8      | 2     | -     | 10  |
|            |         | Normal  | 1      | 51    | 1     | 53             | 2      | 111   | 6     | 119 |
|            |         | Upper   | -      | 2     | 1     | 3              | -      | -     | 1     | 1   |
|            |         | Total   | 2      | 55    | 2     | 59             | 10     | 113   | 7     | 130 |
|            | Week 28 | Lower   | 1      | 1     | -     | 2              | 8      | 2     | -     | 10  |
|            |         | Normal  | 1      | 51    | 1     | 53             | 2      | 107   | 4     | 113 |
|            |         | Upper   | -      | 3     | 1     | 4              | -      | 1     | 3     | 4   |
|            |         | Total   | 2      | 55    | 2     | 59             | 10     | 110   | 7     | 127 |
|            | Week 32 | Lower   | 2      | 1     | -     | 3              | 6      | -     | -     | 6   |
|            |         | Normal  | -      | 51    | 1     | 52             | 4      | 107   | 4     | 115 |
|            |         | Upper   | -      | 2     | 1     | 3              | -      | 2     | 3     | 5   |
|            |         | Total   | 2      | 54    | 2     | 58             | 10     | 109   | 7     | 126 |
|            | Week 36 | Lower   | 2      | 2     | -     | 4              | 5      | 1     | -     | 6   |
|            |         | Normal  | -      | 52    | 1     | 53             | 5      | 104   | 3     | 112 |
|            |         | Upper   | -      | -     | -     | -              | -      | 2     | 4     | 6   |
|            |         | Total   | 2      | 54    | 1     | 57             | 10     | 107   | 7     | 124 |
|            | Week 40 | Lower   | 1      | -     | -     | 1              | 5      | 1     | -     | 6   |
|            |         | Normal  | 1      | 53    | -     | 54             | 4      | 101   | 4     | 109 |
|            |         | Upper   | -      | 1     | 2     | 3              | -      | 4     | 3     | 7   |
|            |         | Total   | 2      | 54    | 2     | 58             | 9      | 106   | 7     | 122 |
|            | Week 44 | Lower   | 1      | -     | -     | 1              | 4      | 2     | -     | 6   |
|            |         | Normal  | 1      | 54    | -     | 55             | 4      | 100   | 2     | 106 |
|            |         | Upper   | -      | -     | 2     | 2              | -      | 3     | 4     | 7   |
|            |         | Total   | 2      | 54    | 2     | 58             | 8      | 105   | 6     | 119 |
| Week 48    | Lower   | 1       | -      | -     | 1     | 5              | -      | -     | 5     |     |
|            | Normal  | 1       | 53     | -     | 54    | 3              | 103    | 4     | 110   |     |
|            | Upper   | -       | -      | 2     | 2     | -              | 2      | 2     | 4     |     |
|            | Total   | 2       | 53     | 2     | 57    | 8              | 105    | 6     | 119   |     |
| Week 52    | Lower   | 1       | 1      | -     | 2     | 3              | 2      | -     | 5     |     |
|            | Normal  | 1       | 51     | -     | 52    | 5              | 99     | 3     | 107   |     |
|            | Upper   | -       | 1      | 2     | 3     | -              | 3      | 3     | 6     |     |
|            | Total   | 2       | 53     | 2     | 57    | 8              | 104    | 6     | 118   |     |

表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表 (CCT-002 試験) (2/11)

|            |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|            |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|            |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Hematocrit | Week 2  | Lower   | -      | 1     | -     | 1              | 13     | 2     | -     | 15  |
|            |         | Normal  | 4      | 59    | -     | 63             | 3      | 118   | 4     | 125 |
|            |         | Upper   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|            |         | Total   | 4      | 61    | -     | 65             | 16     | 121   | 4     | 141 |
|            | Week 4  | Lower   | 1      | 1     | -     | 2              | 14     | -     | -     | 14  |
|            |         | Normal  | 3      | 56    | -     | 59             | 1      | 120   | 3     | 124 |
|            |         | Upper   | -      | 2     | -     | 2              | -      | -     | -     | -   |
|            |         | Total   | 4      | 59    | -     | 63             | 15     | 120   | 3     | 138 |
|            | Week 8  | Lower   | 1      | -     | -     | 1              | 12     | 1     | -     | 13  |
|            |         | Normal  | 3      | 57    | -     | 60             | 3      | 117   | 2     | 122 |
|            |         | Upper   | -      | 1     | -     | 1              | -      | 1     | 1     | 2   |
|            |         | Total   | 4      | 58    | -     | 62             | 15     | 119   | 3     | 137 |
|            | Week 12 | Lower   | 1      | 2     | -     | 3              | 9      | 1     | -     | 10  |
|            |         | Normal  | 3      | 51    | -     | 54             | 6      | 117   | 3     | 126 |
|            |         | Upper   | -      | 2     | -     | 2              | -      | 1     | -     | 1   |
|            |         | Total   | 4      | 55    | -     | 59             | 15     | 119   | 3     | 137 |
|            | Week 16 | Lower   | 1      | 1     | -     | 2              | 11     | 2     | -     | 13  |
|            |         | Normal  | 3      | 52    | -     | 55             | 3      | 114   | 2     | 119 |
|            |         | Upper   | -      | 2     | -     | 2              | -      | 1     | 1     | 2   |
|            |         | Total   | 4      | 55    | -     | 59             | 14     | 117   | 3     | 134 |
|            | Week 20 | Lower   | 1      | 2     | -     | 3              | 8      | 2     | -     | 10  |
|            |         | Normal  | 3      | 50    | -     | 53             | 6      | 112   | 3     | 121 |
|            |         | Upper   | -      | 3     | -     | 3              | -      | 2     | -     | 2   |
|            |         | Total   | 4      | 55    | -     | 59             | 14     | 116   | 3     | 133 |
|            | Week 24 | Lower   | 1      | 4     | -     | 5              | 9      | 2     | -     | 11  |
|            |         | Normal  | 3      | 48    | -     | 51             | 5      | 111   | 2     | 118 |
|            |         | Upper   | -      | 3     | -     | 3              | -      | -     | 1     | 1   |
|            |         | Total   | 4      | 55    | -     | 59             | 14     | 113   | 3     | 130 |
|            | Week 28 | Lower   | 1      | 2     | -     | 3              | 10     | 2     | -     | 12  |
|            |         | Normal  | 3      | 50    | -     | 53             | 4      | 107   | 2     | 113 |
|            |         | Upper   | -      | 3     | -     | 3              | -      | 1     | 1     | 2   |
|            |         | Total   | 4      | 55    | -     | 59             | 14     | 110   | 3     | 127 |
|            | Week 32 | Lower   | 1      | 2     | -     | 3              | 10     | 1     | -     | 11  |
|            |         | Normal  | 3      | 51    | -     | 54             | 4      | 107   | 3     | 114 |
|            |         | Upper   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|            |         | Total   | 4      | 54    | -     | 58             | 14     | 109   | 3     | 126 |
|            | Week 36 | Lower   | 2      | 2     | -     | 4              | 6      | 2     | -     | 8   |
|            |         | Normal  | 2      | 51    | -     | 53             | 8      | 103   | 3     | 114 |
|            |         | Upper   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|            |         | Total   | 4      | 54    | -     | 58             | 14     | 107   | 3     | 124 |
|            | Week 40 | Lower   | -      | 1     | -     | 1              | 6      | 1     | -     | 7   |
|            |         | Normal  | 4      | 52    | -     | 56             | 7      | 105   | 2     | 114 |
|            |         | Upper   | -      | 1     | -     | 1              | -      | -     | 1     | 1   |
|            |         | Total   | 4      | 54    | -     | 58             | 13     | 106   | 3     | 122 |
|            | Week 44 | Lower   | 1      | 2     | -     | 3              | 7      | 1     | -     | 8   |
|            |         | Normal  | 3      | 51    | -     | 54             | 5      | 103   | 3     | 111 |
|            |         | Upper   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|            |         | Total   | 4      | 54    | -     | 58             | 12     | 104   | 3     | 119 |
| Week 48    | Lower   | -       | 3      | -     | 3     | 5              | 1      | -     | 6     |     |
|            | Normal  | 4       | 49     | -     | 53    | 7              | 102    | 3     | 112   |     |
|            | Upper   | -       | 1      | -     | 1     | -              | 1      | -     | 1     |     |
|            | Total   | 4       | 53     | -     | 57    | 12             | 104    | 3     | 119   |     |
| Week 52    | Lower   | 1       | 2      | -     | 3     | 5              | 2      | -     | 7     |     |
|            | Normal  | 3       | 49     | -     | 52    | 7              | 100    | 2     | 109   |     |
|            | Upper   | -       | 2      | -     | 2     | -              | 1      | 1     | 2     |     |
|            | Total   | 4       | 53     | -     | 57    | 12             | 103    | 3     | 118   |     |



表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表 (CCT-002 試験) (3/11)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| RBC     | Week 2  | Lower   | 2      | -     | -     | 2              | 8      | 3     | -     | 11  |
|         |         | Normal  | 3      | 58    | -     | 61             | 1      | 122   | 3     | 126 |
|         |         | Upper   | -      | 1     | 1     | 2              | -      | -     | 4     | 4   |
|         |         | Total   | 5      | 59    | 1     | 65             | 9      | 125   | 7     | 141 |
|         | Week 4  | Lower   | 2      | -     | -     | 2              | 8      | 3     | -     | 11  |
|         |         | Normal  | 3      | 56    | -     | 59             | -      | 121   | 3     | 124 |
|         |         | Upper   | -      | 1     | 1     | 2              | -      | -     | 3     | 3   |
|         |         | Total   | 5      | 57    | 1     | 63             | 8      | 124   | 6     | 138 |
|         | Week 8  | Lower   | 2      | -     | -     | 2              | 7      | 2     | -     | 9   |
|         |         | Normal  | 3      | 56    | -     | 59             | 1      | 119   | 3     | 123 |
|         |         | Upper   | -      | -     | 1     | 1              | -      | 2     | 3     | 5   |
|         |         | Total   | 5      | 56    | 1     | 62             | 8      | 123   | 6     | 137 |
|         | Week 12 | Lower   | 1      | -     | -     | 1              | 6      | 2     | -     | 8   |
|         |         | Normal  | 3      | 53    | 1     | 57             | 2      | 120   | 3     | 125 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | 1     | 3     | 4   |
|         |         | Total   | 4      | 54    | 1     | 59             | 8      | 123   | 6     | 137 |
|         | Week 16 | Lower   | 2      | 1     | -     | 3              | 6      | 3     | -     | 9   |
|         |         | Normal  | 2      | 51    | -     | 53             | 2      | 117   | 4     | 123 |
|         |         | Upper   | -      | 2     | 1     | 3              | -      | -     | 2     | 2   |
|         |         | Total   | 4      | 54    | 1     | 59             | 8      | 120   | 6     | 134 |
|         | Week 20 | Lower   | 2      | 1     | -     | 3              | 7      | 2     | -     | 9   |
|         |         | Normal  | 2      | 50    | -     | 52             | 1      | 116   | 3     | 120 |
|         |         | Upper   | -      | 3     | 1     | 4              | -      | 1     | 3     | 4   |
|         |         | Total   | 4      | 54    | 1     | 59             | 8      | 119   | 6     | 133 |
|         | Week 24 | Lower   | 3      | 1     | -     | 4              | 6      | -     | -     | 6   |
|         |         | Normal  | 1      | 50    | -     | 51             | 2      | 116   | 5     | 123 |
|         |         | Upper   | -      | 3     | 1     | 4              | -      | -     | 1     | 1   |
|         |         | Total   | 4      | 54    | 1     | 59             | 8      | 116   | 6     | 130 |
|         | Week 28 | Lower   | 1      | 2     | -     | 3              | 7      | 3     | -     | 10  |
|         |         | Normal  | 3      | 50    | -     | 53             | 1      | 109   | 5     | 115 |
|         |         | Upper   | -      | 2     | 1     | 3              | -      | 1     | 1     | 2   |
|         |         | Total   | 4      | 54    | 1     | 59             | 8      | 113   | 6     | 127 |
|         | Week 32 | Lower   | 2      | 2     | -     | 4              | 7      | 3     | -     | 10  |
|         |         | Normal  | 2      | 49    | -     | 51             | 1      | 109   | 5     | 115 |
|         |         | Upper   | -      | 2     | 1     | 3              | -      | -     | 1     | 1   |
|         |         | Total   | 4      | 53    | 1     | 58             | 8      | 112   | 6     | 126 |
|         | Week 36 | Lower   | 1      | 2     | -     | 3              | 5      | 3     | -     | 8   |
|         |         | Normal  | 3      | 50    | -     | 53             | 3      | 107   | 3     | 113 |
|         |         | Upper   | -      | 1     | 1     | 2              | -      | 1     | 2     | 3   |
|         |         | Total   | 4      | 53    | 1     | 58             | 8      | 111   | 5     | 124 |
|         | Week 40 | Lower   | 3      | 1     | -     | 4              | 5      | 4     | -     | 9   |
|         |         | Normal  | 1      | 50    | 1     | 52             | 3      | 105   | 2     | 110 |
|         |         | Upper   | -      | 2     | -     | 2              | -      | -     | 3     | 3   |
|         |         | Total   | 4      | 53    | 1     | 58             | 8      | 109   | 5     | 122 |
|         | Week 44 | Lower   | 2      | 1     | -     | 3              | 5      | 3     | -     | 8   |
|         |         | Normal  | 2      | 51    | -     | 53             | 2      | 103   | 3     | 108 |
|         |         | Upper   | -      | 1     | 1     | 2              | -      | 1     | 2     | 3   |
|         |         | Total   | 4      | 53    | 1     | 58             | 7      | 107   | 5     | 119 |
| Week 48 | Lower   | 3       | 2      | -     | 5     | 6              | 2      | -     | 8     |     |
|         | Normal  | 1       | 48     | -     | 49    | 1              | 103    | 3     | 107   |     |
|         | Upper   | -       | 2      | 1     | 3     | -              | 2      | 2     | 4     |     |
|         | Total   | 4       | 52     | 1     | 57    | 7              | 107    | 5     | 119   |     |
| Week 52 | Lower   | 3       | 2      | -     | 5     | 4              | 4      | -     | 8     |     |
|         | Normal  | 1       | 47     | -     | 48    | 3              | 101    | 3     | 107   |     |
|         | Upper   | -       | 3      | 1     | 4     | -              | 1      | 2     | 3     |     |
|         | Total   | 4       | 52     | 1     | 57    | 7              | 106    | 5     | 118   |     |

表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表 (CCT-002 試験) (4/11)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| WBC     | Week 2  | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 52    | 3     | 55             | -      | 121   | 2     | 123 |
|         |         | Upper   | -      | 3     | 7     | 10             | -      | 8     | 9     | 17  |
|         |         | Total   | -      | 55    | 10    | 65             | 1      | 129   | 11    | 141 |
|         | Week 4  | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 49    | 4     | 53             | -      | 115   | 1     | 116 |
|         |         | Upper   | -      | 4     | 6     | 10             | -      | 12    | 9     | 21  |
|         |         | Total   | -      | 53    | 10    | 63             | 1      | 127   | 10    | 138 |
|         | Week 8  | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 48    | 7     | 55             | -      | 113   | 3     | 116 |
|         |         | Upper   | -      | 4     | 3     | 7              | -      | 13    | 7     | 20  |
|         |         | Total   | -      | 52    | 10    | 62             | 1      | 126   | 10    | 137 |
|         | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 48    | 4     | 52             | 1      | 111   | 3     | 115 |
|         |         | Upper   | -      | 3     | 4     | 7              | -      | 15    | 7     | 22  |
|         |         | Total   | -      | 51    | 8     | 59             | 1      | 126   | 10    | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 47    | 4     | 51             | 1      | 113   | 4     | 118 |
|         |         | Upper   | -      | 4     | 4     | 8              | -      | 10    | 6     | 16  |
|         |         | Total   | -      | 51    | 8     | 59             | 1      | 123   | 10    | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 45    | 4     | 49             | 1      | 115   | 3     | 119 |
|         |         | Upper   | -      | 6     | 4     | 10             | -      | 7     | 7     | 14  |
|         |         | Total   | -      | 51    | 8     | 59             | 1      | 122   | 10    | 133 |
|         | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 47    | 5     | 52             | 1      | 107   | 4     | 112 |
|         |         | Upper   | -      | 4     | 3     | 7              | -      | 12    | 6     | 18  |
|         |         | Total   | -      | 51    | 8     | 59             | 1      | 119   | 10    | 130 |
|         | Week 28 | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 47    | 4     | 51             | -      | 106   | 3     | 109 |
|         |         | Upper   | -      | 4     | 4     | 8              | -      | 10    | 7     | 17  |
|         |         | Total   | -      | 51    | 8     | 59             | 1      | 116   | 10    | 127 |
|         | Week 32 | Lower   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|         |         | Normal  | -      | 48    | 4     | 52             | 1      | 108   | 5     | 114 |
|         |         | Upper   | -      | 1     | 4     | 5              | -      | 7     | 5     | 12  |
|         |         | Total   | -      | 50    | 8     | 58             | 1      | 115   | 10    | 126 |
|         | Week 36 | Lower   | -      | -     | -     | -              | 1      | 1     | -     | 2   |
|         |         | Normal  | -      | 48    | 4     | 52             | -      | 105   | 2     | 107 |
|         |         | Upper   | -      | 2     | 4     | 6              | -      | 7     | 8     | 15  |
|         |         | Total   | -      | 50    | 8     | 58             | 1      | 113   | 10    | 124 |
|         | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 48    | 6     | 54             | 1      | 102   | 3     | 106 |
|         |         | Upper   | -      | 2     | 2     | 4              | -      | 9     | 7     | 16  |
|         |         | Total   | -      | 50    | 8     | 58             | 1      | 111   | 10    | 122 |
|         | Week 44 | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 47    | 5     | 52             | -      | 102   | 5     | 107 |
|         |         | Upper   | -      | 3     | 3     | 6              | -      | 6     | 5     | 11  |
|         |         | Total   | -      | 50    | 8     | 58             | 1      | 108   | 10    | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | 1              | -      | -     | 1     |     |
|         | Normal  | -       | 46     | 6     | 52    | -              | 103    | 3     | 106   |     |
|         | Upper   | -       | 3      | 2     | 5     | -              | 5      | 7     | 12    |     |
|         | Total   | -       | 49     | 8     | 57    | 1              | 108    | 10    | 119   |     |
| Week 52 | Lower   | -       | 1      | -     | 1     | 1              | 1      | -     | 2     |     |
|         | Normal  | -       | 47     | 6     | 53    | -              | 98     | 4     | 102   |     |
|         | Upper   | -       | 1      | 2     | 3     | -              | 8      | 6     | 14    |     |
|         | Total   | -       | 49     | 8     | 57    | 1              | 107    | 10    | 118   |     |

表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表 (CCT-002 試験) (5/11)

|             |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|             |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|             |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Neutrophils | Week 2  | Lower   | -      | 1     | -     | 1              | 1      | 5     | -     | 6   |
|             |         | Normal  | -      | 62    | 1     | 63             | 3      | 132   | -     | 135 |
|             |         | Upper   | -      | -     | 1     | 1              | -      | -     | -     | -   |
|             |         | Total   | -      | 63    | 2     | 65             | 4      | 137   | -     | 141 |
|             | Week 4  | Lower   | -      | 1     | -     | 1              | 3      | 1     | -     | 4   |
|             |         | Normal  | -      | 59    | 2     | 61             | 1      | 132   | -     | 133 |
|             |         | Upper   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|             |         | Total   | -      | 61    | 2     | 63             | 4      | 134   | -     | 138 |
|             | Week 8  | Lower   | -      | -     | -     | -              | 1      | 1     | -     | 2   |
|             |         | Normal  | -      | 60    | 2     | 62             | 3      | 131   | -     | 134 |
|             |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|             |         | Total   | -      | 60    | 2     | 62             | 4      | 133   | -     | 137 |
|             | Week 12 | Lower   | -      | 1     | -     | 1              | 1      | 1     | -     | 2   |
|             |         | Normal  | -      | 56    | 2     | 58             | 3      | 130   | -     | 133 |
|             |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|             |         | Total   | -      | 57    | 2     | 59             | 4      | 133   | -     | 137 |
|             | Week 16 | Lower   | -      | 2     | -     | 2              | 1      | 2     | -     | 3   |
|             |         | Normal  | -      | 55    | 2     | 57             | 3      | 128   | -     | 131 |
|             |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Total   | -      | 57    | 2     | 59             | 4      | 130   | -     | 134 |
|             | Week 20 | Lower   | -      | 1     | -     | 1              | -      | 3     | -     | 3   |
|             |         | Normal  | -      | 56    | 2     | 58             | 4      | 126   | -     | 130 |
|             |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Total   | -      | 57    | 2     | 59             | 4      | 129   | -     | 133 |
|             | Week 24 | Lower   | -      | -     | -     | -              | 1      | 2     | -     | 3   |
|             |         | Normal  | -      | 57    | 2     | 59             | 3      | 124   | -     | 127 |
|             |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Total   | -      | 57    | 2     | 59             | 4      | 126   | -     | 130 |
|             | Week 28 | Lower   | -      | -     | -     | -              | 1      | 2     | -     | 3   |
|             |         | Normal  | -      | 57    | 2     | 59             | 3      | 120   | -     | 123 |
|             |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|             |         | Total   | -      | 57    | 2     | 59             | 4      | 123   | -     | 127 |
|             | Week 32 | Lower   | -      | -     | -     | -              | 2      | 2     | -     | 4   |
|             |         | Normal  | -      | 56    | 2     | 58             | 2      | 120   | -     | 122 |
|             |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Total   | -      | 56    | 2     | 58             | 4      | 122   | -     | 126 |
|             | Week 36 | Lower   | -      | 1     | -     | 1              | 2      | 1     | -     | 3   |
|             |         | Normal  | -      | 55    | 2     | 57             | 2      | 119   | -     | 121 |
|             |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Total   | -      | 56    | 2     | 58             | 4      | 120   | -     | 124 |
|             | Week 40 | Lower   | -      | 4     | -     | 4              | -      | 3     | -     | 3   |
|             |         | Normal  | -      | 52    | 2     | 54             | 4      | 114   | -     | 118 |
|             |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|             |         | Total   | -      | 56    | 2     | 58             | 4      | 118   | -     | 122 |
|             | Week 44 | Lower   | -      | 2     | -     | 2              | 1      | 5     | -     | 6   |
|             |         | Normal  | -      | 54    | 2     | 56             | 3      | 110   | -     | 113 |
|             |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Total   | -      | 56    | 2     | 58             | 4      | 115   | -     | 119 |
| Week 48     | Lower   | -       | -      | -     | -     | 1              | 1      | -     | 2     |     |
|             | Normal  | -       | 55     | 2     | 57    | 3              | 114    | -     | 117   |     |
|             | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|             | Total   | -       | 55     | 2     | 57    | 4              | 115    | -     | 119   |     |
| Week 52     | Lower   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|             | Normal  | -       | 55     | 2     | 57    | 3              | 112    | -     | 115   |     |
|             | Upper   | -       | -      | -     | -     | -              | 2      | -     | 2     |     |
|             | Total   | -       | 55     | 2     | 57    | 3              | 115    | -     | 118   |     |

表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表 (CCT-002 試験) (6/11)

|             |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|             |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|             |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Lymphocytes | Week 2  | Lower   | 1      | 2     | -     | 3              | -      | 1     | -     | 1   |
|             |         | Normal  | 3      | 59    | -     | 62             | 1      | 131   | 3     | 135 |
|             |         | Upper   | -      | -     | -     | -              | -      | 3     | 2     | 5   |
|             |         | Total   | 4      | 61    | -     | 65             | 1      | 135   | 5     | 141 |
|             | Week 4  | Lower   | 1      | 2     | -     | 3              | -      | 2     | -     | 2   |
|             |         | Normal  | 3      | 57    | -     | 60             | -      | 128   | 3     | 131 |
|             |         | Upper   | -      | -     | -     | -              | -      | 3     | 2     | 5   |
|             |         | Total   | 4      | 59    | -     | 63             | -      | 133   | 5     | 138 |
|             | Week 8  | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|             |         | Normal  | 4      | 58    | -     | 62             | -      | 130   | 3     | 133 |
|             |         | Upper   | -      | -     | -     | -              | -      | 1     | 2     | 3   |
|             |         | Total   | 4      | 58    | -     | 62             | -      | 132   | 5     | 137 |
|             | Week 12 | Lower   | -      | -     | -     | -              | -      | 3     | -     | 3   |
|             |         | Normal  | 4      | 55    | -     | 59             | -      | 126   | 3     | 129 |
|             |         | Upper   | -      | -     | -     | -              | -      | 3     | 2     | 5   |
|             |         | Total   | 4      | 55    | -     | 59             | -      | 132   | 5     | 137 |
|             | Week 16 | Lower   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|             |         | Normal  | 4      | 53    | -     | 57             | -      | 127   | 3     | 130 |
|             |         | Upper   | -      | 1     | -     | 1              | -      | 2     | 2     | 4   |
|             |         | Total   | 4      | 55    | -     | 59             | -      | 129   | 5     | 134 |
|             | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | 4      | 55    | -     | 59             | -      | 126   | 4     | 130 |
|             |         | Upper   | -      | -     | -     | -              | -      | 2     | 1     | 3   |
|             |         | Total   | 4      | 55    | -     | 59             | -      | 128   | 5     | 133 |
|             | Week 24 | Lower   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|             |         | Normal  | 4      | 54    | -     | 58             | -      | 121   | 4     | 125 |
|             |         | Upper   | -      | -     | -     | -              | -      | 2     | 1     | 3   |
|             |         | Total   | 4      | 55    | -     | 59             | -      | 125   | 5     | 130 |
|             | Week 28 | Lower   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|             |         | Normal  | 4      | 54    | -     | 58             | -      | 116   | 4     | 120 |
|             |         | Upper   | -      | 1     | -     | 1              | -      | 4     | 1     | 5   |
|             |         | Total   | 4      | 55    | -     | 59             | -      | 122   | 5     | 127 |
|             | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | 4      | 54    | -     | 58             | -      | 119   | 3     | 122 |
|             |         | Upper   | -      | -     | -     | -              | -      | 2     | 2     | 4   |
|             |         | Total   | 4      | 54    | -     | 58             | -      | 121   | 5     | 126 |
|             | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | 4      | 54    | -     | 58             | -      | 116   | 4     | 120 |
|             |         | Upper   | -      | -     | -     | -              | -      | 3     | 1     | 4   |
|             |         | Total   | 4      | 54    | -     | 58             | -      | 119   | 5     | 124 |
|             | Week 40 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|             |         | Normal  | 4      | 49    | -     | 53             | -      | 112   | 5     | 117 |
|             |         | Upper   | -      | 5     | -     | 5              | -      | 4     | -     | 4   |
|             |         | Total   | 4      | 54    | -     | 58             | -      | 117   | 5     | 122 |
|             | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | 4      | 52    | -     | 56             | -      | 111   | 2     | 113 |
|             |         | Upper   | -      | 2     | -     | 2              | -      | 3     | 3     | 6   |
|             |         | Total   | 4      | 54    | -     | 58             | -      | 114   | 5     | 119 |
| Week 48     | Lower   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|             | Normal  | 4       | 53     | -     | 57    | -              | 111    | 4     | 115   |     |
|             | Upper   | -       | -      | -     | -     | -              | 2      | 1     | 3     |     |
|             | Total   | 4       | 53     | -     | 57    | -              | 114    | 5     | 119   |     |
| Week 52     | Lower   | -       | 1      | -     | 1     | -              | 4      | -     | 4     |     |
|             | Normal  | 4       | 52     | -     | 56    | -              | 109    | 4     | 113   |     |
|             | Upper   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|             | Total   | 4       | 53     | -     | 57    | -              | 114    | 4     | 118   |     |

表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表 (CCT-002 試験) (7/11)

|           |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-----------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|           |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|           |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Monocytes | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|           |         | Normal  | -      | 64    | 1     | 65             | -      | 139   | -     | 139 |
|           |         | Upper   | -      | -     | -     | -              | -      | 1     | 1     | 2   |
|           |         | Total   | -      | 64    | 1     | 65             | -      | 140   | 1     | 141 |
|           | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 62    | 1     | 63             | -      | 135   | -     | 135 |
|           |         | Upper   | -      | -     | -     | -              | -      | 3     | -     | 3   |
|           |         | Total   | -      | 62    | 1     | 63             | -      | 138   | -     | 138 |
|           | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 61    | 1     | 62             | -      | 135   | -     | 135 |
|           |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|           |         | Total   | -      | 61    | 1     | 62             | -      | 137   | -     | 137 |
|           | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 56    | 1     | 57             | -      | 135   | -     | 135 |
|           |         | Upper   | -      | 2     | -     | 2              | -      | 2     | -     | 2   |
|           |         | Total   | -      | 58    | 1     | 59             | -      | 137   | -     | 137 |
|           | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 58    | 1     | 59             | -      | 133   | -     | 133 |
|           |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Total   | -      | 58    | 1     | 59             | -      | 134   | -     | 134 |
|           | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 58    | 1     | 59             | -      | 131   | -     | 131 |
|           |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|           |         | Total   | -      | 58    | 1     | 59             | -      | 133   | -     | 133 |
|           | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 58    | 1     | 59             | -      | 129   | -     | 129 |
|           |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Total   | -      | 58    | 1     | 59             | -      | 130   | -     | 130 |
|           | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 58    | 1     | 59             | -      | 127   | -     | 127 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 58    | 1     | 59             | -      | 127   | -     | 127 |
|           | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 56    | 1     | 57             | -      | 126   | -     | 126 |
|           |         | Upper   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|           |         | Total   | -      | 57    | 1     | 58             | -      | 126   | -     | 126 |
|           | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 57    | 1     | 58             | -      | 123   | -     | 123 |
|           |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Total   | -      | 57    | 1     | 58             | -      | 124   | -     | 124 |
|           | Week 40 | Lower   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|           |         | Normal  | -      | 56    | 1     | 57             | -      | 122   | -     | 122 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 57    | 1     | 58             | -      | 122   | -     | 122 |
|           | Week 44 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Normal  | -      | 55    | 1     | 56             | -      | 115   | -     | 115 |
|           |         | Upper   | -      | 2     | -     | 2              | -      | 3     | -     | 3   |
|           |         | Total   | -      | 57    | 1     | 58             | -      | 119   | -     | 119 |
| Week 48   | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|           | Normal  | -       | 53     | 1     | 54    | -              | 118    | -     | 118   |     |
|           | Upper   | -       | 3      | -     | 3     | -              | 1      | -     | 1     |     |
|           | Total   | -       | 56     | 1     | 57    | -              | 119    | -     | 119   |     |
| Week 52   | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|           | Normal  | -       | 54     | 1     | 55    | -              | 117    | -     | 117   |     |
|           | Upper   | -       | 2      | -     | 2     | -              | 1      | -     | 1     |     |
|           | Total   | -       | 56     | 1     | 57    | -              | 118    | -     | 118   |     |

表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表 (CCT-002 試験) (8/11)

|             |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|             |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|             |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Eosinophils | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|             |         | Normal  | -      | 63    | 1     | 64             | -      | 138   | -     | 138 |
|             |         | Upper   | -      | -     | 1     | 1              | -      | 2     | 1     | 3   |
|             |         | Total   | -      | 63    | 2     | 65             | -      | 140   | 1     | 141 |
|             | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 60    | 2     | 62             | -      | 135   | -     | 135 |
|             |         | Upper   | -      | 1     | -     | 1              | -      | 2     | 1     | 3   |
|             |         | Total   | -      | 61    | 2     | 63             | -      | 137   | 1     | 138 |
|             | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 59    | 2     | 61             | -      | 134   | -     | 134 |
|             |         | Upper   | -      | 1     | -     | 1              | -      | 2     | 1     | 3   |
|             |         | Total   | -      | 60    | 2     | 62             | -      | 136   | 1     | 137 |
|             | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 56    | 2     | 58             | -      | 132   | -     | 132 |
|             |         | Upper   | -      | 1     | -     | 1              | -      | 4     | 1     | 5   |
|             |         | Total   | -      | 57    | 2     | 59             | -      | 136   | 1     | 137 |
|             | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 57    | 2     | 59             | -      | 134   | -     | 134 |
|             |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Total   | -      | 57    | 2     | 59             | -      | 134   | -     | 134 |
|             | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 56    | 2     | 58             | -      | 133   | -     | 133 |
|             |         | Upper   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|             |         | Total   | -      | 57    | 2     | 59             | -      | 133   | -     | 133 |
|             | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 57    | 2     | 59             | -      | 130   | -     | 130 |
|             |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Total   | -      | 57    | 2     | 59             | -      | 130   | -     | 130 |
|             | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 57    | 2     | 59             | -      | 125   | -     | 125 |
|             |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|             |         | Total   | -      | 57    | 2     | 59             | -      | 127   | -     | 127 |
|             | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 56    | 1     | 57             | -      | 125   | -     | 125 |
|             |         | Upper   | -      | -     | 1     | 1              | -      | 1     | -     | 1   |
|             |         | Total   | -      | 56    | 2     | 58             | -      | 126   | -     | 126 |
|             | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 55    | 2     | 57             | -      | 123   | -     | 123 |
|             |         | Upper   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|             |         | Total   | -      | 56    | 2     | 58             | -      | 124   | -     | 124 |
|             | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 56    | 2     | 58             | -      | 121   | -     | 121 |
|             |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|             |         | Total   | -      | 56    | 2     | 58             | -      | 122   | -     | 122 |
|             | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|             |         | Normal  | -      | 55    | 2     | 57             | -      | 116   | -     | 116 |
|             |         | Upper   | -      | 1     | -     | 1              | -      | 3     | -     | 3   |
|             |         | Total   | -      | 56    | 2     | 58             | -      | 119   | -     | 119 |
| Week 48     | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|             | Normal  | -       | 54     | 2     | 56    | -              | 117    | -     | 117   |     |
|             | Upper   | -       | 1      | -     | 1     | -              | 2      | -     | 2     |     |
|             | Total   | -       | 55     | 2     | 57    | -              | 119    | -     | 119   |     |
| Week 52     | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|             | Normal  | -       | 56     | -     | 56    | -              | 117    | -     | 117   |     |
|             | Upper   | -       | -      | 1     | 1     | -              | 1      | -     | 1     |     |
|             | Total   | -       | 56     | 1     | 57    | -              | 118    | -     | 118   |     |

表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表 (CCT-002 試験) (9/11)

|           |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-----------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|           |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|           |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Basophils | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|           |         | Normal  | -      | 65    | -     | 65             | -      | 141   | -     | 141 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 65    | -     | 65             | -      | 141   | -     | 141 |
|           | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 63    | -     | 63             | -      | 136   | -     | 136 |
|           |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|           |         | Total   | -      | 63    | -     | 63             | -      | 138   | -     | 138 |
|           | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 61    | -     | 61             | -      | 137   | -     | 137 |
|           |         | Upper   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|           |         | Total   | -      | 62    | -     | 62             | -      | 137   | -     | 137 |
|           | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 59    | -     | 59             | -      | 137   | -     | 137 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 59    | -     | 59             | -      | 137   | -     | 137 |
|           | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 59    | -     | 59             | -      | 134   | -     | 134 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 59    | -     | 59             | -      | 134   | -     | 134 |
|           | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 59    | -     | 59             | -      | 133   | -     | 133 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 59    | -     | 59             | -      | 133   | -     | 133 |
|           | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 59    | -     | 59             | -      | 130   | -     | 130 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 59    | -     | 59             | -      | 130   | -     | 130 |
|           | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 59    | -     | 59             | -      | 127   | -     | 127 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 59    | -     | 59             | -      | 127   | -     | 127 |
|           | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 58    | -     | 58             | -      | 126   | -     | 126 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 58    | -     | 58             | -      | 126   | -     | 126 |
|           | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 58    | -     | 58             | -      | 123   | -     | 123 |
|           |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Total   | -      | 58    | -     | 58             | -      | 124   | -     | 124 |
|           | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 58    | -     | 58             | -      | 121   | -     | 121 |
|           |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Total   | -      | 58    | -     | 58             | -      | 122   | -     | 122 |
|           | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 58    | -     | 58             | -      | 119   | -     | 119 |
|           |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Total   | -      | 58    | -     | 58             | -      | 119   | -     | 119 |
| Week 48   | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|           | Normal  | -       | 57     | -     | 57    | -              | 117    | -     | 117   |     |
|           | Upper   | -       | -      | -     | -     | -              | 2      | -     | 2     |     |
|           | Total   | -       | 57     | -     | 57    | -              | 119    | -     | 119   |     |
| Week 52   | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|           | Normal  | -       | 57     | -     | 57    | -              | 118    | -     | 118   |     |
|           | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|           | Total   | -       | 57     | -     | 57    | -              | 118    | -     | 118   |     |

表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表(CCT-002 試験)(10/11)

|                |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|----------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|                |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|                |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Platelet Count | Week 2  | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|                |         | Normal  | 2      | 55    | 1     | 58             | -      | 121   | 3     | 124 |
|                |         | Upper   | -      | 4     | 3     | 7              | -      | 5     | 8     | 13  |
|                |         | Total   | 2      | 59    | 4     | 65             | 1      | 126   | 11    | 138 |
|                | Week 4  | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|                |         | Normal  | 2      | 52    | -     | 54             | -      | 121   | 1     | 122 |
|                |         | Upper   | -      | 5     | 4     | 9              | -      | 3     | 10    | 13  |
|                |         | Total   | 2      | 57    | 4     | 63             | 1      | 124   | 11    | 136 |
|                | Week 8  | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|                |         | Normal  | 2      | 54    | 1     | 57             | -      | 116   | 2     | 118 |
|                |         | Upper   | -      | 1     | 3     | 4              | -      | 5     | 9     | 14  |
|                |         | Total   | 2      | 55    | 4     | 61             | 1      | 121   | 11    | 133 |
|                | Week 12 | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|                |         | Normal  | 2      | 50    | -     | 52             | -      | 121   | 3     | 124 |
|                |         | Upper   | -      | 2     | 4     | 6              | -      | 1     | 8     | 9   |
|                |         | Total   | 2      | 52    | 4     | 58             | 1      | 122   | 11    | 134 |
|                | Week 16 | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|                |         | Normal  | 2      | 49    | -     | 51             | -      | 116   | 2     | 118 |
|                |         | Upper   | -      | 2     | 4     | 6              | -      | 3     | 9     | 12  |
|                |         | Total   | 2      | 51    | 4     | 57             | 1      | 119   | 11    | 131 |
|                | Week 20 | Lower   | 1      | -     | -     | 1              | 1      | -     | -     | 1   |
|                |         | Normal  | 1      | 52    | -     | 53             | -      | 112   | 5     | 117 |
|                |         | Upper   | -      | 1     | 4     | 5              | -      | 5     | 6     | 11  |
|                |         | Total   | 2      | 53    | 4     | 59             | 1      | 117   | 11    | 129 |
|                | Week 24 | Lower   | 2      | -     | -     | 2              | 1      | 1     | -     | 2   |
|                |         | Normal  | -      | 53    | 1     | 54             | -      | 109   | 4     | 113 |
|                |         | Upper   | -      | -     | 3     | 3              | -      | 3     | 6     | 9   |
|                |         | Total   | 2      | 53    | 4     | 59             | 1      | 113   | 10    | 124 |
|                | Week 28 | Lower   | 2      | -     | -     | 2              | 1      | -     | -     | 1   |
|                |         | Normal  | -      | 52    | 2     | 54             | -      | 109   | 3     | 112 |
|                |         | Upper   | -      | 1     | 2     | 3              | -      | 3     | 8     | 11  |
|                |         | Total   | 2      | 53    | 4     | 59             | 1      | 112   | 11    | 124 |
|                | Week 32 | Lower   | 1      | 1     | -     | 2              | 1      | -     | -     | 1   |
|                |         | Normal  | 1      | 50    | 2     | 53             | -      | 111   | 6     | 117 |
|                |         | Upper   | -      | 1     | 2     | 3              | -      | 1     | 5     | 6   |
|                |         | Total   | 2      | 52    | 4     | 58             | 1      | 112   | 11    | 124 |
|                | Week 36 | Lower   | 2      | -     | -     | 2              | 1      | -     | -     | 1   |
|                |         | Normal  | -      | 49    | 3     | 52             | -      | 110   | 7     | 117 |
|                |         | Upper   | -      | 2     | 1     | 3              | -      | 1     | 3     | 4   |
|                |         | Total   | 2      | 51    | 4     | 57             | 1      | 111   | 10    | 122 |
|                | Week 40 | Lower   | 2      | 1     | -     | 3              | 1      | 1     | -     | 2   |
|                |         | Normal  | -      | 49    | 3     | 52             | -      | 108   | 5     | 113 |
|                |         | Upper   | -      | 1     | 1     | 2              | -      | -     | 5     | 5   |
|                |         | Total   | 2      | 51    | 4     | 57             | 1      | 109   | 10    | 120 |
|                | Week 44 | Lower   | 2      | 1     | -     | 3              | 1      | -     | -     | 1   |
|                |         | Normal  | -      | 50    | 1     | 51             | -      | 105   | 4     | 109 |
|                |         | Upper   | -      | 1     | 3     | 4              | -      | 2     | 5     | 7   |
|                |         | Total   | 2      | 52    | 4     | 58             | 1      | 107   | 9     | 117 |
| Week 48        | Lower   | 2       | -      | -     | 2     | -              | -      | -     | -     |     |
|                | Normal  | -       | 52     | 2     | 54    | 1              | 103    | 6     | 110   |     |
|                | Upper   | -       | -      | 1     | 1     | -              | 4      | 3     | 7     |     |
|                | Total   | 2       | 52     | 3     | 57    | 1              | 107    | 9     | 117   |     |
| Week 52        | Lower   | 2       | 1      | -     | 3     | 1              | 1      | -     | 2     |     |
|                | Normal  | -       | 49     | 2     | 51    | -              | 101    | 5     | 106   |     |
|                | Upper   | -       | 2      | 1     | 3     | -              | 3      | 4     | 7     |     |
|                | Total   | 2       | 52     | 3     | 57    | 1              | 105    | 9     | 115   |     |



表 2.7.4.7-付録-19 血液学的検査の評価時点別正異判定クロス表(CCT-002 試験)(11/11)

|                  |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|------------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|                  |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|                  |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Prothrombin Time | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|                  |         | Normal  | -      | 59    | 1     | 60             | -      | 133   | 3     | 136 |
|                  |         | Upper   | -      | 2     | 3     | 5              | -      | 2     | 3     | 5   |
|                  |         | Total   | -      | 61    | 4     | 65             | -      | 135   | 6     | 141 |
|                  | Week 4  | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|                  |         | Normal  | -      | 57    | 2     | 59             | -      | 127   | 5     | 132 |
|                  |         | Upper   | -      | 2     | 2     | 4              | -      | 4     | 1     | 5   |
|                  |         | Total   | -      | 59    | 4     | 63             | -      | 132   | 6     | 138 |
|                  | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 57    | 1     | 58             | -      | 124   | 2     | 126 |
|                  |         | Upper   | -      | 1     | 3     | 4              | -      | 7     | 4     | 11  |
|                  |         | Total   | -      | 58    | 4     | 62             | -      | 131   | 6     | 137 |
|                  | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 54    | 1     | 55             | -      | 117   | 4     | 121 |
|                  |         | Upper   | -      | 2     | 2     | 4              | -      | 14    | 2     | 16  |
|                  |         | Total   | -      | 56    | 3     | 59             | -      | 131   | 6     | 137 |
|                  | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 54    | 1     | 55             | -      | 113   | 4     | 117 |
|                  |         | Upper   | -      | 2     | 2     | 4              | -      | 15    | 2     | 17  |
|                  |         | Total   | -      | 56    | 3     | 59             | -      | 128   | 6     | 134 |
|                  | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 51    | 1     | 52             | -      | 113   | 1     | 114 |
|                  |         | Upper   | -      | 5     | 2     | 7              | -      | 14    | 5     | 19  |
|                  |         | Total   | -      | 56    | 3     | 59             | -      | 127   | 6     | 133 |
|                  | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 51    | 2     | 53             | -      | 111   | 3     | 114 |
|                  |         | Upper   | -      | 5     | 1     | 6              | -      | 13    | 3     | 16  |
|                  |         | Total   | -      | 56    | 3     | 59             | -      | 124   | 6     | 130 |
|                  | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 52    | 1     | 53             | -      | 108   | 2     | 110 |
|                  |         | Upper   | -      | 4     | 2     | 6              | -      | 13    | 4     | 17  |
|                  |         | Total   | -      | 56    | 3     | 59             | -      | 121   | 6     | 127 |
|                  | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 50    | 1     | 51             | -      | 110   | 3     | 113 |
|                  |         | Upper   | -      | 5     | 2     | 7              | -      | 11    | 3     | 14  |
|                  |         | Total   | -      | 55    | 3     | 58             | -      | 121   | 6     | 127 |
|                  | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 51    | 1     | 52             | -      | 109   | 3     | 112 |
|                  |         | Upper   | -      | 3     | 2     | 5              | -      | 9     | 3     | 12  |
|                  |         | Total   | -      | 54    | 3     | 57             | -      | 118   | 6     | 124 |
|                  | Week 40 | Lower   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 51    | 2     | 53             | -      | 113   | 5     | 118 |
|                  |         | Upper   | -      | 3     | 1     | 4              | -      | 3     | 1     | 4   |
|                  |         | Total   | -      | 55    | 3     | 58             | -      | 116   | 6     | 122 |
|                  | Week 44 | Lower   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 52    | 2     | 54             | -      | 108   | 4     | 112 |
|                  |         | Upper   | -      | 2     | 1     | 3              | -      | 5     | 2     | 7   |
|                  |         | Total   | -      | 55    | 3     | 58             | -      | 113   | 6     | 119 |
| Week 48          | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|                  | Normal  | -       | 53     | 2     | 55    | -              | 106    | 5     | 111   |     |
|                  | Upper   | -       | 2      | 1     | 3     | -              | 7      | 1     | 8     |     |
|                  | Total   | -       | 55     | 3     | 58    | -              | 113    | 6     | 119   |     |
| Week 52          | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|                  | Normal  | -       | 52     | 1     | 53    | -              | 105    | 2     | 107   |     |
|                  | Upper   | -       | 3      | 2     | 5     | -              | 7      | 4     | 11    |     |
|                  | Total   | -       | 55     | 3     | 58    | -              | 112    | 6     | 118   |     |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (1/12)

| Total Protein (g/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|----------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit                | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2               | Placebo        | 65                 | -0.05 | 0.289 | -0.7 | -0.20 | -0.10  | 0.20 | 0.7 |
|                      | ATL-962 120 mg | 141                | -0.05 | 0.334 | -1.3 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 4               | Placebo        | 63                 | 0.01  | 0.329 | -0.8 | -0.20 | 0.00   | 0.20 | 0.9 |
|                      | ATL-962 120 mg | 138                | -0.01 | 0.336 | -0.8 | -0.30 | 0.00   | 0.20 | 1.0 |
| Week 8               | Placebo        | 62                 | -0.05 | 0.320 | -0.8 | -0.20 | -0.10  | 0.10 | 0.9 |
|                      | ATL-962 120 mg | 137                | -0.08 | 0.335 | -1.3 | -0.20 | -0.10  | 0.10 | 1.0 |
| Week 12              | Placebo        | 59                 | -0.07 | 0.388 | -1.1 | -0.30 | -0.10  | 0.10 | 0.9 |
|                      | ATL-962 120 mg | 137                | -0.07 | 0.317 | -0.9 | -0.30 | -0.10  | 0.10 | 0.7 |
| Week 16              | Placebo        | 59                 | -0.04 | 0.342 | -0.9 | -0.30 | 0.00   | 0.20 | 0.6 |
|                      | ATL-962 120 mg | 134                | -0.09 | 0.347 | -1.5 | -0.30 | -0.10  | 0.20 | 0.7 |
| Week 20              | Placebo        | 59                 | -0.06 | 0.375 | -0.8 | -0.30 | -0.10  | 0.10 | 1.1 |
|                      | ATL-962 120 mg | 133                | -0.10 | 0.328 | -1.0 | -0.30 | -0.10  | 0.10 | 1.0 |
| Week 24              | Placebo        | 59                 | -0.12 | 0.339 | -1.0 | -0.30 | -0.10  | 0.10 | 0.8 |
|                      | ATL-962 120 mg | 130                | -0.14 | 0.337 | -1.3 | -0.40 | -0.20  | 0.10 | 0.9 |
| Week 28              | Placebo        | 59                 | -0.07 | 0.356 | -1.1 | -0.20 | -0.10  | 0.10 | 0.7 |
|                      | ATL-962 120 mg | 127                | -0.12 | 0.316 | -1.3 | -0.30 | -0.10  | 0.10 | 0.8 |
| Week 32              | Placebo        | 58                 | -0.05 | 0.362 | -0.9 | -0.20 | -0.10  | 0.20 | 0.8 |
|                      | ATL-962 120 mg | 127                | -0.08 | 0.356 | -1.4 | -0.30 | -0.10  | 0.10 | 0.9 |
| Week 36              | Placebo        | 58                 | -0.14 | 0.321 | -0.9 | -0.30 | -0.20  | 0.10 | 0.6 |
|                      | ATL-962 120 mg | 124                | -0.09 | 0.312 | -1.0 | -0.30 | -0.10  | 0.10 | 0.9 |
| Week 40              | Placebo        | 58                 | -0.09 | 0.288 | -0.8 | -0.30 | -0.10  | 0.10 | 0.6 |
|                      | ATL-962 120 mg | 122                | -0.07 | 0.418 | -1.4 | -0.30 | -0.10  | 0.20 | 1.3 |
| Week 44              | Placebo        | 58                 | -0.03 | 0.295 | -0.7 | -0.20 | 0.00   | 0.10 | 0.6 |
|                      | ATL-962 120 mg | 119                | -0.08 | 0.362 | -1.2 | -0.30 | -0.10  | 0.20 | 0.8 |
| Week 48              | Placebo        | 58                 | -0.06 | 0.338 | -1.0 | -0.30 | -0.10  | 0.20 | 1.0 |
|                      | ATL-962 120 mg | 119                | -0.11 | 0.375 | -1.1 | -0.30 | -0.10  | 0.20 | 1.0 |
| Week 52              | Placebo        | 58                 | -0.09 | 0.337 | -1.2 | -0.30 | -0.10  | 0.20 | 0.5 |
|                      | ATL-962 120 mg | 118                | -0.10 | 0.405 | -1.0 | -0.40 | -0.10  | 0.20 | 1.3 |

| Albumin (g/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|----------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit          | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2         | Placebo        | 65                 | 0.01  | 0.178 | -0.4 | -0.10 | 0.00   | 0.10 | 0.5 |
|                | ATL-962 120 mg | 141                | -0.02 | 0.204 | -0.7 | -0.10 | 0.00   | 0.10 | 0.4 |
| Week 4         | Placebo        | 63                 | 0.02  | 0.218 | -0.4 | -0.10 | 0.00   | 0.10 | 0.7 |
|                | ATL-962 120 mg | 138                | -0.01 | 0.210 | -0.5 | -0.20 | 0.00   | 0.10 | 0.7 |
| Week 8         | Placebo        | 62                 | -0.02 | 0.201 | -0.4 | -0.20 | 0.00   | 0.10 | 0.5 |
|                | ATL-962 120 mg | 137                | -0.04 | 0.214 | -0.6 | -0.20 | 0.00   | 0.10 | 0.5 |
| Week 12        | Placebo        | 59                 | -0.05 | 0.258 | -0.7 | -0.20 | 0.00   | 0.10 | 0.5 |
|                | ATL-962 120 mg | 137                | -0.08 | 0.208 | -0.6 | -0.20 | -0.10  | 0.10 | 0.6 |
| Week 16        | Placebo        | 59                 | -0.05 | 0.218 | -0.6 | -0.20 | -0.10  | 0.10 | 0.4 |
|                | ATL-962 120 mg | 134                | -0.10 | 0.223 | -0.8 | -0.20 | -0.10  | 0.10 | 0.5 |
| Week 20        | Placebo        | 59                 | -0.04 | 0.226 | -0.5 | -0.20 | -0.10  | 0.10 | 0.6 |
|                | ATL-962 120 mg | 133                | -0.08 | 0.206 | -0.5 | -0.20 | -0.10  | 0.10 | 0.6 |
| Week 24        | Placebo        | 59                 | -0.09 | 0.223 | -0.6 | -0.30 | -0.10  | 0.00 | 0.6 |
|                | ATL-962 120 mg | 130                | -0.10 | 0.216 | -0.7 | -0.30 | -0.10  | 0.10 | 0.5 |
| Week 28        | Placebo        | 59                 | -0.04 | 0.240 | -0.6 | -0.20 | 0.00   | 0.10 | 0.5 |
|                | ATL-962 120 mg | 127                | -0.09 | 0.208 | -0.7 | -0.20 | -0.10  | 0.10 | 0.4 |
| Week 32        | Placebo        | 58                 | -0.03 | 0.228 | -0.5 | -0.20 | -0.10  | 0.10 | 0.5 |
|                | ATL-962 120 mg | 127                | -0.05 | 0.215 | -0.6 | -0.20 | -0.10  | 0.10 | 0.5 |
| Week 36        | Placebo        | 58                 | -0.07 | 0.213 | -0.7 | -0.20 | -0.05  | 0.00 | 0.5 |
|                | ATL-962 120 mg | 124                | -0.06 | 0.207 | -0.7 | -0.20 | -0.10  | 0.10 | 0.4 |
| Week 40        | Placebo        | 58                 | -0.03 | 0.228 | -0.6 | -0.20 | 0.00   | 0.20 | 0.4 |
|                | ATL-962 120 mg | 122                | -0.03 | 0.271 | -0.7 | -0.20 | 0.00   | 0.10 | 0.8 |
| Week 44        | Placebo        | 58                 | 0.02  | 0.201 | -0.4 | -0.10 | 0.00   | 0.20 | 0.6 |
|                | ATL-962 120 mg | 119                | -0.04 | 0.234 | -0.8 | -0.20 | 0.00   | 0.10 | 0.5 |
| Week 48        | Placebo        | 58                 | 0.02  | 0.214 | -0.4 | -0.10 | 0.00   | 0.20 | 0.6 |
|                | ATL-962 120 mg | 119                | -0.03 | 0.235 | -0.6 | -0.20 | 0.00   | 0.10 | 0.6 |
| Week 52        | Placebo        | 58                 | -0.01 | 0.244 | -0.7 | -0.20 | 0.00   | 0.20 | 0.5 |
|                | ATL-962 120 mg | 118                | -0.04 | 0.255 | -0.6 | -0.20 | -0.10  | 0.10 | 0.8 |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (2/12)

| ALT (GPT) (IU/L) |                | Summary Statistics |      |       |     |       |        |     |     |
|------------------|----------------|--------------------|------|-------|-----|-------|--------|-----|-----|
| Visit            | Treatment      | N                  | Mean | SD    | Min | Q1    | Median | Q3  | Max |
| Week 2           | Placebo        | 65                 | -1.1 | 10.68 | -38 | -5.0  | 0.0    | 3.0 | 28  |
|                  | ATL-962 120 mg | 141                | 3.0  | 13.94 | -28 | -4.0  | 3.0    | 9.0 | 80  |
| Week 4           | Placebo        | 63                 | -2.2 | 9.40  | -26 | -6.0  | -1.0   | 3.0 | 23  |
|                  | ATL-962 120 mg | 138                | 2.5  | 13.59 | -39 | -4.0  | 1.0    | 8.0 | 57  |
| Week 8           | Placebo        | 62                 | -2.9 | 10.65 | -32 | -6.0  | -1.0   | 3.0 | 20  |
|                  | ATL-962 120 mg | 137                | 0.1  | 15.19 | -40 | -9.0  | -1.0   | 7.0 | 58  |
| Week 12          | Placebo        | 59                 | -1.9 | 12.39 | -39 | -8.0  | -1.0   | 4.0 | 21  |
|                  | ATL-962 120 mg | 137                | -1.0 | 15.73 | -46 | -9.0  | -2.0   | 8.0 | 48  |
| Week 16          | Placebo        | 59                 | -2.9 | 12.12 | -38 | -9.0  | -3.0   | 3.0 | 24  |
|                  | ATL-962 120 mg | 134                | -1.2 | 17.95 | -51 | -9.0  | -0.5   | 7.0 | 71  |
| Week 20          | Placebo        | 59                 | -1.7 | 14.62 | -46 | -8.0  | -2.0   | 8.0 | 43  |
|                  | ATL-962 120 mg | 133                | -2.1 | 18.84 | -50 | -12.0 | -2.0   | 8.0 | 97  |
| Week 24          | Placebo        | 59                 | -2.9 | 15.03 | -39 | -9.0  | -2.0   | 5.0 | 34  |
|                  | ATL-962 120 mg | 130                | -0.4 | 27.86 | -47 | -13.0 | -1.0   | 9.0 | 226 |
| Week 28          | Placebo        | 59                 | -1.1 | 15.99 | -35 | -8.0  | 1.0    | 7.0 | 67  |
|                  | ATL-962 120 mg | 127                | -3.4 | 17.84 | -58 | -14.0 | -2.0   | 7.0 | 45  |
| Week 32          | Placebo        | 58                 | -2.1 | 14.37 | -46 | -8.0  | -1.5   | 6.0 | 37  |
|                  | ATL-962 120 mg | 127                | -3.4 | 18.11 | -45 | -13.0 | -2.0   | 7.0 | 56  |
| Week 36          | Placebo        | 58                 | -2.1 | 16.56 | -56 | -8.0  | -1.0   | 4.0 | 51  |
|                  | ATL-962 120 mg | 124                | -0.7 | 33.56 | -47 | -12.0 | -2.0   | 8.5 | 315 |
| Week 40          | Placebo        | 58                 | -1.9 | 15.47 | -55 | -8.0  | -1.0   | 7.0 | 39  |
|                  | ATL-962 120 mg | 122                | -3.0 | 17.56 | -45 | -12.0 | -1.0   | 7.0 | 55  |
| Week 44          | Placebo        | 58                 | 0.1  | 14.00 | -49 | -6.0  | 0.0    | 8.0 | 31  |
|                  | ATL-962 120 mg | 119                | -3.5 | 17.57 | -54 | -13.0 | -2.0   | 7.0 | 39  |
| Week 48          | Placebo        | 58                 | -1.9 | 14.29 | -57 | -8.0  | -1.0   | 5.0 | 40  |
|                  | ATL-962 120 mg | 119                | -3.5 | 18.32 | -57 | -14.0 | -3.0   | 8.0 | 52  |
| Week 52          | Placebo        | 58                 | -1.7 | 15.52 | -51 | -9.0  | -2.0   | 7.0 | 42  |
|                  | ATL-962 120 mg | 118                | -3.8 | 20.44 | -58 | -17.0 | -2.0   | 7.0 | 63  |

| AST (GOT) (IU/L) |                | Summary Statistics |      |       |     |      |        |     |     |
|------------------|----------------|--------------------|------|-------|-----|------|--------|-----|-----|
| Visit            | Treatment      | N                  | Mean | SD    | Min | Q1   | Median | Q3  | Max |
| Week 2           | Placebo        | 65                 | -1.3 | 8.62  | -31 | -3.0 | -1.0   | 3.0 | 17  |
|                  | ATL-962 120 mg | 141                | 0.5  | 8.18  | -24 | -3.0 | 0.0    | 4.0 | 25  |
| Week 4           | Placebo        | 63                 | -0.7 | 8.29  | -26 | -3.0 | 0.0    | 3.0 | 39  |
|                  | ATL-962 120 mg | 138                | 0.1  | 8.10  | -31 | -3.0 | 0.0    | 4.0 | 33  |
| Week 8           | Placebo        | 62                 | -1.5 | 9.20  | -46 | -3.0 | 0.0    | 3.0 | 16  |
|                  | ATL-962 120 mg | 137                | -0.8 | 8.74  | -25 | -5.0 | -1.0   | 4.0 | 29  |
| Week 12          | Placebo        | 59                 | -0.4 | 8.33  | -29 | -4.0 | 0.0    | 5.0 | 15  |
|                  | ATL-962 120 mg | 137                | -1.0 | 11.06 | -23 | -6.0 | -1.0   | 3.0 | 83  |
| Week 16          | Placebo        | 59                 | -1.3 | 9.10  | -30 | -5.0 | -1.0   | 2.0 | 21  |
|                  | ATL-962 120 mg | 134                | -2.3 | 9.66  | -31 | -6.0 | -1.0   | 3.0 | 39  |
| Week 20          | Placebo        | 59                 | 0.2  | 9.79  | -26 | -4.0 | 0.0    | 4.0 | 27  |
|                  | ATL-962 120 mg | 133                | -2.8 | 9.22  | -30 | -6.0 | -2.0   | 3.0 | 29  |
| Week 24          | Placebo        | 59                 | -0.7 | 11.56 | -25 | -6.0 | 0.0    | 3.0 | 47  |
|                  | ATL-962 120 mg | 130                | -1.5 | 12.35 | -33 | -7.0 | -1.0   | 3.0 | 78  |
| Week 28          | Placebo        | 59                 | 0.3  | 10.72 | -26 | -4.0 | 1.0    | 5.0 | 52  |
|                  | ATL-962 120 mg | 127                | -2.7 | 10.39 | -34 | -7.0 | -1.0   | 3.0 | 27  |
| Week 32          | Placebo        | 58                 | -0.1 | 11.35 | -28 | -5.0 | -1.0   | 3.0 | 51  |
|                  | ATL-962 120 mg | 127                | -2.8 | 10.60 | -32 | -8.0 | -2.0   | 3.0 | 41  |
| Week 36          | Placebo        | 58                 | -0.6 | 10.63 | -41 | -5.0 | -0.5   | 3.0 | 38  |
|                  | ATL-962 120 mg | 124                | -0.0 | 22.73 | -32 | -7.0 | -1.0   | 4.0 | 224 |
| Week 40          | Placebo        | 58                 | -0.5 | 7.91  | -28 | -5.0 | -1.0   | 4.0 | 22  |
|                  | ATL-962 120 mg | 122                | -2.1 | 11.31 | -32 | -6.0 | -2.0   | 3.0 | 64  |
| Week 44          | Placebo        | 58                 | 0.8  | 8.29  | -23 | -4.0 | 0.0    | 5.0 | 16  |
|                  | ATL-962 120 mg | 119                | -1.5 | 13.88 | -31 | -7.0 | -1.0   | 4.0 | 106 |
| Week 48          | Placebo        | 58                 | 0.2  | 8.87  | -22 | -4.0 | -1.0   | 4.0 | 29  |
|                  | ATL-962 120 mg | 119                | -2.0 | 11.58 | -32 | -9.0 | -2.0   | 4.0 | 44  |
| Week 52          | Placebo        | 58                 | -0.3 | 9.87  | -29 | -4.0 | -1.0   | 5.0 | 38  |
|                  | ATL-962 120 mg | 118                | -2.2 | 11.43 | -32 | -7.0 | -3.0   | 3.0 | 57  |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (3/12)

| γ-GTP (IU/L) |                | Summary Statistics |       |       |      |       |        |     |     |
|--------------|----------------|--------------------|-------|-------|------|-------|--------|-----|-----|
| Visit        | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3  | Max |
| Week 2       | Placebo        | 65                 | -4.7  | 20.61 | -141 | -6.0  | -2.0   | 2.0 | 26  |
|              | ATL-962 120 mg | 141                | -1.4  | 21.83 | -111 | -6.0  | -2.0   | 2.0 | 164 |
| Week 4       | Placebo        | 63                 | -5.4  | 26.53 | -180 | -4.0  | -2.0   | 3.0 | 24  |
|              | ATL-962 120 mg | 138                | -3.5  | 18.15 | -80  | -9.0  | -2.0   | 2.0 | 74  |
| Week 8       | Placebo        | 62                 | -1.1  | 31.85 | -72  | -8.0  | -2.5   | 1.0 | 205 |
|              | ATL-962 120 mg | 137                | -3.4  | 20.97 | -84  | -11.0 | -4.0   | 3.0 | 131 |
| Week 12      | Placebo        | 59                 | -4.3  | 33.17 | -193 | -8.0  | -2.0   | 2.0 | 96  |
|              | ATL-962 120 mg | 137                | -3.1  | 20.41 | -98  | -12.0 | -3.0   | 5.0 | 82  |
| Week 16      | Placebo        | 59                 | -5.5  | 35.63 | -228 | -7.0  | -2.0   | 3.0 | 112 |
|              | ATL-962 120 mg | 134                | -4.7  | 19.89 | -110 | -11.0 | -4.0   | 4.0 | 46  |
| Week 20      | Placebo        | 59                 | -2.0  | 34.72 | -152 | -9.0  | -4.0   | 5.0 | 132 |
|              | ATL-962 120 mg | 133                | -5.4  | 24.79 | -120 | -15.0 | -3.0   | 3.0 | 104 |
| Week 24      | Placebo        | 59                 | -1.4  | 36.50 | -142 | -10.0 | -3.0   | 5.0 | 157 |
|              | ATL-962 120 mg | 130                | -4.8  | 25.63 | -114 | -13.0 | -3.5   | 5.0 | 109 |
| Week 28      | Placebo        | 59                 | 1.1   | 33.79 | -121 | -9.0  | -1.0   | 6.0 | 126 |
|              | ATL-962 120 mg | 127                | -6.1  | 32.82 | -165 | -14.0 | -4.0   | 5.0 | 202 |
| Week 32      | Placebo        | 58                 | -0.2  | 25.74 | -49  | -12.0 | -3.0   | 2.0 | 144 |
|              | ATL-962 120 mg | 127                | -6.6  | 26.52 | -131 | -15.0 | -4.0   | 4.0 | 83  |
| Week 36      | Placebo        | 58                 | -2.7  | 16.73 | -55  | -11.0 | -4.5   | 3.0 | 61  |
|              | ATL-962 120 mg | 124                | -4.3  | 28.09 | -128 | -17.0 | -4.0   | 4.0 | 126 |
| Week 40      | Placebo        | 58                 | -2.8  | 21.13 | -65  | -10.0 | -3.0   | 3.0 | 74  |
|              | ATL-962 120 mg | 122                | -7.1  | 26.72 | -119 | -15.0 | -4.0   | 3.0 | 73  |
| Week 44      | Placebo        | 58                 | -0.8  | 21.21 | -68  | -8.0  | -1.5   | 6.0 | 97  |
|              | ATL-962 120 mg | 119                | -9.8  | 31.31 | -137 | -17.0 | -5.0   | 2.0 | 121 |
| Week 48      | Placebo        | 58                 | -2.0  | 21.66 | -71  | -11.0 | -2.0   | 5.0 | 86  |
|              | ATL-962 120 mg | 119                | -10.2 | 31.04 | -130 | -18.0 | -7.0   | 2.0 | 103 |
| Week 52      | Placebo        | 58                 | 0.3   | 31.68 | -61  | -11.0 | -1.0   | 5.0 | 181 |
|              | ATL-962 120 mg | 118                | -9.1  | 28.82 | -156 | -17.0 | -6.5   | 2.0 | 82  |

| ALP (IU/L) |                | Summary Statistics |       |       |      |       |        |      |     |
|------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit      | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2     | Placebo        | 65                 | -8.7  | 37.54 | -137 | -22.0 | -8.0   | 6.0  | 143 |
|            | ATL-962 120 mg | 141                | -0.1  | 27.77 | -103 | -12.0 | 1.0    | 15.0 | 81  |
| Week 4     | Placebo        | 63                 | -10.9 | 36.36 | -144 | -26.0 | -7.0   | 12.0 | 58  |
|            | ATL-962 120 mg | 138                | 0.8   | 26.76 | -126 | -13.0 | 1.5    | 19.0 | 55  |
| Week 8     | Placebo        | 62                 | -15.1 | 28.88 | -118 | -32.0 | -14.5  | 3.0  | 61  |
|            | ATL-962 120 mg | 137                | -4.4  | 31.18 | -132 | -21.0 | -3.0   | 13.0 | 72  |
| Week 12    | Placebo        | 59                 | -19.5 | 33.67 | -138 | -33.0 | -16.0  | -2.0 | 67  |
|            | ATL-962 120 mg | 137                | -2.8  | 33.58 | -126 | -21.0 | -1.0   | 15.0 | 81  |
| Week 16    | Placebo        | 59                 | -14.5 | 34.98 | -138 | -35.0 | -16.0  | 11.0 | 66  |
|            | ATL-962 120 mg | 134                | -5.0  | 34.73 | -148 | -20.0 | -3.5   | 16.0 | 84  |
| Week 20    | Placebo        | 59                 | -18.7 | 33.27 | -134 | -29.0 | -17.0  | -3.0 | 81  |
|            | ATL-962 120 mg | 133                | -4.5  | 37.03 | -171 | -25.0 | -2.0   | 16.0 | 101 |
| Week 24    | Placebo        | 59                 | -16.1 | 44.84 | -135 | -36.0 | -17.0  | 3.0  | 104 |
|            | ATL-962 120 mg | 130                | -2.6  | 39.61 | -113 | -23.0 | -2.5   | 19.0 | 112 |
| Week 28    | Placebo        | 59                 | -17.2 | 39.37 | -142 | -36.0 | -18.0  | 7.0  | 97  |
|            | ATL-962 120 mg | 127                | -5.1  | 37.99 | -171 | -23.0 | 0.0    | 16.0 | 72  |
| Week 32    | Placebo        | 58                 | -19.8 | 42.62 | -154 | -37.0 | -19.5  | 1.0  | 102 |
|            | ATL-962 120 mg | 127                | -1.4  | 42.98 | -184 | -23.0 | 1.0    | 22.0 | 157 |
| Week 36    | Placebo        | 58                 | -18.8 | 55.71 | -158 | -39.0 | -18.5  | 3.0  | 209 |
|            | ATL-962 120 mg | 124                | 0.3   | 37.15 | -152 | -18.0 | 4.0    | 19.0 | 105 |
| Week 40    | Placebo        | 58                 | -19.6 | 46.54 | -155 | -40.0 | -14.0  | 4.0  | 65  |
|            | ATL-962 120 mg | 122                | -1.1  | 43.44 | -145 | -24.0 | 6.0    | 20.0 | 120 |
| Week 44    | Placebo        | 58                 | -12.5 | 59.62 | -159 | -32.0 | -10.5  | 13.0 | 283 |
|            | ATL-962 120 mg | 119                | -1.1  | 40.18 | -133 | -21.0 | 4.0    | 19.0 | 136 |
| Week 48    | Placebo        | 58                 | -12.8 | 46.69 | -139 | -36.0 | -13.5  | 9.0  | 116 |
|            | ATL-962 120 mg | 119                | 1.3   | 39.95 | -129 | -24.0 | 2.0    | 28.0 | 159 |
| Week 52    | Placebo        | 58                 | -12.2 | 39.60 | -137 | -30.0 | -11.5  | 7.0  | 131 |
|            | ATL-962 120 mg | 118                | 4.3   | 43.84 | -118 | -23.0 | 7.5    | 29.0 | 153 |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (4/12)

| LDH (IU/L) |                | Summary Statistics |      |       |      |       |        |      |     |
|------------|----------------|--------------------|------|-------|------|-------|--------|------|-----|
| Visit      | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2     | Placebo        | 65                 | -2.9 | 17.74 | -48  | -15.0 | -3.0   | 11.0 | 42  |
|            | ATL-962 120 mg | 141                | -4.6 | 18.52 | -83  | -14.0 | -4.0   | 4.0  | 74  |
| Week 4     | Placebo        | 63                 | 2.2  | 20.74 | -45  | -13.0 | 2.0    | 12.0 | 91  |
|            | ATL-962 120 mg | 138                | -1.6 | 17.02 | -92  | -12.0 | -2.0   | 9.0  | 38  |
| Week 8     | Placebo        | 62                 | 1.2  | 18.26 | -63  | -9.0  | -1.0   | 15.0 | 45  |
|            | ATL-962 120 mg | 137                | -2.2 | 18.84 | -108 | -13.0 | -2.0   | 10.0 | 44  |
| Week 12    | Placebo        | 59                 | 2.7  | 21.38 | -38  | -10.0 | 0.0    | 16.0 | 89  |
|            | ATL-962 120 mg | 137                | -2.8 | 19.78 | -108 | -16.0 | -2.0   | 9.0  | 44  |
| Week 16    | Placebo        | 59                 | -0.7 | 15.37 | -32  | -13.0 | 0.0    | 12.0 | 37  |
|            | ATL-962 120 mg | 134                | -2.8 | 21.08 | -93  | -13.0 | -2.0   | 8.0  | 93  |
| Week 20    | Placebo        | 59                 | 3.6  | 22.12 | -30  | -12.0 | 1.0    | 17.0 | 65  |
|            | ATL-962 120 mg | 133                | -4.6 | 20.73 | -125 | -17.0 | -5.0   | 10.0 | 46  |
| Week 24    | Placebo        | 59                 | 1.2  | 20.61 | -36  | -11.0 | -4.0   | 14.0 | 65  |
|            | ATL-962 120 mg | 130                | -1.6 | 27.45 | -120 | -14.0 | -4.0   | 12.0 | 173 |
| Week 28    | Placebo        | 59                 | 2.5  | 16.66 | -50  | -12.0 | 4.0    | 15.0 | 35  |
|            | ATL-962 120 mg | 127                | -1.2 | 23.57 | -125 | -15.0 | 0.0    | 13.0 | 64  |
| Week 32    | Placebo        | 58                 | -2.2 | 18.24 | -42  | -14.0 | -3.5   | 9.0  | 50  |
|            | ATL-962 120 mg | 127                | -5.5 | 20.95 | -108 | -16.0 | -6.0   | 8.0  | 59  |
| Week 36    | Placebo        | 58                 | 1.1  | 26.27 | -52  | -12.0 | 0.5    | 9.0  | 138 |
|            | ATL-962 120 mg | 124                | -2.0 | 27.57 | -138 | -14.5 | -4.0   | 9.0  | 161 |
| Week 40    | Placebo        | 58                 | -0.3 | 15.85 | -47  | -10.0 | -0.5   | 8.0  | 45  |
|            | ATL-962 120 mg | 122                | -2.3 | 21.56 | -102 | -15.0 | -3.0   | 12.0 | 62  |
| Week 44    | Placebo        | 58                 | 3.6  | 20.08 | -60  | -7.0  | 3.0    | 15.0 | 55  |
|            | ATL-962 120 mg | 119                | 2.4  | 48.34 | -118 | -14.0 | -1.0   | 12.0 | 470 |
| Week 48    | Placebo        | 58                 | 9.2  | 25.09 | -57  | -6.0  | 6.0    | 24.0 | 115 |
|            | ATL-962 120 mg | 119                | 2.3  | 26.17 | -71  | -12.0 | 0.0    | 17.0 | 126 |
| Week 52    | Placebo        | 58                 | 7.5  | 21.04 | -72  | -4.0  | 8.0    | 17.0 | 74  |
|            | ATL-962 120 mg | 118                | 2.1  | 22.32 | -61  | -10.0 | 3.0    | 13.0 | 86  |

| CPK (IU/L) |                | Summary Statistics |       |         |      |       |        |      |       |
|------------|----------------|--------------------|-------|---------|------|-------|--------|------|-------|
| Visit      | Treatment      | N                  | Mean  | SD      | Min  | Q1    | Median | Q3   | Max   |
| Week 2     | Placebo        | 65                 | 6.6   | 54.83   | -133 | -12.0 | 3.0    | 16.0 | 342   |
|            | ATL-962 120 mg | 141                | -17.6 | 76.99   | -586 | -18.0 | -3.0   | 6.0  | 124   |
| Week 4     | Placebo        | 63                 | 3.9   | 32.19   | -99  | -8.0  | 2.0    | 18.0 | 94    |
|            | ATL-962 120 mg | 138                | -13.5 | 84.01   | -459 | -24.0 | -3.0   | 15.0 | 382   |
| Week 8     | Placebo        | 62                 | 15.0  | 76.45   | -79  | -9.0  | 4.0    | 21.0 | 556   |
|            | ATL-962 120 mg | 137                | -6.3  | 86.86   | -564 | -18.0 | 4.0    | 19.0 | 203   |
| Week 12    | Placebo        | 59                 | 13.5  | 50.04   | -81  | -11.0 | 11.0   | 32.0 | 264   |
|            | ATL-962 120 mg | 137                | -4.2  | 101.29  | -624 | -17.0 | 0.0    | 16.0 | 604   |
| Week 16    | Placebo        | 59                 | 21.2  | 97.23   | -65  | -6.0  | 9.0    | 21.0 | 714   |
|            | ATL-962 120 mg | 134                | -11.0 | 85.63   | -604 | -14.0 | 3.0    | 23.0 | 127   |
| Week 20    | Placebo        | 59                 | 34.8  | 99.17   | -111 | -2.0  | 14.0   | 46.0 | 514   |
|            | ATL-962 120 mg | 133                | -12.4 | 95.73   | -673 | -26.0 | 1.0    | 21.0 | 284   |
| Week 24    | Placebo        | 59                 | 25.4  | 103.52  | -86  | -9.0  | 5.0    | 34.0 | 630   |
|            | ATL-962 120 mg | 130                | 26.4  | 444.30  | -602 | -18.0 | 2.0    | 20.0 | 4938  |
| Week 28    | Placebo        | 59                 | 8.0   | 44.33   | -142 | -12.0 | 4.0    | 19.0 | 144   |
|            | ATL-962 120 mg | 127                | -8.7  | 94.53   | -547 | -26.0 | -1.0   | 25.0 | 254   |
| Week 32    | Placebo        | 58                 | 10.2  | 42.97   | -104 | -10.0 | 3.5    | 32.0 | 124   |
|            | ATL-962 120 mg | 127                | -9.8  | 90.85   | -487 | -25.0 | 2.0    | 18.0 | 313   |
| Week 36    | Placebo        | 58                 | 10.1  | 60.84   | -136 | -14.0 | 6.5    | 33.0 | 287   |
|            | ATL-962 120 mg | 124                | 3.4   | 105.82  | -586 | -19.5 | 1.5    | 33.0 | 293   |
| Week 40    | Placebo        | 58                 | 2.8   | 37.16   | -128 | -13.0 | 3.0    | 18.0 | 112   |
|            | ATL-962 120 mg | 122                | -9.5  | 99.99   | -656 | -21.0 | 1.0    | 25.0 | 257   |
| Week 44    | Placebo        | 58                 | 11.1  | 75.61   | -120 | -14.0 | 4.5    | 27.0 | 475   |
|            | ATL-962 120 mg | 119                | 117.8 | 1206.18 | -659 | -18.0 | 6.0    | 34.0 | 13084 |
| Week 48    | Placebo        | 58                 | 15.4  | 61.80   | -76  | -14.0 | 6.5    | 28.0 | 333   |
|            | ATL-962 120 mg | 119                | -5.7  | 101.59  | -673 | -20.0 | 0.0    | 22.0 | 253   |
| Week 52    | Placebo        | 58                 | 11.3  | 54.43   | -109 | -7.0  | 11.5   | 30.0 | 269   |
|            | ATL-962 120 mg | 118                | 12.9  | 227.98  | -440 | -25.0 | -2.0   | 21.0 | 2252  |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (5/12)

| Uric Acid (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2            | Placebo        | 65                 | 0.04  | 0.715 | -1.5 | -0.30 | 0.00   | 0.40 | 2.8 |
|                   | ATL-962 120 mg | 141                | 0.12  | 0.632 | -1.9 | -0.30 | 0.10   | 0.40 | 2.1 |
| Week 4            | Placebo        | 63                 | 0.13  | 0.707 | -1.7 | -0.30 | 0.10   | 0.60 | 1.9 |
|                   | ATL-962 120 mg | 138                | 0.05  | 0.625 | -2.1 | -0.20 | 0.00   | 0.50 | 1.7 |
| Week 8            | Placebo        | 62                 | 0.15  | 0.688 | -1.3 | -0.30 | 0.00   | 0.60 | 2.4 |
|                   | ATL-962 120 mg | 137                | 0.15  | 0.746 | -1.5 | -0.40 | 0.00   | 0.50 | 3.5 |
| Week 12           | Placebo        | 59                 | 0.20  | 0.875 | -1.3 | -0.40 | 0.10   | 0.70 | 3.4 |
|                   | ATL-962 120 mg | 137                | 0.15  | 0.782 | -1.9 | -0.30 | 0.00   | 0.60 | 3.0 |
| Week 16           | Placebo        | 59                 | 0.23  | 0.756 | -1.1 | -0.30 | 0.10   | 0.70 | 2.2 |
|                   | ATL-962 120 mg | 134                | 0.04  | 0.783 | -2.4 | -0.40 | -0.10  | 0.40 | 2.5 |
| Week 20           | Placebo        | 59                 | 0.32  | 0.870 | -1.9 | -0.20 | 0.20   | 0.80 | 2.8 |
|                   | ATL-962 120 mg | 133                | 0.17  | 0.849 | -2.2 | -0.40 | 0.10   | 0.70 | 2.5 |
| Week 24           | Placebo        | 59                 | 0.30  | 0.919 | -1.8 | -0.30 | 0.30   | 0.90 | 2.7 |
|                   | ATL-962 120 mg | 130                | 0.24  | 0.832 | -2.2 | -0.40 | 0.20   | 0.70 | 3.0 |
| Week 28           | Placebo        | 59                 | 0.38  | 0.884 | -1.4 | -0.20 | 0.30   | 1.00 | 2.6 |
|                   | ATL-962 120 mg | 127                | 0.15  | 0.832 | -1.9 | -0.40 | 0.10   | 0.50 | 2.7 |
| Week 32           | Placebo        | 58                 | 0.22  | 0.942 | -1.4 | -0.50 | 0.15   | 0.80 | 2.3 |
|                   | ATL-962 120 mg | 127                | 0.09  | 0.833 | -2.5 | -0.50 | 0.00   | 0.60 | 2.5 |
| Week 36           | Placebo        | 58                 | 0.21  | 0.918 | -1.8 | -0.40 | 0.10   | 0.80 | 2.5 |
|                   | ATL-962 120 mg | 124                | 0.06  | 0.768 | -2.0 | -0.40 | 0.00   | 0.40 | 2.7 |
| Week 40           | Placebo        | 58                 | 0.09  | 0.850 | -1.9 | -0.40 | 0.05   | 0.50 | 2.6 |
|                   | ATL-962 120 mg | 122                | -0.03 | 0.775 | -2.7 | -0.50 | -0.15  | 0.40 | 2.8 |
| Week 44           | Placebo        | 58                 | 0.09  | 0.967 | -1.8 | -0.40 | 0.05   | 0.60 | 3.3 |
|                   | ATL-962 120 mg | 119                | 0.03  | 0.765 | -2.1 | -0.50 | 0.00   | 0.50 | 2.5 |
| Week 48           | Placebo        | 58                 | 0.05  | 0.943 | -2.0 | -0.50 | 0.00   | 0.70 | 2.6 |
|                   | ATL-962 120 mg | 119                | -0.13 | 0.989 | -2.2 | -0.70 | -0.20  | 0.30 | 6.5 |
| Week 52           | Placebo        | 58                 | 0.07  | 1.051 | -2.3 | -0.60 | 0.10   | 0.80 | 3.4 |
|                   | ATL-962 120 mg | 118                | -0.13 | 0.745 | -2.5 | -0.60 | -0.10  | 0.20 | 2.3 |

| Total Bilirubin (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-------------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit                   | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2                  | Placebo        | 65                 | -0.00 | 0.242 | -0.6 | -0.10 | 0.00   | 0.10 | 0.8 |
|                         | ATL-962 120 mg | 141                | -0.01 | 0.201 | -0.6 | -0.10 | 0.00   | 0.10 | 0.6 |
| Week 4                  | Placebo        | 63                 | 0.05  | 0.215 | -0.5 | 0.00  | 0.00   | 0.10 | 0.8 |
|                         | ATL-962 120 mg | 138                | -0.02 | 0.164 | -0.5 | -0.10 | 0.00   | 0.10 | 0.4 |
| Week 8                  | Placebo        | 62                 | 0.08  | 0.257 | -0.7 | 0.00  | 0.10   | 0.20 | 0.7 |
|                         | ATL-962 120 mg | 137                | -0.01 | 0.205 | -0.5 | -0.10 | 0.00   | 0.10 | 0.6 |
| Week 12                 | Placebo        | 59                 | 0.06  | 0.228 | -0.6 | -0.10 | 0.00   | 0.20 | 0.6 |
|                         | ATL-962 120 mg | 137                | 0.00  | 0.216 | -0.5 | -0.10 | 0.00   | 0.10 | 1.1 |
| Week 16                 | Placebo        | 59                 | 0.12  | 0.301 | -0.5 | -0.10 | 0.10   | 0.30 | 1.0 |
|                         | ATL-962 120 mg | 134                | 0.00  | 0.212 | -0.6 | -0.10 | 0.00   | 0.10 | 0.9 |
| Week 20                 | Placebo        | 59                 | 0.13  | 0.252 | -0.5 | 0.00  | 0.10   | 0.30 | 0.7 |
|                         | ATL-962 120 mg | 133                | 0.02  | 0.203 | -0.4 | -0.10 | 0.00   | 0.10 | 0.5 |
| Week 24                 | Placebo        | 59                 | 0.07  | 0.236 | -0.4 | -0.10 | 0.10   | 0.20 | 0.6 |
|                         | ATL-962 120 mg | 130                | 0.04  | 0.222 | -0.6 | -0.10 | 0.00   | 0.10 | 0.9 |
| Week 28                 | Placebo        | 59                 | 0.13  | 0.246 | -0.5 | 0.00  | 0.10   | 0.30 | 0.9 |
|                         | ATL-962 120 mg | 127                | 0.03  | 0.239 | -0.4 | -0.10 | 0.00   | 0.10 | 1.0 |
| Week 32                 | Placebo        | 58                 | 0.12  | 0.271 | -0.5 | -0.10 | 0.10   | 0.30 | 1.0 |
|                         | ATL-962 120 mg | 127                | 0.04  | 0.232 | -0.6 | -0.10 | 0.00   | 0.10 | 0.9 |
| Week 36                 | Placebo        | 58                 | 0.05  | 0.282 | -0.7 | -0.10 | 0.00   | 0.30 | 0.7 |
|                         | ATL-962 120 mg | 124                | 0.04  | 0.209 | -0.5 | -0.10 | 0.00   | 0.10 | 0.8 |
| Week 40                 | Placebo        | 58                 | 0.08  | 0.260 | -0.7 | 0.00  | 0.10   | 0.20 | 0.7 |
|                         | ATL-962 120 mg | 122                | 0.04  | 0.256 | -0.4 | -0.10 | 0.00   | 0.10 | 1.5 |
| Week 44                 | Placebo        | 58                 | 0.08  | 0.275 | -0.6 | -0.10 | 0.10   | 0.20 | 0.9 |
|                         | ATL-962 120 mg | 119                | 0.01  | 0.214 | -0.8 | -0.10 | 0.00   | 0.10 | 0.8 |
| Week 48                 | Placebo        | 58                 | 0.04  | 0.267 | -0.5 | -0.10 | 0.00   | 0.20 | 0.8 |
|                         | ATL-962 120 mg | 119                | -0.01 | 0.230 | -0.5 | -0.10 | 0.00   | 0.10 | 1.0 |
| Week 52                 | Placebo        | 58                 | 0.04  | 0.238 | -0.5 | -0.10 | 0.00   | 0.20 | 0.7 |
|                         | ATL-962 120 mg | 118                | -0.01 | 0.216 | -0.6 | -0.10 | 0.00   | 0.10 | 0.6 |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (6/12)

| ChE (U/L) |                | Summary Statistics |       |       |      |       |        |      |     |
|-----------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit     | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2    | Placebo        | 65                 | -5.2  | 21.80 | -69  | -17.0 | -5.0   | 9.0  | 39  |
|           | ATL-962 120 mg | 141                | -13.7 | 26.42 | -114 | -26.0 | -10.0  | -1.0 | 47  |
| Week 4    | Placebo        | 63                 | -1.7  | 23.45 | -66  | -14.0 | -4.0   | 15.0 | 59  |
|           | ATL-962 120 mg | 138                | -13.7 | 30.00 | -88  | -32.0 | -14.5  | 5.0  | 82  |
| Week 8    | Placebo        | 62                 | -1.2  | 26.02 | -70  | -20.0 | 3.0    | 15.0 | 63  |
|           | ATL-962 120 mg | 137                | -17.2 | 31.18 | -143 | -35.0 | -17.0  | 3.0  | 61  |
| Week 12   | Placebo        | 59                 | -4.5  | 33.14 | -89  | -25.0 | -10.0  | 22.0 | 66  |
|           | ATL-962 120 mg | 137                | -17.4 | 33.81 | -117 | -38.0 | -17.0  | 3.0  | 79  |
| Week 16   | Placebo        | 59                 | -5.1  | 28.31 | -62  | -25.0 | -1.0   | 14.0 | 64  |
|           | ATL-962 120 mg | 134                | -20.7 | 35.57 | -179 | -37.0 | -18.5  | 1.0  | 72  |
| Week 20   | Placebo        | 59                 | -8.5  | 34.15 | -126 | -33.0 | -10.0  | 12.0 | 61  |
|           | ATL-962 120 mg | 133                | -25.6 | 38.04 | -169 | -47.0 | -22.0  | -5.0 | 69  |
| Week 24   | Placebo        | 59                 | -16.8 | 33.84 | -121 | -42.0 | -14.0  | 7.0  | 44  |
|           | ATL-962 120 mg | 130                | -29.4 | 38.18 | -185 | -48.0 | -26.0  | -7.0 | 52  |
| Week 28   | Placebo        | 59                 | -13.4 | 34.45 | -79  | -35.0 | -19.0  | 9.0  | 73  |
|           | ATL-962 120 mg | 127                | -28.0 | 39.87 | -210 | -45.0 | -25.0  | -6.0 | 69  |
| Week 32   | Placebo        | 58                 | -13.3 | 34.81 | -97  | -38.0 | -16.5  | 12.0 | 62  |
|           | ATL-962 120 mg | 127                | -22.7 | 43.44 | -218 | -44.0 | -16.0  | 2.0  | 81  |
| Week 36   | Placebo        | 58                 | -13.6 | 32.17 | -110 | -29.0 | -18.5  | 5.0  | 90  |
|           | ATL-962 120 mg | 124                | -17.0 | 43.65 | -179 | -39.5 | -15.0  | 5.5  | 157 |
| Week 40   | Placebo        | 58                 | -6.7  | 28.83 | -77  | -26.0 | -7.5   | 15.0 | 48  |
|           | ATL-962 120 mg | 122                | -17.2 | 46.71 | -149 | -40.0 | -17.0  | 11.0 | 155 |
| Week 44   | Placebo        | 58                 | 0.6   | 31.41 | -92  | -19.0 | -2.0   | 23.0 | 69  |
|           | ATL-962 120 mg | 119                | -15.5 | 44.26 | -187 | -42.0 | -11.0  | 9.0  | 159 |
| Week 48   | Placebo        | 58                 | 0.1   | 33.59 | -91  | -19.0 | 1.5    | 27.0 | 66  |
|           | ATL-962 120 mg | 119                | -18.7 | 43.80 | -185 | -41.0 | -19.0  | 9.0  | 139 |
| Week 52   | Placebo        | 58                 | -1.3  | 35.67 | -132 | -22.0 | 6.5    | 18.0 | 72  |
|           | ATL-962 120 mg | 118                | -13.9 | 46.48 | -189 | -40.0 | -10.5  | 15.0 | 153 |

| Creatinine (mg/dL) |                | Summary Statistics |        |        |       |        |        |       |      |
|--------------------|----------------|--------------------|--------|--------|-------|--------|--------|-------|------|
| Visit              | Treatment      | N                  | Mean   | SD     | Min   | Q1     | Median | Q3    | Max  |
| Week 2             | Placebo        | 65                 | 0.002  | 0.0711 | -0.34 | -0.030 | 0.000  | 0.040 | 0.12 |
|                    | ATL-962 120 mg | 141                | 0.015  | 0.0592 | -0.12 | -0.020 | 0.010  | 0.040 | 0.27 |
| Week 4             | Placebo        | 63                 | -0.000 | 0.0626 | -0.10 | -0.050 | -0.010 | 0.030 | 0.21 |
|                    | ATL-962 120 mg | 138                | 0.013  | 0.0549 | -0.12 | -0.020 | 0.010  | 0.040 | 0.25 |
| Week 8             | Placebo        | 62                 | -0.002 | 0.0636 | -0.11 | -0.050 | 0.000  | 0.040 | 0.25 |
|                    | ATL-962 120 mg | 137                | 0.007  | 0.0584 | -0.15 | -0.030 | 0.010  | 0.040 | 0.24 |
| Week 12            | Placebo        | 59                 | 0.002  | 0.0581 | -0.12 | -0.030 | 0.000  | 0.030 | 0.24 |
|                    | ATL-962 120 mg | 137                | 0.024  | 0.0945 | -0.17 | -0.020 | 0.020  | 0.050 | 0.88 |
| Week 16            | Placebo        | 59                 | 0.018  | 0.0604 | -0.09 | -0.030 | 0.020  | 0.050 | 0.25 |
|                    | ATL-962 120 mg | 134                | 0.012  | 0.0619 | -0.18 | -0.030 | 0.010  | 0.050 | 0.19 |
| Week 20            | Placebo        | 59                 | 0.030  | 0.0625 | -0.09 | -0.010 | 0.020  | 0.070 | 0.24 |
|                    | ATL-962 120 mg | 133                | 0.029  | 0.0653 | -0.28 | -0.010 | 0.030  | 0.060 | 0.30 |
| Week 24            | Placebo        | 59                 | 0.024  | 0.0618 | -0.11 | -0.020 | 0.020  | 0.070 | 0.19 |
|                    | ATL-962 120 mg | 130                | 0.030  | 0.0616 | -0.13 | -0.010 | 0.030  | 0.070 | 0.24 |
| Week 28            | Placebo        | 59                 | 0.035  | 0.0861 | -0.15 | -0.020 | 0.020  | 0.100 | 0.24 |
|                    | ATL-962 120 mg | 127                | 0.032  | 0.0691 | -0.18 | -0.010 | 0.030  | 0.070 | 0.24 |
| Week 32            | Placebo        | 58                 | 0.030  | 0.0765 | -0.10 | -0.030 | 0.030  | 0.070 | 0.35 |
|                    | ATL-962 120 mg | 127                | 0.037  | 0.0638 | -0.13 | 0.000  | 0.030  | 0.080 | 0.29 |
| Week 36            | Placebo        | 58                 | 0.033  | 0.0564 | -0.10 | -0.010 | 0.035  | 0.070 | 0.15 |
|                    | ATL-962 120 mg | 124                | 0.031  | 0.0660 | -0.17 | -0.010 | 0.030  | 0.070 | 0.27 |
| Week 40            | Placebo        | 58                 | 0.026  | 0.0697 | -0.09 | -0.020 | 0.020  | 0.070 | 0.26 |
|                    | ATL-962 120 mg | 122                | 0.026  | 0.0721 | -0.19 | -0.010 | 0.020  | 0.060 | 0.32 |
| Week 44            | Placebo        | 58                 | 0.042  | 0.0767 | -0.13 | -0.010 | 0.040  | 0.070 | 0.31 |
|                    | ATL-962 120 mg | 119                | 0.032  | 0.0802 | -0.20 | -0.020 | 0.020  | 0.070 | 0.42 |
| Week 48            | Placebo        | 58                 | 0.021  | 0.0693 | -0.10 | -0.030 | 0.010  | 0.050 | 0.21 |
|                    | ATL-962 120 mg | 119                | 0.028  | 0.0819 | -0.11 | -0.020 | 0.010  | 0.070 | 0.50 |
| Week 52            | Placebo        | 58                 | 0.025  | 0.0758 | -0.10 | -0.030 | 0.030  | 0.060 | 0.26 |
|                    | ATL-962 120 mg | 118                | 0.023  | 0.0729 | -0.20 | -0.020 | 0.015  | 0.060 | 0.36 |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (7/12)

| BUN (mg/dL) |                | Summary Statistics |      |      |     |      |        |     |     |
|-------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit       | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2      | Placebo        | 65                 | -0.5 | 2.34 | -5  | -2.0 | 0.0    | 1.0 | 6   |
|             | ATL-962 120 mg | 141                | -0.2 | 3.00 | -11 | -2.0 | 0.0    | 1.0 | 9   |
| Week 4      | Placebo        | 63                 | -0.2 | 2.35 | -6  | -1.0 | -1.0   | 1.0 | 7   |
|             | ATL-962 120 mg | 138                | -0.2 | 3.01 | -11 | -2.0 | 0.0    | 2.0 | 8   |
| Week 8      | Placebo        | 62                 | -0.2 | 2.78 | -6  | -2.0 | -1.0   | 1.0 | 8   |
|             | ATL-962 120 mg | 137                | -0.3 | 3.15 | -10 | -2.0 | 0.0    | 1.0 | 12  |
| Week 12     | Placebo        | 59                 | 0.0  | 2.92 | -5  | -2.0 | 0.0    | 2.0 | 11  |
|             | ATL-962 120 mg | 137                | -0.0 | 3.86 | -9  | -2.0 | 0.0    | 2.0 | 22  |
| Week 16     | Placebo        | 59                 | -0.3 | 2.83 | -7  | -2.0 | 0.0    | 2.0 | 9   |
|             | ATL-962 120 mg | 134                | -0.4 | 3.08 | -11 | -2.0 | -1.0   | 1.0 | 8   |
| Week 20     | Placebo        | 59                 | 0.1  | 5.14 | -6  | -2.0 | -1.0   | 2.0 | 33  |
|             | ATL-962 120 mg | 133                | -0.3 | 3.10 | -9  | -2.0 | 0.0    | 2.0 | 8   |
| Week 24     | Placebo        | 59                 | -0.3 | 2.63 | -6  | -2.0 | 0.0    | 1.0 | 7   |
|             | ATL-962 120 mg | 130                | -0.1 | 3.35 | -11 | -2.0 | 0.0    | 2.0 | 8   |
| Week 28     | Placebo        | 59                 | 0.0  | 3.21 | -6  | -2.0 | 0.0    | 2.0 | 8   |
|             | ATL-962 120 mg | 127                | -0.1 | 2.91 | -10 | -2.0 | 0.0    | 2.0 | 7   |
| Week 32     | Placebo        | 58                 | -0.2 | 2.71 | -5  | -2.0 | 0.0    | 1.0 | 9   |
|             | ATL-962 120 mg | 127                | -0.6 | 2.99 | -9  | -2.0 | -1.0   | 1.0 | 7   |
| Week 36     | Placebo        | 58                 | -0.8 | 3.15 | -6  | -3.0 | -1.0   | 1.0 | 9   |
|             | ATL-962 120 mg | 124                | -0.5 | 3.50 | -10 | -2.5 | 0.0    | 2.0 | 14  |
| Week 40     | Placebo        | 58                 | -0.4 | 2.56 | -6  | -2.0 | 0.0    | 1.0 | 6   |
|             | ATL-962 120 mg | 122                | -0.2 | 3.57 | -10 | -2.0 | 0.0    | 2.0 | 9   |
| Week 44     | Placebo        | 58                 | -0.3 | 2.50 | -6  | -2.0 | 0.0    | 1.0 | 7   |
|             | ATL-962 120 mg | 119                | -0.3 | 3.52 | -10 | -2.0 | 0.0    | 2.0 | 10  |
| Week 48     | Placebo        | 58                 | 0.3  | 2.74 | -5  | -1.0 | 0.0    | 1.0 | 8   |
|             | ATL-962 120 mg | 119                | 0.0  | 3.41 | -11 | -2.0 | 0.0    | 2.0 | 13  |
| Week 52     | Placebo        | 58                 | 0.1  | 3.18 | -7  | -2.0 | 0.0    | 2.0 | 7   |
|             | ATL-962 120 mg | 118                | -0.1 | 3.09 | -8  | -2.0 | 0.0    | 2.0 | 7   |

| Na (mEq/L) |                | Summary Statistics |      |      |     |      |        |     |     |
|------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit      | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2     | Placebo        | 65                 | 0.4  | 2.00 | -5  | -1.0 | 0.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 141                | 0.5  | 1.68 | -4  | 0.0  | 1.0    | 2.0 | 5   |
| Week 4     | Placebo        | 63                 | 0.3  | 1.77 | -4  | -1.0 | 0.0    | 2.0 | 5   |
|            | ATL-962 120 mg | 138                | 0.5  | 1.69 | -4  | -1.0 | 0.0    | 1.0 | 6   |
| Week 8     | Placebo        | 62                 | 0.3  | 2.26 | -6  | -1.0 | 0.0    | 1.0 | 6   |
|            | ATL-962 120 mg | 137                | 0.4  | 1.58 | -4  | 0.0  | 1.0    | 1.0 | 4   |
| Week 12    | Placebo        | 59                 | 0.4  | 1.95 | -4  | -1.0 | 0.0    | 2.0 | 5   |
|            | ATL-962 120 mg | 137                | 0.3  | 1.76 | -4  | -1.0 | 1.0    | 2.0 | 4   |
| Week 16    | Placebo        | 59                 | 0.3  | 1.84 | -3  | -1.0 | 0.0    | 1.0 | 4   |
|            | ATL-962 120 mg | 134                | 0.5  | 1.87 | -6  | -1.0 | 0.0    | 2.0 | 5   |
| Week 20    | Placebo        | 59                 | 0.3  | 2.09 | -5  | -1.0 | 0.0    | 2.0 | 4   |
|            | ATL-962 120 mg | 133                | 0.9  | 1.85 | -4  | 0.0  | 1.0    | 2.0 | 5   |
| Week 24    | Placebo        | 59                 | 0.7  | 2.07 | -4  | -1.0 | 1.0    | 2.0 | 5   |
|            | ATL-962 120 mg | 130                | 0.8  | 1.89 | -3  | -1.0 | 1.0    | 2.0 | 6   |
| Week 28    | Placebo        | 59                 | 0.8  | 2.09 | -4  | -1.0 | 1.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 127                | 0.8  | 1.88 | -5  | 0.0  | 1.0    | 2.0 | 5   |
| Week 32    | Placebo        | 58                 | 0.6  | 2.14 | -3  | -1.0 | 0.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 127                | 0.8  | 2.06 | -4  | 0.0  | 1.0    | 2.0 | 6   |
| Week 36    | Placebo        | 58                 | 0.7  | 2.27 | -4  | -1.0 | 1.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 124                | 0.7  | 1.94 | -5  | -1.0 | 1.0    | 2.0 | 5   |
| Week 40    | Placebo        | 58                 | 0.7  | 2.44 | -4  | -1.0 | 1.0    | 2.0 | 5   |
|            | ATL-962 120 mg | 122                | 0.9  | 1.83 | -3  | 0.0  | 1.0    | 2.0 | 6   |
| Week 44    | Placebo        | 58                 | 0.8  | 2.20 | -4  | -1.0 | 0.5    | 2.0 | 6   |
|            | ATL-962 120 mg | 119                | 0.9  | 1.71 | -3  | 0.0  | 1.0    | 2.0 | 5   |
| Week 48    | Placebo        | 58                 | 0.7  | 2.13 | -5  | -1.0 | 1.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 119                | 0.7  | 1.71 | -4  | 0.0  | 1.0    | 2.0 | 5   |
| Week 52    | Placebo        | 58                 | 0.8  | 2.02 | -4  | -1.0 | 1.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 118                | 0.7  | 1.74 | -4  | 0.0  | 1.0    | 2.0 | 6   |



表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (8/12)

| K (mEq/L) |                | Summary Statistics |       |       |      |       |        |      |     |
|-----------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit     | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2    | Placebo        | 65                 | 0.05  | 0.301 | -0.5 | -0.20 | 0.00   | 0.20 | 1.0 |
|           | ATL-962 120 mg | 141                | -0.01 | 0.281 | -0.8 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 4    | Placebo        | 63                 | 0.04  | 0.263 | -0.6 | -0.10 | 0.00   | 0.20 | 0.7 |
|           | ATL-962 120 mg | 138                | -0.01 | 0.256 | -0.9 | -0.10 | 0.00   | 0.10 | 0.7 |
| Week 8    | Placebo        | 62                 | -0.02 | 0.299 | -1.0 | -0.20 | 0.00   | 0.20 | 0.7 |
|           | ATL-962 120 mg | 137                | -0.04 | 0.249 | -0.7 | -0.20 | 0.00   | 0.10 | 0.5 |
| Week 12   | Placebo        | 59                 | 0.01  | 0.301 | -0.6 | -0.20 | 0.00   | 0.20 | 0.9 |
|           | ATL-962 120 mg | 137                | -0.02 | 0.281 | -0.8 | -0.20 | 0.00   | 0.10 | 1.2 |
| Week 16   | Placebo        | 59                 | 0.02  | 0.328 | -0.9 | -0.20 | 0.00   | 0.20 | 1.1 |
|           | ATL-962 120 mg | 134                | -0.01 | 0.272 | -0.8 | -0.20 | 0.00   | 0.10 | 1.1 |
| Week 20   | Placebo        | 59                 | 0.01  | 0.238 | -0.5 | -0.20 | 0.00   | 0.20 | 0.7 |
|           | ATL-962 120 mg | 133                | -0.02 | 0.311 | -0.9 | -0.20 | 0.00   | 0.20 | 0.8 |
| Week 24   | Placebo        | 59                 | -0.07 | 0.260 | -0.7 | -0.20 | -0.10  | 0.10 | 0.6 |
|           | ATL-962 120 mg | 130                | -0.04 | 0.279 | -0.8 | -0.20 | 0.00   | 0.10 | 0.6 |
| Week 28   | Placebo        | 59                 | -0.01 | 0.250 | -0.6 | -0.20 | 0.00   | 0.20 | 0.5 |
|           | ATL-962 120 mg | 127                | -0.05 | 0.310 | -0.8 | -0.30 | 0.00   | 0.10 | 0.8 |
| Week 32   | Placebo        | 58                 | -0.01 | 0.290 | -0.7 | -0.20 | 0.00   | 0.20 | 0.8 |
|           | ATL-962 120 mg | 127                | -0.08 | 0.293 | -0.9 | -0.30 | 0.00   | 0.10 | 0.4 |
| Week 36   | Placebo        | 58                 | -0.06 | 0.306 | -0.9 | -0.20 | -0.10  | 0.10 | 0.9 |
|           | ATL-962 120 mg | 124                | -0.10 | 0.330 | -1.0 | -0.30 | -0.10  | 0.10 | 0.7 |
| Week 40   | Placebo        | 58                 | -0.02 | 0.249 | -0.5 | -0.20 | -0.10  | 0.10 | 0.8 |
|           | ATL-962 120 mg | 122                | -0.03 | 0.299 | -0.9 | -0.20 | 0.00   | 0.20 | 0.8 |
| Week 44   | Placebo        | 58                 | 0.04  | 0.283 | -0.7 | -0.10 | 0.00   | 0.30 | 0.8 |
|           | ATL-962 120 mg | 119                | -0.05 | 0.276 | -0.8 | -0.20 | -0.10  | 0.10 | 0.6 |
| Week 48   | Placebo        | 58                 | 0.02  | 0.263 | -0.6 | -0.20 | 0.00   | 0.20 | 0.6 |
|           | ATL-962 120 mg | 119                | -0.02 | 0.278 | -0.8 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 52   | Placebo        | 58                 | 0.02  | 0.278 | -0.5 | -0.20 | 0.00   | 0.20 | 0.7 |
|           | ATL-962 120 mg | 118                | -0.01 | 0.311 | -0.8 | -0.20 | 0.00   | 0.20 | 1.2 |

| Cl (mEq/L) |                | Summary Statistics |      |      |     |      |        |     |     |
|------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit      | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2     | Placebo        | 65                 | 0.3  | 2.35 | -6  | -1.0 | 0.0    | 2.0 | 5   |
|            | ATL-962 120 mg | 141                | 0.9  | 1.83 | -5  | 0.0  | 1.0    | 2.0 | 5   |
| Week 4     | Placebo        | 63                 | 0.4  | 1.89 | -4  | -1.0 | 0.0    | 2.0 | 4   |
|            | ATL-962 120 mg | 138                | 0.7  | 2.07 | -5  | -1.0 | 1.0    | 2.0 | 5   |
| Week 8     | Placebo        | 62                 | 0.2  | 2.35 | -6  | -1.0 | 0.0    | 1.0 | 6   |
|            | ATL-962 120 mg | 137                | 1.0  | 2.11 | -6  | 0.0  | 1.0    | 2.0 | 7   |
| Week 12    | Placebo        | 59                 | 0.6  | 2.16 | -3  | -1.0 | 0.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 137                | 0.9  | 1.97 | -4  | 0.0  | 1.0    | 2.0 | 6   |
| Week 16    | Placebo        | 59                 | 0.9  | 2.06 | -4  | 0.0  | 1.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 134                | 1.2  | 2.10 | -6  | 0.0  | 1.0    | 3.0 | 7   |
| Week 20    | Placebo        | 59                 | 0.7  | 2.01 | -4  | 0.0  | 1.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 133                | 1.5  | 2.13 | -7  | 0.0  | 1.0    | 3.0 | 7   |
| Week 24    | Placebo        | 59                 | 1.3  | 2.28 | -4  | 0.0  | 1.0    | 3.0 | 7   |
|            | ATL-962 120 mg | 130                | 1.5  | 2.32 | -6  | 0.0  | 2.0    | 3.0 | 6   |
| Week 28    | Placebo        | 59                 | 1.0  | 1.97 | -4  | 0.0  | 1.0    | 2.0 | 7   |
|            | ATL-962 120 mg | 127                | 1.3  | 2.33 | -7  | 0.0  | 1.0    | 3.0 | 8   |
| Week 32    | Placebo        | 58                 | 0.9  | 2.01 | -3  | 0.0  | 1.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 127                | 1.2  | 2.37 | -5  | 0.0  | 1.0    | 3.0 | 7   |
| Week 36    | Placebo        | 58                 | 1.3  | 2.28 | -3  | -1.0 | 1.5    | 3.0 | 6   |
|            | ATL-962 120 mg | 124                | 1.1  | 2.34 | -6  | 0.0  | 1.0    | 2.0 | 9   |
| Week 40    | Placebo        | 58                 | 1.0  | 2.17 | -3  | -1.0 | 1.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 122                | 0.9  | 2.26 | -6  | 0.0  | 1.0    | 2.0 | 8   |
| Week 44    | Placebo        | 58                 | 0.7  | 2.17 | -4  | -1.0 | 1.0    | 2.0 | 5   |
|            | ATL-962 120 mg | 119                | 0.9  | 2.19 | -6  | -1.0 | 1.0    | 2.0 | 7   |
| Week 48    | Placebo        | 58                 | 0.8  | 2.09 | -4  | -1.0 | 1.0    | 2.0 | 6   |
|            | ATL-962 120 mg | 119                | 0.9  | 2.11 | -4  | -1.0 | 1.0    | 2.0 | 8   |
| Week 52    | Placebo        | 58                 | 0.7  | 1.80 | -4  | 0.0  | 0.0    | 2.0 | 5   |
|            | ATL-962 120 mg | 118                | 0.6  | 2.20 | -5  | -1.0 | 0.0    | 2.0 | 7   |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (9/12)

| Ca (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit      | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2     | Placebo        | 65                 | 0.01  | 0.286 | -0.6 | -0.20 | 0.00   | 0.20 | 0.7 |
|            | ATL-962 120 mg | 141                | -0.02 | 0.280 | -0.7 | -0.20 | 0.00   | 0.20 | 0.9 |
| Week 4     | Placebo        | 63                 | 0.01  | 0.256 | -0.6 | -0.20 | 0.00   | 0.20 | 0.5 |
|            | ATL-962 120 mg | 138                | -0.03 | 0.302 | -1.0 | -0.20 | 0.00   | 0.20 | 0.9 |
| Week 8     | Placebo        | 62                 | -0.06 | 0.317 | -0.6 | -0.30 | -0.10  | 0.20 | 1.0 |
|            | ATL-962 120 mg | 137                | -0.07 | 0.327 | -0.9 | -0.30 | 0.00   | 0.10 | 0.7 |
| Week 12    | Placebo        | 59                 | -0.12 | 0.297 | -0.8 | -0.30 | -0.10  | 0.00 | 0.5 |
|            | ATL-962 120 mg | 137                | -0.14 | 0.321 | -1.3 | -0.30 | -0.20  | 0.10 | 0.7 |
| Week 16    | Placebo        | 59                 | -0.09 | 0.330 | -0.7 | -0.30 | -0.10  | 0.10 | 0.7 |
|            | ATL-962 120 mg | 134                | -0.15 | 0.325 | -1.1 | -0.40 | -0.20  | 0.10 | 0.7 |
| Week 20    | Placebo        | 59                 | -0.08 | 0.267 | -0.7 | -0.30 | -0.10  | 0.10 | 0.4 |
|            | ATL-962 120 mg | 133                | -0.13 | 0.320 | -1.2 | -0.30 | -0.10  | 0.10 | 0.7 |
| Week 24    | Placebo        | 59                 | -0.15 | 0.339 | -0.8 | -0.40 | -0.20  | 0.10 | 0.8 |
|            | ATL-962 120 mg | 130                | -0.16 | 0.330 | -1.1 | -0.40 | -0.20  | 0.00 | 0.8 |
| Week 28    | Placebo        | 59                 | -0.09 | 0.429 | -0.9 | -0.40 | -0.10  | 0.20 | 1.0 |
|            | ATL-962 120 mg | 127                | -0.15 | 0.294 | -1.0 | -0.30 | -0.20  | 0.10 | 0.8 |
| Week 32    | Placebo        | 58                 | -0.07 | 0.334 | -0.8 | -0.30 | -0.10  | 0.20 | 0.7 |
|            | ATL-962 120 mg | 127                | -0.14 | 0.308 | -1.1 | -0.30 | -0.10  | 0.00 | 0.7 |
| Week 36    | Placebo        | 58                 | -0.13 | 0.253 | -0.7 | -0.30 | -0.10  | 0.00 | 0.4 |
|            | ATL-962 120 mg | 124                | -0.14 | 0.279 | -0.8 | -0.30 | -0.10  | 0.00 | 0.4 |
| Week 40    | Placebo        | 58                 | -0.09 | 0.302 | -0.8 | -0.30 | -0.10  | 0.10 | 0.7 |
|            | ATL-962 120 mg | 122                | -0.13 | 0.361 | -1.0 | -0.40 | -0.10  | 0.10 | 0.9 |
| Week 44    | Placebo        | 58                 | -0.04 | 0.345 | -0.7 | -0.30 | -0.05  | 0.10 | 1.3 |
|            | ATL-962 120 mg | 119                | -0.12 | 0.318 | -0.9 | -0.30 | -0.10  | 0.10 | 0.7 |
| Week 48    | Placebo        | 58                 | -0.08 | 0.274 | -0.6 | -0.30 | -0.10  | 0.10 | 0.5 |
|            | ATL-962 120 mg | 119                | -0.15 | 0.311 | -1.0 | -0.30 | -0.10  | 0.10 | 0.7 |
| Week 52    | Placebo        | 58                 | -0.10 | 0.356 | -1.2 | -0.30 | -0.10  | 0.10 | 0.7 |
|            | ATL-962 120 mg | 118                | -0.13 | 0.327 | -1.1 | -0.40 | -0.10  | 0.10 | 0.6 |

| P (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-----------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit     | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2    | Placebo        | 65                 | 0.02  | 0.395 | -1.0 | -0.20 | 0.00   | 0.30 | 1.1 |
|           | ATL-962 120 mg | 141                | 0.03  | 0.368 | -1.0 | -0.20 | 0.00   | 0.20 | 1.4 |
| Week 4    | Placebo        | 63                 | -0.00 | 0.322 | -1.0 | -0.10 | 0.00   | 0.20 | 0.5 |
|           | ATL-962 120 mg | 138                | 0.02  | 0.338 | -0.8 | -0.20 | 0.00   | 0.20 | 0.9 |
| Week 8    | Placebo        | 62                 | 0.02  | 0.371 | -0.8 | -0.20 | 0.05   | 0.30 | 0.8 |
|           | ATL-962 120 mg | 137                | -0.01 | 0.331 | -1.0 | -0.20 | 0.00   | 0.20 | 1.0 |
| Week 12   | Placebo        | 59                 | -0.08 | 0.356 | -0.9 | -0.30 | -0.10  | 0.20 | 0.9 |
|           | ATL-962 120 mg | 137                | -0.00 | 0.377 | -1.4 | -0.20 | 0.00   | 0.30 | 0.8 |
| Week 16   | Placebo        | 59                 | 0.01  | 0.318 | -0.8 | -0.20 | 0.00   | 0.30 | 0.7 |
|           | ATL-962 120 mg | 134                | 0.01  | 0.375 | -1.3 | -0.30 | 0.00   | 0.30 | 1.2 |
| Week 20   | Placebo        | 59                 | -0.05 | 0.272 | -0.7 | -0.30 | -0.10  | 0.20 | 0.5 |
|           | ATL-962 120 mg | 133                | 0.05  | 0.416 | -1.0 | -0.20 | 0.10   | 0.30 | 1.2 |
| Week 24   | Placebo        | 59                 | -0.06 | 0.328 | -0.7 | -0.30 | 0.00   | 0.20 | 0.6 |
|           | ATL-962 120 mg | 130                | 0.08  | 0.409 | -1.2 | -0.20 | 0.00   | 0.40 | 1.3 |
| Week 28   | Placebo        | 59                 | 0.04  | 0.364 | -0.9 | -0.20 | 0.00   | 0.30 | 0.8 |
|           | ATL-962 120 mg | 127                | 0.08  | 0.374 | -1.3 | -0.10 | 0.10   | 0.30 | 1.0 |
| Week 32   | Placebo        | 58                 | 0.01  | 0.429 | -0.7 | -0.30 | 0.00   | 0.30 | 1.9 |
|           | ATL-962 120 mg | 127                | 0.07  | 0.400 | -0.8 | -0.20 | 0.00   | 0.30 | 1.4 |
| Week 36   | Placebo        | 58                 | -0.05 | 0.359 | -0.8 | -0.30 | -0.10  | 0.20 | 0.6 |
|           | ATL-962 120 mg | 124                | 0.09  | 0.345 | -0.7 | -0.10 | 0.10   | 0.30 | 1.0 |
| Week 40   | Placebo        | 58                 | -0.07 | 0.320 | -0.8 | -0.30 | -0.05  | 0.20 | 0.6 |
|           | ATL-962 120 mg | 122                | 0.08  | 0.370 | -0.8 | -0.10 | 0.10   | 0.30 | 1.1 |
| Week 44   | Placebo        | 58                 | 0.01  | 0.368 | -0.8 | -0.20 | -0.05  | 0.10 | 1.1 |
|           | ATL-962 120 mg | 119                | 0.07  | 0.346 | -0.9 | -0.20 | 0.10   | 0.30 | 1.3 |
| Week 48   | Placebo        | 58                 | -0.08 | 0.333 | -1.0 | -0.30 | -0.10  | 0.20 | 0.6 |
|           | ATL-962 120 mg | 119                | 0.05  | 0.385 | -0.7 | -0.20 | 0.10   | 0.30 | 1.1 |
| Week 52   | Placebo        | 58                 | -0.02 | 0.326 | -0.9 | -0.20 | 0.00   | 0.20 | 0.6 |
|           | ATL-962 120 mg | 118                | 0.04  | 0.310 | -0.6 | -0.20 | 0.05   | 0.20 | 1.2 |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (10/12)

| Vitamin A (ng/mL) |                | Summary Statistics |       |        |      |       |        |      |     |
|-------------------|----------------|--------------------|-------|--------|------|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean  | SD     | Min  | Q1    | Median | Q3   | Max |
| Week 2            | Placebo        | 65                 | -13.9 | 94.96  | -278 | -62.0 | -7.0   | 38.0 | 162 |
|                   | ATL-962 120 mg | 141                | -6.7  | 112.31 | -683 | -70.0 | -8.0   | 67.0 | 263 |
| Week 4            | Placebo        | 63                 | -15.2 | 86.47  | -325 | -43.0 | 1.0    | 38.0 | 147 |
|                   | ATL-962 120 mg | 138                | -20.8 | 105.06 | -755 | -66.0 | -14.0  | 32.0 | 255 |
| Week 8            | Placebo        | 62                 | -31.5 | 101.32 | -327 | -66.0 | -15.5  | 27.0 | 204 |
|                   | ATL-962 120 mg | 137                | -22.5 | 104.76 | -657 | -70.0 | -14.0  | 34.0 | 209 |
| Week 12           | Placebo        | 59                 | -33.1 | 101.09 | -315 | -91.0 | -26.0  | 9.0  | 229 |
|                   | ATL-962 120 mg | 137                | -20.9 | 108.85 | -609 | -72.0 | -19.0  | 47.0 | 182 |
| Week 16           | Placebo        | 59                 | -34.8 | 107.02 | -466 | -80.0 | -36.0  | 36.0 | 232 |
|                   | ATL-962 120 mg | 134                | -17.8 | 107.12 | -603 | -68.0 | -9.5   | 44.0 | 274 |
| Week 20           | Placebo        | 59                 | -4.7  | 108.02 | -253 | -66.0 | -22.0  | 53.0 | 279 |
|                   | ATL-962 120 mg | 133                | -5.9  | 107.80 | -690 | -51.0 | -2.0   | 60.0 | 204 |
| Week 24           | Placebo        | 59                 | -17.2 | 96.46  | -264 | -75.0 | -12.0  | 38.0 | 204 |
|                   | ATL-962 120 mg | 130                | 0.7   | 115.76 | -616 | -66.0 | -7.5   | 47.0 | 465 |
| Week 28           | Placebo        | 59                 | 8.7   | 129.52 | -261 | -56.0 | -7.0   | 75.0 | 545 |
|                   | ATL-962 120 mg | 127                | 5.6   | 118.82 | -686 | -63.0 | 20.0   | 71.0 | 293 |
| Week 32           | Placebo        | 58                 | -6.2  | 98.60  | -212 | -64.0 | -14.5  | 58.0 | 272 |
|                   | ATL-962 120 mg | 127                | 1.3   | 114.48 | -818 | -49.0 | 15.0   | 64.0 | 205 |
| Week 36           | Placebo        | 58                 | -21.2 | 106.10 | -526 | -54.0 | -11.0  | 44.0 | 169 |
|                   | ATL-962 120 mg | 124                | 0.1   | 116.21 | -643 | -60.5 | 6.0    | 70.0 | 388 |
| Week 40           | Placebo        | 58                 | -5.5  | 104.07 | -318 | -74.0 | -3.5   | 41.0 | 295 |
|                   | ATL-962 120 mg | 122                | 3.9   | 113.52 | -607 | -51.0 | 14.0   | 66.0 | 308 |
| Week 44           | Placebo        | 58                 | -3.5  | 90.32  | -321 | -43.0 | 1.0    | 44.0 | 259 |
|                   | ATL-962 120 mg | 119                | -1.1  | 114.29 | -649 | -54.0 | 9.0    | 61.0 | 286 |
| Week 48           | Placebo        | 58                 | -10.6 | 106.25 | -405 | -52.0 | -16.0  | 52.0 | 271 |
|                   | ATL-962 120 mg | 119                | -9.7  | 107.88 | -583 | -66.0 | -4.0   | 56.0 | 241 |
| Week 52           | Placebo        | 58                 | -11.9 | 127.96 | -568 | -76.0 | -11.5  | 45.0 | 334 |
|                   | ATL-962 120 mg | 118                | -4.9  | 110.85 | -495 | -62.0 | -1.5   | 69.0 | 261 |

| Vitamin D (pg/mL) |                | Summary Statistics |      |       |     |       |        |      |     |
|-------------------|----------------|--------------------|------|-------|-----|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean | SD    | Min | Q1    | Median | Q3   | Max |
| Week 2            | Placebo        | 65                 | -3.1 | 20.71 | -63 | -11.0 | 0.0    | 7.0  | 61  |
|                   | ATL-962 120 mg | 141                | -1.0 | 17.95 | -48 | -13.0 | -1.0   | 11.0 | 50  |
| Week 4            | Placebo        | 63                 | 2.4  | 21.08 | -71 | -12.0 | 3.0    | 16.0 | 56  |
|                   | ATL-962 120 mg | 138                | 1.8  | 19.13 | -46 | -10.0 | 2.0    | 14.0 | 46  |
| Week 8            | Placebo        | 62                 | 5.3  | 22.07 | -66 | -8.0  | 3.5    | 23.0 | 67  |
|                   | ATL-962 120 mg | 137                | 2.0  | 19.99 | -38 | -9.0  | 0.0    | 11.0 | 63  |
| Week 12           | Placebo        | 59                 | 2.6  | 22.88 | -63 | -14.0 | 4.0    | 17.0 | 68  |
|                   | ATL-962 120 mg | 137                | 1.4  | 16.77 | -42 | -10.0 | 2.0    | 10.0 | 53  |
| Week 16           | Placebo        | 59                 | 2.1  | 22.21 | -70 | -8.0  | 3.0    | 15.0 | 46  |
|                   | ATL-962 120 mg | 134                | 0.4  | 18.99 | -46 | -14.0 | 1.0    | 12.0 | 61  |
| Week 20           | Placebo        | 59                 | 2.7  | 24.46 | -67 | -10.0 | 6.0    | 16.0 | 89  |
|                   | ATL-962 120 mg | 133                | -1.5 | 19.94 | -50 | -13.0 | 1.0    | 10.0 | 59  |
| Week 24           | Placebo        | 59                 | -0.0 | 22.40 | -80 | -11.0 | 4.0    | 10.0 | 49  |
|                   | ATL-962 120 mg | 130                | 0.9  | 21.75 | -55 | -11.0 | -1.5   | 12.0 | 84  |
| Week 28           | Placebo        | 59                 | 1.7  | 23.65 | -68 | -9.0  | 4.0    | 16.0 | 66  |
|                   | ATL-962 120 mg | 127                | -1.0 | 20.23 | -51 | -15.0 | -1.0   | 13.0 | 49  |
| Week 32           | Placebo        | 58                 | 0.7  | 25.63 | -77 | -13.0 | 0.5    | 19.0 | 52  |
|                   | ATL-962 120 mg | 127                | 1.5  | 17.35 | -46 | -10.0 | 4.0    | 12.0 | 49  |
| Week 36           | Placebo        | 58                 | 2.4  | 21.33 | -68 | -7.0  | 3.5    | 19.0 | 39  |
|                   | ATL-962 120 mg | 124                | 1.9  | 21.07 | -47 | -12.0 | 2.0    | 15.0 | 86  |
| Week 40           | Placebo        | 58                 | 2.2  | 20.23 | -64 | -8.0  | 6.0    | 16.0 | 40  |
|                   | ATL-962 120 mg | 122                | 4.8  | 20.99 | -66 | -9.0  | 4.0    | 17.0 | 74  |
| Week 44           | Placebo        | 58                 | 5.4  | 22.12 | -55 | -7.0  | 5.5    | 21.0 | 76  |
|                   | ATL-962 120 mg | 119                | 1.7  | 21.47 | -55 | -12.0 | -2.0   | 13.0 | 73  |
| Week 48           | Placebo        | 58                 | 2.7  | 25.12 | -63 | -12.0 | 4.5    | 14.0 | 79  |
|                   | ATL-962 120 mg | 119                | 1.0  | 20.60 | -55 | -11.0 | 2.0    | 14.0 | 60  |
| Week 52           | Placebo        | 58                 | 4.1  | 23.95 | -47 | -12.0 | 3.5    | 18.0 | 87  |
|                   | ATL-962 120 mg | 118                | 3.5  | 21.07 | -51 | -10.0 | 2.0    | 15.0 | 92  |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (11/12)

| Vitamin E (mg/dL) |                | Summary Statistics |        |        |       |        |        |        |      |
|-------------------|----------------|--------------------|--------|--------|-------|--------|--------|--------|------|
| Visit             | Treatment      | N                  | Mean   | SD     | Min   | Q1     | Median | Q3     | Max  |
| Week 2            | Placebo        | 65                 | -0.042 | 0.3204 | -1.30 | -0.230 | -0.060 | 0.120  | 0.89 |
|                   | ATL-962 120 mg | 141                | -0.232 | 0.3189 | -1.29 | -0.430 | -0.220 | -0.050 | 0.68 |
| Week 4            | Placebo        | 63                 | 0.031  | 0.2684 | -0.69 | -0.130 | 0.050  | 0.150  | 0.77 |
|                   | ATL-962 120 mg | 138                | -0.207 | 0.3324 | -1.31 | -0.350 | -0.190 | -0.050 | 1.15 |
| Week 8            | Placebo        | 62                 | 0.008  | 0.3190 | -0.88 | -0.130 | 0.040  | 0.190  | 1.19 |
|                   | ATL-962 120 mg | 137                | -0.246 | 0.2922 | -1.25 | -0.380 | -0.230 | -0.080 | 0.73 |
| Week 12           | Placebo        | 59                 | 0.087  | 0.3592 | -0.67 | -0.150 | 0.080  | 0.260  | 1.21 |
|                   | ATL-962 120 mg | 137                | -0.192 | 0.3032 | -1.37 | -0.370 | -0.190 | -0.030 | 0.92 |
| Week 16           | Placebo        | 59                 | 0.080  | 0.2931 | -0.58 | -0.050 | 0.060  | 0.230  | 0.95 |
|                   | ATL-962 120 mg | 134                | -0.191 | 0.3308 | -1.27 | -0.360 | -0.190 | 0.010  | 1.12 |
| Week 20           | Placebo        | 59                 | 0.067  | 0.3513 | -0.64 | -0.170 | 0.080  | 0.260  | 1.13 |
|                   | ATL-962 120 mg | 133                | -0.196 | 0.3543 | -1.72 | -0.420 | -0.200 | 0.030  | 1.16 |
| Week 24           | Placebo        | 59                 | 0.041  | 0.3297 | -0.91 | -0.140 | 0.030  | 0.240  | 0.92 |
|                   | ATL-962 120 mg | 130                | -0.224 | 0.3824 | -1.49 | -0.410 | -0.255 | 0.000  | 0.99 |
| Week 28           | Placebo        | 59                 | -0.040 | 0.3896 | -1.01 | -0.330 | -0.040 | 0.180  | 1.06 |
|                   | ATL-962 120 mg | 127                | -0.289 | 0.4125 | -1.91 | -0.440 | -0.250 | -0.090 | 1.33 |
| Week 32           | Placebo        | 58                 | -0.056 | 0.3213 | -0.68 | -0.300 | -0.100 | 0.140  | 0.72 |
|                   | ATL-962 120 mg | 127                | -0.290 | 0.4110 | -1.97 | -0.500 | -0.270 | -0.090 | 1.09 |
| Week 36           | Placebo        | 58                 | -0.108 | 0.3666 | -0.72 | -0.360 | -0.115 | 0.080  | 1.33 |
|                   | ATL-962 120 mg | 124                | -0.319 | 0.3738 | -1.69 | -0.500 | -0.330 | -0.080 | 1.15 |
| Week 40           | Placebo        | 58                 | -0.141 | 0.2710 | -0.79 | -0.300 | -0.135 | 0.010  | 0.49 |
|                   | ATL-962 120 mg | 122                | -0.347 | 0.3888 | -1.74 | -0.530 | -0.315 | -0.100 | 0.76 |
| Week 44           | Placebo        | 58                 | -0.143 | 0.3451 | -0.89 | -0.270 | -0.130 | 0.040  | 0.84 |
|                   | ATL-962 120 mg | 119                | -0.367 | 0.3881 | -1.79 | -0.510 | -0.330 | -0.170 | 0.79 |
| Week 48           | Placebo        | 58                 | -0.137 | 0.3112 | -1.11 | -0.290 | -0.155 | 0.030  | 0.53 |
|                   | ATL-962 120 mg | 119                | -0.399 | 0.3707 | -2.18 | -0.570 | -0.350 | -0.190 | 0.41 |
| Week 52           | Placebo        | 58                 | -0.092 | 0.3403 | -0.96 | -0.290 | -0.110 | 0.060  | 1.41 |
|                   | ATL-962 120 mg | 118                | -0.378 | 0.3696 | -1.77 | -0.550 | -0.365 | -0.130 | 0.38 |

| β-Carotene (μg/dL) |                | Summary Statistics |       |        |       |       |        |      |      |
|--------------------|----------------|--------------------|-------|--------|-------|-------|--------|------|------|
| Visit              | Treatment      | N                  | Mean  | SD     | Min   | Q1    | Median | Q3   | Max  |
| Week 2             | Placebo        | 64                 | -0.38 | 12.826 | -84.0 | -1.70 | 0.00   | 2.90 | 27.0 |
|                    | ATL-962 120 mg | 141                | -1.22 | 7.247  | -26.0 | -4.40 | -0.60  | 1.30 | 30.2 |
| Week 4             | Placebo        | 63                 | -0.44 | 11.720 | -72.4 | -1.80 | 0.40   | 4.10 | 12.6 |
|                    | ATL-962 120 mg | 138                | -0.80 | 7.950  | -30.1 | -4.00 | -0.30  | 2.20 | 27.9 |
| Week 8             | Placebo        | 62                 | -0.10 | 15.450 | -77.4 | -3.10 | 0.00   | 6.00 | 43.5 |
|                    | ATL-962 120 mg | 137                | -0.40 | 8.762  | -34.4 | -3.70 | 0.00   | 2.60 | 33.1 |
| Week 12            | Placebo        | 59                 | -0.54 | 17.846 | -73.6 | -1.90 | 0.30   | 3.60 | 46.3 |
|                    | ATL-962 120 mg | 137                | -0.21 | 10.494 | -44.2 | -4.50 | -0.30  | 3.90 | 45.5 |
| Week 16            | Placebo        | 59                 | -0.10 | 16.323 | -71.0 | -1.90 | 1.10   | 5.70 | 54.7 |
|                    | ATL-962 120 mg | 134                | 0.34  | 11.272 | -36.4 | -5.10 | 0.00   | 3.40 | 62.9 |
| Week 20            | Placebo        | 59                 | -0.38 | 16.624 | -72.4 | -2.10 | 0.10   | 3.50 | 38.3 |
|                    | ATL-962 120 mg | 133                | -0.33 | 10.183 | -42.9 | -4.30 | -0.40  | 3.30 | 53.7 |
| Week 24            | Placebo        | 59                 | -0.04 | 15.107 | -72.7 | -3.30 | 1.10   | 5.80 | 36.1 |
|                    | ATL-962 120 mg | 130                | -0.11 | 10.305 | -49.9 | -4.50 | -0.25  | 3.70 | 45.4 |
| Week 28            | Placebo        | 59                 | 2.64  | 15.664 | -72.2 | -2.00 | 1.70   | 6.90 | 43.9 |
|                    | ATL-962 120 mg | 127                | -1.12 | 10.128 | -27.3 | -6.90 | -1.80  | 2.60 | 53.8 |
| Week 32            | Placebo        | 58                 | 3.90  | 18.275 | -73.1 | -1.80 | 2.00   | 7.40 | 60.6 |
|                    | ATL-962 120 mg | 127                | -0.28 | 11.871 | -44.4 | -5.10 | -0.20  | 2.20 | 45.1 |
| Week 36            | Placebo        | 58                 | -1.10 | 14.781 | -72.8 | -2.50 | 0.15   | 3.40 | 27.7 |
|                    | ATL-962 120 mg | 124                | -1.69 | 12.292 | -49.3 | -6.35 | -0.95  | 2.10 | 44.6 |
| Week 40            | Placebo        | 58                 | -1.93 | 13.970 | -76.8 | -3.50 | -0.50  | 3.90 | 30.3 |
|                    | ATL-962 120 mg | 122                | -1.62 | 13.179 | -52.2 | -5.80 | -0.50  | 2.80 | 71.7 |
| Week 44            | Placebo        | 58                 | -1.57 | 14.498 | -65.2 | -3.40 | -0.55  | 3.80 | 30.8 |
|                    | ATL-962 120 mg | 119                | -1.90 | 12.274 | -56.4 | -5.90 | -0.90  | 3.20 | 39.3 |
| Week 48            | Placebo        | 58                 | -1.60 | 14.983 | -77.9 | -2.60 | -0.10  | 3.40 | 20.9 |
|                    | ATL-962 120 mg | 119                | -1.83 | 12.273 | -45.4 | -6.30 | -1.00  | 4.20 | 42.7 |
| Week 52            | Placebo        | 58                 | -2.30 | 13.633 | -77.0 | -2.40 | -0.30  | 2.80 | 15.6 |
|                    | ATL-962 120 mg | 118                | -2.05 | 11.522 | -43.6 | -6.70 | -0.95  | 2.30 | 41.4 |

表 2.7.4.7-付録-20 血液生化学検査の評価時点別の変化量 (CCT-002 試験) (12/12)

| m-AST (IU/L) |                | Summary Statistics |      |      |     |      |        |     |     |
|--------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit        | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2       | Placebo        | 65                 | -0.3 | 1.85 | -10 | -1.0 | 0.0    | 0.0 | 3   |
|              | ATL-962 120 mg | 141                | 0.4  | 1.79 | -4  | -1.0 | 0.0    | 1.0 | 7   |
| Week 4       | Placebo        | 63                 | -0.1 | 2.31 | -9  | -1.0 | 0.0    | 1.0 | 9   |
|              | ATL-962 120 mg | 138                | 0.1  | 1.94 | -6  | -1.0 | 0.0    | 1.0 | 11  |
| Week 8       | Placebo        | 62                 | -0.5 | 2.14 | -10 | -1.0 | 0.0    | 1.0 | 2   |
|              | ATL-962 120 mg | 137                | 0.1  | 2.19 | -7  | -1.0 | 0.0    | 1.0 | 15  |
| Week 12      | Placebo        | 59                 | -0.4 | 1.92 | -10 | -1.0 | 0.0    | 1.0 | 4   |
|              | ATL-962 120 mg | 137                | 0.5  | 6.88 | -6  | -1.0 | 0.0    | 1.0 | 77  |
| Week 16      | Placebo        | 59                 | -0.4 | 2.20 | -9  | -1.0 | 0.0    | 1.0 | 7   |
|              | ATL-962 120 mg | 134                | 0.0  | 2.67 | -7  | -1.0 | 0.0    | 1.0 | 22  |
| Week 20      | Placebo        | 59                 | -0.3 | 1.77 | -7  | -1.0 | 0.0    | 1.0 | 5   |
|              | ATL-962 120 mg | 133                | -0.4 | 1.79 | -6  | -1.0 | 0.0    | 1.0 | 6   |
| Week 24      | Placebo        | 59                 | -0.1 | 2.38 | -9  | -1.0 | 0.0    | 1.0 | 12  |
|              | ATL-962 120 mg | 130                | -0.1 | 1.96 | -7  | -1.0 | 0.0    | 1.0 | 7   |
| Week 28      | Placebo        | 59                 | 0.0  | 2.01 | -10 | -1.0 | 0.0    | 1.0 | 6   |
|              | ATL-962 120 mg | 127                | 0.1  | 1.94 | -6  | -1.0 | 0.0    | 1.0 | 8   |
| Week 32      | Placebo        | 58                 | -0.1 | 2.88 | -8  | -1.0 | 0.0    | 1.0 | 16  |
|              | ATL-962 120 mg | 127                | -0.1 | 2.23 | -8  | -1.0 | 0.0    | 1.0 | 14  |
| Week 36      | Placebo        | 58                 | 0.1  | 1.91 | -7  | -1.0 | 0.0    | 1.0 | 4   |
|              | ATL-962 120 mg | 124                | 0.2  | 2.94 | -7  | -1.0 | 0.0    | 1.0 | 25  |
| Week 40      | Placebo        | 58                 | 0.1  | 1.94 | -8  | -1.0 | 0.0    | 1.0 | 5   |
|              | ATL-962 120 mg | 122                | 0.3  | 3.17 | -6  | -1.0 | 0.0    | 1.0 | 27  |
| Week 44      | Placebo        | 58                 | 0.2  | 1.84 | -5  | -1.0 | 0.0    | 1.0 | 5   |
|              | ATL-962 120 mg | 119                | 0.1  | 1.80 | -7  | -1.0 | 0.0    | 1.0 | 5   |
| Week 48      | Placebo        | 58                 | 0.3  | 2.21 | -9  | 0.0  | 0.0    | 1.0 | 8   |
|              | ATL-962 120 mg | 119                | 0.1  | 2.13 | -5  | -1.0 | 0.0    | 1.0 | 8   |
| Week 52      | Placebo        | 58                 | 0.4  | 2.27 | -7  | -1.0 | 0.0    | 2.0 | 8   |
|              | ATL-962 120 mg | 118                | 0.4  | 2.27 | -6  | -1.0 | 0.0    | 1.0 | 8   |

| CK18 (Segment) (U/L) |                | Summary Statistics |        |         |        |         |        |        |       |
|----------------------|----------------|--------------------|--------|---------|--------|---------|--------|--------|-------|
| Visit                | Treatment      | N                  | Mean   | SD      | Min    | Q1      | Median | Q3     | Max   |
| Week 2               | Placebo        | 65                 | -16.93 | 151.125 | -502.3 | -68.00  | 0.00   | 59.80  | 472.7 |
|                      | ATL-962 120 mg | 141                | -25.67 | 145.097 | -492.6 | -78.20  | -13.70 | 28.60  | 469.8 |
| Week 4               | Placebo        | 63                 | -17.24 | 132.047 | -449.9 | -73.80  | -2.60  | 44.20  | 437.7 |
|                      | ATL-962 120 mg | 138                | -33.51 | 150.838 | -599.9 | -82.50  | -25.20 | 33.60  | 577.8 |
| Week 8               | Placebo        | 62                 | -39.82 | 160.421 | -785.6 | -85.00  | -12.05 | 47.40  | 403.2 |
|                      | ATL-962 120 mg | 137                | -62.28 | 161.660 | -697.5 | -127.80 | -43.40 | 9.60   | 641.2 |
| Week 12              | Placebo        | 59                 | 15.46  | 152.811 | -348.2 | -89.50  | 25.70  | 109.20 | 392.0 |
|                      | ATL-962 120 mg | 137                | -60.16 | 158.202 | -526.1 | -108.10 | -41.10 | 13.30  | 641.2 |
| Week 16              | Placebo        | 59                 | 15.16  | 148.862 | -551.9 | -58.40  | 3.00   | 85.70  | 395.4 |
|                      | ATL-962 120 mg | 134                | -50.70 | 156.115 | -595.6 | -98.30  | -32.70 | 20.60  | 590.0 |
| Week 20              | Placebo        | 59                 | 5.90   | 126.265 | -332.1 | -58.50  | 18.30  | 85.60  | 342.9 |
|                      | ATL-962 120 mg | 133                | -52.17 | 172.792 | -681.4 | -106.50 | -28.40 | 18.30  | 807.3 |
| Week 24              | Placebo        | 59                 | 24.59  | 169.610 | -380.0 | -49.50  | 17.50  | 103.40 | 638.2 |
|                      | ATL-962 120 mg | 130                | -61.54 | 172.148 | -611.5 | -129.50 | -24.25 | 28.00  | 807.3 |
| Week 28              | Placebo        | 59                 | 18.51  | 113.687 | -288.9 | -65.50  | 15.40  | 74.40  | 401.6 |
|                      | ATL-962 120 mg | 127                | -64.49 | 192.862 | -670.3 | -142.50 | -41.50 | 18.80  | 807.3 |
| Week 32              | Placebo        | 58                 | 15.15  | 150.364 | -421.1 | -69.70  | 0.00   | 84.20  | 638.2 |
|                      | ATL-962 120 mg | 127                | -56.90 | 203.261 | -716.0 | -124.60 | -30.70 | 21.60  | 807.3 |
| Week 36              | Placebo        | 58                 | -2.89  | 175.312 | -466.1 | -74.50  | 0.00   | 83.10  | 694.9 |
|                      | ATL-962 120 mg | 124                | -50.63 | 204.538 | -682.2 | -123.00 | -35.00 | 23.90  | 760.0 |
| Week 40              | Placebo        | 58                 | 6.44   | 131.139 | -339.2 | -61.90  | 0.00   | 73.60  | 379.6 |
|                      | ATL-962 120 mg | 122                | -57.23 | 209.797 | -772.6 | -132.20 | -41.15 | 25.40  | 807.3 |
| Week 44              | Placebo        | 58                 | -14.19 | 149.117 | -442.3 | -106.80 | -4.70  | 60.30  | 524.4 |
|                      | ATL-962 120 mg | 119                | -79.52 | 206.743 | -761.0 | -128.10 | -48.40 | 4.00   | 807.3 |
| Week 48              | Placebo        | 58                 | -29.32 | 173.881 | -554.4 | -88.20  | -17.40 | 39.30  | 694.9 |
|                      | ATL-962 120 mg | 119                | -86.09 | 216.485 | -709.5 | -148.70 | -63.50 | -3.80  | 772.9 |
| Week 52              | Placebo        | 58                 | -57.19 | 195.288 | -855.6 | -123.50 | -31.35 | 24.80  | 442.8 |
|                      | ATL-962 120 mg | 118                | -92.77 | 200.426 | -668.6 | -166.10 | -56.10 | -3.30  | 807.3 |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験)(1/24)

|               |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|               |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|               |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Total Protein | Week 2  | Lower   | 1      | -     | -     | 1              | 1      | 2     | -     | 3   |
|               |         | Normal  | 1      | 60    | 2     | 63             | 3      | 133   | 1     | 137 |
|               |         | Upper   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|               |         | Total   | 2      | 61    | 2     | 65             | 4      | 136   | 1     | 141 |
|               | Week 4  | Lower   | 1      | -     | -     | 1              | 1      | 2     | -     | 3   |
|               |         | Normal  | 1      | 59    | -     | 60             | 3      | 127   | -     | 130 |
|               |         | Upper   | -      | -     | 2     | 2              | -      | 4     | 1     | 5   |
|               |         | Total   | 2      | 59    | 2     | 63             | 4      | 133   | 1     | 138 |
|               | Week 8  | Lower   | -      | 1     | -     | 1              | 2      | 2     | -     | 4   |
|               |         | Normal  | 2      | 57    | 2     | 61             | 2      | 128   | 1     | 131 |
|               |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|               |         | Total   | 2      | 58    | 2     | 62             | 4      | 132   | 1     | 137 |
|               | Week 12 | Lower   | 1      | 2     | -     | 3              | 2      | 3     | -     | 5   |
|               |         | Normal  | 1      | 52    | 2     | 55             | 2      | 128   | 1     | 131 |
|               |         | Upper   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|               |         | Total   | 2      | 55    | 2     | 59             | 4      | 132   | 1     | 137 |
|               | Week 16 | Lower   | 1      | -     | -     | 1              | 3      | 7     | -     | 10  |
|               |         | Normal  | 1      | 53    | 2     | 56             | 1      | 122   | -     | 123 |
|               |         | Upper   | -      | 2     | -     | 2              | -      | -     | 1     | 1   |
|               |         | Total   | 2      | 55    | 2     | 59             | 4      | 129   | 1     | 134 |
|               | Week 20 | Lower   | 1      | 1     | -     | 2              | 1      | 5     | -     | 6   |
|               |         | Normal  | 1      | 54    | 2     | 57             | 3      | 123   | 1     | 127 |
|               |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|               |         | Total   | 2      | 55    | 2     | 59             | 4      | 128   | 1     | 133 |
|               | Week 24 | Lower   | 1      | 2     | -     | 3              | 3      | 8     | -     | 11  |
|               |         | Normal  | 1      | 53    | 2     | 56             | 1      | 117   | 1     | 119 |
|               |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|               |         | Total   | 2      | 55    | 2     | 59             | 4      | 125   | 1     | 130 |
|               | Week 28 | Lower   | 2      | 1     | -     | 3              | 2      | 4     | -     | 6   |
|               |         | Normal  | -      | 54    | 2     | 56             | 2      | 117   | 1     | 120 |
|               |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|               |         | Total   | 2      | 55    | 2     | 59             | 4      | 122   | 1     | 127 |
|               | Week 32 | Lower   | 1      | 3     | -     | 4              | 2      | 6     | -     | 8   |
|               |         | Normal  | 1      | 51    | 2     | 54             | 2      | 114   | 1     | 117 |
|               |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|               |         | Total   | 2      | 54    | 2     | 58             | 4      | 122   | 1     | 127 |
|               | Week 36 | Lower   | 1      | 3     | -     | 4              | 2      | 5     | -     | 7   |
|               |         | Normal  | 1      | 51    | 2     | 54             | 2      | 114   | 1     | 117 |
|               |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|               |         | Total   | 2      | 54    | 2     | 58             | 4      | 119   | 1     | 124 |
|               | Week 40 | Lower   | 1      | 2     | -     | 3              | 3      | 6     | -     | 9   |
|               |         | Normal  | 1      | 52    | 2     | 55             | 1      | 110   | 1     | 112 |
|               |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|               |         | Total   | 2      | 54    | 2     | 58             | 4      | 117   | 1     | 122 |
|               | Week 44 | Lower   | 1      | 1     | -     | 2              | 3      | 5     | -     | 8   |
|               |         | Normal  | 1      | 52    | 1     | 54             | 1      | 107   | 1     | 109 |
|               |         | Upper   | -      | 1     | 1     | 2              | -      | 2     | -     | 2   |
|               |         | Total   | 2      | 54    | 2     | 58             | 4      | 114   | 1     | 119 |
| Week 48       | Lower   | 1       | 3      | -     | 4     | 2              | 9      | -     | 11    |     |
|               | Normal  | 1       | 51     | -     | 52    | 2              | 101    | 1     | 104   |     |
|               | Upper   | -       | -      | 2     | 2     | -              | 4      | -     | 4     |     |
|               | Total   | 2       | 54     | 2     | 58    | 4              | 114    | 1     | 119   |     |
| Week 52       | Lower   | 1       | 3      | -     | 4     | 1              | 8      | -     | 9     |     |
|               | Normal  | 1       | 50     | 2     | 53    | 3              | 103    | 1     | 107   |     |
|               | Upper   | -       | 1      | -     | 1     | -              | 2      | -     | 2     |     |
|               | Total   | 2       | 54     | 2     | 58    | 4              | 113    | 1     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験) (2/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Albumin | Week 2  | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 65    | -     | 65             | -      | 140   | -     | 140 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 65    | -     | 65             | -      | 141   | -     | 141 |
|         | Week 4  | Lower   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|         |         | Normal  | -      | 62    | -     | 62             | -      | 138   | -     | 138 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 63    | -     | 63             | -      | 138   | -     | 138 |
|         | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 62    | -     | 62             | -      | 137   | -     | 137 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 62    | -     | 62             | -      | 137   | -     | 137 |
|         | Week 12 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 136   | -     | 136 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 137   | -     | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 132   | -     | 132 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 134   | -     | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 132   | -     | 132 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 133   | -     | 133 |
|         | Week 24 | Lower   | -      | -     | -     | -              | -      | 4     | -     | 4   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 126   | -     | 126 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 130   | -     | 130 |
|         | Week 28 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 126   | -     | 126 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 127   | -     | 127 |
|         | Week 32 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 58    | -     | 58             | -      | 126   | -     | 126 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 127   | -     | 127 |
|         | Week 36 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 58    | -     | 58             | -      | 123   | -     | 123 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 124   | -     | 124 |
| Week 40 | Lower   | -       | 1      | -     | 1     | -              | 1      | -     | 1     |     |
|         | Normal  | -       | 57     | -     | 57    | -              | 121    | -     | 121   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | -       | 58     | -     | 58    | -              | 122    | -     | 122   |     |
| Week 44 | Lower   | -       | -      | -     | -     | -              | 2      | -     | 2     |     |
|         | Normal  | -       | 58     | -     | 58    | -              | 117    | -     | 117   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | -       | 58     | -     | 58    | -              | 119    | -     | 119   |     |
| Week 48 | Lower   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|         | Normal  | -       | 58     | -     | 58    | -              | 117    | -     | 117   |     |
|         | Upper   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|         | Total   | -       | 58     | -     | 58    | -              | 119    | -     | 119   |     |
| Week 52 | Lower   | -       | -      | -     | -     | -              | 2      | -     | 2     |     |
|         | Normal  | -       | 58     | -     | 58    | -              | 115    | -     | 115   |     |
|         | Upper   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|         | Total   | -       | 58     | -     | 58    | -              | 118    | -     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験)(3/24)

|           |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-----------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|           |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|           |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| ALT (GPT) | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|           |         | Normal  | -      | 43    | 3     | 46             | -      | 74    | 12    | 86  |
|           |         | Upper   | -      | 4     | 15    | 19             | -      | 11    | 44    | 55  |
|           |         | Total   | -      | 47    | 18    | 65             | -      | 85    | 56    | 141 |
|           | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 45    | 2     | 47             | -      | 69    | 12    | 81  |
|           |         | Upper   | -      | 1     | 15    | 16             | -      | 14    | 43    | 57  |
|           |         | Total   | -      | 46    | 17    | 63             | -      | 83    | 55    | 138 |
|           | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 42    | 2     | 44             | -      | 69    | 17    | 86  |
|           |         | Upper   | -      | 3     | 15    | 18             | -      | 13    | 38    | 51  |
|           |         | Total   | -      | 45    | 17    | 62             | -      | 82    | 55    | 137 |
|           | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 37    | 3     | 40             | -      | 71    | 21    | 92  |
|           |         | Upper   | -      | 7     | 12    | 19             | -      | 11    | 34    | 45  |
|           |         | Total   | -      | 44    | 15    | 59             | -      | 82    | 55    | 137 |
|           | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 38    | 4     | 42             | -      | 64    | 24    | 88  |
|           |         | Upper   | -      | 6     | 11    | 17             | -      | 15    | 31    | 46  |
|           |         | Total   | -      | 44    | 15    | 59             | -      | 79    | 55    | 134 |
|           | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 36    | 4     | 40             | -      | 64    | 24    | 88  |
|           |         | Upper   | -      | 8     | 11    | 19             | -      | 14    | 31    | 45  |
|           |         | Total   | -      | 44    | 15    | 59             | -      | 78    | 55    | 133 |
|           | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 38    | 5     | 43             | -      | 66    | 21    | 87  |
|           |         | Upper   | -      | 6     | 10    | 16             | -      | 10    | 33    | 43  |
|           |         | Total   | -      | 44    | 15    | 59             | -      | 76    | 54    | 130 |
|           | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 37    | 4     | 41             | -      | 63    | 22    | 85  |
|           |         | Upper   | -      | 7     | 11    | 18             | -      | 12    | 30    | 42  |
|           |         | Total   | -      | 44    | 15    | 59             | -      | 75    | 52    | 127 |
|           | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 37    | 4     | 41             | -      | 65    | 24    | 89  |
|           |         | Upper   | -      | 6     | 11    | 17             | -      | 10    | 28    | 38  |
|           |         | Total   | -      | 43    | 15    | 58             | -      | 75    | 52    | 127 |
|           | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 38    | 2     | 40             | -      | 58    | 24    | 82  |
|           |         | Upper   | -      | 5     | 13    | 18             | -      | 15    | 27    | 42  |
|           |         | Total   | -      | 43    | 15    | 58             | -      | 73    | 51    | 124 |
|           | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 36    | 3     | 39             | -      | 62    | 19    | 81  |
|           |         | Upper   | -      | 7     | 12    | 19             | -      | 9     | 32    | 41  |
|           |         | Total   | -      | 43    | 15    | 58             | -      | 71    | 51    | 122 |
|           | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 36    | 3     | 39             | -      | 57    | 21    | 78  |
|           |         | Upper   | -      | 7     | 12    | 19             | -      | 12    | 29    | 41  |
|           |         | Total   | -      | 43    | 15    | 58             | -      | 69    | 50    | 119 |
| Week 48   | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|           | Normal  | -       | 38     | 2     | 40    | -              | 61     | 27    | 88    |     |
|           | Upper   | -       | 5      | 13    | 18    | -              | 8      | 23    | 31    |     |
|           | Total   | -       | 43     | 15    | 58    | -              | 69     | 50    | 119   |     |
| Week 52   | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|           | Normal  | -       | 38     | 2     | 40    | -              | 64     | 24    | 88    |     |
|           | Upper   | -       | 5      | 13    | 18    | -              | 4      | 26    | 30    |     |
|           | Total   | -       | 43     | 15    | 58    | -              | 68     | 50    | 118   |     |



表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験)(4/24)

|           |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-----------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|           |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|           |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| AST (GOT) | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|           |         | Normal  | -      | 52    | 3     | 55             | -      | 103   | 10    | 113 |
|           |         | Upper   | -      | 1     | 9     | 10             | -      | 13    | 15    | 28  |
|           |         | Total   | -      | 53    | 12    | 65             | -      | 116   | 25    | 141 |
|           | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 48    | 4     | 52             | -      | 106   | 14    | 120 |
|           |         | Upper   | -      | 4     | 7     | 11             | -      | 7     | 11    | 18  |
|           |         | Total   | -      | 52    | 11    | 63             | -      | 113   | 25    | 138 |
|           | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 48    | 4     | 52             | -      | 101   | 16    | 117 |
|           |         | Upper   | -      | 3     | 7     | 10             | -      | 11    | 9     | 20  |
|           |         | Total   | -      | 51    | 11    | 62             | -      | 112   | 25    | 137 |
|           | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 45    | 2     | 47             | -      | 105   | 14    | 119 |
|           |         | Upper   | -      | 5     | 7     | 12             | -      | 7     | 11    | 18  |
|           |         | Total   | -      | 50    | 9     | 59             | -      | 112   | 25    | 137 |
|           | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 44    | 4     | 48             | -      | 104   | 19    | 123 |
|           |         | Upper   | -      | 6     | 5     | 11             | -      | 5     | 6     | 11  |
|           |         | Total   | -      | 50    | 9     | 59             | -      | 109   | 25    | 134 |
|           | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 44    | 2     | 46             | -      | 103   | 19    | 122 |
|           |         | Upper   | -      | 6     | 7     | 13             | -      | 5     | 6     | 11  |
|           |         | Total   | -      | 50    | 9     | 59             | -      | 108   | 25    | 133 |
|           | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 46    | 4     | 50             | -      | 99    | 14    | 113 |
|           |         | Upper   | -      | 4     | 5     | 9              | -      | 7     | 10    | 17  |
|           |         | Total   | -      | 50    | 9     | 59             | -      | 106   | 24    | 130 |
|           | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 44    | 1     | 45             | -      | 97    | 17    | 114 |
|           |         | Upper   | -      | 6     | 8     | 14             | -      | 7     | 6     | 13  |
|           |         | Total   | -      | 50    | 9     | 59             | -      | 104   | 23    | 127 |
|           | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 46    | 3     | 49             | -      | 95    | 17    | 112 |
|           |         | Upper   | -      | 3     | 6     | 9              | -      | 9     | 6     | 15  |
|           |         | Total   | -      | 49    | 9     | 58             | -      | 104   | 23    | 127 |
|           | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 46    | 3     | 49             | -      | 96    | 14    | 110 |
|           |         | Upper   | -      | 3     | 6     | 9              | -      | 6     | 8     | 14  |
|           |         | Total   | -      | 49    | 9     | 58             | -      | 102   | 22    | 124 |
|           | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 47    | 2     | 49             | -      | 95    | 14    | 109 |
|           |         | Upper   | -      | 2     | 7     | 9              | -      | 5     | 8     | 13  |
|           |         | Total   | -      | 49    | 9     | 58             | -      | 100   | 22    | 122 |
|           | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 42    | 1     | 43             | -      | 92    | 15    | 107 |
|           |         | Upper   | -      | 7     | 8     | 15             | -      | 5     | 7     | 12  |
|           |         | Total   | -      | 49    | 9     | 58             | -      | 97    | 22    | 119 |
| Week 48   | Lower   | -       | 1      | -     | 1     | -              | -      | -     | -     |     |
|           | Normal  | -       | 40     | 4     | 44    | -              | 87     | 16    | 103   |     |
|           | Upper   | -       | 8      | 5     | 13    | -              | 10     | 6     | 16    |     |
|           | Total   | -       | 49     | 9     | 58    | -              | 97     | 22    | 119   |     |
| Week 52   | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|           | Normal  | -       | 44     | 5     | 49    | -              | 88     | 12    | 100   |     |
|           | Upper   | -       | 5      | 4     | 9     | -              | 8      | 10    | 18    |     |
|           | Total   | -       | 49     | 9     | 58    | -              | 96     | 22    | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験)(5/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| γ-GTP   | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|         |         | Normal  | -      | 45    | 2     | 47             | -      | 88    | 6     | 94  |
|         |         | Upper   | -      | -     | 18    | 18             | -      | 4     | 43    | 47  |
|         |         | Total   | -      | 45    | 20    | 65             | -      | 92    | 49    | 141 |
|         | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 44    | 2     | 46             | -      | 82    | 11    | 93  |
|         |         | Upper   | -      | 1     | 16    | 17             | -      | 7     | 38    | 45  |
|         |         | Total   | -      | 45    | 18    | 63             | -      | 89    | 49    | 138 |
|         | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 43    | 4     | 47             | -      | 83    | 11    | 94  |
|         |         | Upper   | -      | 1     | 14    | 15             | -      | 5     | 38    | 43  |
|         |         | Total   | -      | 44    | 18    | 62             | -      | 88    | 49    | 137 |
|         | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 41    | 3     | 44             | -      | 83    | 10    | 93  |
|         |         | Upper   | -      | 1     | 14    | 15             | -      | 5     | 39    | 44  |
|         |         | Total   | -      | 42    | 17    | 59             | -      | 88    | 49    | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 41    | 3     | 44             | -      | 82    | 12    | 94  |
|         |         | Upper   | -      | 1     | 14    | 15             | -      | 4     | 36    | 40  |
|         |         | Total   | -      | 42    | 17    | 59             | -      | 86    | 48    | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 41    | 3     | 44             | -      | 79    | 14    | 93  |
|         |         | Upper   | -      | 1     | 14    | 15             | -      | 6     | 34    | 40  |
|         |         | Total   | -      | 42    | 17    | 59             | -      | 85    | 48    | 133 |
|         | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 41    | 4     | 45             | -      | 76    | 14    | 90  |
|         |         | Upper   | -      | 1     | 13    | 14             | -      | 7     | 33    | 40  |
|         |         | Total   | -      | 42    | 17    | 59             | -      | 83    | 47    | 130 |
|         | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 40    | 4     | 44             | -      | 74    | 13    | 87  |
|         |         | Upper   | -      | 2     | 13    | 15             | -      | 7     | 33    | 40  |
|         |         | Total   | -      | 42    | 17    | 59             | -      | 81    | 46    | 127 |
|         | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 41    | 4     | 45             | -      | 75    | 15    | 90  |
|         |         | Upper   | -      | -     | 13    | 13             | -      | 6     | 31    | 37  |
|         |         | Total   | -      | 41    | 17    | 58             | -      | 81    | 46    | 127 |
|         | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 41    | 3     | 44             | -      | 68    | 13    | 81  |
|         |         | Upper   | -      | -     | 14    | 14             | -      | 11    | 32    | 43  |
|         |         | Total   | -      | 41    | 17    | 58             | -      | 79    | 45    | 124 |
|         | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 41    | 4     | 45             | -      | 70    | 10    | 80  |
|         |         | Upper   | -      | -     | 13    | 13             | -      | 7     | 35    | 42  |
|         |         | Total   | -      | 41    | 17    | 58             | -      | 77    | 45    | 122 |
|         | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 38    | 5     | 43             | -      | 68    | 12    | 80  |
|         |         | Upper   | -      | 3     | 12    | 15             | -      | 7     | 32    | 39  |
|         |         | Total   | -      | 41    | 17    | 58             | -      | 75    | 44    | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 38     | 6     | 44    | -              | 68     | 12    | 80    |     |
|         | Upper   | -       | 3      | 11    | 14    | -              | 7      | 32    | 39    |     |
|         | Total   | -       | 41     | 17    | 58    | -              | 75     | 44    | 119   |     |
| Week 52 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 39     | 6     | 45    | -              | 68     | 15    | 83    |     |
|         | Upper   | -       | 2      | 11    | 13    | -              | 6      | 29    | 35    |     |
|         | Total   | -       | 41     | 17    | 58    | -              | 74     | 44    | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験)(6/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| ALP     | Week 2  | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 57    | 2     | 59             | -      | 118   | 6     | 124 |
|         |         | Upper   | -      | 2     | 4     | 6              | -      | 5     | 11    | 16  |
|         |         | Total   | -      | 59    | 6     | 65             | 1      | 123   | 17    | 141 |
|         | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 55    | 2     | 57             | -      | 118   | 4     | 122 |
|         |         | Upper   | -      | 2     | 4     | 6              | -      | 4     | 12    | 16  |
|         |         | Total   | -      | 57    | 6     | 63             | -      | 122   | 16    | 138 |
|         | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 55    | 1     | 56             | -      | 119   | 9     | 128 |
|         |         | Upper   | -      | 1     | 5     | 6              | -      | 2     | 7     | 9   |
|         |         | Total   | -      | 56    | 6     | 62             | -      | 121   | 16    | 137 |
|         | Week 12 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 52    | 3     | 55             | -      | 116   | 9     | 125 |
|         |         | Upper   | -      | 1     | 3     | 4              | -      | 4     | 7     | 11  |
|         |         | Total   | -      | 53    | 6     | 59             | -      | 121   | 16    | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 50    | 4     | 54             | -      | 117   | 8     | 125 |
|         |         | Upper   | -      | 3     | 2     | 5              | -      | 1     | 8     | 9   |
|         |         | Total   | -      | 53    | 6     | 59             | -      | 118   | 16    | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 51    | 2     | 53             | -      | 115   | 9     | 124 |
|         |         | Upper   | -      | 2     | 4     | 6              | -      | 2     | 7     | 9   |
|         |         | Total   | -      | 53    | 6     | 59             | -      | 117   | 16    | 133 |
|         | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 47    | 3     | 50             | -      | 110   | 7     | 117 |
|         |         | Upper   | -      | 6     | 3     | 9              | -      | 4     | 9     | 13  |
|         |         | Total   | -      | 53    | 6     | 59             | -      | 114   | 16    | 130 |
|         | Week 28 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 52    | 2     | 54             | -      | 107   | 5     | 112 |
|         |         | Upper   | -      | 1     | 4     | 5              | -      | 3     | 11    | 14  |
|         |         | Total   | -      | 53    | 6     | 59             | -      | 111   | 16    | 127 |
|         | Week 32 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 48    | 2     | 50             | -      | 105   | 7     | 112 |
|         |         | Upper   | -      | 4     | 4     | 8              | -      | 5     | 9     | 14  |
|         |         | Total   | -      | 52    | 6     | 58             | -      | 111   | 16    | 127 |
|         | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 47    | 3     | 50             | -      | 103   | 6     | 109 |
|         |         | Upper   | -      | 5     | 3     | 8              | -      | 5     | 10    | 15  |
|         |         | Total   | -      | 52    | 6     | 58             | -      | 108   | 16    | 124 |
|         | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 50    | 3     | 53             | -      | 100   | 6     | 106 |
|         |         | Upper   | -      | 2     | 3     | 5              | -      | 7     | 9     | 16  |
|         |         | Total   | -      | 52    | 6     | 58             | -      | 107   | 15    | 122 |
|         | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 49    | 4     | 53             | -      | 100   | 7     | 107 |
|         |         | Upper   | -      | 3     | 2     | 5              | -      | 5     | 7     | 12  |
|         |         | Total   | -      | 52    | 6     | 58             | -      | 105   | 14    | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 47     | 3     | 50    | -              | 101    | 6     | 107   |     |
|         | Upper   | -       | 5      | 3     | 8     | -              | 4      | 8     | 12    |     |
|         | Total   | -       | 52     | 6     | 58    | -              | 105    | 14    | 119   |     |
| Week 52 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 49     | 2     | 51    | -              | 99     | 4     | 103   |     |
|         | Upper   | -       | 3      | 4     | 7     | -              | 6      | 9     | 15    |     |
|         | Total   | -       | 52     | 6     | 58    | -              | 105    | 13    | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験)(7/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| LDH     | Week 2  | Lower   | -      | -     | -     | -              | 4      | 2     | -     | 6   |
|         |         | Normal  | -      | 63    | 1     | 64             | 1      | 121   | 3     | 125 |
|         |         | Upper   | -      | -     | 1     | 1              | -      | 5     | 5     | 10  |
|         |         | Total   | -      | 63    | 2     | 65             | 5      | 128   | 8     | 141 |
|         | Week 4  | Lower   | -      | -     | -     | -              | 4      | 1     | -     | 5   |
|         |         | Normal  | -      | 59    | -     | 59             | 1      | 124   | 1     | 126 |
|         |         | Upper   | -      | 2     | 2     | 4              | -      | -     | 7     | 7   |
|         |         | Total   | -      | 61    | 2     | 63             | 5      | 125   | 8     | 138 |
|         | Week 8  | Lower   | -      | -     | -     | -              | 1      | 2     | -     | 3   |
|         |         | Normal  | -      | 60    | 1     | 61             | 4      | 117   | 4     | 125 |
|         |         | Upper   | -      | -     | 1     | 1              | -      | 5     | 4     | 9   |
|         |         | Total   | -      | 60    | 2     | 62             | 5      | 124   | 8     | 137 |
|         | Week 12 | Lower   | -      | -     | -     | -              | 4      | -     | -     | 4   |
|         |         | Normal  | -      | 57    | -     | 57             | 1      | 118   | 3     | 122 |
|         |         | Upper   | -      | 1     | 1     | 2              | -      | 6     | 5     | 11  |
|         |         | Total   | -      | 58    | 1     | 59             | 5      | 124   | 8     | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | 3      | -     | -     | 3   |
|         |         | Normal  | -      | 58    | -     | 58             | 2      | 117   | 4     | 123 |
|         |         | Upper   | -      | -     | 1     | 1              | -      | 4     | 4     | 8   |
|         |         | Total   | -      | 58    | 1     | 59             | 5      | 121   | 8     | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | 3      | -     | -     | 3   |
|         |         | Normal  | -      | 58    | -     | 58             | 2      | 116   | 5     | 123 |
|         |         | Upper   | -      | -     | 1     | 1              | -      | 4     | 3     | 7   |
|         |         | Total   | -      | 58    | 1     | 59             | 5      | 120   | 8     | 133 |
|         | Week 24 | Lower   | -      | 2     | -     | 2              | 3      | 1     | -     | 4   |
|         |         | Normal  | -      | 55    | -     | 55             | 2      | 113   | 6     | 121 |
|         |         | Upper   | -      | 1     | 1     | 2              | -      | 3     | 2     | 5   |
|         |         | Total   | -      | 58    | 1     | 59             | 5      | 117   | 8     | 130 |
|         | Week 28 | Lower   | -      | -     | -     | -              | 2      | -     | -     | 2   |
|         |         | Normal  | -      | 57    | -     | 57             | 3      | 110   | 6     | 119 |
|         |         | Upper   | -      | 1     | 1     | 2              | -      | 4     | 2     | 6   |
|         |         | Total   | -      | 58    | 1     | 59             | 5      | 114   | 8     | 127 |
|         | Week 32 | Lower   | -      | -     | -     | -              | 5      | -     | -     | 5   |
|         |         | Normal  | -      | 57    | -     | 57             | -      | 111   | 5     | 116 |
|         |         | Upper   | -      | -     | 1     | 1              | -      | 3     | 3     | 6   |
|         |         | Total   | -      | 57    | 1     | 58             | 5      | 114   | 8     | 127 |
|         | Week 36 | Lower   | -      | -     | -     | -              | 1      | 1     | -     | 2   |
|         |         | Normal  | -      | 56    | -     | 56             | 3      | 109   | 3     | 115 |
|         |         | Upper   | -      | 1     | 1     | 2              | -      | 4     | 3     | 7   |
|         |         | Total   | -      | 57    | 1     | 58             | 4      | 114   | 6     | 124 |
|         | Week 40 | Lower   | -      | -     | -     | -              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 57    | -     | 57             | 3      | 107   | 2     | 112 |
|         |         | Upper   | -      | -     | 1     | 1              | -      | 5     | 4     | 9   |
|         |         | Total   | -      | 57    | 1     | 58             | 4      | 112   | 6     | 122 |
|         | Week 44 | Lower   | -      | -     | -     | -              | 2      | -     | -     | 2   |
|         |         | Normal  | -      | 56    | -     | 56             | 1      | 107   | 3     | 111 |
|         |         | Upper   | -      | 1     | 1     | 2              | 1      | 2     | 3     | 6   |
|         |         | Total   | -      | 57    | 1     | 58             | 4      | 109   | 6     | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | 2              | 1      | -     | 3     |     |
|         | Normal  | -       | 54     | -     | 54    | 2              | 103    | 1     | 106   |     |
|         | Upper   | -       | 3      | 1     | 4     | -              | 5      | 5     | 10    |     |
|         | Total   | -       | 57     | 1     | 58    | 4              | 109    | 6     | 119   |     |
| Week 52 | Lower   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|         | Normal  | -       | 57     | -     | 57    | 3              | 107    | 3     | 113   |     |
|         | Upper   | -       | -      | 1     | 1     | -              | 1      | 3     | 4     |     |
|         | Total   | -       | 57     | 1     | 58    | 3              | 109    | 6     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験)(8/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| CPK     | Week 2  | Lower   | 4      | 3     | -     | 7              | 5      | 6     | -     | 11  |
|         |         | Normal  | 2      | 52    | -     | 54             | 3      | 112   | 9     | 124 |
|         |         | Upper   | -      | 1     | 3     | 4              | -      | 1     | 5     | 6   |
|         |         | Total   | 6      | 56    | 3     | 65             | 8      | 119   | 14    | 141 |
|         | Week 4  | Lower   | 6      | 1     | -     | 7              | 4      | 6     | 1     | 11  |
|         |         | Normal  | -      | 52    | -     | 52             | 4      | 107   | 7     | 118 |
|         |         | Upper   | -      | 1     | 3     | 4              | -      | 3     | 6     | 9   |
|         |         | Total   | 6      | 54    | 3     | 63             | 8      | 116   | 14    | 138 |
|         | Week 8  | Lower   | 4      | 2     | -     | 6              | 4      | 4     | -     | 8   |
|         |         | Normal  | 2      | 51    | -     | 53             | 4      | 106   | 9     | 119 |
|         |         | Upper   | -      | -     | 3     | 3              | -      | 5     | 5     | 10  |
|         |         | Total   | 6      | 53    | 3     | 62             | 8      | 115   | 14    | 137 |
|         | Week 12 | Lower   | 4      | 2     | -     | 6              | 5      | 4     | -     | 9   |
|         |         | Normal  | 1      | 48    | 1     | 50             | 3      | 108   | 8     | 119 |
|         |         | Upper   | -      | 1     | 2     | 3              | -      | 3     | 6     | 9   |
|         |         | Total   | 5      | 51    | 3     | 59             | 8      | 115   | 14    | 137 |
|         | Week 16 | Lower   | 1      | 2     | -     | 3              | 4      | 4     | 1     | 9   |
|         |         | Normal  | 4      | 46    | -     | 50             | 4      | 104   | 8     | 116 |
|         |         | Upper   | -      | 3     | 3     | 6              | -      | 4     | 5     | 9   |
|         |         | Total   | 5      | 51    | 3     | 59             | 8      | 112   | 14    | 134 |
|         | Week 20 | Lower   | 2      | 2     | -     | 4              | 2      | 4     | 1     | 7   |
|         |         | Normal  | 3      | 47    | -     | 50             | 6      | 102   | 8     | 116 |
|         |         | Upper   | -      | 2     | 3     | 5              | -      | 5     | 5     | 10  |
|         |         | Total   | 5      | 51    | 3     | 59             | 8      | 111   | 14    | 133 |
|         | Week 24 | Lower   | 4      | 2     | -     | 6              | 3      | 6     | 1     | 10  |
|         |         | Normal  | 1      | 47    | 2     | 50             | 5      | 95    | 11    | 111 |
|         |         | Upper   | -      | 2     | 1     | 3              | -      | 7     | 2     | 9   |
|         |         | Total   | 5      | 51    | 3     | 59             | 8      | 108   | 14    | 130 |
|         | Week 28 | Lower   | 4      | -     | -     | 4              | 3      | 4     | 1     | 8   |
|         |         | Normal  | 1      | 50    | 2     | 53             | 4      | 97    | 9     | 110 |
|         |         | Upper   | -      | 1     | 1     | 2              | 1      | 4     | 4     | 9   |
|         |         | Total   | 5      | 51    | 3     | 59             | 8      | 105   | 14    | 127 |
|         | Week 32 | Lower   | 4      | 1     | -     | 5              | 3      | 3     | -     | 6   |
|         |         | Normal  | 1      | 48    | 1     | 50             | 5      | 97    | 8     | 110 |
|         |         | Upper   | -      | 1     | 2     | 3              | -      | 5     | 6     | 11  |
|         |         | Total   | 5      | 50    | 3     | 58             | 8      | 105   | 14    | 127 |
|         | Week 36 | Lower   | 4      | -     | -     | 4              | 4      | 3     | -     | 7   |
|         |         | Normal  | 1      | 47    | 2     | 50             | 3      | 94    | 8     | 105 |
|         |         | Upper   | -      | 3     | 1     | 4              | -      | 7     | 5     | 12  |
|         |         | Total   | 5      | 50    | 3     | 58             | 7      | 104   | 13    | 124 |
|         | Week 40 | Lower   | 3      | 2     | -     | 5              | 4      | 2     | -     | 6   |
|         |         | Normal  | 2      | 48    | 2     | 52             | 3      | 95    | 7     | 105 |
|         |         | Upper   | -      | -     | 1     | 1              | -      | 6     | 5     | 11  |
|         |         | Total   | 5      | 50    | 3     | 58             | 7      | 103   | 12    | 122 |
|         | Week 44 | Lower   | 2      | 3     | -     | 5              | 3      | 4     | -     | 7   |
|         |         | Normal  | 3      | 45    | 1     | 49             | 4      | 86    | 7     | 97  |
|         |         | Upper   | -      | 2     | 2     | 4              | -      | 10    | 5     | 15  |
|         |         | Total   | 5      | 50    | 3     | 58             | 7      | 100   | 12    | 119 |
| Week 48 | Lower   | 4       | 3      | -     | 7     | 3              | 3      | -     | 6     |     |
|         | Normal  | 1       | 46     | 1     | 48    | 4              | 89     | 7     | 100   |     |
|         | Upper   | -       | 1      | 2     | 3     | -              | 8      | 5     | 13    |     |
|         | Total   | 5       | 50     | 3     | 58    | 7              | 100    | 12    | 119   |     |
| Week 52 | Lower   | 2       | 2      | -     | 4     | 4              | 4      | -     | 8     |     |
|         | Normal  | 3       | 47     | 2     | 52    | 2              | 86     | 5     | 93    |     |
|         | Upper   | -       | 1      | 1     | 2     | -              | 10     | 7     | 17    |     |
|         | Total   | 5       | 50     | 3     | 58    | 6              | 100    | 12    | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験)(9/24)

|           |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-----------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|           |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|           |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Uric Acid | Week 2  | Lower   | -      | 2     | -     | 2              | 8      | -     | -     | 8   |
|           |         | Normal  | 1      | 44    | 4     | 49             | 1      | 105   | 6     | 112 |
|           |         | Upper   | -      | 5     | 9     | 14             | -      | 11    | 10    | 21  |
|           |         | Total   | 1      | 51    | 13    | 65             | 9      | 116   | 16    | 141 |
|           | Week 4  | Lower   | 1      | -     | -     | 1              | 8      | -     | -     | 8   |
|           |         | Normal  | -      | 45    | 4     | 49             | 1      | 106   | 6     | 113 |
|           |         | Upper   | -      | 4     | 9     | 13             | -      | 7     | 10    | 17  |
|           |         | Total   | 1      | 49    | 13    | 63             | 9      | 113   | 16    | 138 |
|           | Week 8  | Lower   | 1      | 1     | -     | 2              | 8      | -     | -     | 8   |
|           |         | Normal  | -      | 42    | 2     | 44             | 1      | 102   | 4     | 107 |
|           |         | Upper   | -      | 5     | 11    | 16             | -      | 10    | 12    | 22  |
|           |         | Total   | 1      | 48    | 13    | 62             | 9      | 112   | 16    | 137 |
|           | Week 12 | Lower   | -      | -     | -     | -              | 8      | -     | -     | 8   |
|           |         | Normal  | 1      | 41    | 3     | 45             | 1      | 97    | 5     | 103 |
|           |         | Upper   | -      | 6     | 8     | 14             | -      | 15    | 11    | 26  |
|           |         | Total   | 1      | 47    | 11    | 59             | 9      | 112   | 16    | 137 |
|           | Week 16 | Lower   | 1      | -     | -     | 1              | 9      | -     | -     | 9   |
|           |         | Normal  | -      | 39    | 1     | 40             | -      | 101   | 7     | 108 |
|           |         | Upper   | -      | 8     | 10    | 18             | -      | 11    | 6     | 17  |
|           |         | Total   | 1      | 47    | 11    | 59             | 9      | 112   | 13    | 134 |
|           | Week 20 | Lower   | 1      | -     | -     | 1              | 6      | -     | -     | 6   |
|           |         | Normal  | -      | 40    | 1     | 41             | 3      | 99    | 6     | 108 |
|           |         | Upper   | -      | 7     | 10    | 17             | -      | 12    | 7     | 19  |
|           |         | Total   | 1      | 47    | 11    | 59             | 9      | 111   | 13    | 133 |
|           | Week 24 | Lower   | 1      | -     | -     | 1              | 7      | -     | -     | 7   |
|           |         | Normal  | -      | 38    | 4     | 42             | 2      | 92    | 7     | 101 |
|           |         | Upper   | -      | 9     | 7     | 16             | -      | 16    | 6     | 22  |
|           |         | Total   | 1      | 47    | 11    | 59             | 9      | 108   | 13    | 130 |
|           | Week 28 | Lower   | 1      | -     | -     | 1              | 7      | -     | -     | 7   |
|           |         | Normal  | -      | 40    | 3     | 43             | 2      | 92    | 5     | 99  |
|           |         | Upper   | -      | 7     | 8     | 15             | -      | 13    | 8     | 21  |
|           |         | Total   | 1      | 47    | 11    | 59             | 9      | 105   | 13    | 127 |
|           | Week 32 | Lower   | 1      | -     | -     | 1              | 9      | -     | -     | 9   |
|           |         | Normal  | -      | 39    | 5     | 44             | -      | 93    | 5     | 98  |
|           |         | Upper   | -      | 7     | 6     | 13             | -      | 12    | 8     | 20  |
|           |         | Total   | 1      | 46    | 11    | 58             | 9      | 105   | 13    | 127 |
|           | Week 36 | Lower   | -      | -     | -     | -              | 8      | -     | -     | 8   |
|           |         | Normal  | 1      | 39    | 6     | 46             | 1      | 96    | 8     | 105 |
|           |         | Upper   | -      | 7     | 5     | 12             | -      | 7     | 4     | 11  |
|           |         | Total   | 1      | 46    | 11    | 58             | 9      | 103   | 12    | 124 |
|           | Week 40 | Lower   | -      | -     | -     | -              | 8      | -     | -     | 8   |
|           |         | Normal  | 1      | 44    | 6     | 51             | 1      | 93    | 7     | 101 |
|           |         | Upper   | -      | 2     | 5     | 7              | -      | 8     | 5     | 13  |
|           |         | Total   | 1      | 46    | 11    | 58             | 9      | 101   | 12    | 122 |
|           | Week 44 | Lower   | 1      | 1     | -     | 2              | 7      | -     | -     | 7   |
|           |         | Normal  | -      | 39    | 6     | 45             | 1      | 89    | 7     | 97  |
|           |         | Upper   | -      | 6     | 5     | 11             | -      | 10    | 5     | 15  |
|           |         | Total   | 1      | 46    | 11    | 58             | 8      | 99    | 12    | 119 |
| Week 48   | Lower   | -       | 1      | -     | 1     | 6              | 1      | -     | 7     |     |
|           | Normal  | 1       | 41     | 7     | 49    | 2              | 89     | 9     | 100   |     |
|           | Upper   | -       | 4      | 4     | 8     | -              | 9      | 3     | 12    |     |
|           | Total   | 1       | 46     | 11    | 58    | 8              | 99     | 12    | 119   |     |
| Week 52   | Lower   | 1       | -      | -     | 1     | 8              | 1      | -     | 9     |     |
|           | Normal  | -       | 40     | 6     | 46    | -              | 89     | 8     | 97    |     |
|           | Upper   | -       | 6      | 5     | 11    | -              | 8      | 4     | 12    |     |
|           | Total   | 1       | 46     | 11    | 58    | 8              | 98     | 12    | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (10/24)

|                 |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-----------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|                 |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|                 |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Total Bilirubin | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|                 |         | Normal  | -      | 60    | -     | 60             | -      | 131   | -     | 131 |
|                 |         | Upper   | -      | 2     | 3     | 5              | -      | 7     | 3     | 10  |
|                 |         | Total   | -      | 62    | 3     | 65             | -      | 138   | 3     | 141 |
|                 | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 57    | -     | 57             | -      | 131   | -     | 131 |
|                 |         | Upper   | -      | 3     | 3     | 6              | -      | 4     | 3     | 7   |
|                 |         | Total   | -      | 60    | 3     | 63             | -      | 135   | 3     | 138 |
|                 | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 54    | 1     | 55             | -      | 130   | -     | 130 |
|                 |         | Upper   | -      | 5     | 2     | 7              | -      | 4     | 3     | 7   |
|                 |         | Total   | -      | 59    | 3     | 62             | -      | 134   | 3     | 137 |
|                 | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 51    | -     | 51             | -      | 130   | 2     | 132 |
|                 |         | Upper   | -      | 5     | 3     | 8              | -      | 4     | 1     | 5   |
|                 |         | Total   | -      | 56    | 3     | 59             | -      | 134   | 3     | 137 |
|                 | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 47    | -     | 47             | -      | 125   | 1     | 126 |
|                 |         | Upper   | -      | 9     | 3     | 12             | -      | 6     | 2     | 8   |
|                 |         | Total   | -      | 56    | 3     | 59             | -      | 131   | 3     | 134 |
|                 | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 48    | -     | 48             | -      | 123   | 1     | 124 |
|                 |         | Upper   | -      | 8     | 3     | 11             | -      | 7     | 2     | 9   |
|                 |         | Total   | -      | 56    | 3     | 59             | -      | 130   | 3     | 133 |
|                 | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 52    | 1     | 53             | -      | 121   | 1     | 122 |
|                 |         | Upper   | -      | 4     | 2     | 6              | -      | 6     | 2     | 8   |
|                 |         | Total   | -      | 56    | 3     | 59             | -      | 127   | 3     | 130 |
|                 | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 49    | -     | 49             | -      | 116   | 1     | 117 |
|                 |         | Upper   | -      | 7     | 3     | 10             | -      | 8     | 2     | 10  |
|                 |         | Total   | -      | 56    | 3     | 59             | -      | 124   | 3     | 127 |
|                 | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 49    | 1     | 50             | -      | 120   | -     | 120 |
|                 |         | Upper   | -      | 6     | 2     | 8              | -      | 4     | 3     | 7   |
|                 |         | Total   | -      | 55    | 3     | 58             | -      | 124   | 3     | 127 |
|                 | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 50    | 1     | 51             | -      | 116   | -     | 116 |
|                 |         | Upper   | -      | 5     | 2     | 7              | -      | 5     | 3     | 8   |
|                 |         | Total   | -      | 55    | 3     | 58             | -      | 121   | 3     | 124 |
|                 | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 51    | 2     | 53             | -      | 112   | 1     | 113 |
|                 |         | Upper   | -      | 4     | 1     | 5              | -      | 7     | 2     | 9   |
|                 |         | Total   | -      | 55    | 3     | 58             | -      | 119   | 3     | 122 |
|                 | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                 |         | Normal  | -      | 51    | 1     | 52             | -      | 114   | 1     | 115 |
|                 |         | Upper   | -      | 4     | 2     | 6              | -      | 2     | 2     | 4   |
|                 |         | Total   | -      | 55    | 3     | 58             | -      | 116   | 3     | 119 |
| Week 48         | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|                 | Normal  | -       | 51     | -     | 51    | -              | 112    | 1     | 113   |     |
|                 | Upper   | -       | 4      | 3     | 7     | -              | 4      | 2     | 6     |     |
|                 | Total   | -       | 55     | 3     | 58    | -              | 116    | 3     | 119   |     |
| Week 52         | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|                 | Normal  | -       | 51     | -     | 51    | -              | 114    | 2     | 116   |     |
|                 | Upper   | -       | 4      | 3     | 7     | -              | 1      | 1     | 2     |     |
|                 | Total   | -       | 55     | 3     | 58    | -              | 115    | 3     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (11/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| ChE     | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|         |         | Normal  | -      | 49    | 3     | 52             | -      | 99    | 11    | 110 |
|         |         | Upper   | -      | 1     | 12    | 13             | -      | 5     | 26    | 31  |
|         |         | Total   | -      | 50    | 15    | 65             | -      | 104   | 37    | 141 |
|         | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 49    | 2     | 51             | -      | 93    | 12    | 105 |
|         |         | Upper   | -      | 1     | 11    | 12             | -      | 8     | 25    | 33  |
|         |         | Total   | -      | 50    | 13    | 63             | -      | 101   | 37    | 138 |
|         | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 43    | 4     | 47             | -      | 99    | 13    | 112 |
|         |         | Upper   | -      | 6     | 9     | 15             | -      | 2     | 23    | 25  |
|         |         | Total   | -      | 49    | 13    | 62             | -      | 101   | 36    | 137 |
|         | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 43    | 3     | 46             | -      | 98    | 17    | 115 |
|         |         | Upper   | -      | 4     | 9     | 13             | -      | 3     | 19    | 22  |
|         |         | Total   | -      | 47    | 12    | 59             | -      | 101   | 36    | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 46    | 2     | 48             | -      | 97    | 16    | 113 |
|         |         | Upper   | -      | 1     | 10    | 11             | -      | 2     | 19    | 21  |
|         |         | Total   | -      | 47    | 12    | 59             | -      | 99    | 35    | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 45    | 3     | 48             | -      | 94    | 22    | 116 |
|         |         | Upper   | -      | 2     | 9     | 11             | -      | 4     | 13    | 17  |
|         |         | Total   | -      | 47    | 12    | 59             | -      | 98    | 35    | 133 |
|         | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 46    | 5     | 51             | -      | 93    | 20    | 113 |
|         |         | Upper   | -      | 1     | 7     | 8              | -      | 3     | 14    | 17  |
|         |         | Total   | -      | 47    | 12    | 59             | -      | 96    | 34    | 130 |
|         | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 43    | 4     | 47             | -      | 95    | 19    | 114 |
|         |         | Upper   | -      | 4     | 8     | 12             | -      | -     | 13    | 13  |
|         |         | Total   | -      | 47    | 12    | 59             | -      | 95    | 32    | 127 |
|         | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 44    | 5     | 49             | -      | 91    | 15    | 106 |
|         |         | Upper   | -      | 2     | 7     | 9              | -      | 4     | 17    | 21  |
|         |         | Total   | -      | 46    | 12    | 58             | -      | 95    | 32    | 127 |
|         | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 43    | 5     | 48             | -      | 86    | 14    | 100 |
|         |         | Upper   | -      | 3     | 7     | 10             | -      | 6     | 18    | 24  |
|         |         | Total   | -      | 46    | 12    | 58             | -      | 92    | 32    | 124 |
|         | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 45    | 2     | 47             | -      | 83    | 14    | 97  |
|         |         | Upper   | -      | 1     | 10    | 11             | -      | 7     | 18    | 25  |
|         |         | Total   | -      | 46    | 12    | 58             | -      | 90    | 32    | 122 |
|         | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 44    | 3     | 47             | -      | 79    | 12    | 91  |
|         |         | Upper   | -      | 2     | 9     | 11             | -      | 8     | 20    | 28  |
|         |         | Total   | -      | 46    | 12    | 58             | -      | 87    | 32    | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 41     | 4     | 45    | -              | 83     | 14    | 97    |     |
|         | Upper   | -       | 5      | 8     | 13    | -              | 4      | 18    | 22    |     |
|         | Total   | -       | 46     | 12    | 58    | -              | 87     | 32    | 119   |     |
| Week 52 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 43     | 2     | 45    | -              | 77     | 12    | 89    |     |
|         | Upper   | -       | 3      | 10    | 13    | -              | 10     | 19    | 29    |     |
|         | Total   | -       | 46     | 12    | 58    | -              | 87     | 31    | 118   |     |



表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (12/24)

|            |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|            |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|            |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Creatinine | Week 2  | Lower   | 9      | 4     | -     | 13             | 22     | 3     | -     | 25  |
|            |         | Normal  | 3      | 42    | 2     | 47             | 8      | 102   | -     | 110 |
|            |         | Upper   | -      | 3     | 2     | 5              | -      | -     | 6     | 6   |
|            |         | Total   | 12     | 49    | 4     | 65             | 30     | 105   | 6     | 141 |
|            | Week 4  | Lower   | 8      | 2     | -     | 10             | 22     | 6     | -     | 28  |
|            |         | Normal  | 2      | 46    | -     | 48             | 8      | 96    | -     | 104 |
|            |         | Upper   | -      | 1     | 4     | 5              | -      | -     | 6     | 6   |
|            |         | Total   | 10     | 49    | 4     | 63             | 30     | 102   | 6     | 138 |
|            | Week 8  | Lower   | 10     | 3     | -     | 13             | 25     | 3     | -     | 28  |
|            |         | Normal  | -      | 44    | -     | 44             | 5      | 98    | 1     | 104 |
|            |         | Upper   | -      | 1     | 4     | 5              | -      | -     | 5     | 5   |
|            |         | Total   | 10     | 48    | 4     | 62             | 30     | 101   | 6     | 137 |
|            | Week 12 | Lower   | 9      | 2     | -     | 11             | 20     | 4     | -     | 24  |
|            |         | Normal  | 1      | 44    | -     | 45             | 10     | 97    | 1     | 108 |
|            |         | Upper   | -      | 1     | 2     | 3              | -      | -     | 5     | 5   |
|            |         | Total   | 10     | 47    | 2     | 59             | 30     | 101   | 6     | 137 |
|            | Week 16 | Lower   | 8      | 1     | -     | 9              | 22     | 5     | -     | 27  |
|            |         | Normal  | 2      | 45    | -     | 47             | 7      | 94    | 1     | 102 |
|            |         | Upper   | -      | 1     | 2     | 3              | -      | 1     | 4     | 5   |
|            |         | Total   | 10     | 47    | 2     | 59             | 29     | 100   | 5     | 134 |
|            | Week 20 | Lower   | 7      | -     | -     | 7              | 19     | 2     | -     | 21  |
|            |         | Normal  | 3      | 46    | -     | 49             | 10     | 96    | -     | 106 |
|            |         | Upper   | -      | 1     | 2     | 3              | -      | 1     | 5     | 6   |
|            |         | Total   | 10     | 47    | 2     | 59             | 29     | 99    | 5     | 133 |
|            | Week 24 | Lower   | 8      | -     | -     | 8              | 19     | 4     | -     | 23  |
|            |         | Normal  | 2      | 45    | -     | 47             | 9      | 93    | 2     | 104 |
|            |         | Upper   | -      | 2     | 2     | 4              | -      | -     | 3     | 3   |
|            |         | Total   | 10     | 47    | 2     | 59             | 28     | 97    | 5     | 130 |
|            | Week 28 | Lower   | 6      | 3     | -     | 9              | 17     | 3     | -     | 20  |
|            |         | Normal  | 4      | 41    | -     | 45             | 10     | 90    | 2     | 102 |
|            |         | Upper   | -      | 3     | 2     | 5              | -      | 2     | 3     | 5   |
|            |         | Total   | 10     | 47    | 2     | 59             | 27     | 95    | 5     | 127 |
|            | Week 32 | Lower   | 7      | 1     | -     | 8              | 18     | -     | -     | 18  |
|            |         | Normal  | 3      | 42    | -     | 45             | 9      | 93    | 1     | 103 |
|            |         | Upper   | -      | 3     | 2     | 5              | -      | 2     | 4     | 6   |
|            |         | Total   | 10     | 46    | 2     | 58             | 27     | 95    | 5     | 127 |
|            | Week 36 | Lower   | 6      | 2     | -     | 8              | 14     | -     | -     | 14  |
|            |         | Normal  | 4      | 42    | -     | 46             | 11     | 93    | 2     | 106 |
|            |         | Upper   | -      | 2     | 2     | 4              | -      | 2     | 2     | 4   |
|            |         | Total   | 10     | 46    | 2     | 58             | 25     | 95    | 4     | 124 |
|            | Week 40 | Lower   | 8      | -     | -     | 8              | 17     | 1     | -     | 18  |
|            |         | Normal  | 2      | 45    | -     | 47             | 7      | 91    | 1     | 99  |
|            |         | Upper   | -      | 1     | 2     | 3              | -      | 2     | 3     | 5   |
|            |         | Total   | 10     | 46    | 2     | 58             | 24     | 94    | 4     | 122 |
|            | Week 44 | Lower   | 6      | -     | -     | 6              | 17     | 1     | -     | 18  |
|            |         | Normal  | 4      | 43    | -     | 47             | 7      | 88    | 2     | 97  |
|            |         | Upper   | -      | 3     | 2     | 5              | -      | 2     | 2     | 4   |
|            |         | Total   | 10     | 46    | 2     | 58             | 24     | 91    | 4     | 119 |
| Week 48    | Lower   | 7       | -      | -     | 7     | 16             | 1      | -     | 17    |     |
|            | Normal  | 3       | 44     | -     | 47    | 7              | 89     | 1     | 97    |     |
|            | Upper   | -       | 2      | 2     | 4     | 1              | 1      | 3     | 5     |     |
|            | Total   | 10      | 46     | 2     | 58    | 24             | 91     | 4     | 119   |     |
| Week 52    | Lower   | 9       | 1      | -     | 10    | 17             | 1      | -     | 18    |     |
|            | Normal  | 1       | 43     | -     | 44    | 6              | 88     | 1     | 95    |     |
|            | Upper   | -       | 2      | 2     | 4     | -              | 2      | 3     | 5     |     |
|            | Total   | 10      | 46     | 2     | 58    | 23             | 91     | 4     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (13/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| BUN     | Week 2  | Lower   | -      | 1     | -     | 1              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 64    | -     | 64             | 3      | 133   | 2     | 138 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 2     | 2   |
|         |         | Total   | -      | 65    | -     | 65             | 4      | 133   | 4     | 141 |
|         | Week 4  | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 63    | -     | 63             | 4      | 129   | 2     | 135 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 2     | 2   |
|         |         | Total   | -      | 63    | -     | 63             | 4      | 130   | 4     | 138 |
|         | Week 8  | Lower   | -      | 1     | -     | 1              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 60    | -     | 60             | 3      | 128   | 1     | 132 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | 1     | 3     | 4   |
|         |         | Total   | -      | 62    | -     | 62             | 4      | 129   | 4     | 137 |
|         | Week 12 | Lower   | -      | 1     | -     | 1              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 57    | -     | 57             | 3      | 129   | 1     | 133 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | -     | 3     | 3   |
|         |         | Total   | -      | 59    | -     | 59             | 4      | 129   | 4     | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | 1      | 2     | -     | 3   |
|         |         | Normal  | -      | 59    | -     | 59             | 3      | 125   | 2     | 130 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 59    | -     | 59             | 4      | 127   | 3     | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | 2      | 1     | -     | 3   |
|         |         | Normal  | -      | 58    | -     | 58             | 2      | 125   | 2     | 129 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 59    | -     | 59             | 4      | 126   | 3     | 133 |
|         | Week 24 | Lower   | -      | 1     | -     | 1              | 2      | 1     | -     | 3   |
|         |         | Normal  | -      | 58    | -     | 58             | 2      | 121   | 2     | 125 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | 1     | 2   |
|         |         | Total   | -      | 59    | -     | 59             | 4      | 123   | 3     | 130 |
|         | Week 28 | Lower   | -      | 2     | -     | 2              | 1      | -     | -     | 1   |
|         |         | Normal  | -      | 56    | -     | 56             | 2      | 121   | 3     | 126 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | 3      | 121   | 3     | 127 |
|         | Week 32 | Lower   | -      | -     | -     | -              | 1      | 2     | -     | 3   |
|         |         | Normal  | -      | 58    | -     | 58             | 2      | 119   | 2     | 123 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 58    | -     | 58             | 3      | 121   | 3     | 127 |
|         | Week 36 | Lower   | -      | 2     | -     | 2              | -      | 2     | -     | 2   |
|         |         | Normal  | -      | 56    | -     | 56             | 3      | 116   | 1     | 120 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | 1     | 2   |
|         |         | Total   | -      | 58    | -     | 58             | 3      | 119   | 2     | 124 |
|         | Week 40 | Lower   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|         |         | Normal  | -      | 57    | -     | 57             | 3      | 116   | 1     | 120 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 58    | -     | 58             | 3      | 117   | 2     | 122 |
|         | Week 44 | Lower   | -      | 1     | -     | 1              | 2      | 1     | -     | 3   |
|         |         | Normal  | -      | 57    | -     | 57             | 1      | 113   | 2     | 116 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 58    | -     | 58             | 3      | 114   | 2     | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | 1              | -      | -     | 1     |     |
|         | Normal  | -       | 57     | -     | 57    | 2              | 113    | 1     | 116   |     |
|         | Upper   | -       | 1      | -     | 1     | -              | 1      | 1     | 2     |     |
|         | Total   | -       | 58     | -     | 58    | 3              | 114    | 2     | 119   |     |
| Week 52 | Lower   | -       | -      | -     | -     | 1              | 2      | -     | 3     |     |
|         | Normal  | -       | 57     | -     | 57    | 2              | 110    | 2     | 114   |     |
|         | Upper   | -       | 1      | -     | 1     | -              | 1      | -     | 1     |     |
|         | Total   | -       | 58     | -     | 58    | 3              | 113    | 2     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (14/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Na      | Week 2  | Lower   | 5      | 4     | -     | 9              | 2      | 6     | -     | 8   |
|         |         | Normal  | 5      | 51    | -     | 56             | 13     | 120   | -     | 133 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 10     | 55    | -     | 65             | 15     | 126   | -     | 141 |
|         | Week 4  | Lower   | 4      | 1     | -     | 5              | 4      | -     | -     | 4   |
|         |         | Normal  | 5      | 53    | -     | 58             | 10     | 124   | -     | 134 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 9      | 54    | -     | 63             | 14     | 124   | -     | 138 |
|         | Week 8  | Lower   | 3      | 2     | -     | 5              | 5      | 5     | -     | 10  |
|         |         | Normal  | 6      | 51    | -     | 57             | 9      | 118   | -     | 127 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 9      | 53    | -     | 62             | 14     | 123   | -     | 137 |
|         | Week 12 | Lower   | 5      | 2     | -     | 7              | 6      | 8     | -     | 14  |
|         |         | Normal  | 3      | 49    | -     | 52             | 8      | 115   | -     | 123 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 51    | -     | 59             | 14     | 123   | -     | 137 |
|         | Week 16 | Lower   | 3      | 2     | -     | 5              | 3      | 5     | -     | 8   |
|         |         | Normal  | 5      | 49    | -     | 54             | 11     | 115   | -     | 126 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 51    | -     | 59             | 14     | 120   | -     | 134 |
|         | Week 20 | Lower   | 2      | 4     | -     | 6              | 2      | 4     | -     | 6   |
|         |         | Normal  | 6      | 47    | -     | 53             | 12     | 115   | -     | 127 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 51    | -     | 59             | 14     | 119   | -     | 133 |
|         | Week 24 | Lower   | 3      | 4     | -     | 7              | 4      | 4     | -     | 8   |
|         |         | Normal  | 5      | 47    | -     | 52             | 10     | 112   | -     | 122 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 51    | -     | 59             | 14     | 116   | -     | 130 |
|         | Week 28 | Lower   | 1      | 1     | -     | 2              | 1      | 2     | -     | 3   |
|         |         | Normal  | 7      | 50    | -     | 57             | 12     | 112   | -     | 124 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 51    | -     | 59             | 13     | 114   | -     | 127 |
|         | Week 32 | Lower   | 4      | 3     | -     | 7              | 3      | 5     | -     | 8   |
|         |         | Normal  | 4      | 47    | -     | 51             | 10     | 109   | -     | 119 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 50    | -     | 58             | 13     | 114   | -     | 127 |
|         | Week 36 | Lower   | 4      | 3     | -     | 7              | 2      | 5     | -     | 7   |
|         |         | Normal  | 4      | 47    | -     | 51             | 11     | 106   | -     | 117 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 50    | -     | 58             | 13     | 111   | -     | 124 |
|         | Week 40 | Lower   | 3      | 4     | -     | 7              | 2      | 3     | -     | 5   |
|         |         | Normal  | 5      | 46    | -     | 51             | 11     | 106   | -     | 117 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 50    | -     | 58             | 13     | 109   | -     | 122 |
|         | Week 44 | Lower   | 3      | 2     | -     | 5              | 3      | 4     | -     | 7   |
|         |         | Normal  | 5      | 48    | -     | 53             | 10     | 102   | -     | 112 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 50    | -     | 58             | 13     | 106   | -     | 119 |
| Week 48 | Lower   | 3       | 3      | -     | 6     | 4              | 4      | -     | 8     |     |
|         | Normal  | 5       | 47     | -     | 52    | 9              | 102    | -     | 111   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | 8       | 50     | -     | 58    | 13             | 106    | -     | 119   |     |
| Week 52 | Lower   | 2       | 1      | -     | 3     | 3              | 5      | -     | 8     |     |
|         | Normal  | 6       | 49     | -     | 55    | 9              | 101    | -     | 110   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | 8       | 50     | -     | 58    | 12             | 106    | -     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (15/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| K       | Week 2  | Lower   | 1      | -     | -     | 1              | -      | 1     | -     | 1   |
|         |         | Normal  | 1      | 63    | -     | 64             | -      | 136   | 2     | 138 |
|         |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|         |         | Total   | 2      | 63    | -     | 65             | -      | 139   | 2     | 141 |
|         | Week 4  | Lower   | 1      | 1     | -     | 2              | -      | 1     | -     | 1   |
|         |         | Normal  | 1      | 60    | -     | 61             | -      | 135   | 2     | 137 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 2      | 61    | -     | 63             | -      | 136   | 2     | 138 |
|         | Week 8  | Lower   | 1      | 1     | -     | 2              | -      | -     | -     | -   |
|         |         | Normal  | 1      | 59    | -     | 60             | -      | 134   | 2     | 136 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Total   | 2      | 60    | -     | 62             | -      | 135   | 2     | 137 |
|         | Week 12 | Lower   | 1      | 2     | -     | 3              | -      | 1     | -     | 1   |
|         |         | Normal  | 1      | 54    | -     | 55             | -      | 132   | 1     | 133 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | 2     | 1     | 3   |
|         |         | Total   | 2      | 57    | -     | 59             | -      | 135   | 2     | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | 2      | 56    | -     | 58             | -      | 130   | 2     | 132 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|         |         | Total   | 2      | 57    | -     | 59             | -      | 132   | 2     | 134 |
|         | Week 20 | Lower   | 1      | 1     | -     | 2              | -      | -     | -     | -   |
|         |         | Normal  | 1      | 56    | -     | 57             | -      | 131   | 2     | 133 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 2      | 57    | -     | 59             | -      | 131   | 2     | 133 |
|         | Week 24 | Lower   | 1      | 1     | -     | 2              | -      | -     | -     | -   |
|         |         | Normal  | 1      | 56    | -     | 57             | -      | 128   | 2     | 130 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 2      | 57    | -     | 59             | -      | 128   | 2     | 130 |
|         | Week 28 | Lower   | 1      | -     | -     | 1              | -      | 1     | -     | 1   |
|         |         | Normal  | 1      | 57    | -     | 58             | -      | 124   | 2     | 126 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 2      | 57    | -     | 59             | -      | 125   | 2     | 127 |
|         | Week 32 | Lower   | 1      | 1     | -     | 2              | -      | 1     | -     | 1   |
|         |         | Normal  | 1      | 55    | -     | 56             | -      | 124   | 2     | 126 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 2      | 56    | -     | 58             | -      | 125   | 2     | 127 |
|         | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | 2      | 56    | -     | 58             | -      | 122   | 2     | 124 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 2      | 56    | -     | 58             | -      | 122   | 2     | 124 |
|         | Week 40 | Lower   | 1      | -     | -     | 1              | -      | -     | -     | -   |
|         |         | Normal  | 1      | 56    | -     | 57             | -      | 120   | 2     | 122 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 2      | 56    | -     | 58             | -      | 120   | 2     | 122 |
|         | Week 44 | Lower   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|         |         | Normal  | 2      | 55    | -     | 57             | -      | 117   | 1     | 118 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Total   | 2      | 56    | -     | 58             | -      | 118   | 1     | 119 |
| Week 48 | Lower   | 1       | -      | -     | 1     | -              | -      | -     | -     |     |
|         | Normal  | 1       | 56     | -     | 57    | -              | 118    | 1     | 119   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | 2       | 56     | -     | 58    | -              | 118    | 1     | 119   |     |
| Week 52 | Lower   | 1       | -      | -     | 1     | -              | 1      | -     | 1     |     |
|         | Normal  | 1       | 56     | -     | 57    | -              | 116    | 1     | 117   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | 2       | 56     | -     | 58    | -              | 117    | 1     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (16/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| CI      | Week 2  | Lower   | 1      | 1     | -     | 2              | -      | 4     | -     | 4   |
|         |         | Normal  | 7      | 56    | -     | 63             | 9      | 128   | -     | 137 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 8      | 57    | -     | 65             | 9      | 132   | -     | 141 |
|         | Week 4  | Lower   | 4      | -     | -     | 4              | 4      | 1     | -     | 5   |
|         |         | Normal  | 3      | 56    | -     | 59             | 4      | 129   | -     | 133 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 56    | -     | 63             | 8      | 130   | -     | 138 |
|         | Week 8  | Lower   | 3      | 5     | -     | 8              | 1      | 2     | -     | 3   |
|         |         | Normal  | 4      | 50    | -     | 54             | 7      | 127   | -     | 134 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 55    | -     | 62             | 8      | 129   | -     | 137 |
|         | Week 12 | Lower   | 2      | 3     | -     | 5              | 3      | -     | -     | 3   |
|         |         | Normal  | 5      | 49    | -     | 54             | 5      | 129   | -     | 134 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 52    | -     | 59             | 8      | 129   | -     | 137 |
|         | Week 16 | Lower   | 2      | 1     | -     | 3              | 2      | 1     | -     | 3   |
|         |         | Normal  | 5      | 51    | -     | 56             | 6      | 124   | -     | 130 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Total   | 7      | 52    | -     | 59             | 8      | 126   | -     | 134 |
|         | Week 20 | Lower   | 2      | 2     | -     | 4              | -      | -     | -     | -   |
|         |         | Normal  | 5      | 50    | -     | 55             | 8      | 125   | -     | 133 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 52    | -     | 59             | 8      | 125   | -     | 133 |
|         | Week 24 | Lower   | 2      | 1     | -     | 3              | 1      | -     | -     | 1   |
|         |         | Normal  | 5      | 51    | -     | 56             | 7      | 122   | -     | 129 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 52    | -     | 59             | 8      | 122   | -     | 130 |
|         | Week 28 | Lower   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|         |         | Normal  | 7      | 51    | -     | 58             | 8      | 119   | -     | 127 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 52    | -     | 59             | 8      | 119   | -     | 127 |
|         | Week 32 | Lower   | 3      | -     | -     | 3              | -      | -     | -     | -   |
|         |         | Normal  | 4      | 51    | -     | 55             | 8      | 119   | -     | 127 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 51    | -     | 58             | 8      | 119   | -     | 127 |
|         | Week 36 | Lower   | 2      | 1     | -     | 3              | 1      | 2     | -     | 3   |
|         |         | Normal  | 5      | 50    | -     | 55             | 7      | 114   | -     | 121 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 51    | -     | 58             | 8      | 116   | -     | 124 |
|         | Week 40 | Lower   | 1      | -     | -     | 1              | 2      | 1     | -     | 3   |
|         |         | Normal  | 6      | 51    | -     | 57             | 6      | 113   | -     | 119 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 51    | -     | 58             | 8      | 114   | -     | 122 |
|         | Week 44 | Lower   | 1      | 1     | -     | 2              | -      | 1     | -     | 1   |
|         |         | Normal  | 6      | 50    | -     | 56             | 8      | 110   | -     | 118 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 7      | 51    | -     | 58             | 8      | 111   | -     | 119 |
| Week 48 | Lower   | 3       | 1      | -     | 4     | 1              | 2      | -     | 3     |     |
|         | Normal  | 4       | 50     | -     | 54    | 7              | 108    | -     | 115   |     |
|         | Upper   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|         | Total   | 7       | 51     | -     | 58    | 8              | 111    | -     | 119   |     |
| Week 52 | Lower   | 3       | -      | -     | 3     | 3              | 2      | -     | 5     |     |
|         | Normal  | 4       | 51     | -     | 55    | 5              | 108    | -     | 113   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | 7       | 51     | -     | 58    | 8              | 110    | -     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (17/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Ca      | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|         |         | Normal  | -      | 65    | -     | 65             | -      | 140   | -     | 140 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 65    | -     | 65             | -      | 140   | 1     | 141 |
|         | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 63    | -     | 63             | -      | 137   | -     | 137 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 63    | -     | 63             | -      | 137   | 1     | 138 |
|         | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 62    | -     | 62             | -      | 136   | -     | 136 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 62    | -     | 62             | -      | 136   | 1     | 137 |
|         | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 135   | -     | 135 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | 1     | 2   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 136   | 1     | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 133   | -     | 133 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 133   | 1     | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 132   | -     | 132 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 132   | 1     | 133 |
|         | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 129   | 1     | 130 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 129   | 1     | 130 |
|         | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 126   | -     | 126 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 126   | 1     | 127 |
|         | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 58    | -     | 58             | -      | 126   | -     | 126 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | 1     | 1   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 126   | 1     | 127 |
|         | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 58    | -     | 58             | -      | 124   | -     | 124 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 124   | -     | 124 |
|         | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 58    | -     | 58             | -      | 121   | -     | 121 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 122   | -     | 122 |
|         | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 57    | -     | 57             | -      | 119   | -     | 119 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 119   | -     | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 58     | -     | 58    | -              | 118    | -     | 118   |     |
|         | Upper   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|         | Total   | -       | 58     | -     | 58    | -              | 119    | -     | 119   |     |
| Week 52 | Lower   | -       | 1      | -     | 1     | -              | -      | -     | -     |     |
|         | Normal  | -       | 57     | -     | 57    | -              | 118    | -     | 118   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | -       | 58     | -     | 58    | -              | 118    | -     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (18/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| P       | Week 2  | Lower   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|         |         | Normal  | 1      | 63    | -     | 64             | -      | 138   | -     | 138 |
|         |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|         |         | Total   | 1      | 64    | -     | 65             | -      | 141   | -     | 141 |
|         | Week 4  | Lower   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|         |         | Normal  | 1      | 61    | -     | 62             | -      | 136   | -     | 136 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 1      | 62    | -     | 63             | -      | 138   | -     | 138 |
|         | Week 8  | Lower   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|         |         | Normal  | 1      | 60    | -     | 61             | -      | 133   | -     | 133 |
|         |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|         |         | Total   | 1      | 61    | -     | 62             | -      | 137   | -     | 137 |
|         | Week 12 | Lower   | -      | 2     | -     | 2              | -      | 5     | -     | 5   |
|         |         | Normal  | 1      | 56    | -     | 57             | -      | 131   | -     | 131 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Total   | 1      | 58    | -     | 59             | -      | 137   | -     | 137 |
|         | Week 16 | Lower   | -      | 1     | -     | 1              | -      | 4     | -     | 4   |
|         |         | Normal  | 1      | 57    | -     | 58             | -      | 129   | -     | 129 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Total   | 1      | 58    | -     | 59             | -      | 134   | -     | 134 |
|         | Week 20 | Lower   | -      | 2     | -     | 2              | -      | 3     | -     | 3   |
|         |         | Normal  | 1      | 56    | -     | 57             | -      | 130   | -     | 130 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 1      | 58    | -     | 59             | -      | 133   | -     | 133 |
|         | Week 24 | Lower   | -      | 2     | -     | 2              | -      | 2     | -     | 2   |
|         |         | Normal  | 1      | 56    | -     | 57             | -      | 126   | -     | 126 |
|         |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|         |         | Total   | 1      | 58    | -     | 59             | -      | 130   | -     | 130 |
|         | Week 28 | Lower   | -      | 2     | -     | 2              | -      | 2     | -     | 2   |
|         |         | Normal  | 1      | 56    | -     | 57             | -      | 125   | -     | 125 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | 1      | 58    | -     | 59             | -      | 127   | -     | 127 |
|         | Week 32 | Lower   | -      | 2     | -     | 2              | -      | 2     | -     | 2   |
|         |         | Normal  | 1      | 54    | -     | 55             | -      | 125   | -     | 125 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|         |         | Total   | 1      | 57    | -     | 58             | -      | 127   | -     | 127 |
|         | Week 36 | Lower   | 1      | 1     | -     | 2              | -      | 2     | -     | 2   |
|         |         | Normal  | -      | 56    | -     | 56             | -      | 121   | -     | 121 |
|         |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|         |         | Total   | 1      | 57    | -     | 58             | -      | 124   | -     | 124 |
|         | Week 40 | Lower   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|         |         | Normal  | 1      | 56    | -     | 57             | -      | 119   | -     | 119 |
|         |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|         |         | Total   | 1      | 57    | -     | 58             | -      | 122   | -     | 122 |
|         | Week 44 | Lower   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|         |         | Normal  | 1      | 56    | -     | 57             | -      | 116   | -     | 116 |
|         |         | Upper   | -      | -     | -     | -              | -      | 3     | -     | 3   |
|         |         | Total   | 1      | 57    | -     | 58             | -      | 119   | -     | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | -              | 3      | -     | 3     |     |
|         | Normal  | 1       | 57     | -     | 58    | -              | 116    | -     | 116   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | 1       | 57     | -     | 58    | -              | 119    | -     | 119   |     |
| Week 52 | Lower   | 1       | -      | -     | 1     | -              | 1      | -     | 1     |     |
|         | Normal  | -       | 57     | -     | 57    | -              | 117    | -     | 117   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | 1       | 57     | -     | 58    | -              | 118    | -     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験) (19/24)

|           |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-----------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|           |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|           |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Vitamin A | Week 2  | Lower   | 8      | 4     | -     | 12             | 13     | 16    | -     | 29  |
|           |         | Normal  | 2      | 47    | 1     | 50             | 10     | 98    | 2     | 110 |
|           |         | Upper   | -      | 2     | 1     | 3              | -      | -     | 2     | 2   |
|           |         | Total   | 10     | 53    | 2     | 65             | 23     | 114   | 4     | 141 |
|           | Week 4  | Lower   | 7      | 2     | -     | 9              | 15     | 8     | -     | 23  |
|           |         | Normal  | 3      | 49    | 2     | 54             | 8      | 102   | 4     | 114 |
|           |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Total   | 10     | 51    | 2     | 63             | 23     | 111   | 4     | 138 |
|           | Week 8  | Lower   | 9      | 5     | -     | 14             | 19     | 11    | -     | 30  |
|           |         | Normal  | 1      | 45    | 2     | 48             | 4      | 98    | 4     | 106 |
|           |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Total   | 10     | 50    | 2     | 62             | 23     | 110   | 4     | 137 |
|           | Week 12 | Lower   | 9      | 4     | -     | 13             | 15     | 13    | -     | 28  |
|           |         | Normal  | -      | 44    | -     | 44             | 8      | 96    | 3     | 107 |
|           |         | Upper   | -      | 1     | 1     | 2              | -      | 1     | 1     | 2   |
|           |         | Total   | 9      | 49    | 1     | 59             | 23     | 110   | 4     | 137 |
|           | Week 16 | Lower   | 8      | 6     | -     | 14             | 19     | 7     | -     | 26  |
|           |         | Normal  | 1      | 42    | 1     | 44             | 4      | 99    | 3     | 106 |
|           |         | Upper   | -      | 1     | -     | 1              | -      | 1     | 1     | 2   |
|           |         | Total   | 9      | 49    | 1     | 59             | 23     | 107   | 4     | 134 |
|           | Week 20 | Lower   | 8      | 3     | -     | 11             | 18     | 9     | -     | 27  |
|           |         | Normal  | 1      | 45    | 1     | 47             | 5      | 96    | 3     | 104 |
|           |         | Upper   | -      | 1     | -     | 1              | -      | 1     | 1     | 2   |
|           |         | Total   | 9      | 49    | 1     | 59             | 23     | 106   | 4     | 133 |
|           | Week 24 | Lower   | 7      | 5     | -     | 12             | 17     | 7     | -     | 24  |
|           |         | Normal  | 2      | 44    | 1     | 47             | 6      | 94    | 3     | 103 |
|           |         | Upper   | -      | -     | -     | -              | -      | 2     | 1     | 3   |
|           |         | Total   | 9      | 49    | 1     | 59             | 23     | 103   | 4     | 130 |
|           | Week 28 | Lower   | 8      | 6     | -     | 14             | 15     | 9     | -     | 24  |
|           |         | Normal  | 1      | 42    | 1     | 44             | 7      | 91    | 3     | 101 |
|           |         | Upper   | -      | 1     | -     | 1              | -      | 1     | 1     | 2   |
|           |         | Total   | 9      | 49    | 1     | 59             | 22     | 101   | 4     | 127 |
|           | Week 32 | Lower   | 7      | 4     | -     | 11             | 12     | 5     | 1     | 18  |
|           |         | Normal  | 2      | 43    | 1     | 46             | 10     | 96    | 2     | 108 |
|           |         | Upper   | -      | 1     | -     | 1              | -      | -     | 1     | 1   |
|           |         | Total   | 9      | 48    | 1     | 58             | 22     | 101   | 4     | 127 |
|           | Week 36 | Lower   | 8      | 3     | -     | 11             | 13     | 5     | -     | 18  |
|           |         | Normal  | 1      | 44    | 1     | 46             | 9      | 91    | 3     | 103 |
|           |         | Upper   | -      | 1     | -     | 1              | -      | 2     | 1     | 3   |
|           |         | Total   | 9      | 48    | 1     | 58             | 22     | 98    | 4     | 124 |
|           | Week 40 | Lower   | 7      | 4     | -     | 11             | 14     | 5     | -     | 19  |
|           |         | Normal  | 2      | 42    | 1     | 45             | 7      | 91    | 2     | 100 |
|           |         | Upper   | -      | 2     | -     | 2              | -      | 2     | 1     | 3   |
|           |         | Total   | 9      | 48    | 1     | 58             | 21     | 98    | 3     | 122 |
|           | Week 44 | Lower   | 6      | 5     | -     | 11             | 13     | 6     | -     | 19  |
|           |         | Normal  | 3      | 42    | 1     | 46             | 8      | 88    | 2     | 98  |
|           |         | Upper   | -      | 1     | -     | 1              | -      | 1     | 1     | 2   |
|           |         | Total   | 9      | 48    | 1     | 58             | 21     | 95    | 3     | 119 |
| Week 48   | Lower   | 7       | 4      | -     | 11    | 15             | 10     | -     | 25    |     |
|           | Normal  | 2       | 43     | 1     | 46    | 6              | 84     | 2     | 92    |     |
|           | Upper   | -       | 1      | -     | 1     | -              | 1      | 1     | 2     |     |
|           | Total   | 9       | 48     | 1     | 58    | 21             | 95     | 3     | 119   |     |
| Week 52   | Lower   | 6       | 5      | -     | 11    | 15             | 9      | -     | 24    |     |
|           | Normal  | 3       | 42     | 1     | 46    | 6              | 84     | 2     | 92    |     |
|           | Upper   | -       | 1      | -     | 1     | -              | 1      | 1     | 2     |     |
|           | Total   | 9       | 48     | 1     | 58    | 21             | 94     | 3     | 118   |     |



表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表(CCT-002 試験) (20/24)

|           |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-----------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|           |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|           |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Vitamin D | Week 2  | Lower   | -      | 1     | -     | 1              | -      | 4     | -     | 4   |
|           |         | Normal  | -      | 41    | 11    | 52             | 1      | 76    | 22    | 99  |
|           |         | Upper   | -      | 6     | 6     | 12             | -      | 16    | 22    | 38  |
|           |         | Total   | -      | 48    | 17    | 65             | 1      | 96    | 44    | 141 |
|           | Week 4  | Lower   | -      | 1     | -     | 1              | -      | 4     | 1     | 5   |
|           |         | Normal  | -      | 37    | 7     | 44             | 1      | 71    | 17    | 89  |
|           |         | Upper   | -      | 9     | 9     | 18             | -      | 19    | 25    | 44  |
|           |         | Total   | -      | 47    | 16    | 63             | 1      | 94    | 43    | 138 |
|           | Week 8  | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Normal  | -      | 35    | 4     | 39             | 1      | 74    | 19    | 94  |
|           |         | Upper   | -      | 11    | 12    | 23             | -      | 18    | 24    | 42  |
|           |         | Total   | -      | 46    | 16    | 62             | 1      | 93    | 43    | 137 |
|           | Week 12 | Lower   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|           |         | Normal  | -      | 31    | 7     | 38             | 1      | 72    | 17    | 90  |
|           |         | Upper   | -      | 12    | 8     | 20             | -      | 20    | 26    | 46  |
|           |         | Total   | -      | 44    | 15    | 59             | 1      | 93    | 43    | 137 |
|           | Week 16 | Lower   | -      | -     | -     | -              | -      | 4     | -     | 4   |
|           |         | Normal  | -      | 36    | 8     | 44             | 1      | 70    | 23    | 94  |
|           |         | Upper   | -      | 8     | 7     | 15             | -      | 16    | 20    | 36  |
|           |         | Total   | -      | 44    | 15    | 59             | 1      | 90    | 43    | 134 |
|           | Week 20 | Lower   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|           |         | Normal  | -      | 36    | 10    | 46             | 1      | 72    | 27    | 100 |
|           |         | Upper   | -      | 8     | 5     | 13             | -      | 15    | 16    | 31  |
|           |         | Total   | -      | 44    | 15    | 59             | 1      | 89    | 43    | 133 |
|           | Week 24 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Normal  | -      | 35    | 10    | 45             | 1      | 69    | 20    | 90  |
|           |         | Upper   | -      | 9     | 5     | 14             | -      | 17    | 22    | 39  |
|           |         | Total   | -      | 44    | 15    | 59             | 1      | 87    | 42    | 130 |
|           | Week 28 | Lower   | -      | 1     | -     | 1              | -      | 2     | 1     | 3   |
|           |         | Normal  | -      | 33    | 11    | 44             | 1      | 68    | 25    | 94  |
|           |         | Upper   | -      | 10    | 4     | 14             | -      | 16    | 14    | 30  |
|           |         | Total   | -      | 44    | 15    | 59             | 1      | 86    | 40    | 127 |
|           | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 32    | 7     | 39             | 1      | 69    | 18    | 88  |
|           |         | Upper   | -      | 11    | 8     | 19             | -      | 17    | 22    | 39  |
|           |         | Total   | -      | 43    | 15    | 58             | 1      | 86    | 40    | 127 |
|           | Week 36 | Lower   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|           |         | Normal  | -      | 31    | 8     | 39             | 1      | 64    | 20    | 85  |
|           |         | Upper   | -      | 11    | 7     | 18             | -      | 19    | 18    | 37  |
|           |         | Total   | -      | 43    | 15    | 58             | 1      | 85    | 38    | 124 |
|           | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|           |         | Normal  | -      | 30    | 8     | 38             | 1      | 66    | 16    | 83  |
|           |         | Upper   | -      | 13    | 7     | 20             | -      | 17    | 22    | 39  |
|           |         | Total   | -      | 43    | 15    | 58             | 1      | 83    | 38    | 122 |
|           | Week 44 | Lower   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|           |         | Normal  | -      | 30    | 5     | 35             | 1      | 60    | 22    | 83  |
|           |         | Upper   | -      | 13    | 10    | 23             | -      | 18    | 16    | 34  |
|           |         | Total   | -      | 43    | 15    | 58             | 1      | 80    | 38    | 119 |
| Week 48   | Lower   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|           | Normal  | -       | 33     | 9     | 42    | 1              | 63     | 15    | 79    |     |
|           | Upper   | -       | 10     | 6     | 16    | -              | 16     | 23    | 39    |     |
|           | Total   | -       | 43     | 15    | 58    | 1              | 80     | 38    | 119   |     |
| Week 52   | Lower   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|           | Normal  | -       | 33     | 7     | 40    | 1              | 59     | 17    | 77    |     |
|           | Upper   | -       | 10     | 8     | 18    | -              | 19     | 21    | 40    |     |
|           | Total   | -       | 43     | 15    | 58    | 1              | 79     | 38    | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (21/24)

|           |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|-----------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|           |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|           |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Vitamin E | Week 2  | Lower   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|           |         | Normal  | -      | 20    | 10    | 30             | 1      | 49    | 39    | 89  |
|           |         | Upper   | -      | 7     | 27    | 34             | -      | 9     | 42    | 51  |
|           |         | Total   | -      | 28    | 37    | 65             | 1      | 59    | 81    | 141 |
|           | Week 4  | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Normal  | -      | 18    | 8     | 26             | 1      | 55    | 32    | 88  |
|           |         | Upper   | -      | 9     | 28    | 37             | -      | 2     | 47    | 49  |
|           |         | Total   | -      | 27    | 36    | 63             | 1      | 58    | 79    | 138 |
|           | Week 8  | Lower   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|           |         | Normal  | -      | 21    | 6     | 27             | 1      | 54    | 30    | 85  |
|           |         | Upper   | -      | 6     | 29    | 35             | -      | 1     | 49    | 50  |
|           |         | Total   | -      | 27    | 35    | 62             | 1      | 57    | 79    | 137 |
|           | Week 12 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Normal  | -      | 16    | 8     | 24             | 1      | 51    | 29    | 81  |
|           |         | Upper   | -      | 8     | 27    | 35             | -      | 5     | 50    | 55  |
|           |         | Total   | -      | 24    | 35    | 59             | 1      | 57    | 79    | 137 |
|           | Week 16 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Normal  | -      | 15    | 4     | 19             | 1      | 45    | 27    | 73  |
|           |         | Upper   | -      | 9     | 31    | 40             | -      | 9     | 51    | 60  |
|           |         | Total   | -      | 24    | 35    | 59             | 1      | 55    | 78    | 134 |
|           | Week 20 | Lower   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|           |         | Normal  | -      | 13    | 6     | 19             | 1      | 49    | 27    | 77  |
|           |         | Upper   | -      | 11    | 29    | 40             | -      | 5     | 50    | 55  |
|           |         | Total   | -      | 24    | 35    | 59             | 1      | 55    | 77    | 133 |
|           | Week 24 | Lower   | -      | 1     | -     | 1              | -      | 3     | -     | 3   |
|           |         | Normal  | -      | 14    | 6     | 20             | 1      | 45    | 28    | 74  |
|           |         | Upper   | -      | 9     | 29    | 38             | -      | 6     | 47    | 53  |
|           |         | Total   | -      | 24    | 35    | 59             | 1      | 54    | 75    | 130 |
|           | Week 28 | Lower   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|           |         | Normal  | -      | 16    | 8     | 24             | 1      | 43    | 39    | 83  |
|           |         | Upper   | -      | 7     | 27    | 34             | -      | 8     | 34    | 42  |
|           |         | Total   | -      | 24    | 35    | 59             | 1      | 53    | 73    | 127 |
|           | Week 32 | Lower   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|           |         | Normal  | -      | 16    | 13    | 29             | 1      | 46    | 38    | 85  |
|           |         | Upper   | -      | 6     | 22    | 28             | -      | 5     | 35    | 40  |
|           |         | Total   | -      | 23    | 35    | 58             | 1      | 53    | 73    | 127 |
|           | Week 36 | Lower   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|           |         | Normal  | -      | 19    | 12    | 31             | 1      | 45    | 36    | 82  |
|           |         | Upper   | -      | 3     | 23    | 26             | -      | 5     | 35    | 40  |
|           |         | Total   | -      | 23    | 35    | 58             | 1      | 52    | 71    | 124 |
|           | Week 40 | Lower   | -      | 1     | -     | 1              | -      | 3     | -     | 3   |
|           |         | Normal  | -      | 19    | 14    | 33             | 1      | 45    | 40    | 86  |
|           |         | Upper   | -      | 3     | 21    | 24             | -      | 3     | 30    | 33  |
|           |         | Total   | -      | 23    | 35    | 58             | 1      | 51    | 70    | 122 |
|           | Week 44 | Lower   | -      | 1     | -     | 1              | -      | 5     | -     | 5   |
|           |         | Normal  | -      | 21    | 13    | 34             | 1      | 40    | 44    | 85  |
|           |         | Upper   | -      | 1     | 22    | 23             | -      | 5     | 24    | 29  |
|           |         | Total   | -      | 23    | 35    | 58             | 1      | 50    | 68    | 119 |
| Week 48   | Lower   | -       | 1      | -     | 1     | -              | 2      | -     | 2     |     |
|           | Normal  | -       | 18     | 13    | 31    | 1              | 46     | 39    | 86    |     |
|           | Upper   | -       | 4      | 22    | 26    | -              | 2      | 29    | 31    |     |
|           | Total   | -       | 23     | 35    | 58    | 1              | 50     | 68    | 119   |     |
| Week 52   | Lower   | -       | 1      | -     | 1     | 1              | 5      | -     | 6     |     |
|           | Normal  | -       | 19     | 14    | 33    | -              | 43     | 40    | 83    |     |
|           | Upper   | -       | 3      | 21    | 24    | -              | 2      | 27    | 29    |     |
|           | Total   | -       | 23     | 35    | 58    | 1              | 50     | 67    | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (22/24)

|            |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|            |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|            |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| β-Carotene | Week 2  | Lower   | 16     | 2     | -     | 18             | 38     | 14    | -     | 52  |
|            |         | Normal  | 4      | 38    | 2     | 44             | 4      | 78    | 1     | 83  |
|            |         | Upper   | -      | -     | 2     | 2              | -      | 2     | 4     | 6   |
|            |         | Total   | 20     | 40    | 4     | 64             | 42     | 94    | 5     | 141 |
|            | Week 4  | Lower   | 13     | 4     | -     | 17             | 39     | 10    | -     | 49  |
|            |         | Normal  | 5      | 37    | 2     | 44             | 3      | 82    | -     | 85  |
|            |         | Upper   | -      | -     | 2     | 2              | -      | -     | 4     | 4   |
|            |         | Total   | 18     | 41    | 4     | 63             | 42     | 92    | 4     | 138 |
|            | Week 8  | Lower   | 15     | 3     | -     | 18             | 37     | 13    | -     | 50  |
|            |         | Normal  | 2      | 37    | 2     | 41             | 4      | 79    | -     | 83  |
|            |         | Upper   | 1      | -     | 2     | 3              | -      | -     | 4     | 4   |
|            |         | Total   | 18     | 40    | 4     | 62             | 41     | 92    | 4     | 137 |
|            | Week 12 | Lower   | 13     | 2     | -     | 15             | 35     | 13    | -     | 48  |
|            |         | Normal  | 5      | 35    | 2     | 42             | 6      | 78    | -     | 84  |
|            |         | Upper   | -      | -     | 2     | 2              | -      | 1     | 4     | 5   |
|            |         | Total   | 18     | 37    | 4     | 59             | 41     | 92    | 4     | 137 |
|            | Week 16 | Lower   | 13     | 5     | -     | 18             | 35     | 15    | -     | 50  |
|            |         | Normal  | 5      | 30    | 2     | 37             | 4      | 76    | -     | 80  |
|            |         | Upper   | -      | 2     | 2     | 4              | -      | -     | 4     | 4   |
|            |         | Total   | 18     | 37    | 4     | 59             | 39     | 91    | 4     | 134 |
|            | Week 20 | Lower   | 11     | 1     | -     | 12             | 34     | 12    | -     | 46  |
|            |         | Normal  | 7      | 36    | 3     | 46             | 5      | 77    | 1     | 83  |
|            |         | Upper   | -      | -     | 1     | 1              | -      | 1     | 3     | 4   |
|            |         | Total   | 18     | 37    | 4     | 59             | 39     | 90    | 4     | 133 |
|            | Week 24 | Lower   | 11     | 3     | -     | 14             | 31     | 16    | -     | 47  |
|            |         | Normal  | 7      | 34    | 2     | 43             | 8      | 70    | -     | 78  |
|            |         | Upper   | -      | -     | 2     | 2              | -      | 2     | 3     | 5   |
|            |         | Total   | 18     | 37    | 4     | 59             | 39     | 88    | 3     | 130 |
|            | Week 28 | Lower   | 12     | 6     | -     | 18             | 30     | 19    | -     | 49  |
|            |         | Normal  | 6      | 28    | 2     | 36             | 8      | 66    | -     | 74  |
|            |         | Upper   | -      | 3     | 2     | 5              | -      | 1     | 3     | 4   |
|            |         | Total   | 18     | 37    | 4     | 59             | 38     | 86    | 3     | 127 |
|            | Week 32 | Lower   | 11     | 7     | -     | 18             | 31     | 16    | -     | 47  |
|            |         | Normal  | 7      | 27    | 2     | 36             | 7      | 69    | 1     | 77  |
|            |         | Upper   | -      | 2     | 2     | 4              | -      | 1     | 2     | 3   |
|            |         | Total   | 18     | 36    | 4     | 58             | 38     | 86    | 3     | 127 |
|            | Week 36 | Lower   | 14     | 6     | -     | 20             | 30     | 15    | -     | 45  |
|            |         | Normal  | 4      | 29    | 2     | 35             | 6      | 68    | 1     | 75  |
|            |         | Upper   | -      | 1     | 2     | 3              | -      | 2     | 2     | 4   |
|            |         | Total   | 18     | 36    | 4     | 58             | 36     | 85    | 3     | 124 |
|            | Week 40 | Lower   | 15     | 6     | -     | 21             | 30     | 17    | -     | 47  |
|            |         | Normal  | 3      | 30    | 2     | 35             | 4      | 64    | 1     | 69  |
|            |         | Upper   | -      | -     | 2     | 2              | -      | 4     | 2     | 6   |
|            |         | Total   | 18     | 36    | 4     | 58             | 34     | 85    | 3     | 122 |
|            | Week 44 | Lower   | 14     | 4     | -     | 18             | 24     | 17    | -     | 41  |
|            |         | Normal  | 4      | 32    | 2     | 38             | 8      | 65    | 2     | 75  |
|            |         | Upper   | -      | -     | 2     | 2              | -      | 2     | 1     | 3   |
|            |         | Total   | 18     | 36    | 4     | 58             | 32     | 84    | 3     | 119 |
| Week 48    | Lower   | 14      | 5      | -     | 19    | 26             | 21     | -     | 47    |     |
|            | Normal  | 4       | 31     | 2     | 37    | 6              | 62     | 2     | 70    |     |
|            | Upper   | -       | -      | 2     | 2     | -              | 1      | 1     | 2     |     |
|            | Total   | 18      | 36     | 4     | 58    | 32             | 84     | 3     | 119   |     |
| Week 52    | Lower   | 12      | 5      | -     | 17    | 25             | 18     | -     | 43    |     |
|            | Normal  | 6       | 31     | 2     | 39    | 7              | 62     | 2     | 71    |     |
|            | Upper   | -       | -      | 2     | 2     | -              | 3      | 1     | 4     |     |
|            | Total   | 18      | 36     | 4     | 58    | 32             | 83     | 3     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (23/24)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| m-AST   | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|         |         | Normal  | -      | 62    | 2     | 64             | -      | 136   | -     | 136 |
|         |         | Upper   | -      | -     | 1     | 1              | -      | 4     | 1     | 5   |
|         |         | Total   | -      | 62    | 3     | 65             | -      | 140   | 1     | 141 |
|         | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | 3     | 62             | -      | 133   | 1     | 134 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | 4     | -     | 4   |
|         |         | Total   | -      | 60    | 3     | 63             | -      | 137   | 1     | 138 |
|         | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | 3     | 62             | -      | 134   | 1     | 135 |
|         |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|         |         | Total   | -      | 59    | 3     | 62             | -      | 136   | 1     | 137 |
|         | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 57    | 2     | 59             | -      | 131   | 1     | 132 |
|         |         | Upper   | -      | -     | -     | -              | -      | 5     | -     | 5   |
|         |         | Total   | -      | 57    | 2     | 59             | -      | 136   | 1     | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 56    | 2     | 58             | -      | 130   | 1     | 131 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | 3     | -     | 3   |
|         |         | Total   | -      | 57    | 2     | 59             | -      | 133   | 1     | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 56    | 2     | 58             | -      | 132   | 1     | 133 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | -     | -     | -   |
|         |         | Total   | -      | 57    | 2     | 59             | -      | 132   | 1     | 133 |
|         | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 55    | 2     | 57             | -      | 127   | 1     | 128 |
|         |         | Upper   | -      | 2     | -     | 2              | -      | 2     | -     | 2   |
|         |         | Total   | -      | 57    | 2     | 59             | -      | 129   | 1     | 130 |
|         | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 56    | 1     | 57             | -      | 123   | 1     | 124 |
|         |         | Upper   | -      | 1     | 1     | 2              | -      | 3     | -     | 3   |
|         |         | Total   | -      | 57    | 2     | 59             | -      | 126   | 1     | 127 |
|         | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 54    | 2     | 56             | -      | 124   | 1     | 125 |
|         |         | Upper   | -      | 2     | -     | 2              | -      | 2     | -     | 2   |
|         |         | Total   | -      | 56    | 2     | 58             | -      | 126   | 1     | 127 |
|         | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 55    | 2     | 57             | -      | 119   | 1     | 120 |
|         |         | Upper   | -      | 1     | -     | 1              | -      | 4     | -     | 4   |
|         |         | Total   | -      | 56    | 2     | 58             | -      | 123   | 1     | 124 |
|         | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 53    | 2     | 55             | -      | 115   | 1     | 116 |
|         |         | Upper   | -      | 3     | -     | 3              | -      | 6     | -     | 6   |
|         |         | Total   | -      | 56    | 2     | 58             | -      | 121   | 1     | 122 |
|         | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 54    | 2     | 56             | -      | 116   | 1     | 117 |
|         |         | Upper   | -      | 2     | -     | 2              | -      | 2     | -     | 2   |
|         |         | Total   | -      | 56    | 2     | 58             | -      | 118   | 1     | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 54     | 2     | 56    | -              | 114    | 1     | 115   |     |
|         | Upper   | -       | 2      | -     | 2     | -              | 4      | -     | 4     |     |
|         | Total   | -       | 56     | 2     | 58    | -              | 118    | 1     | 119   |     |
| Week 52 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 54     | 2     | 56    | -              | 111    | 1     | 112   |     |
|         | Upper   | -       | 2      | -     | 2     | -              | 6      | -     | 6     |     |
|         | Total   | -       | 56     | 2     | 58    | -              | 117    | 1     | 118   |     |

表 2.7.4.7-付録-21 血液生化学検査の評価時点別正異判定クロス表 (CCT-002 試験) (24/24)

|                |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|----------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|                |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|                |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| CK18 (Segment) | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|                |         | Normal  | -      | 46    | 4     | 50             | -      | 101   | 13    | 114 |
|                |         | Upper   | -      | 4     | 11    | 15             | -      | 7     | 20    | 27  |
|                |         | Total   | -      | 50    | 15    | 65             | -      | 108   | 33    | 141 |
|                | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 43    | 6     | 49             | -      | 97    | 15    | 112 |
|                |         | Upper   | -      | 5     | 9     | 14             | -      | 8     | 18    | 26  |
|                |         | Total   | -      | 48    | 15    | 63             | -      | 105   | 33    | 138 |
|                | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 44    | 6     | 50             | -      | 101   | 17    | 118 |
|                |         | Upper   | -      | 3     | 9     | 12             | -      | 3     | 16    | 19  |
|                |         | Total   | -      | 47    | 15    | 62             | -      | 104   | 33    | 137 |
|                | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 35    | 7     | 42             | -      | 97    | 19    | 116 |
|                |         | Upper   | -      | 10    | 7     | 17             | -      | 7     | 14    | 21  |
|                |         | Total   | -      | 45    | 14    | 59             | -      | 104   | 33    | 137 |
|                | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 39    | 6     | 45             | -      | 95    | 19    | 114 |
|                |         | Upper   | -      | 6     | 8     | 14             | -      | 6     | 14    | 20  |
|                |         | Total   | -      | 45    | 14    | 59             | -      | 101   | 33    | 134 |
|                | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 40    | 5     | 45             | -      | 95    | 22    | 117 |
|                |         | Upper   | -      | 5     | 9     | 14             | -      | 5     | 11    | 16  |
|                |         | Total   | -      | 45    | 14    | 59             | -      | 100   | 33    | 133 |
|                | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 37    | 5     | 42             | -      | 92    | 18    | 110 |
|                |         | Upper   | -      | 8     | 9     | 17             | -      | 5     | 15    | 20  |
|                |         | Total   | -      | 45    | 14    | 59             | -      | 97    | 33    | 130 |
|                | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 40    | 3     | 43             | -      | 91    | 20    | 111 |
|                |         | Upper   | -      | 5     | 11    | 16             | -      | 4     | 12    | 16  |
|                |         | Total   | -      | 45    | 14    | 59             | -      | 95    | 32    | 127 |
|                | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 38    | 5     | 43             | -      | 88    | 18    | 106 |
|                |         | Upper   | -      | 6     | 9     | 15             | -      | 7     | 14    | 21  |
|                |         | Total   | -      | 44    | 14    | 58             | -      | 95    | 32    | 127 |
|                | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 40    | 8     | 48             | -      | 85    | 17    | 102 |
|                |         | Upper   | -      | 4     | 6     | 10             | -      | 7     | 15    | 22  |
|                |         | Total   | -      | 44    | 14    | 58             | -      | 92    | 32    | 124 |
|                | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 40    | 4     | 44             | -      | 79    | 20    | 99  |
|                |         | Upper   | -      | 4     | 10    | 14             | -      | 11    | 12    | 23  |
|                |         | Total   | -      | 44    | 14    | 58             | -      | 90    | 32    | 122 |
|                | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                |         | Normal  | -      | 38    | 6     | 44             | -      | 83    | 21    | 104 |
|                |         | Upper   | -      | 6     | 8     | 14             | -      | 4     | 11    | 15  |
|                |         | Total   | -      | 44    | 14    | 58             | -      | 87    | 32    | 119 |
| Week 48        | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|                | Normal  | -       | 39     | 8     | 47    | -              | 81     | 21    | 102   |     |
|                | Upper   | -       | 5      | 6     | 11    | -              | 6      | 11    | 17    |     |
|                | Total   | -       | 44     | 14    | 58    | -              | 87     | 32    | 119   |     |
| Week 52        | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|                | Normal  | -       | 41     | 10    | 51    | -              | 83     | 21    | 104   |     |
|                | Upper   | -       | 3      | 4     | 7     | -              | 3      | 11    | 14    |     |
|                | Total   | -       | 44     | 14    | 58    | -              | 86     | 32    | 118   |     |

表 2.7.4.7-付録-22 尿検査の評価時点別の変化量 (CCT-002 試験)

| Specific Gravity |                | Summary Statistics |         |         |        |         |         |        |       |
|------------------|----------------|--------------------|---------|---------|--------|---------|---------|--------|-------|
| Visit            | Treatment      | N                  | Mean    | SD      | Min    | Q1      | Median  | Q3     | Max   |
| Week 2           | Placebo        | 65                 | -0.0014 | 0.00519 | -0.010 | -0.0050 | 0.0000  | 0.0000 | 0.010 |
|                  | ATL-962 120 mg | 141                | -0.0004 | 0.00670 | -0.020 | -0.0050 | 0.0000  | 0.0050 | 0.015 |
| Week 4           | Placebo        | 63                 | -0.0015 | 0.00603 | -0.020 | -0.0050 | 0.0000  | 0.0020 | 0.010 |
|                  | ATL-962 120 mg | 138                | -0.0004 | 0.00574 | -0.015 | -0.0050 | 0.0000  | 0.0050 | 0.015 |
| Week 8           | Placebo        | 62                 | -0.0007 | 0.00583 | -0.020 | -0.0050 | 0.0000  | 0.0050 | 0.015 |
|                  | ATL-962 120 mg | 137                | -0.0004 | 0.00657 | -0.015 | -0.0050 | 0.0000  | 0.0040 | 0.020 |
| Week 12          | Placebo        | 59                 | 0.0000  | 0.00597 | -0.027 | -0.0030 | 0.0000  | 0.0050 | 0.015 |
|                  | ATL-962 120 mg | 137                | -0.0009 | 0.00628 | -0.015 | -0.0050 | -0.0010 | 0.0040 | 0.020 |
| Week 16          | Placebo        | 59                 | -0.0013 | 0.00680 | -0.018 | -0.0050 | -0.0010 | 0.0030 | 0.021 |
|                  | ATL-962 120 mg | 134                | -0.0011 | 0.00626 | -0.017 | -0.0050 | 0.0000  | 0.0030 | 0.019 |
| Week 20          | Placebo        | 59                 | -0.0008 | 0.00623 | -0.015 | -0.0050 | -0.0010 | 0.0050 | 0.012 |
|                  | ATL-962 120 mg | 133                | -0.0004 | 0.00652 | -0.020 | -0.0050 | -0.0010 | 0.0030 | 0.025 |
| Week 24          | Placebo        | 59                 | -0.0001 | 0.00562 | -0.011 | -0.0040 | 0.0000  | 0.0050 | 0.013 |
|                  | ATL-962 120 mg | 130                | 0.0009  | 0.00643 | -0.018 | -0.0030 | 0.0000  | 0.0050 | 0.020 |
| Week 28          | Placebo        | 59                 | 0.0004  | 0.00562 | -0.014 | -0.0030 | 0.0010  | 0.0050 | 0.012 |
|                  | ATL-962 120 mg | 127                | -0.0003 | 0.00700 | -0.020 | -0.0050 | 0.0000  | 0.0040 | 0.018 |
| Week 32          | Placebo        | 58                 | -0.0002 | 0.00649 | -0.016 | -0.0040 | 0.0000  | 0.0040 | 0.015 |
|                  | ATL-962 120 mg | 127                | -0.0007 | 0.00608 | -0.019 | -0.0050 | -0.0010 | 0.0030 | 0.013 |
| Week 36          | Placebo        | 58                 | -0.0010 | 0.00610 | -0.015 | -0.0030 | -0.0010 | 0.0020 | 0.012 |
|                  | ATL-962 120 mg | 124                | -0.0010 | 0.00581 | -0.015 | -0.0050 | -0.0010 | 0.0020 | 0.020 |
| Week 40          | Placebo        | 58                 | -0.0016 | 0.00490 | -0.015 | -0.0050 | -0.0010 | 0.0010 | 0.011 |
|                  | ATL-962 120 mg | 122                | -0.0011 | 0.00551 | -0.016 | -0.0050 | -0.0010 | 0.0020 | 0.013 |
| Week 44          | Placebo        | 58                 | -0.0015 | 0.00588 | -0.020 | -0.0050 | -0.0010 | 0.0020 | 0.017 |
|                  | ATL-962 120 mg | 119                | -0.0015 | 0.00633 | -0.020 | -0.0050 | -0.0010 | 0.0030 | 0.015 |
| Week 48          | Placebo        | 58                 | -0.0012 | 0.00668 | -0.026 | -0.0040 | -0.0005 | 0.0030 | 0.012 |
|                  | ATL-962 120 mg | 119                | -0.0013 | 0.00689 | -0.020 | -0.0060 | -0.0010 | 0.0030 | 0.015 |
| Week 52          | Placebo        | 58                 | -0.0017 | 0.00583 | -0.025 | -0.0050 | -0.0005 | 0.0020 | 0.015 |
|                  | ATL-962 120 mg | 118                | -0.0007 | 0.00710 | -0.020 | -0.0050 | 0.0000  | 0.0040 | 0.015 |

表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (1/10)

|                  |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|------------------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|                  |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|                  |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| Specific Gravity | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|                  |         | Normal  | -      | 65    | -     | 65             | -      | 141   | -     | 141 |
|                  |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Total   | -      | 65    | -     | 65             | -      | 141   | -     | 141 |
|                  | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 63    | -     | 63             | -      | 138   | -     | 138 |
|                  |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Total   | -      | 63    | -     | 63             | -      | 138   | -     | 138 |
|                  | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 62    | -     | 62             | -      | 137   | -     | 137 |
|                  |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Total   | -      | 62    | -     | 62             | -      | 137   | -     | 137 |
|                  | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 59    | -     | 59             | -      | 136   | -     | 136 |
|                  |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|                  |         | Total   | -      | 59    | -     | 59             | -      | 137   | -     | 137 |
|                  | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 59    | -     | 59             | -      | 134   | -     | 134 |
|                  |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Total   | -      | 59    | -     | 59             | -      | 134   | -     | 134 |
|                  | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 59    | -     | 59             | -      | 132   | -     | 132 |
|                  |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|                  |         | Total   | -      | 59    | -     | 59             | -      | 133   | -     | 133 |
|                  | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 58    | -     | 58             | -      | 128   | -     | 128 |
|                  |         | Upper   | -      | 1     | -     | 1              | -      | 2     | -     | 2   |
|                  |         | Total   | -      | 59    | -     | 59             | -      | 130   | -     | 130 |
|                  | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 59    | -     | 59             | -      | 126   | -     | 126 |
|                  |         | Upper   | -      | -     | -     | -              | -      | 1     | -     | 1   |
|                  |         | Total   | -      | 59    | -     | 59             | -      | 127   | -     | 127 |
|                  | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 57    | -     | 57             | -      | 126   | -     | 126 |
|                  |         | Upper   | -      | 1     | -     | 1              | -      | 1     | -     | 1   |
|                  |         | Total   | -      | 58    | -     | 58             | -      | 127   | -     | 127 |
|                  | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 58    | -     | 58             | -      | 124   | -     | 124 |
|                  |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Total   | -      | 58    | -     | 58             | -      | 124   | -     | 124 |
|                  | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 58    | -     | 58             | -      | 120   | -     | 120 |
|                  |         | Upper   | -      | -     | -     | -              | -      | 2     | -     | 2   |
|                  |         | Total   | -      | 58    | -     | 58             | -      | 122   | -     | 122 |
|                  | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Normal  | -      | 58    | -     | 58             | -      | 119   | -     | 119 |
|                  |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|                  |         | Total   | -      | 58    | -     | 58             | -      | 119   | -     | 119 |
| Week 48          | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|                  | Normal  | -       | 58     | -     | 58    | -              | 118    | -     | 118   |     |
|                  | Upper   | -       | -      | -     | -     | -              | 1      | -     | 1     |     |
|                  | Total   | -       | 58     | -     | 58    | -              | 119    | -     | 119   |     |
| Week 52          | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|                  | Normal  | -       | 57     | -     | 57    | -              | 117    | -     | 117   |     |
|                  | Upper   | -       | 1      | -     | 1     | -              | 1      | -     | 1     |     |
|                  | Total   | -       | 58     | -     | 58    | -              | 118    | -     | 118   |     |

表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (2/10)

|         |         | Placebo |        |       |       | ATL-962 120 mg |        |       |       |     |
|---------|---------|---------|--------|-------|-------|----------------|--------|-------|-------|-----|
|         |         | Week 0  |        |       |       | Week 0         |        |       |       |     |
|         |         | Lower   | Normal | Upper | Total | Lower          | Normal | Upper | Total |     |
| pH      | Week 2  | Lower   | -      | -     | -     | -              | -      | -     | -     |     |
|         |         | Normal  | -      | 65    | -     | 65             | -      | 141   | -     | 141 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 65    | -     | 65             | -      | 141   | -     | 141 |
|         | Week 4  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 63    | -     | 63             | -      | 138   | -     | 138 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 63    | -     | 63             | -      | 138   | -     | 138 |
|         | Week 8  | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 62    | -     | 62             | -      | 137   | -     | 137 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 62    | -     | 62             | -      | 137   | -     | 137 |
|         | Week 12 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 137   | -     | 137 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 137   | -     | 137 |
|         | Week 16 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 134   | -     | 134 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 134   | -     | 134 |
|         | Week 20 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 133   | -     | 133 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 133   | -     | 133 |
|         | Week 24 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 130   | -     | 130 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 130   | -     | 130 |
|         | Week 28 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 59    | -     | 59             | -      | 127   | -     | 127 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 59    | -     | 59             | -      | 127   | -     | 127 |
|         | Week 32 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 58    | -     | 58             | -      | 127   | -     | 127 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 127   | -     | 127 |
|         | Week 36 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 58    | -     | 58             | -      | 124   | -     | 124 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 124   | -     | 124 |
|         | Week 40 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 58    | -     | 58             | -      | 122   | -     | 122 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 122   | -     | 122 |
|         | Week 44 | Lower   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Normal  | -      | 58    | -     | 58             | -      | 119   | -     | 119 |
|         |         | Upper   | -      | -     | -     | -              | -      | -     | -     | -   |
|         |         | Total   | -      | 58    | -     | 58             | -      | 119   | -     | 119 |
| Week 48 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 58     | -     | 58    | -              | 119    | -     | 119   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | -       | 58     | -     | 58    | -              | 119    | -     | 119   |     |
| Week 52 | Lower   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Normal  | -       | 58     | -     | 58    | -              | 118    | -     | 118   |     |
|         | Upper   | -       | -      | -     | -     | -              | -      | -     | -     |     |
|         | Total   | -       | 58     | -     | 58    | -              | 118    | -     | 118   |     |



表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (3/10)

Glucose

| Treatment | Placebo | Frequency |          | Week 0 |          |       |
|-----------|---------|-----------|----------|--------|----------|-------|
|           |         |           |          | Normal | Abnormal | Total |
|           |         | Week 2    | Normal   | 33     | 8        | 41    |
|           |         |           | Abnormal | 6      | 18       | 24    |
|           |         |           | Total    | 39     | 26       | 65    |
|           |         | Week 4    | Normal   | 36     | 8        | 44    |
|           |         |           | Abnormal | 3      | 16       | 19    |
|           |         |           | Total    | 39     | 24       | 63    |
|           |         | Week 8    | Normal   | 32     | 7        | 39    |
|           |         |           | Abnormal | 7      | 16       | 23    |
|           |         |           | Total    | 39     | 23       | 62    |
|           |         | Week 12   | Normal   | 30     | 7        | 37    |
|           |         |           | Abnormal | 7      | 15       | 22    |
|           |         |           | Total    | 37     | 22       | 59    |
|           |         | Week 16   | Normal   | 32     | 11       | 43    |
|           |         |           | Abnormal | 5      | 11       | 16    |
|           |         |           | Total    | 37     | 22       | 59    |
|           |         | Week 20   | Normal   | 32     | 6        | 38    |
|           |         |           | Abnormal | 5      | 16       | 21    |
|           |         |           | Total    | 37     | 22       | 59    |
|           |         | Week 24   | Normal   | 31     | 13       | 44    |
|           |         |           | Abnormal | 6      | 9        | 15    |
|           |         |           | Total    | 37     | 22       | 59    |
|           |         | Week 28   | Normal   | 30     | 12       | 42    |
|           |         |           | Abnormal | 7      | 10       | 17    |
|           |         |           | Total    | 37     | 22       | 59    |
|           |         | Week 32   | Normal   | 32     | 8        | 40    |
|           |         |           | Abnormal | 4      | 14       | 18    |
|           |         |           | Total    | 36     | 22       | 58    |
|           |         | Week 36   | Normal   | 30     | 8        | 38    |
|           |         |           | Abnormal | 6      | 14       | 20    |
|           |         |           | Total    | 36     | 22       | 58    |
|           |         | Week 40   | Normal   | 32     | 8        | 40    |
|           |         |           | Abnormal | 4      | 14       | 18    |
|           |         |           | Total    | 36     | 22       | 58    |
|           |         | Week 44   | Normal   | 32     | 10       | 42    |
|           |         |           | Abnormal | 4      | 12       | 16    |
|           |         |           | Total    | 36     | 22       | 58    |
|           |         | Week 48   | Normal   | 30     | 8        | 38    |
|           |         |           | Abnormal | 6      | 14       | 20    |
|           |         |           | Total    | 36     | 22       | 58    |
|           |         | Week 52   | Normal   | 35     | 7        | 42    |
|           |         |           | Abnormal | 1      | 15       | 16    |
|           |         |           | Total    | 36     | 22       | 58    |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (4/10)

Glucose

| Treatment | ATL-962 120 mg | Frequency | Week 0   |          |       |     |
|-----------|----------------|-----------|----------|----------|-------|-----|
|           |                |           | Normal   | Abnormal | Total |     |
|           |                | Week 2    | Normal   | 86       | 20    | 106 |
|           |                |           | Abnormal | 1        | 34    | 35  |
|           |                |           | Total    | 87       | 54    | 141 |
|           |                | Week 4    | Normal   | 81       | 22    | 103 |
|           |                |           | Abnormal | 4        | 31    | 35  |
|           |                |           | Total    | 85       | 53    | 138 |
|           |                | Week 8    | Normal   | 82       | 20    | 102 |
|           |                |           | Abnormal | 2        | 33    | 35  |
|           |                |           | Total    | 84       | 53    | 137 |
|           |                | Week 12   | Normal   | 78       | 27    | 105 |
|           |                |           | Abnormal | 6        | 26    | 32  |
|           |                |           | Total    | 84       | 53    | 137 |
|           |                | Week 16   | Normal   | 77       | 30    | 107 |
|           |                |           | Abnormal | 4        | 23    | 27  |
|           |                |           | Total    | 81       | 53    | 134 |
|           |                | Week 20   | Normal   | 76       | 25    | 101 |
|           |                |           | Abnormal | 5        | 27    | 32  |
|           |                |           | Total    | 81       | 52    | 133 |
|           |                | Week 24   | Normal   | 75       | 20    | 95  |
|           |                |           | Abnormal | 4        | 31    | 35  |
|           |                |           | Total    | 79       | 51    | 130 |
|           |                | Week 28   | Normal   | 75       | 25    | 100 |
|           |                |           | Abnormal | 3        | 24    | 27  |
|           |                |           | Total    | 78       | 49    | 127 |
|           |                | Week 32   | Normal   | 72       | 22    | 94  |
|           |                |           | Abnormal | 6        | 27    | 33  |
|           |                |           | Total    | 78       | 49    | 127 |
|           |                | Week 36   | Normal   | 69       | 24    | 93  |
|           |                |           | Abnormal | 6        | 25    | 31  |
|           |                |           | Total    | 75       | 49    | 124 |
|           |                | Week 40   | Normal   | 72       | 28    | 100 |
|           |                |           | Abnormal | 2        | 20    | 22  |
|           |                |           | Total    | 74       | 48    | 122 |
|           |                | Week 44   | Normal   | 67       | 29    | 96  |
|           |                |           | Abnormal | 4        | 19    | 23  |
|           |                |           | Total    | 71       | 48    | 119 |
|           |                | Week 48   | Normal   | 66       | 27    | 93  |
|           |                |           | Abnormal | 5        | 21    | 26  |
|           |                |           | Total    | 71       | 48    | 119 |
|           |                | Week 52   | Normal   | 58       | 25    | 83  |
|           |                |           | Abnormal | 13       | 22    | 35  |
|           |                |           | Total    | 71       | 47    | 118 |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (5/10)

| Protein   |         | Frequency |          | Week 0 |          |       |
|-----------|---------|-----------|----------|--------|----------|-------|
| Treatment | Placebo |           |          | Normal | Abnormal | Total |
|           |         | Week 2    | Normal   | 40     | 5        | 45    |
|           |         |           | Abnormal | 5      | 15       | 20    |
|           |         |           | Total    | 45     | 20       | 65    |
|           |         | Week 4    | Normal   | 38     | 3        | 41    |
|           |         |           | Abnormal | 5      | 17       | 22    |
|           |         |           | Total    | 43     | 20       | 63    |
|           |         | Week 8    | Normal   | 35     | 6        | 41    |
|           |         |           | Abnormal | 7      | 14       | 21    |
|           |         |           | Total    | 42     | 20       | 62    |
|           |         | Week 12   | Normal   | 34     | 6        | 40    |
|           |         |           | Abnormal | 8      | 11       | 19    |
|           |         |           | Total    | 42     | 17       | 59    |
|           |         | Week 16   | Normal   | 34     | 6        | 40    |
|           |         |           | Abnormal | 8      | 11       | 19    |
|           |         |           | Total    | 42     | 17       | 59    |
|           |         | Week 20   | Normal   | 31     | 6        | 37    |
|           |         |           | Abnormal | 11     | 11       | 22    |
|           |         |           | Total    | 42     | 17       | 59    |
|           |         | Week 24   | Normal   | 30     | 3        | 33    |
|           |         |           | Abnormal | 12     | 14       | 26    |
|           |         |           | Total    | 42     | 17       | 59    |
|           |         | Week 28   | Normal   | 30     | 3        | 33    |
|           |         |           | Abnormal | 12     | 14       | 26    |
|           |         |           | Total    | 42     | 17       | 59    |
|           |         | Week 32   | Normal   | 33     | 4        | 37    |
|           |         |           | Abnormal | 9      | 12       | 21    |
|           |         |           | Total    | 42     | 16       | 58    |
|           |         | Week 36   | Normal   | 35     | 5        | 40    |
|           |         |           | Abnormal | 7      | 11       | 18    |
|           |         |           | Total    | 42     | 16       | 58    |
|           |         | Week 40   | Normal   | 36     | 3        | 39    |
|           |         |           | Abnormal | 6      | 13       | 19    |
|           |         |           | Total    | 42     | 16       | 58    |
|           |         | Week 44   | Normal   | 35     | 3        | 38    |
|           |         |           | Abnormal | 7      | 13       | 20    |
|           |         |           | Total    | 42     | 16       | 58    |
|           |         | Week 48   | Normal   | 36     | 3        | 39    |
|           |         |           | Abnormal | 6      | 13       | 19    |
|           |         |           | Total    | 42     | 16       | 58    |
|           |         | Week 52   | Normal   | 37     | 7        | 44    |
|           |         |           | Abnormal | 5      | 9        | 14    |
|           |         |           | Total    | 42     | 16       | 58    |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (6/10)

| Protein   |                | Frequency |          | Week 0 |          |       |
|-----------|----------------|-----------|----------|--------|----------|-------|
|           |                |           |          | Normal | Abnormal | Total |
| Treatment | ATL-962 120 mg | Week 2    | Normal   | 74     | 20       | 94    |
|           |                |           | Abnormal | 13     | 34       | 47    |
|           |                |           | Total    | 87     | 54       | 141   |
|           |                | Week 4    | Normal   | 73     | 16       | 89    |
|           |                |           | Abnormal | 13     | 36       | 49    |
|           |                |           | Total    | 86     | 52       | 138   |
|           |                | Week 8    | Normal   | 79     | 17       | 96    |
|           |                |           | Abnormal | 7      | 34       | 41    |
|           |                |           | Total    | 86     | 51       | 137   |
|           |                | Week 12   | Normal   | 72     | 15       | 87    |
|           |                |           | Abnormal | 14     | 36       | 50    |
|           |                |           | Total    | 86     | 51       | 137   |
|           |                | Week 16   | Normal   | 72     | 20       | 92    |
|           |                |           | Abnormal | 12     | 30       | 42    |
|           |                |           | Total    | 84     | 50       | 134   |
|           |                | Week 20   | Normal   | 72     | 20       | 92    |
|           |                |           | Abnormal | 11     | 30       | 41    |
|           |                |           | Total    | 83     | 50       | 133   |
|           |                | Week 24   | Normal   | 69     | 14       | 83    |
|           |                |           | Abnormal | 13     | 34       | 47    |
|           |                |           | Total    | 82     | 48       | 130   |
|           |                | Week 28   | Normal   | 69     | 18       | 87    |
|           |                |           | Abnormal | 11     | 29       | 40    |
|           |                |           | Total    | 80     | 47       | 127   |
|           |                | Week 32   | Normal   | 70     | 21       | 91    |
|           |                |           | Abnormal | 10     | 26       | 36    |
|           |                |           | Total    | 80     | 47       | 127   |
|           |                | Week 36   | Normal   | 72     | 15       | 87    |
|           |                |           | Abnormal | 7      | 30       | 37    |
|           |                |           | Total    | 79     | 45       | 124   |
|           |                | Week 40   | Normal   | 71     | 18       | 89    |
|           |                |           | Abnormal | 7      | 26       | 33    |
|           |                |           | Total    | 78     | 44       | 122   |
|           |                | Week 44   | Normal   | 67     | 17       | 84    |
|           |                |           | Abnormal | 9      | 26       | 35    |
|           |                |           | Total    | 76     | 43       | 119   |
|           |                | Week 48   | Normal   | 62     | 20       | 82    |
|           |                |           | Abnormal | 14     | 23       | 37    |
|           |                |           | Total    | 76     | 43       | 119   |
|           |                | Week 52   | Normal   | 63     | 12       | 75    |
|           |                |           | Abnormal | 13     | 30       | 43    |
|           |                |           | Total    | 76     | 42       | 118   |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (7/10)

Ketones

| Treatment | Placebo | Frequency | Week 0   |          |       |    |
|-----------|---------|-----------|----------|----------|-------|----|
|           |         |           | Normal   | Abnormal | Total |    |
|           |         | Week 2    | Normal   | 56       | 3     | 59 |
|           |         |           | Abnormal | 4        | 2     | 6  |
|           |         |           | Total    | 60       | 5     | 65 |
|           |         | Week 4    | Normal   | 53       | 5     | 58 |
|           |         |           | Abnormal | 5        | -     | 5  |
|           |         |           | Total    | 58       | 5     | 63 |
|           |         | Week 8    | Normal   | 53       | 3     | 56 |
|           |         |           | Abnormal | 4        | 2     | 6  |
|           |         |           | Total    | 57       | 5     | 62 |
|           |         | Week 12   | Normal   | 52       | 3     | 55 |
|           |         |           | Abnormal | 3        | 1     | 4  |
|           |         |           | Total    | 55       | 4     | 59 |
|           |         | Week 16   | Normal   | 44       | 2     | 46 |
|           |         |           | Abnormal | 11       | 2     | 13 |
|           |         |           | Total    | 55       | 4     | 59 |
|           |         | Week 20   | Normal   | 46       | -     | 46 |
|           |         |           | Abnormal | 9        | 4     | 13 |
|           |         |           | Total    | 55       | 4     | 59 |
|           |         | Week 24   | Normal   | 50       | 2     | 52 |
|           |         |           | Abnormal | 5        | 2     | 7  |
|           |         |           | Total    | 55       | 4     | 59 |
|           |         | Week 28   | Normal   | 47       | 1     | 48 |
|           |         |           | Abnormal | 8        | 3     | 11 |
|           |         |           | Total    | 55       | 4     | 59 |
|           |         | Week 32   | Normal   | 47       | 2     | 49 |
|           |         |           | Abnormal | 7        | 2     | 9  |
|           |         |           | Total    | 54       | 4     | 58 |
|           |         | Week 36   | Normal   | 48       | 2     | 50 |
|           |         |           | Abnormal | 6        | 2     | 8  |
|           |         |           | Total    | 54       | 4     | 58 |
|           |         | Week 40   | Normal   | 51       | 4     | 55 |
|           |         |           | Abnormal | 3        | -     | 3  |
|           |         |           | Total    | 54       | 4     | 58 |
|           |         | Week 44   | Normal   | 48       | 2     | 50 |
|           |         |           | Abnormal | 6        | 2     | 8  |
|           |         |           | Total    | 54       | 4     | 58 |
|           |         | Week 48   | Normal   | 49       | 4     | 53 |
|           |         |           | Abnormal | 5        | -     | 5  |
|           |         |           | Total    | 54       | 4     | 58 |
|           |         | Week 52   | Normal   | 50       | 3     | 53 |
|           |         |           | Abnormal | 4        | 1     | 5  |
|           |         |           | Total    | 54       | 4     | 58 |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (8/10)

Ketones

| Treatment | ATL-962 120 mg | Frequency | Week 0   |          |       |     |
|-----------|----------------|-----------|----------|----------|-------|-----|
|           |                |           | Normal   | Abnormal | Total |     |
|           |                | Week 2    | Normal   | 116      | 13    | 129 |
|           |                |           | Abnormal | 6        | 6     | 12  |
|           |                |           | Total    | 122      | 19    | 141 |
|           |                | Week 4    | Normal   | 116      | 14    | 130 |
|           |                |           | Abnormal | 5        | 3     | 8   |
|           |                |           | Total    | 121      | 17    | 138 |
|           |                | Week 8    | Normal   | 115      | 12    | 127 |
|           |                |           | Abnormal | 6        | 4     | 10  |
|           |                |           | Total    | 121      | 16    | 137 |
|           |                | Week 12   | Normal   | 116      | 14    | 130 |
|           |                |           | Abnormal | 5        | 2     | 7   |
|           |                |           | Total    | 121      | 16    | 137 |
|           |                | Week 16   | Normal   | 110      | 13    | 123 |
|           |                |           | Abnormal | 8        | 3     | 11  |
|           |                |           | Total    | 118      | 16    | 134 |
|           |                | Week 20   | Normal   | 111      | 12    | 123 |
|           |                |           | Abnormal | 6        | 4     | 10  |
|           |                |           | Total    | 117      | 16    | 133 |
|           |                | Week 24   | Normal   | 106      | 11    | 117 |
|           |                |           | Abnormal | 8        | 5     | 13  |
|           |                |           | Total    | 114      | 16    | 130 |
|           |                | Week 28   | Normal   | 105      | 13    | 118 |
|           |                |           | Abnormal | 6        | 3     | 9   |
|           |                |           | Total    | 111      | 16    | 127 |
|           |                | Week 32   | Normal   | 101      | 13    | 114 |
|           |                |           | Abnormal | 10       | 3     | 13  |
|           |                |           | Total    | 111      | 16    | 127 |
|           |                | Week 36   | Normal   | 104      | 13    | 117 |
|           |                |           | Abnormal | 4        | 3     | 7   |
|           |                |           | Total    | 108      | 16    | 124 |
|           |                | Week 40   | Normal   | 102      | 13    | 115 |
|           |                |           | Abnormal | 5        | 2     | 7   |
|           |                |           | Total    | 107      | 15    | 122 |
|           |                | Week 44   | Normal   | 99       | 12    | 111 |
|           |                |           | Abnormal | 5        | 3     | 8   |
|           |                |           | Total    | 104      | 15    | 119 |
|           |                | Week 48   | Normal   | 98       | 12    | 110 |
|           |                |           | Abnormal | 6        | 3     | 9   |
|           |                |           | Total    | 104      | 15    | 119 |
|           |                | Week 52   | Normal   | 100      | 13    | 113 |
|           |                |           | Abnormal | 3        | 2     | 5   |
|           |                |           | Total    | 103      | 15    | 118 |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (9/10)

Occult Blood

| Treatment | Placebo | Frequency | Week 0   |          |       |    |
|-----------|---------|-----------|----------|----------|-------|----|
|           |         |           | Normal   | Abnormal | Total |    |
|           |         | Week 2    | Normal   | 56       | 3     | 59 |
|           |         |           | Abnormal | 1        | 5     | 6  |
|           |         |           | Total    | 57       | 8     | 65 |
|           |         | Week 4    | Normal   | 51       | 2     | 53 |
|           |         |           | Abnormal | 4        | 6     | 10 |
|           |         |           | Total    | 55       | 8     | 63 |
|           |         | Week 8    | Normal   | 53       | 3     | 56 |
|           |         |           | Abnormal | 1        | 5     | 6  |
|           |         |           | Total    | 54       | 8     | 62 |
|           |         | Week 12   | Normal   | 51       | 3     | 54 |
|           |         |           | Abnormal | 2        | 3     | 5  |
|           |         |           | Total    | 53       | 6     | 59 |
|           |         | Week 16   | Normal   | 52       | 1     | 53 |
|           |         |           | Abnormal | 1        | 5     | 6  |
|           |         |           | Total    | 53       | 6     | 59 |
|           |         | Week 20   | Normal   | 52       | 3     | 55 |
|           |         |           | Abnormal | 1        | 3     | 4  |
|           |         |           | Total    | 53       | 6     | 59 |
|           |         | Week 24   | Normal   | 49       | 5     | 54 |
|           |         |           | Abnormal | 4        | 1     | 5  |
|           |         |           | Total    | 53       | 6     | 59 |
|           |         | Week 28   | Normal   | 50       | 3     | 53 |
|           |         |           | Abnormal | 3        | 3     | 6  |
|           |         |           | Total    | 53       | 6     | 59 |
|           |         | Week 32   | Normal   | 50       | 3     | 53 |
|           |         |           | Abnormal | 2        | 3     | 5  |
|           |         |           | Total    | 52       | 6     | 58 |
|           |         | Week 36   | Normal   | 49       | 3     | 52 |
|           |         |           | Abnormal | 3        | 3     | 6  |
|           |         |           | Total    | 52       | 6     | 58 |
|           |         | Week 40   | Normal   | 50       | 3     | 53 |
|           |         |           | Abnormal | 2        | 3     | 5  |
|           |         |           | Total    | 52       | 6     | 58 |
|           |         | Week 44   | Normal   | 50       | 3     | 53 |
|           |         |           | Abnormal | 2        | 3     | 5  |
|           |         |           | Total    | 52       | 6     | 58 |
|           |         | Week 48   | Normal   | 51       | 1     | 52 |
|           |         |           | Abnormal | 1        | 5     | 6  |
|           |         |           | Total    | 52       | 6     | 58 |
|           |         | Week 52   | Normal   | 48       | 4     | 52 |
|           |         |           | Abnormal | 4        | 2     | 6  |
|           |         |           | Total    | 52       | 6     | 58 |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-23 尿検査の評価時点別クロス表 (CCT-002 試験) (10/10)

Occult Blood

| Treatment | ATL-962 120 mg | Frequency | Week 0   |          |       |     |
|-----------|----------------|-----------|----------|----------|-------|-----|
|           |                |           | Normal   | Abnormal | Total |     |
|           |                | Week 2    | Normal   | 112      | 6     | 118 |
|           |                |           | Abnormal | 10       | 13    | 23  |
|           |                |           | Total    | 122      | 19    | 141 |
|           |                | Week 4    | Normal   | 112      | 7     | 119 |
|           |                |           | Abnormal | 7        | 12    | 19  |
|           |                |           | Total    | 119      | 19    | 138 |
|           |                | Week 8    | Normal   | 114      | 8     | 122 |
|           |                |           | Abnormal | 4        | 11    | 15  |
|           |                |           | Total    | 118      | 19    | 137 |
|           |                | Week 12   | Normal   | 113      | 7     | 120 |
|           |                |           | Abnormal | 5        | 12    | 17  |
|           |                |           | Total    | 118      | 19    | 137 |
|           |                | Week 16   | Normal   | 111      | 9     | 120 |
|           |                |           | Abnormal | 6        | 8     | 14  |
|           |                |           | Total    | 117      | 17    | 134 |
|           |                | Week 20   | Normal   | 113      | 6     | 119 |
|           |                |           | Abnormal | 4        | 10    | 14  |
|           |                |           | Total    | 117      | 16    | 133 |
|           |                | Week 24   | Normal   | 110      | 6     | 116 |
|           |                |           | Abnormal | 4        | 10    | 14  |
|           |                |           | Total    | 114      | 16    | 130 |
|           |                | Week 28   | Normal   | 108      | 4     | 112 |
|           |                |           | Abnormal | 4        | 11    | 15  |
|           |                |           | Total    | 112      | 15    | 127 |
|           |                | Week 32   | Normal   | 108      | 9     | 117 |
|           |                |           | Abnormal | 4        | 6     | 10  |
|           |                |           | Total    | 112      | 15    | 127 |
|           |                | Week 36   | Normal   | 107      | 5     | 112 |
|           |                |           | Abnormal | 4        | 8     | 12  |
|           |                |           | Total    | 111      | 13    | 124 |
|           |                | Week 40   | Normal   | 108      | 4     | 112 |
|           |                |           | Abnormal | 2        | 8     | 10  |
|           |                |           | Total    | 110      | 12    | 122 |
|           |                | Week 44   | Normal   | 102      | 4     | 106 |
|           |                |           | Abnormal | 6        | 7     | 13  |
|           |                |           | Total    | 108      | 11    | 119 |
|           |                | Week 48   | Normal   | 101      | 3     | 104 |
|           |                |           | Abnormal | 7        | 8     | 15  |
|           |                |           | Total    | 108      | 11    | 119 |
|           |                | Week 52   | Normal   | 102      | 4     | 106 |
|           |                |           | Abnormal | 5        | 7     | 12  |
|           |                |           | Total    | 107      | 11    | 118 |

"Abnormal" is used for this summary table in place of "Upper".



表 2.7.4.7-付録-24 血液学的検査の評価時点別の変化量 (OCT-001 試験) (1/3)

| Hemoglobin (g/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2            | ATL-962 120 mg | 127                | -0.07 | 0.516 | -1.6 | -0.40 | 0.00   | 0.30 | 1.8 |
| Week 4            | ATL-962 120 mg | 126                | -0.07 | 0.574 | -1.3 | -0.50 | -0.10  | 0.30 | 1.8 |
| Week 8            | ATL-962 120 mg | 126                | 0.07  | 0.635 | -2.0 | -0.40 | 0.10   | 0.50 | 1.3 |
| Week 12           | ATL-962 120 mg | 124                | 0.14  | 0.688 | -1.5 | -0.30 | 0.10   | 0.50 | 2.2 |
| Week 16           | ATL-962 120 mg | 123                | 0.22  | 0.701 | -2.4 | -0.20 | 0.20   | 0.70 | 2.4 |
| Week 20           | ATL-962 120 mg | 123                | 0.29  | 0.630 | -1.3 | -0.10 | 0.30   | 0.70 | 2.5 |
| Week 24           | ATL-962 120 mg | 122                | 0.30  | 0.754 | -1.9 | -0.20 | 0.30   | 0.70 | 2.5 |

  

| Hematocrit (%) |                | Summary Statistics |       |       |      |       |        |      |     |
|----------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit          | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2         | ATL-962 120 mg | 127                | -0.26 | 1.610 | -4.4 | -1.20 | -0.10  | 0.70 | 6.9 |
| Week 4         | ATL-962 120 mg | 126                | -0.22 | 1.882 | -4.3 | -1.50 | -0.40  | 0.90 | 6.5 |
| Week 8         | ATL-962 120 mg | 126                | 0.10  | 1.931 | -5.4 | -1.20 | -0.05  | 1.40 | 5.6 |
| Week 12        | ATL-962 120 mg | 124                | 0.24  | 2.151 | -5.0 | -1.05 | 0.40   | 1.40 | 6.1 |
| Week 16        | ATL-962 120 mg | 123                | 0.20  | 2.165 | -6.9 | -1.10 | 0.10   | 1.70 | 7.4 |
| Week 20        | ATL-962 120 mg | 123                | 0.42  | 1.978 | -4.8 | -0.80 | 0.60   | 1.80 | 6.1 |
| Week 24        | ATL-962 120 mg | 122                | 0.35  | 2.234 | -5.8 | -1.00 | 0.45   | 1.60 | 6.0 |

  

| RBC (x10 <sup>4</sup> /μL) |                | Summary Statistics |      |       |     |       |        |      |     |
|----------------------------|----------------|--------------------|------|-------|-----|-------|--------|------|-----|
| Visit                      | Treatment      | N                  | Mean | SD    | Min | Q1    | Median | Q3   | Max |
| Week 2                     | ATL-962 120 mg | 127                | -4.6 | 16.34 | -58 | -15.0 | -3.0   | 6.0  | 66  |
| Week 4                     | ATL-962 120 mg | 126                | -5.7 | 19.87 | -58 | -19.0 | -7.5   | 7.0  | 75  |
| Week 8                     | ATL-962 120 mg | 126                | -3.2 | 20.60 | -73 | -17.0 | -3.5   | 10.0 | 49  |
| Week 12                    | ATL-962 120 mg | 124                | -2.7 | 23.23 | -67 | -17.0 | -1.0   | 10.0 | 68  |
| Week 16                    | ATL-962 120 mg | 123                | -2.8 | 22.72 | -78 | -18.0 | -3.0   | 13.0 | 71  |
| Week 20                    | ATL-962 120 mg | 123                | -0.8 | 21.81 | -53 | -15.0 | 1.0    | 15.0 | 72  |
| Week 24                    | ATL-962 120 mg | 122                | -0.9 | 24.94 | -86 | -14.0 | -1.5   | 12.0 | 67  |

  

| WBC (/μL) |                | Summary Statistics |       |         |       |        |        |        |      |
|-----------|----------------|--------------------|-------|---------|-------|--------|--------|--------|------|
| Visit     | Treatment      | N                  | Mean  | SD      | Min   | Q1     | Median | Q3     | Max  |
| Week 2    | ATL-962 120 mg | 127                | 318.9 | 957.32  | -2700 | -200.0 | 400.0  | 800.0  | 3100 |
| Week 4    | ATL-962 120 mg | 126                | 385.7 | 1346.50 | -4800 | -300.0 | 400.0  | 1000.0 | 7300 |
| Week 8    | ATL-962 120 mg | 125                | 312.0 | 1050.24 | -2700 | -300.0 | 200.0  | 1000.0 | 2800 |
| Week 12   | ATL-962 120 mg | 124                | 131.5 | 1245.37 | -5700 | -500.0 | 200.0  | 700.0  | 4800 |
| Week 16   | ATL-962 120 mg | 122                | 291.0 | 1194.41 | -3500 | -400.0 | 200.0  | 900.0  | 4300 |
| Week 20   | ATL-962 120 mg | 123                | 248.0 | 1171.12 | -2700 | -500.0 | 100.0  | 900.0  | 3600 |
| Week 24   | ATL-962 120 mg | 121                | 165.3 | 1174.57 | -3500 | -500.0 | 400.0  | 800.0  | 2600 |

  

| Neutrophils (%) |                | Summary Statistics |      |       |       |       |        |      |      |
|-----------------|----------------|--------------------|------|-------|-------|-------|--------|------|------|
| Visit           | Treatment      | N                  | Mean | SD    | Min   | Q1    | Median | Q3   | Max  |
| Week 2          | ATL-962 120 mg | 127                | 1.09 | 5.278 | -14.8 | -2.90 | 1.30   | 4.20 | 18.2 |
| Week 4          | ATL-962 120 mg | 126                | 1.43 | 6.069 | -16.0 | -2.00 | 0.90   | 4.60 | 25.3 |
| Week 8          | ATL-962 120 mg | 125                | 1.01 | 4.978 | -16.1 | -2.70 | 1.40   | 3.90 | 17.6 |
| Week 12         | ATL-962 120 mg | 124                | 1.50 | 6.195 | -14.0 | -2.40 | 2.15   | 5.25 | 22.1 |
| Week 16         | ATL-962 120 mg | 122                | 1.81 | 5.974 | -16.5 | -2.10 | 1.40   | 6.00 | 14.9 |
| Week 20         | ATL-962 120 mg | 123                | 1.59 | 5.241 | -16.8 | -1.60 | 1.10   | 5.90 | 14.0 |
| Week 24         | ATL-962 120 mg | 121                | 1.70 | 5.903 | -11.1 | -2.60 | 2.00   | 6.10 | 14.7 |

表 2.7.4.7-付録-24 血液学的検査の評価時点別の変化量 (OCT-001 試験) (2/3)

| Lymphocytes (%) |                | Summary Statistics |       |       |       |       |        |      |      |
|-----------------|----------------|--------------------|-------|-------|-------|-------|--------|------|------|
| Visit           | Treatment      | N                  | Mean  | SD    | Min   | Q1    | Median | Q3   | Max  |
| Week 2          | ATL-962 120 mg | 127                | -0.93 | 4.866 | -19.0 | -3.70 | -1.10  | 2.30 | 15.2 |
| Week 4          | ATL-962 120 mg | 126                | -1.35 | 5.418 | -22.8 | -4.30 | -0.90  | 1.30 | 13.9 |
| Week 8          | ATL-962 120 mg | 125                | -0.62 | 4.266 | -11.9 | -3.60 | -0.90  | 2.50 | 14.0 |
| Week 12         | ATL-962 120 mg | 124                | -1.46 | 5.186 | -21.7 | -4.60 | -1.70  | 2.20 | 12.0 |
| Week 16         | ATL-962 120 mg | 122                | -1.66 | 5.131 | -13.6 | -5.00 | -1.30  | 1.60 | 13.4 |
| Week 20         | ATL-962 120 mg | 123                | -1.38 | 4.653 | -13.5 | -4.80 | -1.00  | 1.80 | 15.2 |
| Week 24         | ATL-962 120 mg | 121                | -2.00 | 5.123 | -15.5 | -5.40 | -1.70  | 1.70 | 9.6  |

  

| Monocytes (%) |                | Summary Statistics |       |       |      |       |        |      |     |
|---------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit         | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2        | ATL-962 120 mg | 127                | -0.12 | 1.345 | -5.6 | -0.80 | 0.00   | 0.70 | 3.8 |
| Week 4        | ATL-962 120 mg | 126                | -0.04 | 1.246 | -6.1 | -0.70 | 0.00   | 0.60 | 4.3 |
| Week 8        | ATL-962 120 mg | 125                | -0.22 | 1.458 | -5.1 | -1.10 | -0.30  | 0.80 | 2.8 |
| Week 12       | ATL-962 120 mg | 124                | 0.03  | 1.651 | -4.9 | -0.85 | 0.10   | 0.60 | 7.5 |
| Week 16       | ATL-962 120 mg | 122                | -0.14 | 1.344 | -5.9 | -1.00 | 0.00   | 0.70 | 3.1 |
| Week 20       | ATL-962 120 mg | 123                | -0.12 | 1.345 | -5.3 | -0.80 | 0.00   | 0.60 | 3.0 |
| Week 24       | ATL-962 120 mg | 121                | 0.12  | 1.393 | -3.5 | -0.70 | 0.10   | 0.80 | 3.9 |

  

| Eosinophils (%) |                | Summary Statistics |       |       |      |       |        |      |      |
|-----------------|----------------|--------------------|-------|-------|------|-------|--------|------|------|
| Visit           | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max  |
| Week 2          | ATL-962 120 mg | 127                | 0.00  | 1.135 | -4.5 | -0.50 | -0.10  | 0.50 | 4.1  |
| Week 4          | ATL-962 120 mg | 126                | -0.03 | 1.531 | -5.3 | -0.60 | 0.00   | 0.50 | 8.4  |
| Week 8          | ATL-962 120 mg | 125                | -0.12 | 1.242 | -4.8 | -0.60 | -0.10  | 0.40 | 5.9  |
| Week 12         | ATL-962 120 mg | 124                | -0.02 | 1.321 | -3.8 | -0.50 | 0.00   | 0.50 | 7.4  |
| Week 16         | ATL-962 120 mg | 122                | 0.01  | 1.416 | -5.5 | -0.60 | 0.00   | 0.60 | 6.6  |
| Week 20         | ATL-962 120 mg | 123                | -0.04 | 1.454 | -4.8 | -0.60 | 0.00   | 0.40 | 8.2  |
| Week 24         | ATL-962 120 mg | 121                | 0.21  | 1.948 | -3.0 | -0.50 | 0.00   | 0.50 | 13.8 |

  

| Basophils (%) |                | Summary Statistics |       |       |      |       |        |      |     |
|---------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit         | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2        | ATL-962 120 mg | 127                | -0.04 | 0.400 | -1.8 | -0.20 | 0.00   | 0.20 | 1.1 |
| Week 4        | ATL-962 120 mg | 126                | -0.01 | 0.334 | -1.0 | -0.20 | 0.00   | 0.20 | 1.0 |
| Week 8        | ATL-962 120 mg | 125                | -0.05 | 0.482 | -2.3 | -0.30 | 0.00   | 0.20 | 1.7 |
| Week 12       | ATL-962 120 mg | 124                | -0.05 | 0.405 | -2.3 | -0.20 | 0.00   | 0.10 | 1.5 |
| Week 16       | ATL-962 120 mg | 122                | -0.02 | 0.463 | -1.7 | -0.20 | 0.00   | 0.20 | 1.6 |
| Week 20       | ATL-962 120 mg | 123                | -0.04 | 0.460 | -2.2 | -0.30 | 0.00   | 0.20 | 1.3 |
| Week 24       | ATL-962 120 mg | 121                | -0.04 | 0.460 | -2.5 | -0.20 | 0.00   | 0.20 | 1.0 |

  

| Platelet Count (x10 <sup>4</sup> /μL) |                | Summary Statistics |       |       |       |       |        |      |     |
|---------------------------------------|----------------|--------------------|-------|-------|-------|-------|--------|------|-----|
| Visit                                 | Treatment      | N                  | Mean  | SD    | Min   | Q1    | Median | Q3   | Max |
| Week 2                                | ATL-962 120 mg | 123                | 0.47  | 2.224 | -11.8 | -0.90 | 0.50   | 1.70 | 7.8 |
| Week 4                                | ATL-962 120 mg | 124                | 0.24  | 2.060 | -6.6  | -0.90 | 0.25   | 1.55 | 6.0 |
| Week 8                                | ATL-962 120 mg | 122                | -0.06 | 2.285 | -10.0 | -1.40 | -0.15  | 1.40 | 6.5 |
| Week 12                               | ATL-962 120 mg | 122                | -0.66 | 2.416 | -10.0 | -1.90 | -0.75  | 1.10 | 6.3 |
| Week 16                               | ATL-962 120 mg | 121                | -0.70 | 2.538 | -8.8  | -2.20 | -0.60  | 0.50 | 7.5 |
| Week 20                               | ATL-962 120 mg | 120                | -0.49 | 2.576 | -9.7  | -2.00 | -0.70  | 0.75 | 8.7 |
| Week 24                               | ATL-962 120 mg | 120                | -0.87 | 2.424 | -7.5  | -2.25 | -0.70  | 0.95 | 4.2 |

表 2.7.4.7-付録-24 血液学的検査の評価時点別の変化量 (OCT-001 試験) (3/3)

| Prothrombin Time (sec) |                | Summary Statistics |       |       |      |       |        |      |      |
|------------------------|----------------|--------------------|-------|-------|------|-------|--------|------|------|
| Visit                  | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max  |
| Week 2                 | ATL-962 120 mg | 126                | 0.13  | 0.685 | -1.7 | -0.30 | 0.10   | 0.50 | 3.8  |
| Week 4                 | ATL-962 120 mg | 126                | 0.18  | 1.038 | -1.8 | -0.30 | 0.10   | 0.50 | 9.4  |
| Week 8                 | ATL-962 120 mg | 126                | 0.20  | 1.479 | -1.7 | -0.30 | 0.00   | 0.40 | 15.0 |
| Week 12                | ATL-962 120 mg | 124                | 0.17  | 1.661 | -1.3 | -0.40 | 0.00   | 0.45 | 15.8 |
| Week 16                | ATL-962 120 mg | 124                | -0.01 | 2.029 | -2.1 | -0.60 | -0.20  | 0.30 | 21.2 |
| Week 20                | ATL-962 120 mg | 123                | -0.15 | 0.673 | -1.4 | -0.60 | -0.10  | 0.20 | 1.9  |
| Week 24                | ATL-962 120 mg | 122                | -0.07 | 1.318 | -2.3 | -0.50 | -0.20  | 0.20 | 10.7 |

表 2.7.4.7-付録-25 血液学的検査の評価時点別正異判定クロス表(OCT-001 試験) (1/6)

|            |         |        | ATL-962 120 mg |        |       |       |
|------------|---------|--------|----------------|--------|-------|-------|
|            |         |        | Week 0         |        |       |       |
|            |         |        | Lower          | Normal | Upper | Total |
| Hemoglobin | Week 2  | Lower  | 4              | 1      | -     | 5     |
|            |         | Normal | 1              | 118    | 2     | 121   |
|            |         | Upper  | -              | -      | 1     | 1     |
|            |         | Total  | 5              | 119    | 3     | 127   |
|            | Week 4  | Lower  | 3              | 2      | -     | 5     |
|            |         | Normal | 2              | 116    | 1     | 119   |
|            |         | Upper  | -              | -      | 2     | 2     |
|            |         | Total  | 5              | 118    | 3     | 126   |
|            | Week 8  | Lower  | 3              | 1      | -     | 4     |
|            |         | Normal | 2              | 114    | 1     | 117   |
|            |         | Upper  | -              | 3      | 2     | 5     |
|            |         | Total  | 5              | 118    | 3     | 126   |
|            | Week 12 | Lower  | 1              | 1      | -     | 2     |
|            |         | Normal | 4              | 113    | 1     | 118   |
|            |         | Upper  | -              | 2      | 2     | 4     |
|            |         | Total  | 5              | 116    | 3     | 124   |
|            | Week 16 | Lower  | 1              | -      | -     | 1     |
|            |         | Normal | 4              | 110    | 1     | 115   |
|            |         | Upper  | -              | 5      | 2     | 7     |
|            |         | Total  | 5              | 115    | 3     | 123   |
|            | Week 20 | Lower  | 1              | 1      | -     | 2     |
|            |         | Normal | 4              | 113    | 1     | 118   |
|            |         | Upper  | -              | 1      | 2     | 3     |
|            |         | Total  | 5              | 115    | 3     | 123   |
| Week 24    | Lower   | -      | 1              | -      | 1     |       |
|            | Normal  | 5      | 110            | -      | 115   |       |
|            | Upper   | -      | 3              | 3      | 6     |       |
|            | Total   | 5      | 114            | 3      | 122   |       |
| Hematocrit | Week 2  | Lower  | 5              | 2      | -     | 7     |
|            |         | Normal | -              | 117    | 2     | 119   |
|            |         | Upper  | -              | -      | 1     | 1     |
|            |         | Total  | 5              | 119    | 3     | 127   |
|            | Week 4  | Lower  | 3              | 3      | -     | 6     |
|            |         | Normal | 2              | 115    | 2     | 119   |
|            |         | Upper  | -              | -      | 1     | 1     |
|            |         | Total  | 5              | 118    | 3     | 126   |
|            | Week 8  | Lower  | 2              | 2      | -     | 4     |
|            |         | Normal | 3              | 116    | 1     | 120   |
|            |         | Upper  | -              | -      | 2     | 2     |
|            |         | Total  | 5              | 118    | 3     | 126   |
|            | Week 12 | Lower  | 2              | 2      | -     | 4     |
|            |         | Normal | 3              | 113    | 2     | 118   |
|            |         | Upper  | -              | 1      | 1     | 2     |
|            |         | Total  | 5              | 116    | 3     | 124   |
|            | Week 16 | Lower  | 2              | -      | -     | 2     |
|            |         | Normal | 3              | 114    | 3     | 120   |
|            |         | Upper  | -              | 1      | -     | 1     |
|            |         | Total  | 5              | 115    | 3     | 123   |
|            | Week 20 | Lower  | 1              | 1      | -     | 2     |
|            |         | Normal | 4              | 114    | 3     | 121   |
|            |         | Upper  | -              | -      | -     | -     |
|            |         | Total  | 5              | 115    | 3     | 123   |
| Week 24    | Lower   | 1      | 1              | -      | 2     |       |
|            | Normal  | 4      | 112            | 2      | 118   |       |
|            | Upper   | -      | 1              | 1      | 2     |       |
|            | Total   | 5      | 114            | 3      | 122   |       |

表 2.7.4.7-付録-25 血液学的検査の評価時点別正異判定クロス表(OCT-001 試験) (2/6)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| RBC     | Week 2  | Lower  | 5              | 3      | -     | 8     |
|         |         | Normal | 1              | 113    | 2     | 116   |
|         |         | Upper  | -              | -      | 3     | 3     |
|         |         | Total  | 6              | 116    | 5     | 127   |
|         | Week 4  | Lower  | 4              | 3      | -     | 7     |
|         |         | Normal | 2              | 112    | 2     | 116   |
|         |         | Upper  | -              | -      | 3     | 3     |
|         |         | Total  | 6              | 115    | 5     | 126   |
|         | Week 8  | Lower  | 4              | 3      | -     | 7     |
|         |         | Normal | 2              | 110    | 2     | 114   |
|         |         | Upper  | -              | 2      | 3     | 5     |
|         |         | Total  | 6              | 115    | 5     | 126   |
|         | Week 12 | Lower  | 4              | 5      | -     | 9     |
|         |         | Normal | 2              | 106    | 4     | 112   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 6              | 113    | 5     | 124   |
|         | Week 16 | Lower  | 3              | 3      | -     | 6     |
|         |         | Normal | 3              | 107    | 4     | 114   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 6              | 112    | 5     | 123   |
|         | Week 20 | Lower  | 5              | 4      | -     | 9     |
|         |         | Normal | 1              | 108    | 3     | 112   |
|         |         | Upper  | -              | -      | 2     | 2     |
|         |         | Total  | 6              | 112    | 5     | 123   |
| Week 24 | Lower   | 5      | 3              | -      | 8     |       |
|         | Normal  | 1      | 107            | 3      | 111   |       |
|         | Upper   | -      | 1              | 2      | 3     |       |
|         | Total   | 6      | 111            | 5      | 122   |       |
| WBC     | Week 2  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 105    | 5     | 110   |
|         |         | Upper  | -              | 6      | 11    | 17    |
|         |         | Total  | -              | 111    | 16    | 127   |
|         | Week 4  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 102    | 6     | 108   |
|         |         | Upper  | -              | 8      | 10    | 18    |
|         |         | Total  | -              | 110    | 16    | 126   |
|         | Week 8  | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | -              | 102    | 6     | 108   |
|         |         | Upper  | -              | 6      | 10    | 16    |
|         |         | Total  | -              | 109    | 16    | 125   |
|         | Week 12 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 103    | 6     | 109   |
|         |         | Upper  | -              | 5      | 10    | 15    |
|         |         | Total  | -              | 108    | 16    | 124   |
|         | Week 16 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 100    | 6     | 106   |
|         |         | Upper  | -              | 6      | 10    | 16    |
|         |         | Total  | -              | 106    | 16    | 122   |
|         | Week 20 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 99     | 5     | 104   |
|         |         | Upper  | -              | 8      | 11    | 19    |
|         |         | Total  | -              | 107    | 16    | 123   |
| Week 24 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 100            | 8      | 108   |       |
|         | Upper   | -      | 5              | 8      | 13    |       |
|         | Total   | -      | 105            | 16     | 121   |       |

表 2.7.4.7-付録-25 血液学的検査の評価時点別正異判定クロス表(OCT-001 試験) (3/6)

|             |         |        | ATL-962 120 mg |        |       |       |
|-------------|---------|--------|----------------|--------|-------|-------|
|             |         |        | Week 0         |        |       |       |
|             |         |        | Lower          | Normal | Upper | Total |
| Neutrophils | Week 2  | Lower  | 2              | 1      | -     | 3     |
|             |         | Normal | 1              | 123    | -     | 124   |
|             |         | Upper  | -              | -      | -     | -     |
|             |         | Total  | 3              | 124    | -     | 127   |
|             | Week 4  | Lower  | 1              | 2      | -     | 3     |
|             |         | Normal | 2              | 119    | -     | 121   |
|             |         | Upper  | -              | 2      | -     | 2     |
|             |         | Total  | 3              | 123    | -     | 126   |
|             | Week 8  | Lower  | 3              | 2      | -     | 5     |
|             |         | Normal | -              | 120    | -     | 120   |
|             |         | Upper  | -              | -      | -     | -     |
|             |         | Total  | 3              | 122    | -     | 125   |
|             | Week 12 | Lower  | 2              | -      | -     | 2     |
|             |         | Normal | 1              | 120    | -     | 121   |
|             |         | Upper  | -              | 1      | -     | 1     |
|             |         | Total  | 3              | 121    | -     | 124   |
|             | Week 16 | Lower  | 1              | -      | -     | 1     |
|             |         | Normal | 2              | 117    | -     | 119   |
|             |         | Upper  | -              | 2      | -     | 2     |
|             |         | Total  | 3              | 119    | -     | 122   |
|             | Week 20 | Lower  | 2              | 1      | -     | 3     |
|             |         | Normal | 1              | 119    | -     | 120   |
|             |         | Upper  | -              | -      | -     | -     |
|             |         | Total  | 3              | 120    | -     | 123   |
| Week 24     | Lower   | 1      | 1              | -      | 2     |       |
|             | Normal  | 2      | 117            | -      | 119   |       |
|             | Upper   | -      | -              | -      | -     |       |
|             | Total   | 3      | 118            | -      | 121   |       |
| Lymphocytes | Week 2  | Lower  | -              | 2      | -     | 2     |
|             |         | Normal | -              | 120    | 3     | 123   |
|             |         | Upper  | -              | 1      | 1     | 2     |
|             |         | Total  | -              | 123    | 4     | 127   |
|             | Week 4  | Lower  | -              | 2      | -     | 2     |
|             |         | Normal | -              | 120    | 2     | 122   |
|             |         | Upper  | -              | -      | 2     | 2     |
|             |         | Total  | -              | 122    | 4     | 126   |
|             | Week 8  | Lower  | -              | 1      | -     | 1     |
|             |         | Normal | -              | 119    | 2     | 121   |
|             |         | Upper  | -              | 1      | 2     | 3     |
|             |         | Total  | -              | 121    | 4     | 125   |
|             | Week 12 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 119    | 2     | 121   |
|             |         | Upper  | -              | 1      | 2     | 3     |
|             |         | Total  | -              | 120    | 4     | 124   |
|             | Week 16 | Lower  | -              | 2      | -     | 2     |
|             |         | Normal | -              | 116    | 2     | 118   |
|             |         | Upper  | -              | -      | 2     | 2     |
|             |         | Total  | -              | 118    | 4     | 122   |
|             | Week 20 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 119    | -     | 119   |
|             |         | Upper  | -              | -      | 4     | 4     |
|             |         | Total  | -              | 119    | 4     | 123   |
| Week 24     | Lower   | -      | -              | -      | -     |       |
|             | Normal  | -      | 117            | 2      | 119   |       |
|             | Upper   | -      | -              | 2      | 2     |       |
|             | Total   | -      | 117            | 4      | 121   |       |

表 2.7.4.7-付録-25 血液学的検査の評価時点別正異判定クロス表(OCT-001 試験) (4/6)

|             |         |        | ATL-962 120 mg |        |       |       |
|-------------|---------|--------|----------------|--------|-------|-------|
|             |         |        | Week 0         |        |       |       |
|             |         |        | Lower          | Normal | Upper | Total |
| Monocytes   | Week 2  | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 126    | 1     | 127   |
|             |         | Upper  | -              | -      | -     | -     |
|             |         | Total  | -              | 126    | 1     | 127   |
|             | Week 4  | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 125    | 1     | 126   |
|             |         | Upper  | -              | -      | -     | -     |
|             |         | Total  | -              | 125    | 1     | 126   |
|             | Week 8  | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 123    | 1     | 124   |
|             |         | Upper  | -              | 1      | -     | 1     |
|             |         | Total  | -              | 124    | 1     | 125   |
|             | Week 12 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 121    | 1     | 122   |
|             |         | Upper  | -              | 2      | -     | 2     |
|             |         | Total  | -              | 123    | 1     | 124   |
|             | Week 16 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 120    | 1     | 121   |
|             |         | Upper  | -              | 1      | -     | 1     |
|             |         | Total  | -              | 121    | 1     | 122   |
| Week 20     | Lower   | -      | -              | -      | -     |       |
|             | Normal  | -      | 122            | 1      | 123   |       |
|             | Upper   | -      | -              | -      | -     |       |
|             | Total   | -      | 122            | 1      | 123   |       |
| Week 24     | Lower   | -      | -              | -      | -     |       |
|             | Normal  | -      | 120            | 1      | 121   |       |
|             | Upper   | -      | -              | -      | -     |       |
|             | Total   | -      | 120            | 1      | 121   |       |
| Eosinophils | Week 2  | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 121    | 2     | 123   |
|             |         | Upper  | -              | 1      | 3     | 4     |
|             |         | Total  | -              | 122    | 5     | 127   |
|             | Week 4  | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 120    | 2     | 122   |
|             |         | Upper  | -              | 1      | 3     | 4     |
|             |         | Total  | -              | 121    | 5     | 126   |
|             | Week 8  | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 120    | 3     | 123   |
|             |         | Upper  | -              | -      | 2     | 2     |
|             |         | Total  | -              | 120    | 5     | 125   |
|             | Week 12 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 119    | 2     | 121   |
|             |         | Upper  | -              | -      | 3     | 3     |
|             |         | Total  | -              | 119    | 5     | 124   |
|             | Week 16 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 114    | 2     | 116   |
|             |         | Upper  | -              | 3      | 3     | 6     |
|             |         | Total  | -              | 117    | 5     | 122   |
| Week 20     | Lower   | -      | -              | -      | -     |       |
|             | Normal  | -      | 116            | 4      | 120   |       |
|             | Upper   | -      | 2              | 1      | 3     |       |
|             | Total   | -      | 118            | 5      | 123   |       |
| Week 24     | Lower   | -      | -              | -      | -     |       |
|             | Normal  | -      | 114            | 1      | 115   |       |
|             | Upper   | -      | 2              | 4      | 6     |       |
|             | Total   | -      | 116            | 5      | 121   |       |

表 2.7.4.7-付録-25 血液学的検査の評価時点別正異判定クロス表(OCT-001 試験) (5/6)

|                |         |        | ATL-962 120 mg |        |       |       |
|----------------|---------|--------|----------------|--------|-------|-------|
|                |         |        | Week 0         |        |       |       |
|                |         |        | Lower          | Normal | Upper | Total |
| Basophils      | Week 2  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 126    | 1     | 127   |
|                |         | Upper  | -              | -      | -     | -     |
|                |         | Total  | -              | 126    | 1     | 127   |
|                | Week 4  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 125    | -     | 125   |
|                |         | Upper  | -              | -      | 1     | 1     |
|                |         | Total  | -              | 125    | 1     | 126   |
|                | Week 8  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 123    | 1     | 124   |
|                |         | Upper  | -              | 1      | -     | 1     |
|                |         | Total  | -              | 124    | 1     | 125   |
|                | Week 12 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 123    | 1     | 124   |
|                |         | Upper  | -              | -      | -     | -     |
|                |         | Total  | -              | 123    | 1     | 124   |
|                | Week 16 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 121    | 1     | 122   |
|                |         | Upper  | -              | -      | -     | -     |
|                |         | Total  | -              | 121    | 1     | 122   |
| Week 20        | Lower   | -      | -              | -      | -     |       |
|                | Normal  | -      | 122            | 1      | 123   |       |
|                | Upper   | -      | -              | -      | -     |       |
|                | Total   | -      | 122            | 1      | 123   |       |
| Week 24        | Lower   | -      | -              | -      | -     |       |
|                | Normal  | -      | 120            | 1      | 121   |       |
|                | Upper   | -      | -              | -      | -     |       |
|                | Total   | -      | 120            | 1      | 121   |       |
| Platelet Count | Week 2  | Lower  | -              | 1      | -     | 1     |
|                |         | Normal | 2              | 111    | -     | 113   |
|                |         | Upper  | -              | -      | 9     | 9     |
|                |         | Total  | 2              | 112    | 9     | 123   |
|                | Week 4  | Lower  | 1              | -      | -     | 1     |
|                |         | Normal | 1              | 112    | 1     | 114   |
|                |         | Upper  | -              | 1      | 8     | 9     |
|                |         | Total  | 2              | 113    | 9     | 124   |
|                | Week 8  | Lower  | 1              | -      | -     | 1     |
|                |         | Normal | 1              | 110    | 4     | 115   |
|                |         | Upper  | -              | 1      | 5     | 6     |
|                |         | Total  | 2              | 111    | 9     | 122   |
|                | Week 12 | Lower  | -              | 1      | -     | 1     |
|                |         | Normal | 2              | 109    | 4     | 115   |
|                |         | Upper  | -              | 1      | 5     | 6     |
|                |         | Total  | 2              | 111    | 9     | 122   |
|                | Week 16 | Lower  | 1              | -      | -     | 1     |
|                |         | Normal | 1              | 110    | 4     | 115   |
|                |         | Upper  | -              | -      | 5     | 5     |
|                |         | Total  | 2              | 110    | 9     | 121   |
| Week 20        | Lower   | -      | -              | -      | -     |       |
|                | Normal  | 2      | 108            | 4      | 114   |       |
|                | Upper   | -      | 1              | 5      | 6     |       |
|                | Total   | 2      | 109            | 9      | 120   |       |
| Week 24        | Lower   | -      | 2              | -      | 2     |       |
|                | Normal  | 2      | 106            | 2      | 110   |       |
|                | Upper   | -      | 1              | 7      | 8     |       |
|                | Total   | 2      | 109            | 9      | 120   |       |



表 2.7.4.7-付録-25 血液学的検査の評価時点別正異判定クロス表(OCT-001 試験) (6/6)

|                  |         |        | ATL-962 120 mg |        |       |       |
|------------------|---------|--------|----------------|--------|-------|-------|
|                  |         |        | Week 0         |        |       |       |
|                  |         |        | Lower          | Normal | Upper | Total |
| Prothrombin Time | Week 2  | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 112    | 4     | 116   |
|                  |         | Upper  | -              | 4      | 6     | 10    |
|                  |         | Total  | -              | 116    | 10    | 126   |
|                  | Week 4  | Lower  | -              | 1      | -     | 1     |
|                  |         | Normal | -              | 107    | 5     | 112   |
|                  |         | Upper  | -              | 8      | 5     | 13    |
|                  |         | Total  | -              | 116    | 10    | 126   |
|                  | Week 8  | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 112    | 3     | 115   |
|                  |         | Upper  | -              | 4      | 7     | 11    |
|                  |         | Total  | -              | 116    | 10    | 126   |
|                  | Week 12 | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 111    | 5     | 116   |
|                  |         | Upper  | -              | 3      | 5     | 8     |
|                  |         | Total  | -              | 114    | 10    | 124   |
|                  | Week 16 | Lower  | -              | 1      | -     | 1     |
|                  |         | Normal | -              | 113    | 4     | 117   |
|                  |         | Upper  | -              | -      | 6     | 6     |
|                  |         | Total  | -              | 114    | 10    | 124   |
|                  | Week 20 | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 109    | 7     | 116   |
|                  |         | Upper  | -              | 4      | 3     | 7     |
|                  |         | Total  | -              | 113    | 10    | 123   |
|                  | Week 24 | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 110    | 6     | 116   |
|                  |         | Upper  | -              | 2      | 4     | 6     |
|                  |         | Total  | -              | 112    | 10    | 122   |

表 2.7.4.7-付録-26 血液生化学検査の評価時点別の変化量 (OCT-001 試験) (1/5)

| Total Protein (g/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|----------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit                | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2               | ATL-962 120 mg | 127                | -0.01 | 0.274 | -0.7 | -0.20 | 0.00   | 0.20 | 0.6 |
| Week 4               | ATL-962 120 mg | 126                | -0.04 | 0.300 | -0.6 | -0.30 | 0.00   | 0.20 | 0.7 |
| Week 8               | ATL-962 120 mg | 126                | 0.01  | 0.320 | -0.9 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 12              | ATL-962 120 mg | 124                | -0.01 | 0.310 | -1.0 | -0.20 | 0.00   | 0.20 | 0.8 |
| Week 16              | ATL-962 120 mg | 124                | 0.04  | 0.305 | -0.8 | -0.20 | 0.00   | 0.20 | 1.2 |
| Week 20              | ATL-962 120 mg | 123                | 0.03  | 0.314 | -0.9 | -0.10 | 0.00   | 0.20 | 1.1 |
| Week 24              | ATL-962 120 mg | 122                | 0.05  | 0.355 | -0.9 | -0.20 | 0.05   | 0.20 | 1.3 |

  

| Albumin (g/dL) |                | Summary Statistics |      |       |      |       |        |      |     |
|----------------|----------------|--------------------|------|-------|------|-------|--------|------|-----|
| Visit          | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2         | ATL-962 120 mg | 127                | 0.03 | 0.175 | -0.4 | -0.10 | 0.00   | 0.20 | 0.5 |
| Week 4         | ATL-962 120 mg | 126                | 0.01 | 0.191 | -0.5 | -0.10 | 0.00   | 0.20 | 0.4 |
| Week 8         | ATL-962 120 mg | 126                | 0.05 | 0.210 | -0.4 | -0.10 | 0.00   | 0.20 | 0.6 |
| Week 12        | ATL-962 120 mg | 124                | 0.00 | 0.189 | -0.5 | -0.10 | 0.00   | 0.10 | 0.5 |
| Week 16        | ATL-962 120 mg | 124                | 0.05 | 0.209 | -0.5 | -0.10 | 0.10   | 0.20 | 0.6 |
| Week 20        | ATL-962 120 mg | 123                | 0.08 | 0.203 | -0.3 | -0.10 | 0.10   | 0.20 | 0.7 |
| Week 24        | ATL-962 120 mg | 122                | 0.07 | 0.221 | -0.5 | -0.10 | 0.10   | 0.20 | 0.8 |

  

| ALT (GPT) (IU/L) |                | Summary Statistics |      |       |     |      |        |     |     |
|------------------|----------------|--------------------|------|-------|-----|------|--------|-----|-----|
| Visit            | Treatment      | N                  | Mean | SD    | Min | Q1   | Median | Q3  | Max |
| Week 2           | ATL-962 120 mg | 127                | 2.6  | 11.59 | -36 | -2.0 | 2.0    | 6.0 | 41  |
| Week 4           | ATL-962 120 mg | 126                | 4.6  | 17.04 | -28 | -3.0 | 1.5    | 8.0 | 92  |
| Week 8           | ATL-962 120 mg | 126                | 2.2  | 23.87 | -38 | -5.0 | 0.5    | 6.0 | 234 |
| Week 12          | ATL-962 120 mg | 124                | 1.1  | 14.70 | -42 | -6.0 | 1.0    | 9.0 | 64  |
| Week 16          | ATL-962 120 mg | 124                | 0.3  | 15.72 | -46 | -7.0 | 0.5    | 6.5 | 69  |
| Week 20          | ATL-962 120 mg | 123                | 0.1  | 19.59 | -51 | -7.0 | 0.0    | 6.0 | 138 |
| Week 24          | ATL-962 120 mg | 122                | -0.9 | 14.85 | -62 | -6.0 | -0.5   | 6.0 | 56  |

  

| AST (GOT) (IU/L) |                | Summary Statistics |      |       |     |      |        |     |     |
|------------------|----------------|--------------------|------|-------|-----|------|--------|-----|-----|
| Visit            | Treatment      | N                  | Mean | SD    | Min | Q1   | Median | Q3  | Max |
| Week 2           | ATL-962 120 mg | 127                | 1.1  | 9.96  | -41 | -2.0 | 0.0    | 4.0 | 34  |
| Week 4           | ATL-962 120 mg | 126                | 1.7  | 11.62 | -30 | -2.0 | 1.0    | 4.0 | 58  |
| Week 8           | ATL-962 120 mg | 126                | 0.4  | 12.81 | -49 | -4.0 | 0.0    | 4.0 | 86  |
| Week 12          | ATL-962 120 mg | 124                | -0.0 | 12.17 | -39 | -5.0 | 0.0    | 4.0 | 72  |
| Week 16          | ATL-962 120 mg | 124                | -1.2 | 10.70 | -42 | -6.0 | 0.0    | 3.0 | 34  |
| Week 20          | ATL-962 120 mg | 123                | -0.5 | 13.78 | -42 | -5.0 | 0.0    | 3.0 | 81  |
| Week 24          | ATL-962 120 mg | 122                | -1.5 | 11.25 | -51 | -6.0 | 0.0    | 4.0 | 50  |

  

| γ-GTP (IU/L) |                | Summary Statistics |      |       |      |       |        |     |     |
|--------------|----------------|--------------------|------|-------|------|-------|--------|-----|-----|
| Visit        | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3  | Max |
| Week 2       | ATL-962 120 mg | 127                | -3.6 | 22.78 | -213 | -7.0  | -2.0   | 3.0 | 69  |
| Week 4       | ATL-962 120 mg | 126                | -3.1 | 32.09 | -280 | -7.0  | -2.0   | 4.0 | 122 |
| Week 8       | ATL-962 120 mg | 126                | -3.2 | 45.47 | -312 | -7.0  | -1.5   | 3.0 | 352 |
| Week 12      | ATL-962 120 mg | 124                | -1.1 | 29.26 | -84  | -8.5  | -2.0   | 6.0 | 233 |
| Week 16      | ATL-962 120 mg | 124                | -5.3 | 34.62 | -275 | -9.5  | -3.5   | 5.0 | 129 |
| Week 20      | ATL-962 120 mg | 123                | -4.9 | 22.60 | -96  | -12.0 | -3.0   | 4.0 | 102 |
| Week 24      | ATL-962 120 mg | 122                | -4.9 | 37.69 | -248 | -13.0 | -3.0   | 4.0 | 195 |

表 2.7.4.7-付録-26 血液生化学検査の評価時点別の変化量 (OCT-001 試験) (2/5)

| ALP (IU/L) |                | Summary Statistics |      |       |      |       |        |      |     |
|------------|----------------|--------------------|------|-------|------|-------|--------|------|-----|
| Visit      | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2     | ATL-962 120 mg | 127                | -2.5 | 24.74 | -123 | -11.0 | 0.0    | 13.0 | 74  |
| Week 4     | ATL-962 120 mg | 126                | 2.3  | 29.33 | -111 | -12.0 | 3.5    | 17.0 | 122 |
| Week 8     | ATL-962 120 mg | 126                | 3.3  | 33.91 | -84  | -10.0 | 3.0    | 18.0 | 227 |
| Week 12    | ATL-962 120 mg | 124                | 3.7  | 32.41 | -133 | -9.0  | 5.0    | 23.0 | 102 |
| Week 16    | ATL-962 120 mg | 124                | 9.1  | 34.69 | -110 | -9.0  | 7.0    | 27.0 | 110 |
| Week 20    | ATL-962 120 mg | 123                | 7.9  | 31.39 | -102 | -11.0 | 8.0    | 23.0 | 88  |
| Week 24    | ATL-962 120 mg | 122                | 9.7  | 33.56 | -89  | -9.0  | 10.0   | 25.0 | 173 |

  

| LDH (IU/L) |                | Summary Statistics |      |       |     |       |        |      |     |
|------------|----------------|--------------------|------|-------|-----|-------|--------|------|-----|
| Visit      | Treatment      | N                  | Mean | SD    | Min | Q1    | Median | Q3   | Max |
| Week 2     | ATL-962 120 mg | 127                | -0.6 | 16.43 | -54 | -10.0 | 1.0    | 10.0 | 43  |
| Week 4     | ATL-962 120 mg | 126                | -2.9 | 16.36 | -55 | -12.0 | -3.0   | 6.0  | 66  |
| Week 8     | ATL-962 120 mg | 126                | 0.3  | 18.43 | -51 | -12.0 | 0.0    | 11.0 | 61  |
| Week 12    | ATL-962 120 mg | 124                | 0.1  | 21.05 | -53 | -11.5 | -1.0   | 8.5  | 87  |
| Week 16    | ATL-962 120 mg | 124                | -2.9 | 19.20 | -55 | -13.0 | -2.0   | 10.0 | 77  |
| Week 20    | ATL-962 120 mg | 123                | 2.0  | 21.70 | -60 | -9.0  | 2.0    | 13.0 | 119 |
| Week 24    | ATL-962 120 mg | 122                | 1.4  | 19.35 | -45 | -10.0 | 3.5    | 15.0 | 51  |

  

| CPK (IU/L) |                | Summary Statistics |      |        |      |       |        |      |      |
|------------|----------------|--------------------|------|--------|------|-------|--------|------|------|
| Visit      | Treatment      | N                  | Mean | SD     | Min  | Q1    | Median | Q3   | Max  |
| Week 2     | ATL-962 120 mg | 127                | 9.1  | 190.90 | -484 | -17.0 | -3.0   | 20.0 | 2026 |
| Week 4     | ATL-962 120 mg | 126                | -5.3 | 66.77  | -537 | -17.0 | -1.0   | 14.0 | 222  |
| Week 8     | ATL-962 120 mg | 126                | 9.6  | 99.95  | -533 | -20.0 | 1.5    | 24.0 | 818  |
| Week 12    | ATL-962 120 mg | 124                | -0.8 | 70.12  | -547 | -12.0 | 2.0    | 22.0 | 326  |
| Week 16    | ATL-962 120 mg | 124                | -6.1 | 64.58  | -537 | -17.5 | 0.0    | 19.0 | 124  |
| Week 20    | ATL-962 120 mg | 123                | 26.4 | 266.91 | -539 | -11.0 | 6.0    | 23.0 | 2873 |
| Week 24    | ATL-962 120 mg | 122                | 10.1 | 68.23  | -412 | -14.0 | 6.0    | 28.0 | 332  |

  

| Uric Acid (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2            | ATL-962 120 mg | 127                | 0.04  | 0.712 | -2.4 | -0.30 | 0.10   | 0.50 | 1.6 |
| Week 4            | ATL-962 120 mg | 126                | 0.03  | 0.697 | -1.9 | -0.40 | 0.05   | 0.50 | 1.6 |
| Week 8            | ATL-962 120 mg | 126                | -0.04 | 0.761 | -2.5 | -0.50 | 0.00   | 0.40 | 2.3 |
| Week 12           | ATL-962 120 mg | 124                | -0.14 | 0.687 | -2.4 | -0.50 | -0.10  | 0.40 | 1.5 |
| Week 16           | ATL-962 120 mg | 124                | -0.16 | 0.725 | -3.1 | -0.60 | -0.10  | 0.30 | 1.2 |
| Week 20           | ATL-962 120 mg | 123                | -0.22 | 0.718 | -2.4 | -0.60 | -0.10  | 0.30 | 1.7 |
| Week 24           | ATL-962 120 mg | 122                | -0.19 | 0.742 | -3.4 | -0.60 | -0.20  | 0.20 | 1.5 |

  

| Total Bilirubin (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-------------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit                   | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2                  | ATL-962 120 mg | 127                | -0.01 | 0.217 | -0.9 | -0.10 | 0.00   | 0.10 | 0.8 |
| Week 4                  | ATL-962 120 mg | 126                | -0.02 | 0.193 | -0.5 | -0.10 | 0.00   | 0.10 | 0.6 |
| Week 8                  | ATL-962 120 mg | 126                | -0.00 | 0.176 | -0.6 | -0.10 | 0.00   | 0.10 | 0.5 |
| Week 12                 | ATL-962 120 mg | 124                | -0.02 | 0.205 | -0.7 | -0.20 | 0.00   | 0.10 | 0.7 |
| Week 16                 | ATL-962 120 mg | 124                | -0.04 | 0.174 | -0.5 | -0.20 | 0.00   | 0.10 | 0.4 |
| Week 20                 | ATL-962 120 mg | 123                | -0.01 | 0.197 | -0.5 | -0.20 | 0.00   | 0.10 | 0.7 |
| Week 24                 | ATL-962 120 mg | 122                | -0.04 | 0.211 | -0.6 | -0.20 | 0.00   | 0.10 | 0.6 |

表 2.7.4.7-付録-26 血液生化学検査の評価時点別の変化量 (OCT-001 試験) (3/5)

| ChE (U/L) |                | Summary Statistics |       |       |      |       |        |      |     |
|-----------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit     | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2    | ATL-962 120 mg | 127                | -11.3 | 22.92 | -73  | -27.0 | -9.0   | 3.0  | 45  |
| Week 4    | ATL-962 120 mg | 126                | -15.1 | 27.57 | -102 | -31.0 | -14.0  | 3.0  | 51  |
| Week 8    | ATL-962 120 mg | 126                | -14.5 | 27.62 | -100 | -35.0 | -12.0  | 2.0  | 50  |
| Week 12   | ATL-962 120 mg | 124                | -10.6 | 33.16 | -120 | -28.0 | -12.0  | 8.0  | 127 |
| Week 16   | ATL-962 120 mg | 124                | -6.8  | 33.08 | -119 | -26.5 | -6.5   | 11.0 | 94  |
| Week 20   | ATL-962 120 mg | 123                | -4.0  | 32.20 | -168 | -22.0 | -4.0   | 18.0 | 65  |
| Week 24   | ATL-962 120 mg | 122                | 1.0   | 41.67 | -226 | -20.0 | 0.0    | 26.0 | 105 |

  

| Creatinine (mg/dL) |                | Summary Statistics |        |        |       |        |        |       |      |
|--------------------|----------------|--------------------|--------|--------|-------|--------|--------|-------|------|
| Visit              | Treatment      | N                  | Mean   | SD     | Min   | Q1     | Median | Q3    | Max  |
| Week 2             | ATL-962 120 mg | 127                | 0.013  | 0.0627 | -0.16 | -0.030 | 0.020  | 0.050 | 0.24 |
| Week 4             | ATL-962 120 mg | 126                | 0.011  | 0.0620 | -0.13 | -0.030 | 0.020  | 0.050 | 0.16 |
| Week 8             | ATL-962 120 mg | 126                | 0.017  | 0.0597 | -0.18 | -0.010 | 0.015  | 0.050 | 0.27 |
| Week 12            | ATL-962 120 mg | 124                | 0.008  | 0.0649 | -0.18 | -0.030 | 0.010  | 0.050 | 0.22 |
| Week 16            | ATL-962 120 mg | 124                | 0.009  | 0.0678 | -0.20 | -0.030 | 0.010  | 0.050 | 0.18 |
| Week 20            | ATL-962 120 mg | 123                | 0.005  | 0.0687 | -0.31 | -0.020 | 0.010  | 0.050 | 0.15 |
| Week 24            | ATL-962 120 mg | 122                | -0.001 | 0.0664 | -0.23 | -0.040 | 0.000  | 0.040 | 0.17 |

  

| BUN (mg/dL) |                | Summary Statistics |      |      |     |      |        |     |     |
|-------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit       | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2      | ATL-962 120 mg | 127                | 0.1  | 3.31 | -9  | -1.0 | 0.0    | 2.0 | 8   |
| Week 4      | ATL-962 120 mg | 126                | -0.2 | 2.94 | -9  | -2.0 | 0.0    | 2.0 | 8   |
| Week 8      | ATL-962 120 mg | 126                | -0.1 | 3.20 | -11 | -2.0 | 0.0    | 2.0 | 9   |
| Week 12     | ATL-962 120 mg | 124                | -0.7 | 2.92 | -11 | -2.0 | -1.0   | 1.0 | 10  |
| Week 16     | ATL-962 120 mg | 124                | -0.1 | 2.73 | -9  | -1.0 | 0.0    | 1.0 | 10  |
| Week 20     | ATL-962 120 mg | 123                | -0.3 | 3.11 | -11 | -2.0 | 0.0    | 2.0 | 9   |
| Week 24     | ATL-962 120 mg | 122                | 0.3  | 2.98 | -7  | -2.0 | 0.0    | 2.0 | 9   |

  

| Na (mEq/L) |                | Summary Statistics |      |      |     |      |        |     |     |
|------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit      | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2     | ATL-962 120 mg | 127                | 0.1  | 1.62 | -4  | -1.0 | 0.0    | 1.0 | 5   |
| Week 4     | ATL-962 120 mg | 126                | 0.2  | 1.55 | -5  | -1.0 | 0.0    | 1.0 | 3   |
| Week 8     | ATL-962 120 mg | 126                | -0.1 | 1.70 | -4  | -1.0 | 0.0    | 1.0 | 4   |
| Week 12    | ATL-962 120 mg | 124                | -0.1 | 1.63 | -5  | -1.0 | 0.0    | 1.0 | 5   |
| Week 16    | ATL-962 120 mg | 124                | 0.0  | 1.48 | -3  | -1.0 | 0.0    | 1.0 | 4   |
| Week 20    | ATL-962 120 mg | 123                | -0.0 | 1.70 | -4  | -1.0 | 0.0    | 1.0 | 7   |
| Week 24    | ATL-962 120 mg | 122                | -0.0 | 1.58 | -5  | -1.0 | 0.0    | 1.0 | 4   |

  

| K (mEq/L) |                | Summary Statistics |       |       |      |       |        |      |     |
|-----------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit     | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2    | ATL-962 120 mg | 127                | -0.04 | 0.245 | -0.6 | -0.20 | 0.00   | 0.10 | 0.8 |
| Week 4    | ATL-962 120 mg | 126                | -0.03 | 0.231 | -0.8 | -0.20 | 0.00   | 0.10 | 0.8 |
| Week 8    | ATL-962 120 mg | 126                | -0.05 | 0.252 | -0.7 | -0.20 | 0.00   | 0.10 | 0.6 |
| Week 12   | ATL-962 120 mg | 124                | -0.03 | 0.255 | -0.6 | -0.20 | 0.00   | 0.10 | 0.5 |
| Week 16   | ATL-962 120 mg | 124                | 0.02  | 0.303 | -0.8 | -0.20 | 0.00   | 0.20 | 1.0 |
| Week 20   | ATL-962 120 mg | 123                | 0.03  | 0.299 | -0.8 | -0.20 | 0.00   | 0.20 | 1.0 |
| Week 24   | ATL-962 120 mg | 122                | 0.02  | 0.298 | -0.7 | -0.20 | 0.00   | 0.20 | 0.9 |

表 2.7.4.7-付録-26 血液生化学検査の評価時点別の変化量 (OCT-001 試験) (4/5)

| Cl (mEq/L) |                | Summary Statistics |      |      |     |      |        |     |     |
|------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit      | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2     | ATL-962 120 mg | 127                | 0.5  | 1.81 | -4  | -1.0 | 1.0    | 2.0 | 4   |
| Week 4     | ATL-962 120 mg | 126                | 0.4  | 1.80 | -3  | -1.0 | 0.0    | 1.0 | 6   |
| Week 8     | ATL-962 120 mg | 126                | 0.2  | 2.02 | -6  | -1.0 | 0.0    | 2.0 | 5   |
| Week 12    | ATL-962 120 mg | 124                | 0.0  | 2.18 | -8  | -1.0 | 0.0    | 1.0 | 5   |
| Week 16    | ATL-962 120 mg | 124                | -0.1 | 1.93 | -4  | -1.0 | 0.0    | 1.0 | 6   |
| Week 20    | ATL-962 120 mg | 123                | -0.2 | 1.88 | -6  | -1.0 | 0.0    | 1.0 | 4   |
| Week 24    | ATL-962 120 mg | 122                | -0.3 | 2.08 | -6  | -2.0 | 0.0    | 1.0 | 4   |

  

| Ca (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit      | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2     | ATL-962 120 mg | 127                | -0.01 | 0.272 | -0.7 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 4     | ATL-962 120 mg | 126                | -0.02 | 0.295 | -0.9 | -0.20 | 0.00   | 0.20 | 0.8 |
| Week 8     | ATL-962 120 mg | 126                | -0.03 | 0.319 | -0.9 | -0.30 | 0.00   | 0.20 | 0.7 |
| Week 12    | ATL-962 120 mg | 124                | 0.00  | 0.276 | -0.8 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 16    | ATL-962 120 mg | 124                | 0.01  | 0.293 | -0.7 | -0.20 | 0.00   | 0.20 | 0.6 |
| Week 20    | ATL-962 120 mg | 123                | 0.02  | 0.293 | -0.6 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 24    | ATL-962 120 mg | 122                | 0.01  | 0.326 | -0.7 | -0.20 | 0.00   | 0.20 | 1.0 |

  

| P (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-----------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit     | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2    | ATL-962 120 mg | 127                | 0.03  | 0.399 | -1.2 | -0.20 | 0.00   | 0.30 | 1.2 |
| Week 4    | ATL-962 120 mg | 126                | 0.02  | 0.406 | -1.0 | -0.20 | 0.00   | 0.30 | 1.1 |
| Week 8    | ATL-962 120 mg | 126                | 0.06  | 0.384 | -1.0 | -0.20 | 0.00   | 0.30 | 1.8 |
| Week 12   | ATL-962 120 mg | 124                | -0.00 | 0.340 | -0.8 | -0.25 | 0.00   | 0.20 | 0.7 |
| Week 16   | ATL-962 120 mg | 124                | 0.07  | 0.381 | -1.0 | -0.20 | 0.00   | 0.30 | 1.2 |
| Week 20   | ATL-962 120 mg | 123                | 0.05  | 0.364 | -1.1 | -0.20 | 0.10   | 0.30 | 1.0 |
| Week 24   | ATL-962 120 mg | 122                | 0.07  | 0.345 | -1.0 | -0.10 | 0.00   | 0.30 | 0.9 |

  

| Vitamin A (ng/mL) |                | Summary Statistics |       |        |      |       |        |      |     |
|-------------------|----------------|--------------------|-------|--------|------|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean  | SD     | Min  | Q1    | Median | Q3   | Max |
| Week 2            | ATL-962 120 mg | 127                | -19.6 | 103.73 | -421 | -69.0 | -16.0  | 42.0 | 338 |
| Week 4            | ATL-962 120 mg | 126                | -14.9 | 105.50 | -684 | -71.0 | -8.0   | 40.0 | 350 |
| Week 8            | ATL-962 120 mg | 126                | -21.1 | 91.04  | -603 | -64.0 | -16.5  | 36.0 | 172 |
| Week 12           | ATL-962 120 mg | 124                | -14.8 | 94.51  | -586 | -54.5 | -3.0   | 39.5 | 183 |
| Week 16           | ATL-962 120 mg | 124                | -22.3 | 112.66 | -700 | -68.0 | -9.5   | 37.0 | 281 |
| Week 20           | ATL-962 120 mg | 123                | -18.4 | 128.42 | -839 | -66.0 | -30.0  | 60.0 | 503 |
| Week 24           | ATL-962 120 mg | 122                | -17.8 | 114.65 | -660 | -84.0 | -12.5  | 43.0 | 451 |

  

| Vitamin D (pg/mL) |                | Summary Statistics |      |       |     |       |        |      |     |
|-------------------|----------------|--------------------|------|-------|-----|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean | SD    | Min | Q1    | Median | Q3   | Max |
| Week 2            | ATL-962 120 mg | 127                | -2.2 | 18.46 | -50 | -13.0 | -4.0   | 9.0  | 55  |
| Week 4            | ATL-962 120 mg | 126                | -3.6 | 20.21 | -59 | -16.0 | -4.0   | 7.0  | 56  |
| Week 8            | ATL-962 120 mg | 126                | -1.8 | 19.18 | -52 | -14.0 | -0.5   | 11.0 | 44  |
| Week 12           | ATL-962 120 mg | 124                | -0.9 | 19.27 | -66 | -13.0 | 0.0    | 13.0 | 44  |
| Week 16           | ATL-962 120 mg | 124                | -2.8 | 19.74 | -56 | -16.0 | -2.0   | 11.0 | 53  |
| Week 20           | ATL-962 120 mg | 123                | -0.8 | 19.26 | -71 | -10.0 | -1.0   | 12.0 | 46  |
| Week 24           | ATL-962 120 mg | 122                | -1.6 | 20.67 | -87 | -12.0 | -2.0   | 9.0  | 59  |

表 2.7.4.7-付録-26 血液生化学検査の評価時点別の変化量 (OCT-001 試験) (5/5)

| Vitamin E (mg/dL) |                | Summary Statistics |        |        |       |        |        |        |      |
|-------------------|----------------|--------------------|--------|--------|-------|--------|--------|--------|------|
| Visit             | Treatment      | N                  | Mean   | SD     | Min   | Q1     | Median | Q3     | Max  |
| Week 2            | ATL-962 120 mg | 127                | -0.232 | 0.3301 | -1.82 | -0.410 | -0.190 | -0.050 | 0.86 |
| Week 4            | ATL-962 120 mg | 126                | -0.246 | 0.3258 | -1.51 | -0.380 | -0.225 | -0.030 | 1.29 |
| Week 8            | ATL-962 120 mg | 126                | -0.301 | 0.3303 | -1.94 | -0.470 | -0.255 | -0.100 | 0.56 |
| Week 12           | ATL-962 120 mg | 124                | -0.320 | 0.3627 | -1.59 | -0.505 | -0.310 | -0.140 | 0.75 |
| Week 16           | ATL-962 120 mg | 124                | -0.381 | 0.3051 | -1.27 | -0.520 | -0.375 | -0.160 | 0.30 |
| Week 20           | ATL-962 120 mg | 123                | -0.368 | 0.3403 | -1.62 | -0.540 | -0.310 | -0.160 | 0.61 |
| Week 24           | ATL-962 120 mg | 122                | -0.379 | 0.3376 | -1.56 | -0.550 | -0.320 | -0.160 | 0.63 |

  

| β-Carotene (μg/dL) |                | Summary Statistics |       |        |        |       |        |      |      |
|--------------------|----------------|--------------------|-------|--------|--------|-------|--------|------|------|
| Visit              | Treatment      | N                  | Mean  | SD     | Min    | Q1    | Median | Q3   | Max  |
| Week 2             | ATL-962 120 mg | 127                | -2.23 | 6.882  | -41.5  | -4.40 | -1.20  | 0.70 | 18.9 |
| Week 4             | ATL-962 120 mg | 126                | -2.16 | 8.324  | -38.5  | -4.90 | -1.25  | 2.20 | 20.8 |
| Week 8             | ATL-962 120 mg | 126                | -3.29 | 8.846  | -29.5  | -5.50 | -1.65  | 1.10 | 17.0 |
| Week 12            | ATL-962 120 mg | 124                | -3.89 | 11.797 | -86.7  | -6.10 | -1.80  | 0.90 | 17.7 |
| Week 16            | ATL-962 120 mg | 124                | -4.96 | 13.653 | -106.6 | -8.00 | -2.00  | 0.85 | 19.8 |
| Week 20            | ATL-962 120 mg | 123                | -4.07 | 13.482 | -111.0 | -6.80 | -2.40  | 2.40 | 27.1 |
| Week 24            | ATL-962 120 mg | 122                | -4.45 | 13.885 | -114.5 | -5.60 | -1.80  | 1.10 | 18.4 |

  

| m-AST (IU/L) |                | Summary Statistics |      |      |     |      |        |     |     |
|--------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit        | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2       | ATL-962 120 mg | 127                | 0.4  | 2.36 | -6  | -1.0 | 0.0    | 1.0 | 15  |
| Week 4       | ATL-962 120 mg | 126                | 0.4  | 2.81 | -6  | -1.0 | 0.0    | 1.0 | 19  |
| Week 8       | ATL-962 120 mg | 126                | 0.4  | 2.65 | -9  | -1.0 | 0.0    | 1.0 | 13  |
| Week 12      | ATL-962 120 mg | 124                | 0.6  | 2.83 | -6  | -1.0 | 0.0    | 1.0 | 17  |
| Week 16      | ATL-962 120 mg | 124                | -0.1 | 2.06 | -7  | -1.0 | 0.0    | 1.0 | 11  |
| Week 20      | ATL-962 120 mg | 123                | 0.1  | 2.49 | -7  | -1.0 | 0.0    | 1.0 | 18  |
| Week 24      | ATL-962 120 mg | 122                | 0.1  | 2.46 | -9  | -1.0 | 0.0    | 1.0 | 14  |

  

| CK18 (Segment) (U/L) |                | Summary Statistics |        |         |        |         |        |        |       |
|----------------------|----------------|--------------------|--------|---------|--------|---------|--------|--------|-------|
| Visit                | Treatment      | N                  | Mean   | SD      | Min    | Q1      | Median | Q3     | Max   |
| Week 2               | ATL-962 120 mg | 127                | -13.01 | 138.565 | -575.5 | -59.90  | -7.60  | 27.20  | 560.9 |
| Week 4               | ATL-962 120 mg | 126                | -15.72 | 147.595 | -472.0 | -76.00  | -14.70 | 44.20  | 583.1 |
| Week 8               | ATL-962 120 mg | 126                | -24.08 | 159.816 | -570.5 | -85.70  | -16.65 | 28.60  | 702.7 |
| Week 12              | ATL-962 120 mg | 124                | -41.50 | 148.319 | -663.4 | -91.10  | -30.40 | 17.50  | 812.6 |
| Week 16              | ATL-962 120 mg | 124                | -52.61 | 161.079 | -635.0 | -96.40  | -45.85 | 10.35  | 564.7 |
| Week 20              | ATL-962 120 mg | 123                | -67.64 | 164.434 | -651.7 | -133.00 | -57.60 | 3.90   | 687.5 |
| Week 24              | ATL-962 120 mg | 122                | -91.31 | 170.836 | -789.5 | -142.40 | -58.75 | -14.20 | 812.6 |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表(OCT-001 試験)(1/12)

|               |         |        | ATL-962 120 mg |        |       |       |
|---------------|---------|--------|----------------|--------|-------|-------|
|               |         |        | Week 0         |        |       |       |
|               |         |        | Lower          | Normal | Upper | Total |
| Total Protein | Week 2  | Lower  | 7              | 4      | -     | 11    |
|               |         | Normal | 3              | 112    | -     | 115   |
|               |         | Upper  | -              | 1      | -     | 1     |
|               |         | Total  | 10             | 117    | -     | 127   |
|               | Week 4  | Lower  | 5              | 6      | -     | 11    |
|               |         | Normal | 5              | 108    | -     | 113   |
|               |         | Upper  | -              | 2      | -     | 2     |
|               |         | Total  | 10             | 116    | -     | 126   |
|               | Week 8  | Lower  | 5              | 2      | -     | 7     |
|               |         | Normal | 5              | 113    | -     | 118   |
|               |         | Upper  | -              | 1      | -     | 1     |
|               |         | Total  | 10             | 116    | -     | 126   |
|               | Week 12 | Lower  | 8              | 1      | -     | 9     |
|               |         | Normal | 2              | 113    | -     | 115   |
|               |         | Upper  | -              | -      | -     | -     |
|               |         | Total  | 10             | 114    | -     | 124   |
|               | Week 16 | Lower  | 5              | 1      | -     | 6     |
|               |         | Normal | 5              | 112    | -     | 117   |
|               |         | Upper  | -              | 1      | -     | 1     |
|               |         | Total  | 10             | 114    | -     | 124   |
|               | Week 20 | Lower  | 5              | 4      | -     | 9     |
|               |         | Normal | 5              | 108    | -     | 113   |
|               |         | Upper  | -              | 1      | -     | 1     |
|               |         | Total  | 10             | 113    | -     | 123   |
| Week 24       | Lower   | 8      | 4              | -      | 12    |       |
|               | Normal  | 2      | 104            | -      | 106   |       |
|               | Upper   | -      | 4              | -      | 4     |       |
|               | Total   | 10     | 112            | -      | 122   |       |
| Albumin       | Week 2  | Lower  | 1              | -      | -     | 1     |
|               |         | Normal | 1              | 125    | -     | 126   |
|               |         | Upper  | -              | -      | -     | -     |
|               |         | Total  | 2              | 125    | -     | 127   |
|               | Week 4  | Lower  | -              | -      | -     | -     |
|               |         | Normal | 2              | 124    | -     | 126   |
|               |         | Upper  | -              | -      | -     | -     |
|               |         | Total  | 2              | 124    | -     | 126   |
|               | Week 8  | Lower  | 2              | 1      | -     | 3     |
|               |         | Normal | -              | 123    | -     | 123   |
|               |         | Upper  | -              | -      | -     | -     |
|               |         | Total  | 2              | 124    | -     | 126   |
|               | Week 12 | Lower  | 1              | -      | -     | 1     |
|               |         | Normal | 1              | 122    | -     | 123   |
|               |         | Upper  | -              | -      | -     | -     |
|               |         | Total  | 2              | 122    | -     | 124   |
|               | Week 16 | Lower  | -              | -      | -     | -     |
|               |         | Normal | 2              | 122    | -     | 124   |
|               |         | Upper  | -              | -      | -     | -     |
|               |         | Total  | 2              | 122    | -     | 124   |
|               | Week 20 | Lower  | -              | -      | -     | -     |
|               |         | Normal | 2              | 120    | -     | 122   |
|               |         | Upper  | -              | 1      | -     | 1     |
|               |         | Total  | 2              | 121    | -     | 123   |
| Week 24       | Lower   | 1      | 2              | -      | 3     |       |
|               | Normal  | 1      | 117            | -      | 118   |       |
|               | Upper   | -      | 1              | -      | 1     |       |
|               | Total   | 2      | 120            | -      | 122   |       |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表 (OCT-001 試験) (2/12)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| ALT (GPT) | Week 2  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 82     | 5     | 87    |
|           |         | Upper  | -              | 10     | 30    | 40    |
|           |         | Total  | -              | 92     | 35    | 127   |
|           | Week 4  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 78     | 8     | 86    |
|           |         | Upper  | -              | 13     | 27    | 40    |
|           |         | Total  | -              | 91     | 35    | 126   |
|           | Week 8  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 81     | 10    | 91    |
|           |         | Upper  | -              | 10     | 25    | 35    |
|           |         | Total  | -              | 91     | 35    | 126   |
|           | Week 12 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 76     | 13    | 89    |
|           |         | Upper  | -              | 13     | 22    | 35    |
|           |         | Total  | -              | 89     | 35    | 124   |
|           | Week 16 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 77     | 15    | 92    |
|           |         | Upper  | -              | 12     | 20    | 32    |
|           |         | Total  | -              | 89     | 35    | 124   |
| Week 20   | Lower   | -      | -              | -      | -     |       |
|           | Normal  | -      | 82             | 14     | 96    |       |
|           | Upper   | -      | 6              | 21     | 27    |       |
|           | Total   | -      | 88             | 35     | 123   |       |
| Week 24   | Lower   | -      | -              | -      | -     |       |
|           | Normal  | -      | 79             | 13     | 92    |       |
|           | Upper   | -      | 8              | 22     | 30    |       |
|           | Total   | -      | 87             | 35     | 122   |       |
| AST (GOT) | Week 2  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 96     | 7     | 103   |
|           |         | Upper  | -              | 9      | 15    | 24    |
|           |         | Total  | -              | 105    | 22    | 127   |
|           | Week 4  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 94     | 8     | 102   |
|           |         | Upper  | -              | 10     | 14    | 24    |
|           |         | Total  | -              | 104    | 22    | 126   |
|           | Week 8  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 97     | 7     | 104   |
|           |         | Upper  | -              | 7      | 15    | 22    |
|           |         | Total  | -              | 104    | 22    | 126   |
|           | Week 12 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 98     | 9     | 107   |
|           |         | Upper  | -              | 4      | 13    | 17    |
|           |         | Total  | -              | 102    | 22    | 124   |
|           | Week 16 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 96     | 10    | 106   |
|           |         | Upper  | -              | 6      | 12    | 18    |
|           |         | Total  | -              | 102    | 22    | 124   |
| Week 20   | Lower   | -      | -              | -      | -     |       |
|           | Normal  | -      | 96             | 10     | 106   |       |
|           | Upper   | -      | 5              | 12     | 17    |       |
|           | Total   | -      | 101            | 22     | 123   |       |
| Week 24   | Lower   | -      | -              | -      | -     |       |
|           | Normal  | -      | 96             | 14     | 110   |       |
|           | Upper   | -      | 4              | 8      | 12    |       |
|           | Total   | -      | 100            | 22     | 122   |       |



表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表 (OCT-001 試験) (3/12)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| γ-GTP   | Week 2  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 80     | 1     | 81    |
|         |         | Upper  | -              | 2      | 44    | 46    |
|         |         | Total  | -              | 82     | 45    | 127   |
|         | Week 4  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 78     | 5     | 83    |
|         |         | Upper  | -              | 4      | 39    | 43    |
|         |         | Total  | -              | 82     | 44    | 126   |
|         | Week 8  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 77     | 6     | 83    |
|         |         | Upper  | -              | 5      | 38    | 43    |
|         |         | Total  | -              | 82     | 44    | 126   |
|         | Week 12 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 78     | 7     | 85    |
|         |         | Upper  | -              | 4      | 35    | 39    |
|         |         | Total  | -              | 82     | 42    | 124   |
|         | Week 16 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 77     | 7     | 84    |
|         |         | Upper  | -              | 5      | 35    | 40    |
|         |         | Total  | -              | 82     | 42    | 124   |
| Week 20 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 79             | 6      | 85    |       |
|         | Upper   | -      | 2              | 36     | 38    |       |
|         | Total   | -      | 81             | 42     | 123   |       |
| Week 24 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 76             | 9      | 85    |       |
|         | Upper   | -      | 4              | 33     | 37    |       |
|         | Total   | -      | 80             | 42     | 122   |       |
| ALP     | Week 2  | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | -              | 112    | 3     | 115   |
|         |         | Upper  | -              | -      | 11    | 11    |
|         |         | Total  | 1              | 112    | 14    | 127   |
|         | Week 4  | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | -              | 109    | 2     | 111   |
|         |         | Upper  | -              | 2      | 12    | 14    |
|         |         | Total  | 1              | 111    | 14    | 126   |
|         | Week 8  | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | -              | 109    | 2     | 111   |
|         |         | Upper  | -              | 2      | 12    | 14    |
|         |         | Total  | 1              | 111    | 14    | 126   |
|         | Week 12 | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | -              | 108    | 4     | 112   |
|         |         | Upper  | -              | 1      | 10    | 11    |
|         |         | Total  | 1              | 109    | 14    | 124   |
|         | Week 16 | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | -              | 109    | 3     | 112   |
|         |         | Upper  | -              | -      | 11    | 11    |
|         |         | Total  | 1              | 109    | 14    | 124   |
| Week 20 | Lower   | 1      | -              | -      | 1     |       |
|         | Normal  | -      | 106            | 2      | 108   |       |
|         | Upper   | -      | 2              | 12     | 14    |       |
|         | Total   | 1      | 108            | 14     | 123   |       |
| Week 24 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | 1      | 107            | 2      | 110   |       |
|         | Upper   | -      | -              | 12     | 12    |       |
|         | Total   | 1      | 107            | 14     | 122   |       |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表 (OCT-001 試験) (4/12)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| LDH     | Week 2  | Lower  | 2              | -      | -     | 2     |
|         |         | Normal | 1              | 114    | 4     | 119   |
|         |         | Upper  | -              | 2      | 4     | 6     |
|         |         | Total  | 3              | 116    | 8     | 127   |
|         | Week 4  | Lower  | 3              | -      | -     | 3     |
|         |         | Normal | -              | 113    | 5     | 118   |
|         |         | Upper  | -              | 2      | 3     | 5     |
|         |         | Total  | 3              | 115    | 8     | 126   |
|         | Week 8  | Lower  | 2              | -      | -     | 2     |
|         |         | Normal | 1              | 113    | 3     | 117   |
|         |         | Upper  | -              | 2      | 5     | 7     |
|         |         | Total  | 3              | 115    | 8     | 126   |
|         | Week 12 | Lower  | 2              | -      | -     | 2     |
|         |         | Normal | 1              | 110    | 4     | 115   |
|         |         | Upper  | -              | 4      | 3     | 7     |
|         |         | Total  | 3              | 114    | 7     | 124   |
|         | Week 16 | Lower  | 3              | 1      | -     | 4     |
|         |         | Normal | -              | 112    | 4     | 116   |
|         |         | Upper  | -              | 1      | 3     | 4     |
|         |         | Total  | 3              | 114    | 7     | 124   |
|         | Week 20 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 3              | 109    | 2     | 114   |
|         |         | Upper  | -              | 3      | 5     | 8     |
|         |         | Total  | 3              | 113    | 7     | 123   |
| Week 24 | Lower   | 1      | -              | -      | 1     |       |
|         | Normal  | 2      | 111            | 4      | 117   |       |
|         | Upper   | -      | 1              | 3      | 4     |       |
|         | Total   | 3      | 112            | 7      | 122   |       |
| CPK     | Week 2  | Lower  | 4              | 4      | -     | 8     |
|         |         | Normal | 2              | 106    | 5     | 113   |
|         |         | Upper  | -              | 1      | 5     | 6     |
|         |         | Total  | 6              | 111    | 10    | 127   |
|         | Week 4  | Lower  | 3              | 1      | -     | 4     |
|         |         | Normal | 3              | 106    | 6     | 115   |
|         |         | Upper  | -              | 3      | 4     | 7     |
|         |         | Total  | 6              | 110    | 10    | 126   |
|         | Week 8  | Lower  | 4              | 2      | -     | 6     |
|         |         | Normal | 2              | 101    | 4     | 107   |
|         |         | Upper  | -              | 7      | 6     | 13    |
|         |         | Total  | 6              | 110    | 10    | 126   |
|         | Week 12 | Lower  | 2              | 4      | -     | 6     |
|         |         | Normal | 4              | 100    | 3     | 107   |
|         |         | Upper  | -              | 5      | 6     | 11    |
|         |         | Total  | 6              | 109    | 9     | 124   |
|         | Week 16 | Lower  | 4              | 3      | -     | 7     |
|         |         | Normal | 2              | 104    | 4     | 110   |
|         |         | Upper  | -              | 2      | 5     | 7     |
|         |         | Total  | 6              | 109    | 9     | 124   |
|         | Week 20 | Lower  | 3              | 1      | -     | 4     |
|         |         | Normal | 3              | 103    | 2     | 108   |
|         |         | Upper  | -              | 4      | 7     | 11    |
|         |         | Total  | 6              | 108    | 9     | 123   |
| Week 24 | Lower   | 2      | 1              | -      | 3     |       |
|         | Normal  | 4      | 99             | 2      | 105   |       |
|         | Upper   | -      | 7              | 7      | 14    |       |
|         | Total   | 6      | 107            | 9      | 122   |       |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表 (OCT-001 試験) (5/12)

|                 |         |        | ATL-962 120 mg |        |       |       |
|-----------------|---------|--------|----------------|--------|-------|-------|
|                 |         |        | Week 0         |        |       |       |
|                 |         |        | Lower          | Normal | Upper | Total |
| Uric Acid       | Week 2  | Lower  | 2              | -      | -     | 2     |
|                 |         | Normal | 3              | 85     | 9     | 97    |
|                 |         | Upper  | -              | 7      | 21    | 28    |
|                 |         | Total  | 5              | 92     | 30    | 127   |
|                 | Week 4  | Lower  | 3              | 1      | -     | 4     |
|                 |         | Normal | 2              | 83     | 11    | 96    |
|                 |         | Upper  | -              | 7      | 19    | 26    |
|                 |         | Total  | 5              | 91     | 30    | 126   |
|                 | Week 8  | Lower  | 2              | 3      | -     | 5     |
|                 |         | Normal | 3              | 83     | 11    | 97    |
|                 |         | Upper  | -              | 5      | 19    | 24    |
|                 |         | Total  | 5              | 91     | 30    | 126   |
|                 | Week 12 | Lower  | 2              | 1      | -     | 3     |
|                 |         | Normal | 3              | 82     | 16    | 101   |
|                 |         | Upper  | -              | 6      | 14    | 20    |
|                 |         | Total  | 5              | 89     | 30    | 124   |
|                 | Week 16 | Lower  | 3              | 2      | -     | 5     |
|                 |         | Normal | 2              | 82     | 13    | 97    |
|                 |         | Upper  | -              | 5      | 17    | 22    |
|                 |         | Total  | 5              | 89     | 30    | 124   |
|                 | Week 20 | Lower  | 3              | 3      | -     | 6     |
|                 |         | Normal | 2              | 84     | 14    | 100   |
|                 |         | Upper  | -              | 1      | 16    | 17    |
|                 |         | Total  | 5              | 88     | 30    | 123   |
| Week 24         | Lower   | 3      | 3              | -      | 6     |       |
|                 | Normal  | 2      | 79             | 13     | 94    |       |
|                 | Upper   | -      | 5              | 17     | 22    |       |
|                 | Total   | 5      | 87             | 30     | 122   |       |
| Total Bilirubin | Week 2  | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 116    | 3     | 119   |
|                 |         | Upper  | -              | 3      | 5     | 8     |
|                 |         | Total  | -              | 119    | 8     | 127   |
|                 | Week 4  | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 116    | 2     | 118   |
|                 |         | Upper  | -              | 2      | 6     | 8     |
|                 |         | Total  | -              | 118    | 8     | 126   |
|                 | Week 8  | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 116    | 4     | 120   |
|                 |         | Upper  | -              | 2      | 4     | 6     |
|                 |         | Total  | -              | 118    | 8     | 126   |
|                 | Week 12 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 116    | 5     | 121   |
|                 |         | Upper  | -              | -      | 3     | 3     |
|                 |         | Total  | -              | 116    | 8     | 124   |
|                 | Week 16 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 116    | 3     | 119   |
|                 |         | Upper  | -              | -      | 5     | 5     |
|                 |         | Total  | -              | 116    | 8     | 124   |
|                 | Week 20 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 113    | 3     | 116   |
|                 |         | Upper  | -              | 2      | 5     | 7     |
|                 |         | Total  | -              | 115    | 8     | 123   |
| Week 24         | Lower   | -      | -              | -      | -     |       |
|                 | Normal  | -      | 112            | 3      | 115   |       |
|                 | Upper   | -      | 2              | 5      | 7     |       |
|                 | Total   | -      | 114            | 8      | 122   |       |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表 (OCT-001 試験) (6/12)

|            |         |        | ATL-962 120 mg |        |       |       |
|------------|---------|--------|----------------|--------|-------|-------|
|            |         |        | Week 0         |        |       |       |
|            |         |        | Lower          | Normal | Upper | Total |
| ChE        | Week 2  | Lower  | -              | 1      | -     | 1     |
|            |         | Normal | -              | 100    | 10    | 110   |
|            |         | Upper  | -              | 1      | 15    | 16    |
|            |         | Total  | -              | 102    | 25    | 127   |
|            | Week 4  | Lower  | -              | -      | -     | -     |
|            |         | Normal | -              | 99     | 11    | 110   |
|            |         | Upper  | -              | 3      | 13    | 16    |
|            |         | Total  | -              | 102    | 24    | 126   |
|            | Week 8  | Lower  | -              | -      | -     | -     |
|            |         | Normal | -              | 99     | 9     | 108   |
|            |         | Upper  | -              | 3      | 15    | 18    |
|            |         | Total  | -              | 102    | 24    | 126   |
|            | Week 12 | Lower  | -              | -      | -     | -     |
|            |         | Normal | -              | 95     | 12    | 107   |
|            |         | Upper  | -              | 6      | 11    | 17    |
|            |         | Total  | -              | 101    | 23    | 124   |
|            | Week 16 | Lower  | -              | -      | -     | -     |
|            |         | Normal | -              | 94     | 8     | 102   |
|            |         | Upper  | -              | 7      | 15    | 22    |
|            |         | Total  | -              | 101    | 23    | 124   |
| Week 20    | Lower   | -      | -              | -      | -     |       |
|            | Normal  | -      | 92             | 7      | 99    |       |
|            | Upper   | -      | 8              | 16     | 24    |       |
|            | Total   | -      | 100            | 23     | 123   |       |
| Week 24    | Lower   | -      | -              | -      | -     |       |
|            | Normal  | -      | 90             | 10     | 100   |       |
|            | Upper   | -      | 9              | 13     | 22    |       |
|            | Total   | -      | 99             | 23     | 122   |       |
| Creatinine | Week 2  | Lower  | 7              | 5      | -     | 12    |
|            |         | Normal | 5              | 96     | 2     | 103   |
|            |         | Upper  | -              | 2      | 10    | 12    |
|            |         | Total  | 12             | 103    | 12    | 127   |
|            | Week 4  | Lower  | 10             | 6      | -     | 16    |
|            |         | Normal | 2              | 95     | 5     | 102   |
|            |         | Upper  | -              | 1      | 7     | 8     |
|            |         | Total  | 12             | 102    | 12    | 126   |
|            | Week 8  | Lower  | 7              | 2      | -     | 9     |
|            |         | Normal | 5              | 94     | 3     | 102   |
|            |         | Upper  | -              | 6      | 9     | 15    |
|            |         | Total  | 12             | 102    | 12    | 126   |
|            | Week 12 | Lower  | 9              | 4      | -     | 13    |
|            |         | Normal | 2              | 95     | 5     | 102   |
|            |         | Upper  | -              | 2      | 7     | 9     |
|            |         | Total  | 11             | 101    | 12    | 124   |
|            | Week 16 | Lower  | 7              | 2      | -     | 9     |
|            |         | Normal | 4              | 98     | 6     | 108   |
|            |         | Upper  | -              | 1      | 6     | 7     |
|            |         | Total  | 11             | 101    | 12    | 124   |
| Week 20    | Lower   | 8      | 5              | -      | 13    |       |
|            | Normal  | 3      | 95             | 6      | 104   |       |
|            | Upper   | -      | -              | 6      | 6     |       |
|            | Total   | 11     | 100            | 12     | 123   |       |
| Week 24    | Lower   | 7      | 6              | -      | 13    |       |
|            | Normal  | 4      | 92             | 5      | 101   |       |
|            | Upper   | -      | 1              | 7      | 8     |       |
|            | Total   | 11     | 99             | 12     | 122   |       |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表 (OCT-001 試験) (7/12)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| BUN     | Week 2  | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | 2              | 116    | 3     | 121   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 3              | 120    | 4     | 127   |
|         | Week 4  | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | 2              | 119    | 4     | 125   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 3              | 119    | 4     | 126   |
|         | Week 8  | Lower  | 2              | 2      | -     | 4     |
|         |         | Normal | 1              | 114    | 3     | 118   |
|         |         | Upper  | -              | 3      | 1     | 4     |
|         |         | Total  | 3              | 119    | 4     | 126   |
|         | Week 12 | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | 2              | 115    | 4     | 121   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 3              | 117    | 4     | 124   |
|         | Week 16 | Lower  | -              | 4      | -     | 4     |
|         |         | Normal | 3              | 112    | 3     | 118   |
|         |         | Upper  | -              | 1      | 1     | 2     |
|         |         | Total  | 3              | 117    | 4     | 124   |
|         | Week 20 | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | 2              | 113    | 3     | 118   |
|         |         | Upper  | -              | 1      | 1     | 2     |
|         |         | Total  | 3              | 116    | 4     | 123   |
| Week 24 | Lower   | -      | 3              | -      | 3     |       |
|         | Normal  | 3      | 111            | 4      | 118   |       |
|         | Upper   | -      | 1              | -      | 1     |       |
|         | Total   | 3      | 115            | 4      | 122   |       |
| Na      | Week 2  | Lower  | 1              | 3      | -     | 4     |
|         |         | Normal | 4              | 119    | -     | 123   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 5              | 122    | -     | 127   |
|         | Week 4  | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | 4              | 119    | -     | 123   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 5              | 121    | -     | 126   |
|         | Week 8  | Lower  | 1              | 8      | -     | 9     |
|         |         | Normal | 4              | 113    | -     | 117   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 5              | 121    | -     | 126   |
|         | Week 12 | Lower  | 1              | 6      | -     | 7     |
|         |         | Normal | 4              | 113    | -     | 117   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 5              | 119    | -     | 124   |
|         | Week 16 | Lower  | -              | 4      | -     | 4     |
|         |         | Normal | 5              | 115    | -     | 120   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 5              | 119    | -     | 124   |
|         | Week 20 | Lower  | 2              | 5      | -     | 7     |
|         |         | Normal | 3              | 113    | -     | 116   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 5              | 118    | -     | 123   |
| Week 24 | Lower   | 2      | 3              | -      | 5     |       |
|         | Normal  | 3      | 114            | -      | 117   |       |
|         | Upper   | -      | -              | -      | -     |       |
|         | Total   | 5      | 117            | -      | 122   |       |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表 (OCT-001 試験) (8/12)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| K       | Week 2  | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | -              | 122    | 1     | 123   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 1              | 125    | 1     | 127   |
|         | Week 4  | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | -              | 122    | -     | 122   |
|         |         | Upper  | -              | 1      | 1     | 2     |
|         |         | Total  | 1              | 124    | 1     | 126   |
|         | Week 8  | Lower  | -              | 2      | -     | 2     |
|         |         | Normal | 1              | 122    | 1     | 124   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 1              | 124    | 1     | 126   |
|         | Week 12 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 1              | 120    | 1     | 122   |
|         |         | Upper  | -              | 2      | -     | 2     |
|         |         | Total  | 1              | 122    | 1     | 124   |
|         | Week 16 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 1              | 118    | 1     | 120   |
|         |         | Upper  | -              | 4      | -     | 4     |
|         |         | Total  | 1              | 122    | 1     | 124   |
| Week 20 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | 1      | 119            | -      | 120   |       |
|         | Upper   | -      | 2              | 1      | 3     |       |
|         | Total   | 1      | 121            | 1      | 123   |       |
| Week 24 | Lower   | 1      | -              | -      | 1     |       |
|         | Normal  | -      | 119            | -      | 119   |       |
|         | Upper   | -      | 1              | 1      | 2     |       |
|         | Total   | 1      | 120            | 1      | 122   |       |
| CI      | Week 2  | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 2              | 124    | -     | 126   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 2              | 125    | -     | 127   |
|         | Week 4  | Lower  | -              | -      | -     | -     |
|         |         | Normal | 2              | 124    | -     | 126   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 2              | 124    | -     | 126   |
|         | Week 8  | Lower  | -              | -      | -     | -     |
|         |         | Normal | 2              | 124    | -     | 126   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 2              | 124    | -     | 126   |
|         | Week 12 | Lower  | -              | 2      | -     | 2     |
|         |         | Normal | 2              | 120    | -     | 122   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 2              | 122    | -     | 124   |
|         | Week 16 | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | 1              | 121    | -     | 122   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 2              | 122    | -     | 124   |
| Week 20 | Lower   | -      | 1              | -      | 1     |       |
|         | Normal  | 2      | 120            | -      | 122   |       |
|         | Upper   | -      | -              | -      | -     |       |
|         | Total   | 2      | 121            | -      | 123   |       |
| Week 24 | Lower   | 1      | 2              | -      | 3     |       |
|         | Normal  | 1      | 118            | -      | 119   |       |
|         | Upper   | -      | -              | -      | -     |       |
|         | Total   | 2      | 120            | -      | 122   |       |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表 (OCT-001 試験) (9/12)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| Ca      | Week 2  | Lower  | -              | -      | -     | -     |
|         |         | Normal | 1              | 126    | -     | 127   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 1              | 126    | -     | 127   |
|         | Week 4  | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | -              | 125    | -     | 125   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 1              | 125    | -     | 126   |
|         | Week 8  | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 1              | 124    | -     | 125   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 1              | 125    | -     | 126   |
|         | Week 12 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 1              | 122    | -     | 123   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 1              | 123    | -     | 124   |
|         | Week 16 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 1              | 122    | -     | 123   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 1              | 123    | -     | 124   |
|         | Week 20 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 1              | 121    | -     | 122   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 1              | 122    | -     | 123   |
| Week 24 | Lower   | -      | 1              | -      | 1     |       |
|         | Normal  | 1      | 118            | -      | 119   |       |
|         | Upper   | -      | 2              | -      | 2     |       |
|         | Total   | 1      | 121            | -      | 122   |       |
| P       | Week 2  | Lower  | -              | 2      | -     | 2     |
|         |         | Normal | 3              | 121    | 1     | 125   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 3              | 123    | 1     | 127   |
|         | Week 4  | Lower  | 1              | 4      | -     | 5     |
|         |         | Normal | 2              | 116    | 1     | 119   |
|         |         | Upper  | -              | 2      | -     | 2     |
|         |         | Total  | 3              | 122    | 1     | 126   |
|         | Week 8  | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 3              | 120    | -     | 123   |
|         |         | Upper  | -              | 1      | 1     | 2     |
|         |         | Total  | 3              | 122    | 1     | 126   |
|         | Week 12 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 3              | 119    | -     | 122   |
|         |         | Upper  | -              | -      | 1     | 1     |
|         |         | Total  | 3              | 120    | 1     | 124   |
|         | Week 16 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 3              | 119    | 1     | 123   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 3              | 120    | 1     | 124   |
|         | Week 20 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 3              | 118    | 1     | 122   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 3              | 119    | 1     | 123   |
| Week 24 | Lower   | -      | 1              | -      | 1     |       |
|         | Normal  | 3      | 117            | 1      | 121   |       |
|         | Upper   | -      | -              | -      | -     |       |
|         | Total   | 3      | 118            | 1      | 122   |       |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表(OCT-001 試験) (10/12)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| Vitamin A | Week 2  | Lower  | 10             | 10     | -     | 20    |
|           |         | Normal | 4              | 100    | 1     | 105   |
|           |         | Upper  | -              | -      | 2     | 2     |
|           |         | Total  | 14             | 110    | 3     | 127   |
|           | Week 4  | Lower  | 9              | 3      | -     | 12    |
|           |         | Normal | 5              | 106    | -     | 111   |
|           |         | Upper  | -              | -      | 3     | 3     |
|           |         | Total  | 14             | 109    | 3     | 126   |
|           | Week 8  | Lower  | 11             | 5      | -     | 16    |
|           |         | Normal | 3              | 103    | -     | 106   |
|           |         | Upper  | -              | 1      | 3     | 4     |
|           |         | Total  | 14             | 109    | 3     | 126   |
|           | Week 12 | Lower  | 10             | 6      | -     | 16    |
|           |         | Normal | 4              | 101    | -     | 105   |
|           |         | Upper  | -              | -      | 3     | 3     |
|           |         | Total  | 14             | 107    | 3     | 124   |
|           | Week 16 | Lower  | 11             | 4      | -     | 15    |
|           |         | Normal | 3              | 101    | -     | 104   |
|           |         | Upper  | -              | 2      | 3     | 5     |
|           |         | Total  | 14             | 107    | 3     | 124   |
|           | Week 20 | Lower  | 10             | 3      | -     | 13    |
|           |         | Normal | 4              | 103    | 2     | 109   |
|           |         | Upper  | -              | -      | 1     | 1     |
|           |         | Total  | 14             | 106    | 3     | 123   |
| Week 24   | Lower   | 11     | 5              | -      | 16    |       |
|           | Normal  | 3      | 98             | 1      | 102   |       |
|           | Upper   | -      | 2              | 2      | 4     |       |
|           | Total   | 14     | 105            | 3      | 122   |       |
| Vitamin D | Week 2  | Lower  | -              | 3      | -     | 3     |
|           |         | Normal | 2              | 71     | 19    | 92    |
|           |         | Upper  | -              | 12     | 20    | 32    |
|           |         | Total  | 2              | 86     | 39    | 127   |
|           | Week 4  | Lower  | -              | 1      | -     | 1     |
|           |         | Normal | 2              | 72     | 23    | 97    |
|           |         | Upper  | -              | 12     | 16    | 28    |
|           |         | Total  | 2              | 85     | 39    | 126   |
|           | Week 8  | Lower  | -              | 2      | -     | 2     |
|           |         | Normal | 2              | 72     | 17    | 91    |
|           |         | Upper  | -              | 11     | 22    | 33    |
|           |         | Total  | 2              | 85     | 39    | 126   |
|           | Week 12 | Lower  | -              | 2      | -     | 2     |
|           |         | Normal | 2              | 68     | 15    | 85    |
|           |         | Upper  | -              | 14     | 23    | 37    |
|           |         | Total  | 2              | 84     | 38    | 124   |
|           | Week 16 | Lower  | -              | -      | -     | -     |
|           |         | Normal | 2              | 69     | 19    | 90    |
|           |         | Upper  | -              | 15     | 19    | 34    |
|           |         | Total  | 2              | 84     | 38    | 124   |
|           | Week 20 | Lower  | -              | 2      | -     | 2     |
|           |         | Normal | 2              | 63     | 14    | 79    |
|           |         | Upper  | -              | 19     | 23    | 42    |
|           |         | Total  | 2              | 84     | 37    | 123   |
| Week 24   | Lower   | 1      | 1              | -      | 2     |       |
|           | Normal  | 1      | 68             | 12     | 81    |       |
|           | Upper   | -      | 15             | 24     | 39    |       |
|           | Total   | 2      | 84             | 36     | 122   |       |



表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表 (OCT-001 試験) (11/12)

|            |         |        | ATL-962 120 mg |        |       |       |
|------------|---------|--------|----------------|--------|-------|-------|
|            |         |        | Week 0         |        |       |       |
|            |         |        | Lower          | Normal | Upper | Total |
| Vitamin E  | Week 2  | Lower  | -              | 2      | -     | 2     |
|            |         | Normal | -              | 38     | 32    | 70    |
|            |         | Upper  | -              | 7      | 48    | 55    |
|            |         | Total  | -              | 47     | 80    | 127   |
|            | Week 4  | Lower  | -              | 1      | -     | 1     |
|            |         | Normal | -              | 42     | 30    | 72    |
|            |         | Upper  | -              | 4      | 49    | 53    |
|            |         | Total  | -              | 47     | 79    | 126   |
|            | Week 8  | Lower  | -              | 2      | -     | 2     |
|            |         | Normal | -              | 41     | 37    | 78    |
|            |         | Upper  | -              | 4      | 42    | 46    |
|            |         | Total  | -              | 47     | 79    | 126   |
|            | Week 12 | Lower  | -              | 1      | -     | 1     |
|            |         | Normal | -              | 39     | 48    | 87    |
|            |         | Upper  | -              | 6      | 30    | 36    |
|            |         | Total  | -              | 46     | 78    | 124   |
|            | Week 16 | Lower  | -              | 3      | -     | 3     |
|            |         | Normal | -              | 42     | 58    | 100   |
|            |         | Upper  | -              | 1      | 20    | 21    |
|            |         | Total  | -              | 46     | 78    | 124   |
|            | Week 20 | Lower  | -              | 5      | 1     | 6     |
|            |         | Normal | -              | 38     | 52    | 90    |
|            |         | Upper  | -              | 3      | 24    | 27    |
|            |         | Total  | -              | 46     | 77    | 123   |
| Week 24    | Lower   | -      | 4              | 1      | 5     |       |
|            | Normal  | -      | 39             | 49     | 88    |       |
|            | Upper   | -      | 3              | 26     | 29    |       |
|            | Total   | -      | 46             | 76     | 122   |       |
| β-Carotene | Week 2  | Lower  | 29             | 6      | -     | 35    |
|            |         | Normal | 4              | 84     | 1     | 89    |
|            |         | Upper  | -              | 1      | 2     | 3     |
|            |         | Total  | 33             | 91     | 3     | 127   |
|            | Week 4  | Lower  | 24             | 12     | -     | 36    |
|            |         | Normal | 9              | 78     | 1     | 88    |
|            |         | Upper  | -              | -      | 2     | 2     |
|            |         | Total  | 33             | 90     | 3     | 126   |
|            | Week 8  | Lower  | 28             | 11     | -     | 39    |
|            |         | Normal | 5              | 79     | 1     | 85    |
|            |         | Upper  | -              | -      | 2     | 2     |
|            |         | Total  | 33             | 90     | 3     | 126   |
|            | Week 12 | Lower  | 28             | 18     | -     | 46    |
|            |         | Normal | 4              | 71     | 1     | 76    |
|            |         | Upper  | -              | -      | 2     | 2     |
|            |         | Total  | 32             | 89     | 3     | 124   |
|            | Week 16 | Lower  | 28             | 14     | -     | 42    |
|            |         | Normal | 4              | 75     | 2     | 81    |
|            |         | Upper  | -              | -      | 1     | 1     |
|            |         | Total  | 32             | 89     | 3     | 124   |
|            | Week 20 | Lower  | 26             | 13     | -     | 39    |
|            |         | Normal | 6              | 75     | 2     | 83    |
|            |         | Upper  | -              | -      | 1     | 1     |
|            |         | Total  | 32             | 88     | 3     | 123   |
| Week 24    | Lower   | 27     | 15             | -      | 42    |       |
|            | Normal  | 5      | 72             | 3      | 80    |       |
|            | Upper   | -      | -              | -      | -     |       |
|            | Total   | 32     | 87             | 3      | 122   |       |

表 2.7.4.7-付録-27 血液生化学検査の評価時点別正異判定クロス表(OCT-001 試験) (12/12)

|                |         |        | ATL-962 120 mg |        |       |       |
|----------------|---------|--------|----------------|--------|-------|-------|
|                |         |        | Week 0         |        |       |       |
|                |         |        | Lower          | Normal | Upper | Total |
| m-AST          | Week 2  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 118    | 3     | 121   |
|                |         | Upper  | -              | 4      | 2     | 6     |
|                |         | Total  | -              | 122    | 5     | 127   |
|                | Week 4  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 114    | 2     | 116   |
|                |         | Upper  | -              | 7      | 3     | 10    |
|                |         | Total  | -              | 121    | 5     | 126   |
|                | Week 8  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 114    | 3     | 117   |
|                |         | Upper  | -              | 7      | 2     | 9     |
|                |         | Total  | -              | 121    | 5     | 126   |
|                | Week 12 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 111    | 3     | 114   |
|                |         | Upper  | -              | 8      | 2     | 10    |
|                |         | Total  | -              | 119    | 5     | 124   |
|                | Week 16 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 116    | 5     | 121   |
|                |         | Upper  | -              | 3      | -     | 3     |
|                |         | Total  | -              | 119    | 5     | 124   |
|                | Week 20 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 114    | 3     | 117   |
|                |         | Upper  | -              | 4      | 2     | 6     |
|                |         | Total  | -              | 118    | 5     | 123   |
| Week 24        | Lower   | -      | -              | -      | -     |       |
|                | Normal  | -      | 113            | 3      | 116   |       |
|                | Upper   | -      | 4              | 2      | 6     |       |
|                | Total   | -      | 117            | 5      | 122   |       |
| CK18 (Segment) | Week 2  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 88     | 13    | 101   |
|                |         | Upper  | -              | 10     | 16    | 26    |
|                |         | Total  | -              | 98     | 29    | 127   |
|                | Week 4  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 91     | 13    | 104   |
|                |         | Upper  | -              | 6      | 16    | 22    |
|                |         | Total  | -              | 97     | 29    | 126   |
|                | Week 8  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 86     | 14    | 100   |
|                |         | Upper  | -              | 11     | 15    | 26    |
|                |         | Total  | -              | 97     | 29    | 126   |
|                | Week 12 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 88     | 16    | 104   |
|                |         | Upper  | -              | 7      | 13    | 20    |
|                |         | Total  | -              | 95     | 29    | 124   |
|                | Week 16 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 87     | 16    | 103   |
|                |         | Upper  | -              | 8      | 13    | 21    |
|                |         | Total  | -              | 95     | 29    | 124   |
|                | Week 20 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 89     | 18    | 107   |
|                |         | Upper  | -              | 5      | 11    | 16    |
|                |         | Total  | -              | 94     | 29    | 123   |
| Week 24        | Lower   | -      | -              | -      | -     |       |
|                | Normal  | -      | 88             | 23     | 111   |       |
|                | Upper   | -      | 5              | 6      | 11    |       |
|                | Total   | -      | 93             | 29     | 122   |       |

表 2.7.4.7-付録-28 尿検査の評価時点別の変化量 (OCT-001 試験)

| Specific Gravity |                | Summary Statistics |         |         |        |         |         |        |       |
|------------------|----------------|--------------------|---------|---------|--------|---------|---------|--------|-------|
| Visit            | Treatment      | N                  | Mean    | SD      | Min    | Q1      | Median  | Q3     | Max   |
| Week 2           | ATL-962 120 mg | 127                | -0.0006 | 0.00554 | -0.015 | -0.0040 | -0.0010 | 0.0020 | 0.017 |
| Week 4           | ATL-962 120 mg | 126                | -0.0006 | 0.00636 | -0.021 | -0.0050 | -0.0005 | 0.0030 | 0.018 |
| Week 8           | ATL-962 120 mg | 126                | -0.0009 | 0.00613 | -0.016 | -0.0050 | 0.0000  | 0.0030 | 0.017 |
| Week 12          | ATL-962 120 mg | 124                | -0.0014 | 0.00636 | -0.015 | -0.0050 | -0.0020 | 0.0010 | 0.021 |
| Week 16          | ATL-962 120 mg | 124                | -0.0012 | 0.00594 | -0.020 | -0.0050 | -0.0020 | 0.0020 | 0.016 |
| Week 20          | ATL-962 120 mg | 123                | -0.0011 | 0.00526 | -0.017 | -0.0040 | -0.0010 | 0.0020 | 0.014 |
| Week 24          | ATL-962 120 mg | 122                | -0.0016 | 0.00616 | -0.017 | -0.0050 | -0.0020 | 0.0010 | 0.021 |

表 2.7.4.7-付録-29 尿検査の評価時点別クロス表 (OCT-001 試験) (1/4)

|                  |         | ATL-962 120 mg |        |       |       |     |
|------------------|---------|----------------|--------|-------|-------|-----|
|                  |         | Week 0         |        |       |       |     |
|                  |         | Lower          | Normal | Upper | Total |     |
| Specific Gravity | Week 2  | Lower          | -      | -     | -     | -   |
|                  |         | Normal         | -      | 125   | 1     | 126 |
|                  |         | Upper          | -      | 1     | -     | 1   |
|                  |         | Total          | -      | 126   | 1     | 127 |
|                  | Week 4  | Lower          | -      | -     | -     | -   |
|                  |         | Normal         | -      | 125   | 1     | 126 |
|                  |         | Upper          | -      | -     | -     | -   |
|                  |         | Total          | -      | 125   | 1     | 126 |
|                  | Week 8  | Lower          | -      | -     | -     | -   |
|                  |         | Normal         | -      | 124   | 1     | 125 |
|                  |         | Upper          | -      | 1     | -     | 1   |
|                  |         | Total          | -      | 125   | 1     | 126 |
|                  | Week 12 | Lower          | -      | -     | -     | -   |
|                  |         | Normal         | -      | 123   | 1     | 124 |
|                  |         | Upper          | -      | -     | -     | -   |
|                  |         | Total          | -      | 123   | 1     | 124 |
|                  | Week 16 | Lower          | -      | -     | -     | -   |
|                  |         | Normal         | -      | 123   | 1     | 124 |
|                  |         | Upper          | -      | -     | -     | -   |
|                  |         | Total          | -      | 123   | 1     | 124 |
|                  | Week 20 | Lower          | -      | -     | -     | -   |
|                  |         | Normal         | -      | 122   | -     | 122 |
|                  |         | Upper          | -      | -     | 1     | 1   |
|                  |         | Total          | -      | 122   | 1     | 123 |
| Week 24          | Lower   | -              | -      | -     | -     |     |
|                  | Normal  | -              | 121    | 1     | 122   |     |
|                  | Upper   | -              | -      | -     | -     |     |
|                  | Total   | -              | 121    | 1     | 122   |     |

表 2.7.4.7-付録-29 尿検査の評価時点別クロス表 (OCT-001 試験) (2/4)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| pH      | Week 2  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 127    | -     | 127   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 127    | -     | 127   |
|         | Week 4  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 125    | -     | 125   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | -              | 126    | -     | 126   |
|         | Week 8  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 126    | -     | 126   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 126    | -     | 126   |
|         | Week 12 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 124    | -     | 124   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 124    | -     | 124   |
|         | Week 16 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 124    | -     | 124   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 124    | -     | 124   |
|         | Week 20 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 123    | -     | 123   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 123    | -     | 123   |
| Week 24 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 122            | -      | 122   |       |
|         | Upper   | -      | -              | -      | -     |       |
|         | Total   | -      | 122            | -      | 122   |       |

表 2.7.4.7-付録-29 尿検査の評価時点別クロス表 (OCT-001 試験) (3/4)

Glucose

| Frequency |                |         |          | Week 0 |          |       |
|-----------|----------------|---------|----------|--------|----------|-------|
|           |                |         |          | Normal | Abnormal | Total |
| Treatment | ATL-962 120 mg | Week 2  | Normal   | 75     | 13       | 88    |
|           |                |         | Abnormal | 7      | 32       | 39    |
|           |                |         | Total    | 82     | 45       | 127   |
|           |                | Week 4  | Normal   | 75     | 18       | 93    |
|           |                |         | Abnormal | 7      | 26       | 33    |
|           |                |         | Total    | 82     | 44       | 126   |
|           |                | Week 8  | Normal   | 74     | 20       | 94    |
|           |                |         | Abnormal | 8      | 24       | 32    |
|           |                |         | Total    | 82     | 44       | 126   |
|           |                | Week 12 | Normal   | 73     | 15       | 88    |
|           |                |         | Abnormal | 7      | 29       | 36    |
|           |                |         | Total    | 80     | 44       | 124   |
|           |                | Week 16 | Normal   | 72     | 17       | 89    |
|           |                |         | Abnormal | 8      | 27       | 35    |
|           |                |         | Total    | 80     | 44       | 124   |
|           |                | Week 20 | Normal   | 70     | 17       | 87    |
|           |                |         | Abnormal | 10     | 26       | 36    |
|           |                |         | Total    | 80     | 43       | 123   |
|           |                | Week 24 | Normal   | 66     | 16       | 82    |
|           |                |         | Abnormal | 14     | 26       | 40    |
|           |                |         | Total    | 80     | 42       | 122   |

Protein

| Frequency |                |         |          | Week 0 |          |       |
|-----------|----------------|---------|----------|--------|----------|-------|
|           |                |         |          | Normal | Abnormal | Total |
| Treatment | ATL-962 120 mg | Week 2  | Normal   | 71     | 9        | 80    |
|           |                |         | Abnormal | 8      | 39       | 47    |
|           |                |         | Total    | 79     | 48       | 127   |
|           |                | Week 4  | Normal   | 69     | 10       | 79    |
|           |                |         | Abnormal | 10     | 37       | 47    |
|           |                |         | Total    | 79     | 47       | 126   |
|           |                | Week 8  | Normal   | 67     | 9        | 76    |
|           |                |         | Abnormal | 12     | 38       | 50    |
|           |                |         | Total    | 79     | 47       | 126   |
|           |                | Week 12 | Normal   | 63     | 13       | 76    |
|           |                |         | Abnormal | 15     | 33       | 48    |
|           |                |         | Total    | 78     | 46       | 124   |
|           |                | Week 16 | Normal   | 69     | 14       | 83    |
|           |                |         | Abnormal | 9      | 32       | 41    |
|           |                |         | Total    | 78     | 46       | 124   |
|           |                | Week 20 | Normal   | 67     | 10       | 77    |
|           |                |         | Abnormal | 11     | 35       | 46    |
|           |                |         | Total    | 78     | 45       | 123   |
|           |                | Week 24 | Normal   | 69     | 7        | 76    |
|           |                |         | Abnormal | 9      | 37       | 46    |
|           |                |         | Total    | 78     | 44       | 122   |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-29 尿検査の評価時点別クロス表 (OCT-001 試験) (4/4)

Ketones

| Frequency |                |         |          | Week 0 |          |       |
|-----------|----------------|---------|----------|--------|----------|-------|
|           |                |         |          | Normal | Abnormal | Total |
| Treatment | ATL-962 120 mg | Week 2  | Normal   | 104    | 9        | 113   |
|           |                |         | Abnormal | 9      | 5        | 14    |
|           |                |         | Total    | 113    | 14       | 127   |
|           |                | Week 4  | Normal   | 106    | 8        | 114   |
|           |                |         | Abnormal | 6      | 6        | 12    |
|           |                |         | Total    | 112    | 14       | 126   |
|           |                | Week 8  | Normal   | 105    | 9        | 114   |
|           |                |         | Abnormal | 7      | 5        | 12    |
|           |                |         | Total    | 112    | 14       | 126   |
|           |                | Week 12 | Normal   | 104    | 9        | 113   |
|           |                |         | Abnormal | 6      | 5        | 11    |
|           |                |         | Total    | 110    | 14       | 124   |
|           |                | Week 16 | Normal   | 106    | 9        | 115   |
|           |                |         | Abnormal | 4      | 5        | 9     |
|           |                |         | Total    | 110    | 14       | 124   |
|           |                | Week 20 | Normal   | 102    | 8        | 110   |
|           |                |         | Abnormal | 8      | 5        | 13    |
|           |                |         | Total    | 110    | 13       | 123   |
|           |                | Week 24 | Normal   | 103    | 9        | 112   |
|           |                |         | Abnormal | 6      | 4        | 10    |
|           |                |         | Total    | 109    | 13       | 122   |

Occult Blood

| Frequency |                |         |          | Week 0 |          |       |
|-----------|----------------|---------|----------|--------|----------|-------|
|           |                |         |          | Normal | Abnormal | Total |
| Treatment | ATL-962 120 mg | Week 2  | Normal   | 111    | 4        | 115   |
|           |                |         | Abnormal | 4      | 8        | 12    |
|           |                |         | Total    | 115    | 12       | 127   |
|           |                | Week 4  | Normal   | 111    | 3        | 114   |
|           |                |         | Abnormal | 3      | 9        | 12    |
|           |                |         | Total    | 114    | 12       | 126   |
|           |                | Week 8  | Normal   | 111    | 4        | 115   |
|           |                |         | Abnormal | 3      | 8        | 11    |
|           |                |         | Total    | 114    | 12       | 126   |
|           |                | Week 12 | Normal   | 110    | 3        | 113   |
|           |                |         | Abnormal | 3      | 8        | 11    |
|           |                |         | Total    | 113    | 11       | 124   |
|           |                | Week 16 | Normal   | 110    | 6        | 116   |
|           |                |         | Abnormal | 3      | 5        | 8     |
|           |                |         | Total    | 113    | 11       | 124   |
|           |                | Week 20 | Normal   | 105    | 4        | 109   |
|           |                |         | Abnormal | 7      | 7        | 14    |
|           |                |         | Total    | 112    | 11       | 123   |
|           |                | Week 24 | Normal   | 102    | 3        | 105   |
|           |                |         | Abnormal | 9      | 8        | 17    |
|           |                |         | Total    | 111    | 11       | 122   |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-30 血液学的検査の評価時点別の変化量 (OCT-002 試験) (1/4)

| Hemoglobin (g/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2            | ATL-962 120 mg | 503                | -0.08 | 0.453 | -2.2 | -0.40 | -0.10  | 0.20 | 1.3 |
| Week 4            | ATL-962 120 mg | 492                | -0.09 | 0.462 | -1.8 | -0.40 | -0.10  | 0.20 | 1.4 |
| Week 8            | ATL-962 120 mg | 488                | 0.02  | 0.548 | -3.0 | -0.30 | 0.00   | 0.35 | 1.5 |
| Week 12           | ATL-962 120 mg | 478                | 0.07  | 0.569 | -2.0 | -0.30 | 0.10   | 0.40 | 2.1 |
| Week 16           | ATL-962 120 mg | 471                | 0.21  | 0.579 | -1.8 | -0.20 | 0.20   | 0.60 | 2.2 |
| Week 20           | ATL-962 120 mg | 468                | 0.22  | 0.583 | -1.6 | -0.20 | 0.20   | 0.50 | 2.2 |
| Week 24           | ATL-962 120 mg | 461                | 0.14  | 0.627 | -1.4 | -0.30 | 0.10   | 0.50 | 2.3 |
| Week 28           | ATL-962 120 mg | 452                | 0.11  | 0.629 | -2.4 | -0.30 | 0.10   | 0.50 | 2.0 |
| Week 32           | ATL-962 120 mg | 455                | 0.11  | 0.653 | -2.0 | -0.30 | 0.10   | 0.50 | 2.0 |
| Week 36           | ATL-962 120 mg | 451                | -0.02 | 0.701 | -2.6 | -0.40 | 0.00   | 0.40 | 2.1 |
| Week 40           | ATL-962 120 mg | 444                | -0.05 | 0.636 | -2.3 | -0.40 | -0.10  | 0.30 | 1.9 |
| Week 44           | ATL-962 120 mg | 441                | -0.02 | 0.620 | -2.4 | -0.50 | 0.00   | 0.40 | 1.9 |
| Week 48           | ATL-962 120 mg | 438                | 0.00  | 0.618 | -2.2 | -0.40 | 0.00   | 0.40 | 2.4 |
| Week 52           | ATL-962 120 mg | 438                | 0.00  | 0.688 | -3.5 | -0.40 | 0.00   | 0.40 | 2.5 |

| Hematocrit (%) |                | Summary Statistics |       |       |      |       |        |      |     |
|----------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit          | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2         | ATL-962 120 mg | 504                | -0.26 | 1.424 | -6.8 | -1.20 | -0.20  | 0.70 | 4.0 |
| Week 4         | ATL-962 120 mg | 493                | -0.36 | 1.476 | -7.3 | -1.30 | -0.30  | 0.60 | 4.4 |
| Week 8         | ATL-962 120 mg | 489                | -0.30 | 1.654 | -9.3 | -1.30 | -0.30  | 0.80 | 5.0 |
| Week 12        | ATL-962 120 mg | 479                | -0.31 | 1.698 | -5.7 | -1.40 | -0.30  | 0.70 | 5.3 |
| Week 16        | ATL-962 120 mg | 472                | -0.03 | 1.738 | -6.4 | -1.15 | -0.10  | 1.10 | 6.7 |
| Week 20        | ATL-962 120 mg | 469                | 0.07  | 1.715 | -5.5 | -1.00 | -0.10  | 1.10 | 7.4 |
| Week 24        | ATL-962 120 mg | 462                | -0.09 | 1.836 | -4.8 | -1.30 | 0.00   | 1.10 | 6.3 |
| Week 28        | ATL-962 120 mg | 453                | -0.15 | 1.760 | -7.0 | -1.20 | -0.20  | 1.00 | 4.9 |
| Week 32        | ATL-962 120 mg | 456                | 0.02  | 1.870 | -6.2 | -1.20 | 0.10   | 1.20 | 5.8 |
| Week 36        | ATL-962 120 mg | 452                | -0.20 | 1.933 | -6.8 | -1.40 | -0.10  | 0.85 | 6.5 |
| Week 40        | ATL-962 120 mg | 445                | -0.24 | 1.793 | -6.1 | -1.30 | -0.20  | 0.90 | 4.9 |
| Week 44        | ATL-962 120 mg | 442                | -0.17 | 1.776 | -6.1 | -1.30 | -0.20  | 0.90 | 6.5 |
| Week 48        | ATL-962 120 mg | 439                | -0.11 | 1.850 | -7.2 | -1.20 | -0.10  | 1.00 | 8.5 |
| Week 52        | ATL-962 120 mg | 439                | -0.17 | 2.027 | -9.5 | -1.50 | -0.10  | 1.10 | 6.9 |

| RBC (x10 <sup>4</sup> /μL) |                | Summary Statistics |      |       |     |       |        |      |     |
|----------------------------|----------------|--------------------|------|-------|-----|-------|--------|------|-----|
| Visit                      | Treatment      | N                  | Mean | SD    | Min | Q1    | Median | Q3   | Max |
| Week 2                     | ATL-962 120 mg | 504                | -3.2 | 14.88 | -66 | -14.0 | -3.0   | 6.0  | 42  |
| Week 4                     | ATL-962 120 mg | 493                | -4.1 | 15.42 | -66 | -15.0 | -4.0   | 6.0  | 48  |
| Week 8                     | ATL-962 120 mg | 489                | -3.0 | 17.09 | -89 | -13.0 | -3.0   | 8.0  | 53  |
| Week 12                    | ATL-962 120 mg | 479                | -2.0 | 18.13 | -55 | -14.0 | -2.0   | 9.0  | 61  |
| Week 16                    | ATL-962 120 mg | 472                | 1.9  | 18.01 | -65 | -10.0 | 2.0    | 13.0 | 59  |
| Week 20                    | ATL-962 120 mg | 469                | 3.1  | 18.31 | -59 | -9.0  | 3.0    | 15.0 | 66  |
| Week 24                    | ATL-962 120 mg | 462                | 1.4  | 19.57 | -50 | -10.0 | 2.0    | 14.0 | 62  |
| Week 28                    | ATL-962 120 mg | 453                | 1.0  | 19.07 | -72 | -11.0 | 0.0    | 13.0 | 59  |
| Week 32                    | ATL-962 120 mg | 456                | 2.0  | 20.34 | -60 | -11.0 | 3.0    | 15.0 | 67  |
| Week 36                    | ATL-962 120 mg | 452                | -0.6 | 20.69 | -75 | -14.0 | -1.0   | 13.0 | 73  |
| Week 40                    | ATL-962 120 mg | 445                | -1.1 | 19.41 | -64 | -12.0 | -1.0   | 11.0 | 50  |
| Week 44                    | ATL-962 120 mg | 442                | -1.9 | 19.52 | -74 | -15.0 | -2.0   | 10.0 | 70  |
| Week 48                    | ATL-962 120 mg | 439                | -2.4 | 19.32 | -75 | -14.0 | -3.0   | 9.0  | 85  |
| Week 52                    | ATL-962 120 mg | 439                | -4.0 | 20.41 | -72 | -18.0 | -3.0   | 8.0  | 56  |

表 2.7.4.7-付録-30 血液学的検査の評価時点別の変化量 (OCT-002 試験) (2/4)

| WBC (/ $\mu$ L) |                | Summary Statistics |        |         |       |        |        |       |      |
|-----------------|----------------|--------------------|--------|---------|-------|--------|--------|-------|------|
| Visit           | Treatment      | N                  | Mean   | SD      | Min   | Q1     | Median | Q3    | Max  |
| Week 2          | ATL-962 120 mg | 504                | 175.0  | 982.17  | -3400 | -350.0 | 200.0  | 700.0 | 3800 |
| Week 4          | ATL-962 120 mg | 493                | 145.6  | 1078.34 | -3500 | -400.0 | 100.0  | 700.0 | 5700 |
| Week 8          | ATL-962 120 mg | 489                | 86.5   | 1173.94 | -4400 | -500.0 | 0.0    | 600.0 | 7400 |
| Week 12         | ATL-962 120 mg | 479                | 14.2   | 1071.56 | -4200 | -500.0 | 0.0    | 600.0 | 4000 |
| Week 16         | ATL-962 120 mg | 472                | 75.0   | 1027.11 | -4000 | -500.0 | 0.0    | 700.0 | 5200 |
| Week 20         | ATL-962 120 mg | 469                | 79.7   | 1170.98 | -4700 | -500.0 | 100.0  | 700.0 | 6500 |
| Week 24         | ATL-962 120 mg | 462                | 61.5   | 1212.40 | -4300 | -500.0 | 0.0    | 600.0 | 7600 |
| Week 28         | ATL-962 120 mg | 453                | 45.5   | 1092.88 | -3400 | -600.0 | 0.0    | 700.0 | 4900 |
| Week 32         | ATL-962 120 mg | 456                | -5.7   | 1162.19 | -5400 | -700.0 | 0.0    | 600.0 | 4000 |
| Week 36         | ATL-962 120 mg | 452                | -25.2  | 1133.58 | -4000 | -700.0 | -100.0 | 500.0 | 3900 |
| Week 40         | ATL-962 120 mg | 445                | -41.8  | 1141.74 | -3900 | -600.0 | -100.0 | 600.0 | 5300 |
| Week 44         | ATL-962 120 mg | 442                | -20.6  | 1149.11 | -4400 | -700.0 | 0.0    | 600.0 | 3900 |
| Week 48         | ATL-962 120 mg | 439                | 21.6   | 1249.86 | -3900 | -700.0 | 0.0    | 600.0 | 9600 |
| Week 52         | ATL-962 120 mg | 439                | -120.3 | 1167.92 | -5200 | -700.0 | 0.0    | 500.0 | 5600 |

| Neutrophils (%) |                | Summary Statistics |      |       |       |       |        |      |      |
|-----------------|----------------|--------------------|------|-------|-------|-------|--------|------|------|
| Visit           | Treatment      | N                  | Mean | SD    | Min   | Q1    | Median | Q3   | Max  |
| Week 2          | ATL-962 120 mg | 504                | 0.27 | 5.705 | -21.2 | -3.10 | 0.10   | 3.55 | 29.8 |
| Week 4          | ATL-962 120 mg | 493                | 0.43 | 6.202 | -31.8 | -3.10 | 0.10   | 4.00 | 24.3 |
| Week 8          | ATL-962 120 mg | 489                | 0.71 | 6.362 | -28.5 | -2.30 | 0.60   | 4.00 | 39.1 |
| Week 12         | ATL-962 120 mg | 479                | 1.03 | 6.049 | -21.7 | -2.60 | 1.00   | 4.40 | 26.3 |
| Week 16         | ATL-962 120 mg | 472                | 0.71 | 5.991 | -28.5 | -2.95 | 0.90   | 4.25 | 18.0 |
| Week 20         | ATL-962 120 mg | 469                | 1.03 | 6.481 | -17.8 | -2.80 | 0.90   | 4.60 | 34.3 |
| Week 24         | ATL-962 120 mg | 461                | 1.06 | 6.907 | -27.5 | -2.40 | 1.00   | 4.70 | 43.5 |
| Week 28         | ATL-962 120 mg | 453                | 0.86 | 6.411 | -21.3 | -3.00 | 0.80   | 5.00 | 19.3 |
| Week 32         | ATL-962 120 mg | 456                | 0.49 | 6.208 | -18.8 | -3.60 | 0.40   | 4.30 | 20.9 |
| Week 36         | ATL-962 120 mg | 452                | 0.65 | 6.479 | -19.0 | -3.40 | 0.60   | 4.40 | 23.8 |
| Week 40         | ATL-962 120 mg | 445                | 0.19 | 6.844 | -21.0 | -4.00 | -0.10  | 4.30 | 24.6 |
| Week 44         | ATL-962 120 mg | 442                | 0.07 | 6.391 | -25.6 | -3.30 | 0.15   | 3.70 | 25.2 |
| Week 48         | ATL-962 120 mg | 439                | 0.26 | 6.557 | -22.8 | -3.80 | 0.00   | 4.20 | 27.5 |
| Week 52         | ATL-962 120 mg | 439                | 0.04 | 6.458 | -38.0 | -3.60 | 0.00   | 4.20 | 22.7 |

| Lymphocytes (%) |                | Summary Statistics |       |       |       |       |        |      |      |
|-----------------|----------------|--------------------|-------|-------|-------|-------|--------|------|------|
| Visit           | Treatment      | N                  | Mean  | SD    | Min   | Q1    | Median | Q3   | Max  |
| Week 2          | ATL-962 120 mg | 504                | -0.26 | 5.004 | -25.5 | -2.90 | 0.00   | 2.40 | 19.8 |
| Week 4          | ATL-962 120 mg | 493                | -0.56 | 5.374 | -20.2 | -3.80 | -0.20  | 2.40 | 26.9 |
| Week 8          | ATL-962 120 mg | 489                | -0.93 | 5.617 | -34.5 | -4.00 | -0.90  | 2.00 | 24.0 |
| Week 12         | ATL-962 120 mg | 479                | -1.14 | 5.347 | -24.9 | -4.00 | -1.20  | 2.10 | 17.1 |
| Week 16         | ATL-962 120 mg | 472                | -0.95 | 5.188 | -17.0 | -3.95 | -0.85  | 2.35 | 17.8 |
| Week 20         | ATL-962 120 mg | 469                | -1.13 | 5.551 | -27.0 | -4.20 | -1.10  | 2.30 | 16.5 |
| Week 24         | ATL-962 120 mg | 461                | -1.28 | 5.857 | -37.0 | -4.50 | -1.10  | 2.00 | 19.3 |
| Week 28         | ATL-962 120 mg | 453                | -1.07 | 5.621 | -18.9 | -4.50 | -0.80  | 2.50 | 17.0 |
| Week 32         | ATL-962 120 mg | 456                | -0.59 | 5.434 | -21.2 | -3.90 | -0.30  | 3.00 | 17.7 |
| Week 36         | ATL-962 120 mg | 452                | -0.76 | 5.746 | -19.6 | -3.85 | -0.50  | 2.85 | 15.7 |
| Week 40         | ATL-962 120 mg | 445                | -0.31 | 5.917 | -21.4 | -3.50 | -0.30  | 3.70 | 19.6 |
| Week 44         | ATL-962 120 mg | 442                | -0.16 | 5.350 | -22.0 | -3.20 | 0.00   | 2.90 | 16.2 |
| Week 48         | ATL-962 120 mg | 439                | -0.13 | 5.651 | -25.5 | -3.60 | 0.20   | 3.40 | 18.8 |
| Week 52         | ATL-962 120 mg | 439                | -0.06 | 5.511 | -21.0 | -3.40 | -0.20  | 3.20 | 29.6 |



表 2.7.4.7-付録-30 血液学的検査の評価時点別の変化量（OCT-002 試験）（3/4）

| Monocytes (%) |                | Summary Statistics |       |       |      |       |        |      |      |
|---------------|----------------|--------------------|-------|-------|------|-------|--------|------|------|
| Visit         | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max  |
| Week 2        | ATL-962 120 mg | 504                | -0.11 | 1.471 | -4.6 | -0.95 | -0.10  | 0.80 | 5.4  |
| Week 4        | ATL-962 120 mg | 493                | -0.04 | 1.488 | -6.4 | -0.90 | 0.00   | 0.90 | 5.7  |
| Week 8        | ATL-962 120 mg | 489                | 0.06  | 1.519 | -5.0 | -0.80 | 0.00   | 1.00 | 6.0  |
| Week 12       | ATL-962 120 mg | 479                | 0.04  | 1.548 | -6.0 | -0.80 | 0.00   | 1.00 | 8.3  |
| Week 16       | ATL-962 120 mg | 472                | 0.01  | 1.503 | -5.0 | -0.85 | 0.00   | 0.85 | 5.8  |
| Week 20       | ATL-962 120 mg | 469                | 0.04  | 1.561 | -5.5 | -0.80 | 0.00   | 1.00 | 5.4  |
| Week 24       | ATL-962 120 mg | 461                | 0.08  | 1.593 | -5.7 | -0.80 | 0.10   | 1.00 | 6.2  |
| Week 28       | ATL-962 120 mg | 453                | 0.07  | 1.566 | -6.5 | -0.90 | 0.00   | 0.90 | 6.0  |
| Week 32       | ATL-962 120 mg | 456                | -0.03 | 1.553 | -5.4 | -1.00 | 0.00   | 0.95 | 5.3  |
| Week 36       | ATL-962 120 mg | 452                | 0.02  | 1.631 | -5.0 | -1.00 | 0.00   | 1.00 | 11.0 |
| Week 40       | ATL-962 120 mg | 445                | 0.02  | 1.645 | -6.0 | -1.00 | 0.00   | 1.00 | 6.3  |
| Week 44       | ATL-962 120 mg | 442                | -0.02 | 1.573 | -6.6 | -1.00 | 0.00   | 0.90 | 6.9  |
| Week 48       | ATL-962 120 mg | 439                | -0.15 | 1.503 | -6.0 | -1.00 | -0.10  | 0.80 | 4.0  |
| Week 52       | ATL-962 120 mg | 439                | -0.03 | 1.564 | -4.9 | -1.00 | 0.00   | 0.90 | 5.5  |

  

| Eosinophils (%) |                | Summary Statistics |      |       |      |       |        |      |      |
|-----------------|----------------|--------------------|------|-------|------|-------|--------|------|------|
| Visit           | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3   | Max  |
| Week 2          | ATL-962 120 mg | 504                | 0.06 | 1.231 | -5.3 | -0.50 | 0.00   | 0.60 | 11.1 |
| Week 4          | ATL-962 120 mg | 493                | 0.17 | 1.487 | -4.7 | -0.40 | 0.10   | 0.60 | 14.9 |
| Week 8          | ATL-962 120 mg | 489                | 0.16 | 1.324 | -4.4 | -0.50 | 0.00   | 0.70 | 6.3  |
| Week 12         | ATL-962 120 mg | 479                | 0.05 | 1.425 | -6.5 | -0.70 | 0.00   | 0.70 | 10.9 |
| Week 16         | ATL-962 120 mg | 472                | 0.21 | 1.923 | -6.1 | -0.50 | 0.10   | 0.70 | 24.0 |
| Week 20         | ATL-962 120 mg | 469                | 0.06 | 1.513 | -5.9 | -0.60 | 0.00   | 0.70 | 8.9  |
| Week 24         | ATL-962 120 mg | 461                | 0.10 | 1.940 | -5.5 | -0.70 | 0.00   | 0.70 | 20.5 |
| Week 28         | ATL-962 120 mg | 453                | 0.07 | 1.576 | -8.3 | -0.60 | 0.00   | 0.70 | 12.9 |
| Week 32         | ATL-962 120 mg | 456                | 0.14 | 1.614 | -5.6 | -0.50 | 0.10   | 0.70 | 16.4 |
| Week 36         | ATL-962 120 mg | 452                | 0.05 | 1.539 | -8.3 | -0.70 | 0.00   | 0.70 | 8.4  |
| Week 40         | ATL-962 120 mg | 445                | 0.11 | 1.644 | -5.9 | -0.60 | 0.00   | 0.70 | 14.4 |
| Week 44         | ATL-962 120 mg | 442                | 0.10 | 1.913 | -6.4 | -0.70 | 0.00   | 0.70 | 25.4 |
| Week 48         | ATL-962 120 mg | 439                | 0.03 | 1.603 | -6.2 | -0.50 | 0.00   | 0.70 | 15.4 |
| Week 52         | ATL-962 120 mg | 439                | 0.08 | 1.795 | -5.4 | -0.70 | 0.00   | 0.70 | 21.4 |

  

| Basophils (%) |                | Summary Statistics |       |       |      |       |        |      |     |
|---------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit         | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2        | ATL-962 120 mg | 504                | 0.03  | 0.438 | -2.0 | -0.20 | 0.00   | 0.20 | 2.1 |
| Week 4        | ATL-962 120 mg | 493                | 0.01  | 0.401 | -1.5 | -0.20 | 0.00   | 0.20 | 1.5 |
| Week 8        | ATL-962 120 mg | 489                | 0.00  | 0.420 | -2.0 | -0.20 | 0.00   | 0.20 | 2.0 |
| Week 12       | ATL-962 120 mg | 479                | 0.02  | 0.443 | -2.0 | -0.20 | 0.00   | 0.20 | 2.0 |
| Week 16       | ATL-962 120 mg | 472                | 0.02  | 0.411 | -2.0 | -0.20 | 0.00   | 0.20 | 1.6 |
| Week 20       | ATL-962 120 mg | 469                | 0.01  | 0.457 | -2.0 | -0.20 | 0.00   | 0.20 | 2.3 |
| Week 24       | ATL-962 120 mg | 461                | 0.04  | 0.484 | -2.0 | -0.20 | 0.00   | 0.20 | 2.5 |
| Week 28       | ATL-962 120 mg | 453                | 0.07  | 0.485 | -2.0 | -0.20 | 0.00   | 0.30 | 2.4 |
| Week 32       | ATL-962 120 mg | 456                | 0.00  | 0.427 | -1.8 | -0.20 | 0.00   | 0.20 | 1.6 |
| Week 36       | ATL-962 120 mg | 452                | 0.05  | 0.416 | -1.7 | -0.20 | 0.00   | 0.20 | 1.8 |
| Week 40       | ATL-962 120 mg | 445                | 0.00  | 0.399 | -1.7 | -0.20 | 0.00   | 0.20 | 1.6 |
| Week 44       | ATL-962 120 mg | 442                | 0.01  | 0.420 | -1.7 | -0.20 | 0.00   | 0.20 | 2.1 |
| Week 48       | ATL-962 120 mg | 439                | -0.01 | 0.414 | -1.6 | -0.20 | 0.00   | 0.20 | 2.4 |
| Week 52       | ATL-962 120 mg | 439                | -0.02 | 0.395 | -1.6 | -0.20 | 0.00   | 0.20 | 1.7 |

表 2.7.4.7-付録-30 血液学的検査の評価時点別の変化量 (OCT-002 試験) (4/4)

| Platelet Count ( $\times 10^4 / \mu\text{L}$ ) |                | Summary Statistics |       |       |       |       |        |      |      |
|--|----------------|--------------------|-------|-------|-------|-------|--------|------|------|
| Visit  | Treatment      | N                  | Mean  | SD    | Min   | Q1    | Median | Q3   | Max  |
| Week 2   | ATL-962 120 mg | 498                | 0.47  | 2.398 | -9.5  | -0.80 | 0.30   | 1.70 | 15.5 |
| Week 4   | ATL-962 120 mg | 488                | 0.47  | 2.060 | -7.0  | -0.80 | 0.50   | 1.60 | 9.0  |
| Week 8   | ATL-962 120 mg | 485                | 0.56  | 2.241 | -10.3 | -0.80 | 0.50   | 1.90 | 12.5 |
| Week 12  | ATL-962 120 mg | 476                | 0.49  | 2.388 | -8.5  | -1.00 | 0.40   | 1.80 | 11.5 |
| Week 16  | ATL-962 120 mg | 468                | 0.43  | 2.319 | -9.5  | -0.85 | 0.50   | 1.70 | 14.2 |
| Week 20  | ATL-962 120 mg | 465                | 0.63  | 2.665 | -9.5  | -1.00 | 0.40   | 2.00 | 18.0 |
| Week 24  | ATL-962 120 mg | 459                | 0.39  | 2.577 | -8.6  | -1.00 | 0.30   | 1.70 | 13.1 |
| Week 28  | ATL-962 120 mg | 449                | 0.41  | 2.535 | -7.3  | -1.20 | 0.30   | 1.80 | 17.0 |
| Week 32  | ATL-962 120 mg | 450                | 0.30  | 2.520 | -13.3 | -1.20 | 0.20   | 1.90 | 10.7 |
| Week 36  | ATL-962 120 mg | 449                | 0.25  | 2.674 | -9.2  | -1.30 | 0.20   | 1.70 | 17.1 |
| Week 40  | ATL-962 120 mg | 441                | 0.01  | 2.346 | -10.0 | -1.20 | 0.10   | 1.40 | 7.3  |
| Week 44  | ATL-962 120 mg | 438                | -0.03 | 2.436 | -11.5 | -1.50 | 0.10   | 1.30 | 8.8  |
| Week 48  | ATL-962 120 mg | 435                | 0.12  | 2.596 | -10.6 | -1.20 | 0.00   | 1.50 | 10.8 |
| Week 52  | ATL-962 120 mg | 435                | 0.10  | 2.557 | -9.9  | -1.20 | 0.00   | 1.60 | 13.1 |

| Prothrombin Time (sec) |                | Summary Statistics |       |       |       |       |        |      |     |
|------------------------|----------------|--------------------|-------|-------|-------|-------|--------|------|-----|
| Visit                  | Treatment      | N                  | Mean  | SD    | Min   | Q1    | Median | Q3   | Max |
| Week 2                 | ATL-962 120 mg | 502                | 0.14  | 0.775 | -3.9  | -0.20 | 0.10   | 0.40 | 8.3 |
| Week 4                 | ATL-962 120 mg | 493                | 0.08  | 0.582 | -2.7  | -0.20 | 0.10   | 0.40 | 4.7 |
| Week 8                 | ATL-962 120 mg | 487                | 0.05  | 0.618 | -4.7  | -0.30 | 0.10   | 0.40 | 2.5 |
| Week 12                | ATL-962 120 mg | 477                | 0.04  | 0.737 | -7.5  | -0.30 | 0.10   | 0.40 | 5.4 |
| Week 16                | ATL-962 120 mg | 469                | 0.02  | 0.669 | -5.2  | -0.30 | 0.00   | 0.40 | 3.8 |
| Week 20                | ATL-962 120 mg | 467                | -0.01 | 0.823 | -10.6 | -0.30 | 0.00   | 0.30 | 5.4 |
| Week 24                | ATL-962 120 mg | 460                | 0.06  | 0.706 | -4.5  | -0.30 | 0.10   | 0.40 | 7.2 |
| Week 28                | ATL-962 120 mg | 451                | 0.07  | 0.783 | -6.5  | -0.30 | 0.10   | 0.50 | 4.7 |
| Week 32                | ATL-962 120 mg | 454                | 0.26  | 0.843 | -8.8  | -0.10 | 0.30   | 0.60 | 6.6 |
| Week 36                | ATL-962 120 mg | 449                | 0.29  | 0.900 | -7.0  | -0.10 | 0.30   | 0.70 | 6.3 |
| Week 40                | ATL-962 120 mg | 443                | 0.29  | 0.691 | -4.1  | 0.00  | 0.30   | 0.70 | 4.5 |
| Week 44                | ATL-962 120 mg | 440                | 0.24  | 0.808 | -7.7  | -0.10 | 0.30   | 0.60 | 4.7 |
| Week 48                | ATL-962 120 mg | 437                | 0.20  | 0.711 | -3.8  | -0.20 | 0.20   | 0.60 | 6.4 |
| Week 52                | ATL-962 120 mg | 437                | 0.11  | 0.761 | -5.2  | -0.20 | 0.10   | 0.40 | 9.0 |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (1/11)

|            |         |        | ATL-962 120 mg |        |       |       |
|------------|---------|--------|----------------|--------|-------|-------|
|            |         |        | Week 0         |        |       |       |
|            |         |        | Lower          | Normal | Upper | Total |
| Hemoglobin | Week 2  | Lower  | 41             | 17     | -     | 58    |
|            |         | Normal | 7              | 428    | 2     | 437   |
|            |         | Upper  | -              | 2      | 6     | 8     |
|            |         | Total  | 48             | 447    | 8     | 503   |
|            | Week 4  | Lower  | 37             | 18     | -     | 55    |
|            |         | Normal | 10             | 418    | 2     | 430   |
|            |         | Upper  | -              | 1      | 6     | 7     |
|            |         | Total  | 47             | 437    | 8     | 492   |
|            | Week 8  | Lower  | 31             | 13     | -     | 44    |
|            |         | Normal | 15             | 416    | 1     | 432   |
|            |         | Upper  | -              | 5      | 7     | 12    |
|            |         | Total  | 46             | 434    | 8     | 488   |
|            | Week 12 | Lower  | 31             | 11     | -     | 42    |
|            |         | Normal | 14             | 411    | 3     | 428   |
|            |         | Upper  | -              | 3      | 5     | 8     |
|            |         | Total  | 45             | 425    | 8     | 478   |
|            | Week 16 | Lower  | 22             | 11     | -     | 33    |
|            |         | Normal | 22             | 401    | 3     | 426   |
|            |         | Upper  | -              | 7      | 5     | 12    |
|            |         | Total  | 44             | 419    | 8     | 471   |
|            | Week 20 | Lower  | 23             | 9      | -     | 32    |
|            |         | Normal | 20             | 400    | 3     | 423   |
|            |         | Upper  | -              | 8      | 5     | 13    |
|            |         | Total  | 43             | 417    | 8     | 468   |
|            | Week 24 | Lower  | 22             | 12     | -     | 34    |
|            |         | Normal | 20             | 393    | 2     | 415   |
|            |         | Upper  | -              | 6      | 6     | 12    |
|            |         | Total  | 42             | 411    | 8     | 461   |
|            | Week 28 | Lower  | 22             | 13     | -     | 35    |
|            |         | Normal | 18             | 385    | 5     | 408   |
|            |         | Upper  | -              | 6      | 3     | 9     |
|            |         | Total  | 40             | 404    | 8     | 452   |
|            | Week 32 | Lower  | 24             | 17     | -     | 41    |
|            |         | Normal | 17             | 384    | 5     | 406   |
|            |         | Upper  | -              | 5      | 3     | 8     |
|            |         | Total  | 41             | 406    | 8     | 455   |
|            | Week 36 | Lower  | 27             | 26     | -     | 53    |
|            |         | Normal | 14             | 370    | 4     | 388   |
|            |         | Upper  | -              | 6      | 4     | 10    |
|            |         | Total  | 41             | 402    | 8     | 451   |
|            | Week 40 | Lower  | 29             | 26     | -     | 55    |
|            |         | Normal | 9              | 369    | 5     | 383   |
|            |         | Upper  | -              | 3      | 3     | 6     |
|            |         | Total  | 38             | 398    | 8     | 444   |
| Week 44    | Lower   | 29     | 19             | -      | 48    |       |
|            | Normal  | 9      | 373            | 4      | 386   |       |
|            | Upper   | -      | 3              | 4      | 7     |       |
|            | Total   | 38     | 395            | 8      | 441   |       |
| Week 48    | Lower   | 26     | 20             | -      | 46    |       |
|            | Normal  | 12     | 371            | 4      | 387   |       |
|            | Upper   | -      | 1              | 4      | 5     |       |
|            | Total   | 38     | 392            | 8      | 438   |       |
| Week 52    | Lower   | 24     | 21             | -      | 45    |       |
|            | Normal  | 13     | 367            | 3      | 383   |       |
|            | Upper   | -      | 5              | 5      | 10    |       |
|            | Total   | 37     | 393            | 8      | 438   |       |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (2/11)

|            |         |        | ATL-962 120 mg |        |       |       |
|------------|---------|--------|----------------|--------|-------|-------|
|            |         |        | Week 0         |        |       |       |
|            |         |        | Lower          | Normal | Upper | Total |
| Hematocrit | Week 2  | Lower  | 33             | 14     | -     | 47    |
|            |         | Normal | 7              | 441    | 1     | 449   |
|            |         | Upper  | -              | 3      | 5     | 8     |
|            |         | Total  | 40             | 458    | 6     | 504   |
|            | Week 4  | Lower  | 33             | 16     | -     | 49    |
|            |         | Normal | 7              | 429    | 1     | 437   |
|            |         | Upper  | -              | 2      | 5     | 7     |
|            |         | Total  | 40             | 447    | 6     | 493   |
|            | Week 8  | Lower  | 26             | 19     | -     | 45    |
|            |         | Normal | 14             | 420    | 5     | 439   |
|            |         | Upper  | -              | 4      | 1     | 5     |
|            |         | Total  | 40             | 443    | 6     | 489   |
|            | Week 12 | Lower  | 27             | 16     | -     | 43    |
|            |         | Normal | 12             | 416    | 2     | 430   |
|            |         | Upper  | -              | 2      | 4     | 6     |
|            |         | Total  | 39             | 434    | 6     | 479   |
|            | Week 16 | Lower  | 23             | 17     | -     | 40    |
|            |         | Normal | 15             | 407    | 1     | 423   |
|            |         | Upper  | -              | 4      | 5     | 9     |
|            |         | Total  | 38             | 428    | 6     | 472   |
|            | Week 20 | Lower  | 20             | 10     | -     | 30    |
|            |         | Normal | 17             | 414    | 2     | 433   |
|            |         | Upper  | -              | 2      | 4     | 6     |
|            |         | Total  | 37             | 426    | 6     | 469   |
|            | Week 24 | Lower  | 24             | 16     | -     | 40    |
|            |         | Normal | 12             | 401    | -     | 413   |
|            |         | Upper  | -              | 3      | 6     | 9     |
|            |         | Total  | 36             | 420    | 6     | 462   |
|            | Week 28 | Lower  | 23             | 13     | -     | 36    |
|            |         | Normal | 12             | 399    | 4     | 415   |
|            |         | Upper  | -              | -      | 2     | 2     |
|            |         | Total  | 35             | 412    | 6     | 453   |
|            | Week 32 | Lower  | 22             | 18     | -     | 40    |
|            |         | Normal | 13             | 396    | 3     | 412   |
|            |         | Upper  | -              | 1      | 3     | 4     |
|            |         | Total  | 35             | 415    | 6     | 456   |
|            | Week 36 | Lower  | 21             | 20     | -     | 41    |
|            |         | Normal | 14             | 387    | 1     | 402   |
|            |         | Upper  | -              | 4      | 5     | 9     |
|            |         | Total  | 35             | 411    | 6     | 452   |
|            | Week 40 | Lower  | 24             | 22     | -     | 46    |
|            |         | Normal | 9              | 382    | 4     | 395   |
|            |         | Upper  | -              | 2      | 2     | 4     |
|            |         | Total  | 33             | 406    | 6     | 445   |
|            | Week 44 | Lower  | 21             | 15     | -     | 36    |
|            |         | Normal | 12             | 386    | 3     | 401   |
|            |         | Upper  | -              | 2      | 3     | 5     |
|            |         | Total  | 33             | 403    | 6     | 442   |
| Week 48    | Lower   | 19     | 18             | -      | 37    |       |
|            | Normal  | 13     | 382            | 4      | 399   |       |
|            | Upper   | -      | 1              | 2      | 3     |       |
|            | Total   | 32     | 401            | 6      | 439   |       |
| Week 52    | Lower   | 21     | 22             | -      | 43    |       |
|            | Normal  | 10     | 376            | 5      | 391   |       |
|            | Upper   | -      | 4              | 1      | 5     |       |
|            | Total   | 31     | 402            | 6      | 439   |       |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (3/11)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| RBC     | Week 2  | Lower  | 50             | 21     | -     | 71    |
|         |         | Normal | 6              | 417    | 3     | 426   |
|         |         | Upper  | -              | 4      | 3     | 7     |
|         |         | Total  | 56             | 442    | 6     | 504   |
|         | Week 4  | Lower  | 43             | 19     | -     | 62    |
|         |         | Normal | 12             | 411    | 2     | 425   |
|         |         | Upper  | -              | 2      | 4     | 6     |
|         |         | Total  | 55             | 432    | 6     | 493   |
|         | Week 8  | Lower  | 41             | 18     | -     | 59    |
|         |         | Normal | 12             | 408    | 4     | 424   |
|         |         | Upper  | -              | 4      | 2     | 6     |
|         |         | Total  | 53             | 430    | 6     | 489   |
|         | Week 12 | Lower  | 41             | 24     | -     | 65    |
|         |         | Normal | 12             | 394    | 4     | 410   |
|         |         | Upper  | -              | 2      | 2     | 4     |
|         |         | Total  | 53             | 420    | 6     | 479   |
|         | Week 16 | Lower  | 35             | 12     | -     | 47    |
|         |         | Normal | 17             | 396    | 2     | 415   |
|         |         | Upper  | -              | 6      | 4     | 10    |
|         |         | Total  | 52             | 414    | 6     | 472   |
|         | Week 20 | Lower  | 37             | 13     | -     | 50    |
|         |         | Normal | 14             | 397    | 2     | 413   |
|         |         | Upper  | -              | 2      | 4     | 6     |
|         |         | Total  | 51             | 412    | 6     | 469   |
|         | Week 24 | Lower  | 34             | 14     | -     | 48    |
|         |         | Normal | 16             | 388    | 2     | 406   |
|         |         | Upper  | -              | 4      | 4     | 8     |
|         |         | Total  | 50             | 406    | 6     | 462   |
|         | Week 28 | Lower  | 37             | 13     | -     | 50    |
|         |         | Normal | 12             | 382    | 4     | 398   |
|         |         | Upper  | -              | 3      | 2     | 5     |
|         |         | Total  | 49             | 398    | 6     | 453   |
|         | Week 32 | Lower  | 34             | 20     | -     | 54    |
|         |         | Normal | 15             | 379    | 4     | 398   |
|         |         | Upper  | -              | 2      | 2     | 4     |
|         |         | Total  | 49             | 401    | 6     | 456   |
|         | Week 36 | Lower  | 37             | 20     | -     | 57    |
|         |         | Normal | 12             | 373    | 3     | 388   |
|         |         | Upper  | -              | 4      | 3     | 7     |
|         |         | Total  | 49             | 397    | 6     | 452   |
|         | Week 40 | Lower  | 39             | 19     | -     | 58    |
|         |         | Normal | 8              | 369    | 4     | 381   |
|         |         | Upper  | -              | 4      | 2     | 6     |
|         |         | Total  | 47             | 392    | 6     | 445   |
|         | Week 44 | Lower  | 39             | 22     | -     | 61    |
|         |         | Normal | 8              | 364    | 4     | 376   |
|         |         | Upper  | -              | 3      | 2     | 5     |
|         |         | Total  | 47             | 389    | 6     | 442   |
| Week 48 | Lower   | 36     | 18             | -      | 54    |       |
|         | Normal  | 11     | 366            | 3      | 380   |       |
|         | Upper   | -      | 2              | 3      | 5     |       |
|         | Total   | 47     | 386            | 6      | 439   |       |
| Week 52 | Lower   | 39     | 26             | -      | 65    |       |
|         | Normal  | 7      | 360            | 4      | 371   |       |
|         | Upper   | -      | 1              | 2      | 3     |       |
|         | Total   | 46     | 387            | 6      | 439   |       |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (4/11)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| WBC     | Week 2  | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | 1              | 426    | 16    | 443   |
|         |         | Upper  | -              | 22     | 37    | 59    |
|         |         | Total  | 2              | 449    | 53    | 504   |
|         | Week 4  | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | 1              | 421    | 12    | 434   |
|         |         | Upper  | -              | 17     | 41    | 58    |
|         |         | Total  | 2              | 438    | 53    | 493   |
|         | Week 8  | Lower  | -              | 2      | -     | 2     |
|         |         | Normal | 2              | 413    | 20    | 435   |
|         |         | Upper  | -              | 20     | 32    | 52    |
|         |         | Total  | 2              | 435    | 52    | 489   |
|         | Week 12 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 2              | 406    | 16    | 424   |
|         |         | Upper  | -              | 19     | 35    | 54    |
|         |         | Total  | 2              | 426    | 51    | 479   |
|         | Week 16 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 2              | 405    | 19    | 426   |
|         |         | Upper  | -              | 16     | 29    | 45    |
|         |         | Total  | 2              | 422    | 48    | 472   |
|         | Week 20 | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | 1              | 398    | 18    | 417   |
|         |         | Upper  | -              | 21     | 30    | 51    |
|         |         | Total  | 2              | 419    | 48    | 469   |
|         | Week 24 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 2              | 392    | 22    | 416   |
|         |         | Upper  | -              | 20     | 25    | 45    |
|         |         | Total  | 2              | 413    | 47    | 462   |
|         | Week 28 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 2              | 390    | 16    | 408   |
|         |         | Upper  | -              | 14     | 31    | 45    |
|         |         | Total  | 2              | 404    | 47    | 453   |
|         | Week 32 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 2              | 389    | 20    | 411   |
|         |         | Upper  | -              | 18     | 27    | 45    |
|         |         | Total  | 2              | 407    | 47    | 456   |
|         | Week 36 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 2              | 387    | 22    | 411   |
|         |         | Upper  | -              | 16     | 24    | 40    |
|         |         | Total  | 2              | 404    | 46    | 452   |
|         | Week 40 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 2              | 384    | 16    | 402   |
|         |         | Upper  | -              | 13     | 30    | 43    |
|         |         | Total  | 2              | 397    | 46    | 445   |
|         | Week 44 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 2              | 374    | 16    | 392   |
|         |         | Upper  | -              | 19     | 30    | 49    |
|         |         | Total  | 2              | 394    | 46    | 442   |
| Week 48 | Lower   | -      | 1              | -      | 1     |       |
|         | Normal  | 2      | 375            | 20     | 397   |       |
|         | Upper   | -      | 15             | 26     | 41    |       |
|         | Total   | 2      | 391            | 46     | 439   |       |
| Week 52 | Lower   | -      | 3              | -      | 3     |       |
|         | Normal  | 2      | 370            | 19     | 391   |       |
|         | Upper   | -      | 18             | 27     | 45    |       |
|         | Total   | 2      | 391            | 46     | 439   |       |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (5/11)

|             |         |        | ATL-962 120 mg |        |       |       |
|-------------|---------|--------|----------------|--------|-------|-------|
|             |         |        | Week 0         |        |       |       |
|             |         |        | Lower          | Normal | Upper | Total |
| Neutrophils | Week 2  | Lower  | 7              | 5      | -     | 12    |
|             |         | Normal | 5              | 479    | 1     | 485   |
|             |         | Upper  | -              | 4      | 3     | 7     |
|             |         | Total  | 12             | 488    | 4     | 504   |
|             | Week 4  | Lower  | 6              | 9      | -     | 15    |
|             |         | Normal | 5              | 464    | 2     | 471   |
|             |         | Upper  | -              | 5      | 2     | 7     |
|             |         | Total  | 11             | 478    | 4     | 493   |
|             | Week 8  | Lower  | 3              | 5      | -     | 8     |
|             |         | Normal | 8              | 466    | 1     | 475   |
|             |         | Upper  | -              | 3      | 3     | 6     |
|             |         | Total  | 11             | 474    | 4     | 489   |
|             | Week 12 | Lower  | 3              | 5      | -     | 8     |
|             |         | Normal | 8              | 457    | 1     | 466   |
|             |         | Upper  | -              | 2      | 3     | 5     |
|             |         | Total  | 11             | 464    | 4     | 479   |
|             | Week 16 | Lower  | 4              | 6      | -     | 10    |
|             |         | Normal | 7              | 447    | 1     | 455   |
|             |         | Upper  | -              | 4      | 3     | 7     |
|             |         | Total  | 11             | 457    | 4     | 472   |
|             | Week 20 | Lower  | 2              | 4      | -     | 6     |
|             |         | Normal | 9              | 447    | 4     | 460   |
|             |         | Upper  | -              | 3      | -     | 3     |
|             |         | Total  | 11             | 454    | 4     | 469   |
|             | Week 24 | Lower  | 1              | 9      | -     | 10    |
|             |         | Normal | 10             | 433    | 1     | 444   |
|             |         | Upper  | -              | 4      | 3     | 7     |
|             |         | Total  | 11             | 446    | 4     | 461   |
|             | Week 28 | Lower  | 3              | 5      | -     | 8     |
|             |         | Normal | 8              | 426    | 4     | 438   |
|             |         | Upper  | -              | 7      | -     | 7     |
|             |         | Total  | 11             | 438    | 4     | 453   |
|             | Week 32 | Lower  | 5              | 7      | -     | 12    |
|             |         | Normal | 6              | 431    | 3     | 440   |
|             |         | Upper  | -              | 3      | 1     | 4     |
|             |         | Total  | 11             | 441    | 4     | 456   |
|             | Week 36 | Lower  | 7              | 8      | -     | 15    |
|             |         | Normal | 3              | 425    | 3     | 431   |
|             |         | Upper  | -              | 5      | 1     | 6     |
|             |         | Total  | 10             | 438    | 4     | 452   |
|             | Week 40 | Lower  | 2              | 8      | -     | 10    |
|             |         | Normal | 8              | 419    | 4     | 431   |
|             |         | Upper  | -              | 4      | -     | 4     |
|             |         | Total  | 10             | 431    | 4     | 445   |
|             | Week 44 | Lower  | 6              | 5      | -     | 11    |
|             |         | Normal | 4              | 420    | 3     | 427   |
|             |         | Upper  | -              | 3      | 1     | 4     |
|             |         | Total  | 10             | 428    | 4     | 442   |
| Week 48     | Lower   | 5      | 6              | -      | 11    |       |
|             | Normal  | 5      | 417            | 1      | 423   |       |
|             | Upper   | -      | 3              | 2      | 5     |       |
|             | Total   | 10     | 426            | 3      | 439   |       |
| Week 52     | Lower   | 6      | 6              | -      | 12    |       |
|             | Normal  | 4      | 419            | 2      | 425   |       |
|             | Upper   | -      | 1              | 1      | 2     |       |
|             | Total   | 10     | 426            | 3      | 439   |       |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (6/11)

|             |         |        | ATL-962 120 mg |        |       |       |
|-------------|---------|--------|----------------|--------|-------|-------|
|             |         |        | Week 0         |        |       |       |
|             |         |        | Lower          | Normal | Upper | Total |
| Lymphocytes | Week 2  | Lower  | 6              | 3      | -     | 9     |
|             |         | Normal | 3              | 476    | 5     | 484   |
|             |         | Upper  | -              | 6      | 5     | 11    |
|             |         | Total  | 9              | 485    | 10    | 504   |
|             | Week 4  | Lower  | 2              | 10     | -     | 12    |
|             |         | Normal | 7              | 456    | 5     | 468   |
|             |         | Upper  | -              | 8      | 5     | 13    |
|             |         | Total  | 9              | 474    | 10    | 493   |
|             | Week 8  | Lower  | 4              | 11     | -     | 15    |
|             |         | Normal | 5              | 455    | 6     | 466   |
|             |         | Upper  | -              | 4      | 4     | 8     |
|             |         | Total  | 9              | 470    | 10    | 489   |
|             | Week 12 | Lower  | 4              | 5      | -     | 9     |
|             |         | Normal | 5              | 452    | 8     | 465   |
|             |         | Upper  | -              | 3      | 2     | 5     |
|             |         | Total  | 9              | 460    | 10    | 479   |
|             | Week 16 | Lower  | 4              | 4      | -     | 8     |
|             |         | Normal | 5              | 444    | 5     | 454   |
|             |         | Upper  | -              | 5      | 5     | 10    |
|             |         | Total  | 9              | 453    | 10    | 472   |
|             | Week 20 | Lower  | 2              | 4      | -     | 6     |
|             |         | Normal | 7              | 441    | 9     | 457   |
|             |         | Upper  | -              | 5      | 1     | 6     |
|             |         | Total  | 9              | 450    | 10    | 469   |
|             | Week 24 | Lower  | 2              | 8      | -     | 10    |
|             |         | Normal | 7              | 429    | 7     | 443   |
|             |         | Upper  | -              | 5      | 3     | 8     |
|             |         | Total  | 9              | 442    | 10    | 461   |
|             | Week 28 | Lower  | 1              | 9      | -     | 10    |
|             |         | Normal | 8              | 419    | 7     | 434   |
|             |         | Upper  | -              | 6      | 3     | 9     |
|             |         | Total  | 9              | 434    | 10    | 453   |
|             | Week 32 | Lower  | 2              | 5      | -     | 7     |
|             |         | Normal | 7              | 427    | 6     | 440   |
|             |         | Upper  | -              | 5      | 4     | 9     |
|             |         | Total  | 9              | 437    | 10    | 456   |
|             | Week 36 | Lower  | 2              | 9      | -     | 11    |
|             |         | Normal | 7              | 419    | 3     | 429   |
|             |         | Upper  | -              | 5      | 7     | 12    |
|             |         | Total  | 9              | 433    | 10    | 452   |
|             | Week 40 | Lower  | 1              | 6      | -     | 7     |
|             |         | Normal | 8              | 415    | 6     | 429   |
|             |         | Upper  | -              | 6      | 3     | 9     |
|             |         | Total  | 9              | 427    | 9     | 445   |
|             | Week 44 | Lower  | -              | 4      | -     | 4     |
|             |         | Normal | 9              | 415    | 4     | 428   |
|             |         | Upper  | -              | 5      | 5     | 10    |
|             |         | Total  | 9              | 424    | 9     | 442   |
|             | Week 48 | Lower  | 1              | 5      | -     | 6     |
|             |         | Normal | 7              | 411    | 5     | 423   |
|             |         | Upper  | -              | 6      | 4     | 10    |
|             |         | Total  | 8              | 422    | 9     | 439   |
| Week 52     | Lower   | 2      | 1              | -      | 3     |       |
|             | Normal  | 6      | 415            | 4      | 425   |       |
|             | Upper   | -      | 6              | 5      | 11    |       |
|             | Total   | 8      | 422            | 9      | 439   |       |



表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (7/11)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| Monocytes | Week 2  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 473    | 10    | 483   |
|           |         | Upper  | -              | 13     | 8     | 21    |
|           |         | Total  | -              | 486    | 18    | 504   |
|           | Week 4  | Lower  | -              | 1      | -     | 1     |
|           |         | Normal | -              | 467    | 11    | 478   |
|           |         | Upper  | -              | 10     | 4     | 14    |
|           |         | Total  | -              | 478    | 15    | 493   |
|           | Week 8  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 461    | 11    | 472   |
|           |         | Upper  | -              | 13     | 4     | 17    |
|           |         | Total  | -              | 474    | 15    | 489   |
|           | Week 12 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 450    | 9     | 459   |
|           |         | Upper  | -              | 14     | 6     | 20    |
|           |         | Total  | -              | 464    | 15    | 479   |
|           | Week 16 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 446    | 11    | 457   |
|           |         | Upper  | -              | 11     | 4     | 15    |
|           |         | Total  | -              | 457    | 15    | 472   |
|           | Week 20 | Lower  | -              | 1      | -     | 1     |
|           |         | Normal | -              | 443    | 11    | 454   |
|           |         | Upper  | -              | 10     | 4     | 14    |
|           |         | Total  | -              | 454    | 15    | 469   |
|           | Week 24 | Lower  | -              | 1      | -     | 1     |
|           |         | Normal | -              | 432    | 9     | 441   |
|           |         | Upper  | -              | 13     | 6     | 19    |
|           |         | Total  | -              | 446    | 15    | 461   |
|           | Week 28 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 427    | 9     | 436   |
|           |         | Upper  | -              | 11     | 6     | 17    |
|           |         | Total  | -              | 438    | 15    | 453   |
|           | Week 32 | Lower  | -              | 2      | -     | 2     |
|           |         | Normal | -              | 430    | 10    | 440   |
|           |         | Upper  | -              | 9      | 5     | 14    |
|           |         | Total  | -              | 441    | 15    | 456   |
|           | Week 36 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 424    | 9     | 433   |
|           |         | Upper  | -              | 13     | 6     | 19    |
|           |         | Total  | -              | 437    | 15    | 452   |
|           | Week 40 | Lower  | -              | 1      | -     | 1     |
|           |         | Normal | -              | 415    | 11    | 426   |
|           |         | Upper  | -              | 14     | 4     | 18    |
|           |         | Total  | -              | 430    | 15    | 445   |
|           | Week 44 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 422    | 8     | 430   |
|           |         | Upper  | -              | 5      | 7     | 12    |
|           |         | Total  | -              | 427    | 15    | 442   |
|           | Week 48 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 414    | 11    | 425   |
|           |         | Upper  | -              | 10     | 4     | 14    |
|           |         | Total  | -              | 424    | 15    | 439   |
| Week 52   | Lower   | -      | -              | -      | -     |       |
|           | Normal  | -      | 413            | 9      | 422   |       |
|           | Upper   | -      | 11             | 6      | 17    |       |
|           | Total   | -      | 424            | 15     | 439   |       |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (8/11)

|             |         |        | ATL-962 120 mg |        |       |       |
|-------------|---------|--------|----------------|--------|-------|-------|
|             |         |        | Week 0         |        |       |       |
|             |         |        | Lower          | Normal | Upper | Total |
| Eosinophils | Week 2  | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 492    | 4     | 496   |
|             |         | Upper  | -              | 5      | 3     | 8     |
|             |         | Total  | -              | 497    | 7     | 504   |
|             | Week 4  | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 482    | 3     | 485   |
|             |         | Upper  | -              | 4      | 4     | 8     |
|             |         | Total  | -              | 486    | 7     | 493   |
|             | Week 8  | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 479    | 4     | 483   |
|             |         | Upper  | -              | 3      | 3     | 6     |
|             |         | Total  | -              | 482    | 7     | 489   |
|             | Week 12 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 465    | 4     | 469   |
|             |         | Upper  | -              | 7      | 3     | 10    |
|             |         | Total  | -              | 472    | 7     | 479   |
|             | Week 16 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 458    | 4     | 462   |
|             |         | Upper  | -              | 7      | 3     | 10    |
|             |         | Total  | -              | 465    | 7     | 472   |
|             | Week 20 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 459    | 5     | 464   |
|             |         | Upper  | -              | 3      | 2     | 5     |
|             |         | Total  | -              | 462    | 7     | 469   |
|             | Week 24 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 447    | 4     | 451   |
|             |         | Upper  | -              | 7      | 3     | 10    |
|             |         | Total  | -              | 454    | 7     | 461   |
|             | Week 28 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 440    | 5     | 445   |
|             |         | Upper  | -              | 6      | 2     | 8     |
|             |         | Total  | -              | 446    | 7     | 453   |
|             | Week 32 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 442    | 3     | 445   |
|             |         | Upper  | -              | 7      | 4     | 11    |
|             |         | Total  | -              | 449    | 7     | 456   |
|             | Week 36 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 435    | 3     | 438   |
|             |         | Upper  | -              | 10     | 4     | 14    |
|             |         | Total  | -              | 445    | 7     | 452   |
|             | Week 40 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 431    | 4     | 435   |
|             |         | Upper  | -              | 7      | 3     | 10    |
|             |         | Total  | -              | 438    | 7     | 445   |
|             | Week 44 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 432    | 3     | 435   |
|             |         | Upper  | -              | 3      | 4     | 7     |
|             |         | Total  | -              | 435    | 7     | 442   |
|             | Week 48 | Lower  | -              | -      | -     | -     |
|             |         | Normal | -              | 429    | 4     | 433   |
|             |         | Upper  | -              | 3      | 3     | 6     |
|             |         | Total  | -              | 432    | 7     | 439   |
| Week 52     | Lower   | -      | -              | -      | -     |       |
|             | Normal  | -      | 425            | 3      | 428   |       |
|             | Upper   | -      | 7              | 4      | 11    |       |
|             | Total   | -      | 432            | 7      | 439   |       |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (9/11)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| Basophils | Week 2  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 501    | -     | 501   |
|           |         | Upper  | -              | 3      | -     | 3     |
|           |         | Total  | -              | 504    | -     | 504   |
|           | Week 4  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 493    | -     | 493   |
|           |         | Upper  | -              | -      | -     | -     |
|           |         | Total  | -              | 493    | -     | 493   |
|           | Week 8  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 489    | -     | 489   |
|           |         | Upper  | -              | -      | -     | -     |
|           |         | Total  | -              | 489    | -     | 489   |
|           | Week 12 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 476    | -     | 476   |
|           |         | Upper  | -              | 3      | -     | 3     |
|           |         | Total  | -              | 479    | -     | 479   |
|           | Week 16 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 471    | -     | 471   |
|           |         | Upper  | -              | 1      | -     | 1     |
|           |         | Total  | -              | 472    | -     | 472   |
|           | Week 20 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 467    | -     | 467   |
|           |         | Upper  | -              | 2      | -     | 2     |
|           |         | Total  | -              | 469    | -     | 469   |
|           | Week 24 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 457    | -     | 457   |
|           |         | Upper  | -              | 4      | -     | 4     |
|           |         | Total  | -              | 461    | -     | 461   |
|           | Week 28 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 450    | -     | 450   |
|           |         | Upper  | -              | 3      | -     | 3     |
|           |         | Total  | -              | 453    | -     | 453   |
|           | Week 32 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 456    | -     | 456   |
|           |         | Upper  | -              | -      | -     | -     |
|           |         | Total  | -              | 456    | -     | 456   |
|           | Week 36 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 451    | -     | 451   |
|           |         | Upper  | -              | 1      | -     | 1     |
|           |         | Total  | -              | 452    | -     | 452   |
|           | Week 40 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 443    | -     | 443   |
|           |         | Upper  | -              | 2      | -     | 2     |
|           |         | Total  | -              | 445    | -     | 445   |
|           | Week 44 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 441    | -     | 441   |
|           |         | Upper  | -              | 1      | -     | 1     |
|           |         | Total  | -              | 442    | -     | 442   |
|           | Week 48 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 438    | -     | 438   |
|           |         | Upper  | -              | 1      | -     | 1     |
|           |         | Total  | -              | 439    | -     | 439   |
| Week 52   | Lower   | -      | -              | -      | -     |       |
|           | Normal  | -      | 438            | -      | 438   |       |
|           | Upper   | -      | 1              | -      | 1     |       |
|           | Total   | -      | 439            | -      | 439   |       |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (10/11)

|                |         |        | ATL-962 120 mg |        |       |       |
|----------------|---------|--------|----------------|--------|-------|-------|
|                |         |        | Week 0         |        |       |       |
|                |         |        | Lower          | Normal | Upper | Total |
| Platelet Count | Week 2  | Lower  | 15             | 3      | -     | 18    |
|                |         | Normal | 9              | 450    | 6     | 465   |
|                |         | Upper  | -              | 7      | 8     | 15    |
|                |         | Total  | 24             | 460    | 14    | 498   |
|                | Week 4  | Lower  | 15             | 3      | -     | 18    |
|                |         | Normal | 8              | 445    | 5     | 458   |
|                |         | Upper  | -              | 3      | 9     | 12    |
|                |         | Total  | 23             | 451    | 14    | 488   |
|                | Week 8  | Lower  | 15             | 1      | -     | 16    |
|                |         | Normal | 8              | 445    | 4     | 457   |
|                |         | Upper  | -              | 2      | 10    | 12    |
|                |         | Total  | 23             | 448    | 14    | 485   |
|                | Week 12 | Lower  | 14             | 1      | -     | 15    |
|                |         | Normal | 9              | 435    | 4     | 448   |
|                |         | Upper  | -              | 4      | 9     | 13    |
|                |         | Total  | 23             | 440    | 13    | 476   |
|                | Week 16 | Lower  | 15             | 3      | -     | 18    |
|                |         | Normal | 8              | 427    | 3     | 438   |
|                |         | Upper  | -              | 3      | 9     | 12    |
|                |         | Total  | 23             | 433    | 12    | 468   |
|                | Week 20 | Lower  | 10             | 1      | -     | 11    |
|                |         | Normal | 13             | 424    | 7     | 444   |
|                |         | Upper  | -              | 5      | 5     | 10    |
|                |         | Total  | 23             | 430    | 12    | 465   |
|                | Week 24 | Lower  | 13             | 1      | -     | 14    |
|                |         | Normal | 10             | 415    | 6     | 431   |
|                |         | Upper  | -              | 8      | 6     | 14    |
|                |         | Total  | 23             | 424    | 12    | 459   |
|                | Week 28 | Lower  | 14             | 5      | -     | 19    |
|                |         | Normal | 9              | 408    | 5     | 422   |
|                |         | Upper  | -              | 2      | 6     | 8     |
|                |         | Total  | 23             | 415    | 11    | 449   |
|                | Week 32 | Lower  | 13             | 5      | -     | 18    |
|                |         | Normal | 10             | 406    | 3     | 419   |
|                |         | Upper  | -              | 4      | 9     | 13    |
|                |         | Total  | 23             | 415    | 12    | 450   |
|                | Week 36 | Lower  | 14             | 6      | -     | 20    |
|                |         | Normal | 9              | 400    | 4     | 413   |
|                |         | Upper  | -              | 8      | 8     | 16    |
|                |         | Total  | 23             | 414    | 12    | 449   |
|                | Week 40 | Lower  | 13             | 2      | -     | 15    |
|                |         | Normal | 9              | 398    | 4     | 411   |
|                |         | Upper  | -              | 7      | 8     | 15    |
|                |         | Total  | 22             | 407    | 12    | 441   |
|                | Week 44 | Lower  | 17             | 3      | -     | 20    |
|                |         | Normal | 5              | 397    | 5     | 407   |
|                |         | Upper  | -              | 4      | 7     | 11    |
|                |         | Total  | 22             | 404    | 12    | 438   |
| Week 48        | Lower   | 13     | 3              | -      | 16    |       |
|                | Normal  | 8      | 395            | 5      | 408   |       |
|                | Upper   | -      | 4              | 7      | 11    |       |
|                | Total   | 21     | 402            | 12     | 435   |       |
| Week 52        | Lower   | 12     | 4              | -      | 16    |       |
|                | Normal  | 9      | 394            | 5      | 408   |       |
|                | Upper   | -      | 4              | 7      | 11    |       |
|                | Total   | 21     | 402            | 12     | 435   |       |

表 2.7.4.7-付録-31 血液学的検査の評価時点別正異判定クロス表 (OCT-002 試験) (11/11)

|                  |         |        | ATL-962 120 mg |        |       |       |
|------------------|---------|--------|----------------|--------|-------|-------|
|                  |         |        | Week 0         |        |       |       |
|                  |         |        | Lower          | Normal | Upper | Total |
| Prothrombin Time | Week 2  | Lower  | 10             | 19     | -     | 29    |
|                  |         | Normal | 25             | 437    | 1     | 463   |
|                  |         | Upper  | -              | 3      | 7     | 10    |
|                  |         | Total  | 35             | 459    | 8     | 502   |
|                  | Week 4  | Lower  | 11             | 22     | -     | 33    |
|                  |         | Normal | 24             | 427    | 1     | 452   |
|                  |         | Upper  | -              | 1      | 7     | 8     |
|                  |         | Total  | 35             | 450    | 8     | 493   |
|                  | Week 8  | Lower  | 15             | 15     | -     | 30    |
|                  |         | Normal | 20             | 428    | 2     | 450   |
|                  |         | Upper  | -              | 1      | 6     | 7     |
|                  |         | Total  | 35             | 444    | 8     | 487   |
|                  | Week 12 | Lower  | 12             | 18     | -     | 30    |
|                  |         | Normal | 22             | 416    | 1     | 439   |
|                  |         | Upper  | -              | 2      | 6     | 8     |
|                  |         | Total  | 34             | 436    | 7     | 477   |
|                  | Week 16 | Lower  | 16             | 22     | -     | 38    |
|                  |         | Normal | 18             | 404    | 2     | 424   |
|                  |         | Upper  | -              | 1      | 6     | 7     |
|                  |         | Total  | 34             | 427    | 8     | 469   |
|                  | Week 20 | Lower  | 20             | 14     | -     | 34    |
|                  |         | Normal | 14             | 409    | 1     | 424   |
|                  |         | Upper  | -              | 2      | 7     | 9     |
|                  |         | Total  | 34             | 425    | 8     | 467   |
|                  | Week 24 | Lower  | 10             | 19     | -     | 29    |
|                  |         | Normal | 23             | 399    | 1     | 423   |
|                  |         | Upper  | -              | 1      | 7     | 8     |
|                  |         | Total  | 33             | 419    | 8     | 460   |
|                  | Week 28 | Lower  | 12             | 10     | -     | 22    |
|                  |         | Normal | 20             | 400    | 1     | 421   |
|                  |         | Upper  | -              | 1      | 7     | 8     |
|                  |         | Total  | 32             | 411    | 8     | 451   |
|                  | Week 32 | Lower  | 4              | 11     | -     | 15    |
|                  |         | Normal | 29             | 398    | 1     | 428   |
|                  |         | Upper  | -              | 4      | 7     | 11    |
|                  |         | Total  | 33             | 413    | 8     | 454   |
|                  | Week 36 | Lower  | 4              | 3      | -     | 7     |
|                  |         | Normal | 29             | 399    | 2     | 430   |
|                  |         | Upper  | -              | 6      | 6     | 12    |
|                  |         | Total  | 33             | 408    | 8     | 449   |
|                  | Week 40 | Lower  | 2              | 2      | -     | 4     |
|                  |         | Normal | 29             | 398    | 1     | 428   |
|                  |         | Upper  | -              | 4      | 7     | 11    |
|                  |         | Total  | 31             | 404    | 8     | 443   |
|                  | Week 44 | Lower  | 4              | 4      | -     | 8     |
|                  |         | Normal | 27             | 394    | 1     | 422   |
|                  |         | Upper  | -              | 3      | 7     | 10    |
|                  |         | Total  | 31             | 401    | 8     | 440   |
| Week 48          | Lower   | 8      | 5              | -      | 13    |       |
|                  | Normal  | 23     | 389            | 1      | 413   |       |
|                  | Upper   | -      | 4              | 7      | 11    |       |
|                  | Total   | 31     | 398            | 8      | 437   |       |
| Week 52          | Lower   | 8      | 8              | -      | 16    |       |
|                  | Normal  | 22     | 389            | 1      | 412   |       |
|                  | Upper   | -      | 2              | 7      | 9     |       |
|                  | Total   | 30     | 399            | 8      | 437   |       |

表 2.7.4.7-付録-32 血液生化学検査の評価時点別の変化量 (OCT-002 試験) (1/8)

| Total Protein (g/dL) |                | Summary Statistics |       |       |      |       |        |      |     |  |
|----------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|--|
| Visit                | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |  |
| Week 2               | ATL-962 120 mg | 504                | -0.01 | 0.281 | -1.1 | -0.20 | 0.00   | 0.20 | 0.9 |  |
| Week 4               | ATL-962 120 mg | 495                | -0.03 | 0.281 | -1.0 | -0.20 | 0.00   | 0.20 | 0.8 |  |
| Week 8               | ATL-962 120 mg | 489                | 0.02  | 0.303 | -1.1 | -0.20 | 0.00   | 0.20 | 0.9 |  |
| Week 12              | ATL-962 120 mg | 479                | 0.01  | 0.315 | -1.0 | -0.20 | 0.00   | 0.20 | 1.0 |  |
| Week 16              | ATL-962 120 mg | 472                | 0.05  | 0.311 | -0.8 | -0.20 | 0.00   | 0.30 | 1.2 |  |
| Week 20              | ATL-962 120 mg | 469                | 0.06  | 0.299 | -1.0 | -0.10 | 0.10   | 0.20 | 0.9 |  |
| Week 24              | ATL-962 120 mg | 462                | 0.00  | 0.321 | -1.0 | -0.20 | 0.00   | 0.20 | 0.9 |  |
| Week 28              | ATL-962 120 mg | 453                | 0.01  | 0.308 | -0.8 | -0.20 | 0.00   | 0.20 | 1.1 |  |
| Week 32              | ATL-962 120 mg | 456                | 0.03  | 0.318 | -1.2 | -0.20 | 0.00   | 0.20 | 0.9 |  |
| Week 36              | ATL-962 120 mg | 452                | -0.01 | 0.335 | -1.1 | -0.20 | 0.00   | 0.20 | 1.1 |  |
| Week 40              | ATL-962 120 mg | 445                | -0.01 | 0.312 | -1.0 | -0.20 | 0.00   | 0.20 | 1.1 |  |
| Week 44              | ATL-962 120 mg | 442                | 0.00  | 0.306 | -1.0 | -0.20 | 0.00   | 0.20 | 1.0 |  |
| Week 48              | ATL-962 120 mg | 439                | -0.02 | 0.323 | -0.9 | -0.20 | 0.00   | 0.20 | 1.1 |  |
| Week 52              | ATL-962 120 mg | 439                | -0.04 | 0.305 | -1.2 | -0.20 | 0.00   | 0.20 | 1.0 |  |

| Albumin (g/dL) |                | Summary Statistics |       |       |      |       |        |      |     |  |
|----------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|--|
| Visit          | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |  |
| Week 2         | ATL-962 120 mg | 504                | -0.01 | 0.198 | -0.8 | -0.10 | 0.00   | 0.10 | 0.6 |  |
| Week 4         | ATL-962 120 mg | 495                | -0.04 | 0.195 | -0.7 | -0.20 | 0.00   | 0.10 | 0.6 |  |
| Week 8         | ATL-962 120 mg | 489                | -0.01 | 0.213 | -0.9 | -0.10 | 0.00   | 0.10 | 0.6 |  |
| Week 12        | ATL-962 120 mg | 479                | -0.03 | 0.212 | -0.7 | -0.20 | 0.00   | 0.10 | 0.6 |  |
| Week 16        | ATL-962 120 mg | 472                | 0.00  | 0.216 | -0.7 | -0.10 | 0.00   | 0.10 | 0.7 |  |
| Week 20        | ATL-962 120 mg | 469                | 0.00  | 0.212 | -0.8 | -0.10 | 0.00   | 0.20 | 0.8 |  |
| Week 24        | ATL-962 120 mg | 462                | -0.02 | 0.223 | -0.8 | -0.20 | 0.00   | 0.10 | 0.7 |  |
| Week 28        | ATL-962 120 mg | 453                | -0.03 | 0.219 | -0.8 | -0.20 | 0.00   | 0.10 | 0.7 |  |
| Week 32        | ATL-962 120 mg | 456                | -0.01 | 0.226 | -0.7 | -0.20 | 0.00   | 0.10 | 0.6 |  |
| Week 36        | ATL-962 120 mg | 452                | -0.03 | 0.225 | -0.7 | -0.20 | 0.00   | 0.10 | 0.7 |  |
| Week 40        | ATL-962 120 mg | 445                | -0.04 | 0.226 | -0.8 | -0.20 | 0.00   | 0.10 | 0.7 |  |
| Week 44        | ATL-962 120 mg | 442                | -0.04 | 0.217 | -0.7 | -0.20 | 0.00   | 0.10 | 0.6 |  |
| Week 48        | ATL-962 120 mg | 439                | -0.03 | 0.212 | -0.8 | -0.20 | 0.00   | 0.10 | 0.8 |  |
| Week 52        | ATL-962 120 mg | 439                | -0.05 | 0.233 | -0.8 | -0.20 | -0.10  | 0.10 | 0.8 |  |

| ALT (GPT) (IU/L) |                | Summary Statistics |      |       |      |       |        |     |     |  |
|------------------|----------------|--------------------|------|-------|------|-------|--------|-----|-----|--|
| Visit            | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3  | Max |  |
| Week 2           | ATL-962 120 mg | 504                | 1.9  | 10.84 | -58  | -2.0  | 1.0    | 6.0 | 87  |  |
| Week 4           | ATL-962 120 mg | 495                | 2.1  | 16.78 | -80  | -3.0  | 1.0    | 6.0 | 281 |  |
| Week 8           | ATL-962 120 mg | 489                | 1.8  | 13.48 | -64  | -4.0  | 1.0    | 7.0 | 63  |  |
| Week 12          | ATL-962 120 mg | 479                | 0.5  | 14.03 | -56  | -5.0  | 1.0    | 5.0 | 105 |  |
| Week 16          | ATL-962 120 mg | 472                | -0.6 | 14.93 | -89  | -6.0  | 0.0    | 6.0 | 89  |  |
| Week 20          | ATL-962 120 mg | 469                | -2.1 | 15.06 | -77  | -9.0  | -1.0   | 5.0 | 65  |  |
| Week 24          | ATL-962 120 mg | 462                | -3.0 | 16.35 | -102 | -10.0 | -1.0   | 5.0 | 68  |  |
| Week 28          | ATL-962 120 mg | 453                | -3.6 | 19.61 | -91  | -10.0 | -2.0   | 4.0 | 204 |  |
| Week 32          | ATL-962 120 mg | 456                | -4.5 | 16.74 | -99  | -12.0 | -2.5   | 3.0 | 89  |  |
| Week 36          | ATL-962 120 mg | 452                | -4.6 | 20.63 | -107 | -12.5 | -3.0   | 2.5 | 156 |  |
| Week 40          | ATL-962 120 mg | 445                | -5.4 | 17.07 | -109 | -12.0 | -3.0   | 2.0 | 82  |  |
| Week 44          | ATL-962 120 mg | 442                | -4.9 | 17.49 | -111 | -12.0 | -2.0   | 3.0 | 75  |  |
| Week 48          | ATL-962 120 mg | 439                | -5.1 | 17.28 | -121 | -12.0 | -3.0   | 3.0 | 57  |  |
| Week 52          | ATL-962 120 mg | 439                | -5.3 | 17.16 | -124 | -12.0 | -3.0   | 4.0 | 56  |  |

表 2.7.4.7-付録-32 血液生化学検査の評価時点別の変化量 (OCT-002 試験) (2/8)

| AST (GOT) (IU/L) |                | Summary Statistics |      |       |     |      |        |     |     |
|------------------|----------------|--------------------|------|-------|-----|------|--------|-----|-----|
| Visit            | Treatment      | N                  | Mean | SD    | Min | Q1   | Median | Q3  | Max |
| Week 2           | ATL-962 120 mg | 504                | 1.0  | 7.99  | -45 | -2.0 | 1.0    | 4.0 | 35  |
| Week 4           | ATL-962 120 mg | 495                | 1.2  | 11.56 | -57 | -2.0 | 1.0    | 4.0 | 164 |
| Week 8           | ATL-962 120 mg | 489                | 0.8  | 9.24  | -42 | -2.0 | 1.0    | 5.0 | 42  |
| Week 12          | ATL-962 120 mg | 479                | -0.1 | 10.31 | -49 | -3.0 | 1.0    | 4.0 | 108 |
| Week 16          | ATL-962 120 mg | 472                | -1.0 | 9.39  | -43 | -4.0 | 0.0    | 3.0 | 56  |
| Week 20          | ATL-962 120 mg | 469                | -1.7 | 9.80  | -43 | -6.0 | 0.0    | 4.0 | 37  |
| Week 24          | ATL-962 120 mg | 462                | -2.2 | 10.88 | -46 | -6.0 | -1.0   | 3.0 | 39  |
| Week 28          | ATL-962 120 mg | 453                | -2.3 | 14.25 | -42 | -7.0 | -2.0   | 3.0 | 188 |
| Week 32          | ATL-962 120 mg | 456                | -2.9 | 10.86 | -48 | -7.0 | -1.0   | 2.0 | 41  |
| Week 36          | ATL-962 120 mg | 452                | -2.6 | 15.60 | -49 | -7.0 | -2.0   | 2.0 | 183 |
| Week 40          | ATL-962 120 mg | 445                | -3.6 | 10.94 | -46 | -8.0 | -2.0   | 2.0 | 43  |
| Week 44          | ATL-962 120 mg | 442                | -3.1 | 10.85 | -49 | -7.0 | -2.0   | 2.0 | 39  |
| Week 48          | ATL-962 120 mg | 439                | -2.6 | 11.40 | -49 | -8.0 | -2.0   | 2.0 | 57  |
| Week 52          | ATL-962 120 mg | 439                | -2.9 | 10.95 | -48 | -7.0 | -2.0   | 3.0 | 47  |

| γ-GTP (IU/L) |                | Summary Statistics |      |       |      |       |        |     |     |
|--------------|----------------|--------------------|------|-------|------|-------|--------|-----|-----|
| Visit        | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3  | Max |
| Week 2       | ATL-962 120 mg | 504                | 0.6  | 15.73 | -81  | -4.0  | -1.0   | 2.0 | 166 |
| Week 4       | ATL-962 120 mg | 495                | 0.5  | 25.21 | -74  | -5.0  | -1.0   | 3.0 | 451 |
| Week 8       | ATL-962 120 mg | 489                | 0.3  | 20.70 | -118 | -6.0  | -1.0   | 3.0 | 181 |
| Week 12      | ATL-962 120 mg | 479                | -1.8 | 19.94 | -119 | -7.0  | -2.0   | 3.0 | 215 |
| Week 16      | ATL-962 120 mg | 472                | -2.5 | 23.08 | -143 | -8.0  | -2.0   | 2.0 | 187 |
| Week 20      | ATL-962 120 mg | 469                | -5.3 | 20.77 | -196 | -9.0  | -3.0   | 2.0 | 105 |
| Week 24      | ATL-962 120 mg | 462                | -5.2 | 24.74 | -176 | -11.0 | -3.0   | 1.0 | 263 |
| Week 28      | ATL-962 120 mg | 453                | -5.6 | 32.96 | -178 | -11.0 | -4.0   | 0.0 | 394 |
| Week 32      | ATL-962 120 mg | 456                | -6.4 | 26.24 | -199 | -12.0 | -5.0   | 1.0 | 281 |
| Week 36      | ATL-962 120 mg | 452                | -5.5 | 45.75 | -246 | -13.0 | -5.0   | 1.0 | 675 |
| Week 40      | ATL-962 120 mg | 445                | -7.1 | 27.42 | -234 | -12.0 | -5.0   | 1.0 | 229 |
| Week 44      | ATL-962 120 mg | 442                | -8.0 | 24.20 | -212 | -14.0 | -5.0   | 1.0 | 96  |
| Week 48      | ATL-962 120 mg | 439                | -7.2 | 26.71 | -266 | -13.0 | -5.0   | 0.0 | 125 |
| Week 52      | ATL-962 120 mg | 439                | -8.2 | 26.81 | -312 | -13.0 | -5.0   | 0.0 | 124 |

| ALP (IU/L) |                | Summary Statistics |      |       |      |       |        |      |     |
|------------|----------------|--------------------|------|-------|------|-------|--------|------|-----|
| Visit      | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2     | ATL-962 120 mg | 504                | -0.2 | 30.59 | -379 | -10.0 | 1.0    | 12.5 | 145 |
| Week 4     | ATL-962 120 mg | 495                | 1.0  | 31.99 | -385 | -11.0 | 3.0    | 15.0 | 124 |
| Week 8     | ATL-962 120 mg | 489                | 5.2  | 34.36 | -363 | -8.0  | 6.0    | 20.0 | 136 |
| Week 12    | ATL-962 120 mg | 479                | 5.2  | 33.08 | -336 | -8.0  | 7.0    | 21.0 | 159 |
| Week 16    | ATL-962 120 mg | 472                | 8.8  | 33.46 | -320 | -6.0  | 9.0    | 28.0 | 130 |
| Week 20    | ATL-962 120 mg | 469                | 8.1  | 34.71 | -362 | -6.0  | 10.0   | 26.0 | 144 |
| Week 24    | ATL-962 120 mg | 462                | 7.1  | 34.38 | -314 | -11.0 | 8.0    | 26.0 | 120 |
| Week 28    | ATL-962 120 mg | 453                | 7.5  | 37.69 | -333 | -10.0 | 7.0    | 25.0 | 137 |
| Week 32    | ATL-962 120 mg | 456                | 6.0  | 37.66 | -340 | -11.5 | 6.0    | 25.5 | 122 |
| Week 36    | ATL-962 120 mg | 452                | 4.0  | 41.27 | -366 | -16.0 | 1.0    | 22.0 | 302 |
| Week 40    | ATL-962 120 mg | 445                | 4.1  | 38.07 | -351 | -14.0 | 5.0    | 23.0 | 168 |
| Week 44    | ATL-962 120 mg | 442                | 1.3  | 41.40 | -417 | -17.0 | 1.0    | 21.0 | 210 |
| Week 48    | ATL-962 120 mg | 439                | 0.6  | 40.71 | -372 | -17.0 | 1.0    | 22.0 | 148 |
| Week 52    | ATL-962 120 mg | 439                | -0.1 | 39.48 | -347 | -17.0 | 1.0    | 20.0 | 119 |

表 2.7.4.7-付録-32 血液生化学検査の評価時点別の変化量 (OCT-002 試験) (3/8)

| LDH (IU/L) |                | Summary Statistics |      |       |      |       |        |      |     |
|------------|----------------|--------------------|------|-------|------|-------|--------|------|-----|
| Visit      | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2     | ATL-962 120 mg | 504                | -0.3 | 18.27 | -80  | -10.0 | 0.0    | 10.0 | 120 |
| Week 4     | ATL-962 120 mg | 495                | -1.6 | 19.30 | -103 | -12.0 | -1.0   | 9.0  | 80  |
| Week 8     | ATL-962 120 mg | 489                | -0.3 | 19.22 | -96  | -11.0 | 0.0    | 12.0 | 64  |
| Week 12    | ATL-962 120 mg | 479                | -1.9 | 23.48 | -105 | -14.0 | -2.0   | 10.0 | 222 |
| Week 16    | ATL-962 120 mg | 472                | -3.5 | 22.24 | -104 | -16.0 | -4.0   | 9.0  | 132 |
| Week 20    | ATL-962 120 mg | 469                | -3.6 | 22.42 | -108 | -17.0 | -3.0   | 8.0  | 141 |
| Week 24    | ATL-962 120 mg | 462                | -3.5 | 26.44 | -105 | -16.0 | -4.0   | 9.0  | 328 |
| Week 28    | ATL-962 120 mg | 453                | -5.0 | 22.67 | -97  | -18.0 | -5.0   | 8.0  | 158 |
| Week 32    | ATL-962 120 mg | 456                | -4.1 | 23.54 | -106 | -17.0 | -3.0   | 9.0  | 112 |
| Week 36    | ATL-962 120 mg | 452                | -5.6 | 23.70 | -116 | -17.0 | -5.0   | 7.0  | 132 |
| Week 40    | ATL-962 120 mg | 445                | -5.0 | 24.50 | -104 | -17.0 | -6.0   | 6.0  | 234 |
| Week 44    | ATL-962 120 mg | 442                | -5.4 | 23.84 | -99  | -18.0 | -5.0   | 8.0  | 141 |
| Week 48    | ATL-962 120 mg | 439                | -6.8 | 22.41 | -90  | -20.0 | -7.0   | 5.0  | 110 |
| Week 52    | ATL-962 120 mg | 439                | -7.6 | 21.28 | -89  | -19.0 | -6.0   | 6.0  | 83  |

  

| CPK (IU/L) |                | Summary Statistics |      |        |      |       |        |      |      |
|------------|----------------|--------------------|------|--------|------|-------|--------|------|------|
| Visit      | Treatment      | N                  | Mean | SD     | Min  | Q1    | Median | Q3   | Max  |
| Week 2     | ATL-962 120 mg | 504                | 3.1  | 66.21  | -401 | -19.0 | 0.0    | 17.0 | 506  |
| Week 4     | ATL-962 120 mg | 495                | 3.3  | 69.46  | -235 | -17.0 | -2.0   | 15.0 | 792  |
| Week 8     | ATL-962 120 mg | 489                | 5.7  | 59.69  | -240 | -16.0 | 1.0    | 18.0 | 401  |
| Week 12    | ATL-962 120 mg | 479                | 1.1  | 56.43  | -228 | -18.0 | -2.0   | 15.0 | 487  |
| Week 16    | ATL-962 120 mg | 472                | 0.4  | 65.79  | -307 | -23.5 | -1.0   | 17.0 | 626  |
| Week 20    | ATL-962 120 mg | 469                | 4.9  | 78.45  | -272 | -19.0 | -3.0   | 18.0 | 872  |
| Week 24    | ATL-962 120 mg | 462                | 1.1  | 57.90  | -290 | -21.0 | -2.0   | 20.0 | 452  |
| Week 28    | ATL-962 120 mg | 453                | 0.6  | 65.66  | -243 | -23.0 | -5.0   | 15.0 | 448  |
| Week 32    | ATL-962 120 mg | 456                | 5.6  | 90.67  | -223 | -21.0 | -2.0   | 18.0 | 1447 |
| Week 36    | ATL-962 120 mg | 452                | 5.7  | 69.96  | -263 | -20.5 | -1.0   | 18.0 | 775  |
| Week 40    | ATL-962 120 mg | 445                | 24.4 | 377.52 | -301 | -20.0 | 0.0    | 19.0 | 7403 |
| Week 44    | ATL-962 120 mg | 442                | 7.4  | 87.17  | -341 | -20.0 | -0.5   | 20.0 | 834  |
| Week 48    | ATL-962 120 mg | 439                | 8.5  | 106.74 | -350 | -24.0 | -4.0   | 19.0 | 1403 |
| Week 52    | ATL-962 120 mg | 439                | 6.5  | 98.92  | -201 | -22.0 | -1.0   | 17.0 | 1598 |

  

| Uric Acid (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2            | ATL-962 120 mg | 504                | -0.01 | 0.659 | -1.9 | -0.40 | 0.00   | 0.40 | 2.4 |
| Week 4            | ATL-962 120 mg | 495                | -0.04 | 0.689 | -3.2 | -0.40 | 0.00   | 0.40 | 4.1 |
| Week 8            | ATL-962 120 mg | 489                | -0.06 | 0.693 | -2.5 | -0.50 | -0.10  | 0.30 | 2.9 |
| Week 12           | ATL-962 120 mg | 479                | -0.17 | 0.747 | -2.4 | -0.60 | -0.10  | 0.30 | 3.5 |
| Week 16           | ATL-962 120 mg | 472                | -0.19 | 0.778 | -3.0 | -0.70 | -0.20  | 0.20 | 3.6 |
| Week 20           | ATL-962 120 mg | 469                | -0.26 | 0.767 | -3.6 | -0.70 | -0.30  | 0.20 | 2.2 |
| Week 24           | ATL-962 120 mg | 462                | -0.30 | 0.748 | -4.6 | -0.70 | -0.30  | 0.20 | 1.7 |
| Week 28           | ATL-962 120 mg | 453                | -0.28 | 0.739 | -3.3 | -0.80 | -0.20  | 0.20 | 3.0 |
| Week 32           | ATL-962 120 mg | 456                | -0.15 | 0.778 | -3.4 | -0.60 | -0.20  | 0.30 | 2.5 |
| Week 36           | ATL-962 120 mg | 452                | -0.08 | 0.820 | -4.0 | -0.60 | -0.05  | 0.40 | 2.3 |
| Week 40           | ATL-962 120 mg | 445                | -0.09 | 0.827 | -4.1 | -0.60 | -0.10  | 0.40 | 2.6 |
| Week 44           | ATL-962 120 mg | 442                | -0.03 | 0.810 | -3.9 | -0.50 | 0.00   | 0.40 | 2.7 |
| Week 48           | ATL-962 120 mg | 439                | -0.04 | 0.843 | -4.4 | -0.50 | 0.00   | 0.40 | 4.0 |
| Week 52           | ATL-962 120 mg | 439                | -0.03 | 0.784 | -4.7 | -0.40 | 0.00   | 0.40 | 3.7 |



表 2.7.4.7-付録-32 血液生化学検査の評価時点別の変化量 (OCT-002 試験) (4/8)

| Total Bilirubin (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-------------------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit                   | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2                  | ATL-962 120 mg | 504                | -0.03 | 0.211 | -1.2 | -0.20 | 0.00   | 0.10 | 1.0 |
| Week 4                  | ATL-962 120 mg | 495                | -0.05 | 0.226 | -1.4 | -0.10 | -0.10  | 0.10 | 0.9 |
| Week 8                  | ATL-962 120 mg | 489                | -0.04 | 0.205 | -1.3 | -0.10 | 0.00   | 0.10 | 0.5 |
| Week 12                 | ATL-962 120 mg | 479                | -0.05 | 0.212 | -1.1 | -0.20 | -0.10  | 0.00 | 1.3 |
| Week 16                 | ATL-962 120 mg | 472                | -0.05 | 0.205 | -0.9 | -0.10 | 0.00   | 0.10 | 0.8 |
| Week 20                 | ATL-962 120 mg | 469                | -0.05 | 0.217 | -1.3 | -0.20 | -0.10  | 0.10 | 1.0 |
| Week 24                 | ATL-962 120 mg | 462                | -0.05 | 0.226 | -1.2 | -0.20 | -0.10  | 0.10 | 1.0 |
| Week 28                 | ATL-962 120 mg | 453                | -0.05 | 0.224 | -1.7 | -0.20 | 0.00   | 0.10 | 1.0 |
| Week 32                 | ATL-962 120 mg | 456                | -0.04 | 0.234 | -1.2 | -0.10 | 0.00   | 0.10 | 1.1 |
| Week 36                 | ATL-962 120 mg | 452                | -0.03 | 0.209 | -1.3 | -0.10 | 0.00   | 0.10 | 1.2 |
| Week 40                 | ATL-962 120 mg | 445                | -0.03 | 0.229 | -1.6 | -0.10 | 0.00   | 0.10 | 1.0 |
| Week 44                 | ATL-962 120 mg | 442                | -0.01 | 0.217 | -1.5 | -0.10 | 0.00   | 0.10 | 0.8 |
| Week 48                 | ATL-962 120 mg | 439                | -0.01 | 0.225 | -1.4 | -0.10 | 0.00   | 0.10 | 0.9 |
| Week 52                 | ATL-962 120 mg | 439                | -0.02 | 0.209 | -0.9 | -0.10 | 0.00   | 0.10 | 0.6 |

| ChE (U/L) |                | Summary Statistics |       |       |      |       |        |      |     |
|-----------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit     | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2    | ATL-962 120 mg | 504                | -8.7  | 21.37 | -126 | -21.0 | -9.0   | 5.0  | 68  |
| Week 4    | ATL-962 120 mg | 495                | -12.1 | 23.71 | -91  | -27.0 | -11.0  | 4.0  | 59  |
| Week 8    | ATL-962 120 mg | 489                | -9.7  | 28.44 | -144 | -26.0 | -7.0   | 8.0  | 67  |
| Week 12   | ATL-962 120 mg | 479                | -8.8  | 28.95 | -171 | -24.0 | -7.0   | 9.0  | 82  |
| Week 16   | ATL-962 120 mg | 472                | -3.0  | 29.22 | -131 | -18.0 | -3.0   | 15.0 | 90  |
| Week 20   | ATL-962 120 mg | 469                | -4.0  | 29.27 | -182 | -21.0 | -3.0   | 14.0 | 86  |
| Week 24   | ATL-962 120 mg | 462                | -6.8  | 29.77 | -179 | -22.0 | -7.0   | 11.0 | 99  |
| Week 28   | ATL-962 120 mg | 453                | -8.0  | 31.82 | -182 | -25.0 | -7.0   | 10.0 | 104 |
| Week 32   | ATL-962 120 mg | 456                | -7.6  | 33.01 | -139 | -25.0 | -7.5   | 11.0 | 122 |
| Week 36   | ATL-962 120 mg | 452                | -10.9 | 33.51 | -223 | -29.0 | -10.0  | 8.0  | 112 |
| Week 40   | ATL-962 120 mg | 445                | -11.1 | 31.53 | -210 | -29.0 | -10.0  | 7.0  | 122 |
| Week 44   | ATL-962 120 mg | 442                | -10.5 | 33.06 | -212 | -28.0 | -9.0   | 9.0  | 87  |
| Week 48   | ATL-962 120 mg | 439                | -10.4 | 35.77 | -159 | -26.0 | -8.0   | 8.0  | 116 |
| Week 52   | ATL-962 120 mg | 439                | -10.0 | 35.55 | -161 | -28.0 | -8.0   | 12.0 | 102 |

| Creatinine (mg/dL) |                | Summary Statistics |        |        |       |        |        |       |      |
|--------------------|----------------|--------------------|--------|--------|-------|--------|--------|-------|------|
| Visit              | Treatment      | N                  | Mean   | SD     | Min   | Q1     | Median | Q3    | Max  |
| Week 2             | ATL-962 120 mg | 504                | 0.002  | 0.0667 | -0.50 | -0.030 | 0.000  | 0.040 | 0.26 |
| Week 4             | ATL-962 120 mg | 495                | -0.004 | 0.0702 | -0.56 | -0.040 | 0.000  | 0.030 | 0.66 |
| Week 8             | ATL-962 120 mg | 489                | -0.009 | 0.0701 | -0.54 | -0.040 | -0.010 | 0.030 | 0.22 |
| Week 12            | ATL-962 120 mg | 479                | -0.019 | 0.0691 | -0.49 | -0.050 | -0.020 | 0.020 | 0.19 |
| Week 16            | ATL-962 120 mg | 472                | -0.020 | 0.0846 | -0.53 | -0.060 | -0.020 | 0.020 | 0.73 |
| Week 20            | ATL-962 120 mg | 469                | -0.023 | 0.0784 | -0.57 | -0.060 | -0.020 | 0.020 | 0.45 |
| Week 24            | ATL-962 120 mg | 462                | -0.021 | 0.0745 | -0.52 | -0.060 | -0.020 | 0.020 | 0.45 |
| Week 28            | ATL-962 120 mg | 453                | -0.019 | 0.0754 | -0.42 | -0.060 | -0.010 | 0.020 | 0.29 |
| Week 32            | ATL-962 120 mg | 456                | -0.004 | 0.0946 | -0.44 | -0.050 | -0.010 | 0.040 | 0.80 |
| Week 36            | ATL-962 120 mg | 452                | -0.001 | 0.0867 | -0.44 | -0.050 | 0.000  | 0.040 | 0.42 |
| Week 40            | ATL-962 120 mg | 445                | 0.002  | 0.0897 | -0.38 | -0.050 | 0.000  | 0.040 | 0.57 |
| Week 44            | ATL-962 120 mg | 442                | 0.006  | 0.0899 | -0.38 | -0.040 | 0.000  | 0.050 | 0.68 |
| Week 48            | ATL-962 120 mg | 439                | 0.005  | 0.0944 | -0.38 | -0.040 | 0.000  | 0.050 | 0.80 |
| Week 52            | ATL-962 120 mg | 439                | 0.003  | 0.0872 | -0.42 | -0.040 | 0.000  | 0.040 | 0.67 |

表 2.7.4.7-付録-32 血液生化学検査の評価時点別の変化量 (OCT-002 試験) (5/8)

| BUN (mg/dL) |                | Summary Statistics |      |      |     |      |        |     |     |  |
|-------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|--|
| Visit       | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |  |
| Week 2      | ATL-962 120 mg | 504                | -0.3 | 3.31 | -15 | -2.0 | 0.0    | 2.0 | 19  |  |
| Week 4      | ATL-962 120 mg | 495                | -0.3 | 3.36 | -23 | -2.0 | 0.0    | 1.0 | 18  |  |
| Week 8      | ATL-962 120 mg | 489                | -0.4 | 3.18 | -12 | -2.0 | 0.0    | 1.0 | 12  |  |
| Week 12     | ATL-962 120 mg | 479                | -0.4 | 3.47 | -18 | -2.0 | 0.0    | 2.0 | 12  |  |
| Week 16     | ATL-962 120 mg | 472                | -0.4 | 3.46 | -20 | -2.0 | 0.0    | 2.0 | 9   |  |
| Week 20     | ATL-962 120 mg | 469                | -0.5 | 3.35 | -17 | -3.0 | -1.0   | 2.0 | 9   |  |
| Week 24     | ATL-962 120 mg | 462                | -0.3 | 3.60 | -21 | -2.0 | 0.0    | 2.0 | 23  |  |
| Week 28     | ATL-962 120 mg | 453                | -0.2 | 3.14 | -10 | -2.0 | 0.0    | 2.0 | 12  |  |
| Week 32     | ATL-962 120 mg | 456                | -0.2 | 3.46 | -19 | -2.0 | 0.0    | 2.0 | 11  |  |
| Week 36     | ATL-962 120 mg | 452                | 0.0  | 3.55 | -15 | -2.0 | 0.0    | 2.0 | 15  |  |
| Week 40     | ATL-962 120 mg | 445                | 0.0  | 3.38 | -17 | -2.0 | 0.0    | 2.0 | 15  |  |
| Week 44     | ATL-962 120 mg | 442                | 0.2  | 3.48 | -15 | -2.0 | 0.0    | 2.0 | 14  |  |
| Week 48     | ATL-962 120 mg | 439                | -0.1 | 3.41 | -17 | -2.0 | 0.0    | 2.0 | 12  |  |
| Week 52     | ATL-962 120 mg | 439                | 0.0  | 3.26 | -11 | -2.0 | 0.0    | 2.0 | 16  |  |

  

| Na (mEq/L) |                | Summary Statistics |      |      |     |      |        |     |     |  |
|------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|--|
| Visit      | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |  |
| Week 2     | ATL-962 120 mg | 504                | 0.2  | 1.77 | -5  | -1.0 | 0.0    | 1.0 | 6   |  |
| Week 4     | ATL-962 120 mg | 495                | 0.2  | 1.79 | -8  | -1.0 | 0.0    | 1.0 | 6   |  |
| Week 8     | ATL-962 120 mg | 489                | 0.1  | 1.70 | -6  | -1.0 | 0.0    | 1.0 | 6   |  |
| Week 12    | ATL-962 120 mg | 479                | 0.2  | 1.90 | -5  | -1.0 | 0.0    | 1.0 | 5   |  |
| Week 16    | ATL-962 120 mg | 472                | 0.0  | 1.88 | -8  | -1.0 | 0.0    | 1.0 | 7   |  |
| Week 20    | ATL-962 120 mg | 469                | 0.2  | 1.89 | -6  | -1.0 | 0.0    | 1.0 | 6   |  |
| Week 24    | ATL-962 120 mg | 462                | 0.4  | 1.90 | -9  | -1.0 | 0.0    | 2.0 | 6   |  |
| Week 28    | ATL-962 120 mg | 453                | 0.3  | 1.84 | -7  | -1.0 | 0.0    | 1.0 | 6   |  |
| Week 32    | ATL-962 120 mg | 456                | 0.3  | 1.92 | -6  | -1.0 | 0.0    | 2.0 | 7   |  |
| Week 36    | ATL-962 120 mg | 452                | 0.5  | 1.85 | -7  | -1.0 | 1.0    | 2.0 | 6   |  |
| Week 40    | ATL-962 120 mg | 445                | 0.6  | 1.93 | -6  | -1.0 | 1.0    | 2.0 | 6   |  |
| Week 44    | ATL-962 120 mg | 442                | 0.6  | 1.90 | -6  | -1.0 | 1.0    | 2.0 | 6   |  |
| Week 48    | ATL-962 120 mg | 439                | 0.6  | 1.81 | -8  | 0.0  | 1.0    | 2.0 | 7   |  |
| Week 52    | ATL-962 120 mg | 439                | 0.7  | 1.85 | -7  | -1.0 | 1.0    | 2.0 | 7   |  |

  

| K (mEq/L) |                | Summary Statistics |       |       |      |       |        |      |     |  |
|-----------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|--|
| Visit     | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |  |
| Week 2    | ATL-962 120 mg | 504                | -0.02 | 0.283 | -1.1 | -0.20 | 0.00   | 0.20 | 1.1 |  |
| Week 4    | ATL-962 120 mg | 495                | -0.01 | 0.280 | -1.1 | -0.20 | 0.00   | 0.20 | 1.1 |  |
| Week 8    | ATL-962 120 mg | 489                | 0.00  | 0.308 | -2.1 | -0.20 | 0.00   | 0.20 | 1.3 |  |
| Week 12   | ATL-962 120 mg | 479                | 0.00  | 0.343 | -3.0 | -0.20 | 0.00   | 0.20 | 1.3 |  |
| Week 16   | ATL-962 120 mg | 472                | 0.04  | 0.351 | -3.1 | -0.20 | 0.00   | 0.20 | 1.2 |  |
| Week 20   | ATL-962 120 mg | 469                | 0.04  | 0.317 | -1.4 | -0.10 | 0.00   | 0.20 | 1.1 |  |
| Week 24   | ATL-962 120 mg | 462                | 0.03  | 0.329 | -1.2 | -0.20 | 0.00   | 0.20 | 1.7 |  |
| Week 28   | ATL-962 120 mg | 453                | 0.03  | 0.314 | -0.8 | -0.20 | 0.00   | 0.20 | 1.2 |  |
| Week 32   | ATL-962 120 mg | 456                | 0.04  | 0.310 | -1.1 | -0.20 | 0.00   | 0.20 | 1.3 |  |
| Week 36   | ATL-962 120 mg | 452                | 0.02  | 0.314 | -0.9 | -0.20 | 0.00   | 0.20 | 1.2 |  |
| Week 40   | ATL-962 120 mg | 445                | 0.00  | 0.301 | -1.0 | -0.20 | 0.00   | 0.20 | 1.5 |  |
| Week 44   | ATL-962 120 mg | 442                | -0.01 | 0.317 | -1.0 | -0.20 | 0.00   | 0.20 | 1.2 |  |
| Week 48   | ATL-962 120 mg | 439                | -0.03 | 0.307 | -1.2 | -0.20 | 0.00   | 0.10 | 0.9 |  |
| Week 52   | ATL-962 120 mg | 439                | -0.02 | 0.277 | -0.9 | -0.20 | 0.00   | 0.10 | 1.3 |  |

表 2.7.4.7-付録-32 血液生化学検査の評価時点別の変化量 (OCT-002 試験) (6/8)

| Cl (mEq/L) |                | Summary Statistics |      |      |     |      |        |     |     |
|------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit      | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2     | ATL-962 120 mg | 504                | 0.2  | 1.83 | -7  | -1.0 | 0.0    | 1.0 | 6   |
| Week 4     | ATL-962 120 mg | 495                | 0.0  | 2.00 | -11 | -1.0 | 0.0    | 1.0 | 6   |
| Week 8     | ATL-962 120 mg | 489                | -0.1 | 1.83 | -6  | -1.0 | 0.0    | 1.0 | 6   |
| Week 12    | ATL-962 120 mg | 479                | -0.1 | 1.99 | -6  | -1.0 | 0.0    | 1.0 | 5   |
| Week 16    | ATL-962 120 mg | 472                | -0.4 | 2.12 | -9  | -2.0 | 0.0    | 1.0 | 6   |
| Week 20    | ATL-962 120 mg | 469                | -0.2 | 2.16 | -6  | -2.0 | 0.0    | 1.0 | 7   |
| Week 24    | ATL-962 120 mg | 462                | 0.0  | 2.26 | -11 | -1.0 | 0.0    | 2.0 | 6   |
| Week 28    | ATL-962 120 mg | 453                | 0.1  | 2.14 | -7  | -1.0 | 0.0    | 1.0 | 7   |
| Week 32    | ATL-962 120 mg | 456                | 0.2  | 2.15 | -8  | -1.0 | 0.0    | 2.0 | 6   |
| Week 36    | ATL-962 120 mg | 452                | 0.4  | 2.17 | -6  | -1.0 | 0.0    | 2.0 | 6   |
| Week 40    | ATL-962 120 mg | 445                | 0.7  | 2.19 | -5  | -1.0 | 1.0    | 2.0 | 7   |
| Week 44    | ATL-962 120 mg | 442                | 0.7  | 2.23 | -6  | -1.0 | 1.0    | 2.0 | 8   |
| Week 48    | ATL-962 120 mg | 439                | 0.6  | 1.99 | -6  | -1.0 | 1.0    | 2.0 | 7   |
| Week 52    | ATL-962 120 mg | 439                | 0.6  | 2.02 | -6  | -1.0 | 1.0    | 2.0 | 7   |

| Ca (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|------------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit      | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2     | ATL-962 120 mg | 504                | -0.04 | 0.277 | -0.9 | -0.20 | 0.00   | 0.10 | 1.0 |
| Week 4     | ATL-962 120 mg | 495                | -0.05 | 0.280 | -1.3 | -0.20 | 0.00   | 0.10 | 0.6 |
| Week 8     | ATL-962 120 mg | 489                | -0.02 | 0.287 | -1.2 | -0.20 | 0.00   | 0.10 | 1.1 |
| Week 12    | ATL-962 120 mg | 479                | -0.03 | 0.292 | -1.1 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 16    | ATL-962 120 mg | 472                | -0.01 | 0.298 | -1.4 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 20    | ATL-962 120 mg | 469                | -0.01 | 0.295 | -1.2 | -0.20 | 0.00   | 0.20 | 1.1 |
| Week 24    | ATL-962 120 mg | 462                | -0.04 | 0.311 | -1.4 | -0.30 | 0.00   | 0.20 | 1.0 |
| Week 28    | ATL-962 120 mg | 453                | -0.06 | 0.304 | -1.3 | -0.30 | -0.10  | 0.20 | 0.9 |
| Week 32    | ATL-962 120 mg | 456                | -0.04 | 0.310 | -1.9 | -0.20 | 0.00   | 0.20 | 0.7 |
| Week 36    | ATL-962 120 mg | 452                | -0.06 | 0.323 | -1.2 | -0.30 | 0.00   | 0.20 | 1.0 |
| Week 40    | ATL-962 120 mg | 445                | -0.06 | 0.303 | -1.0 | -0.20 | 0.00   | 0.10 | 0.8 |
| Week 44    | ATL-962 120 mg | 442                | -0.04 | 0.322 | -1.3 | -0.30 | 0.00   | 0.10 | 1.1 |
| Week 48    | ATL-962 120 mg | 439                | -0.06 | 0.310 | -1.3 | -0.30 | -0.10  | 0.10 | 1.6 |
| Week 52    | ATL-962 120 mg | 439                | -0.05 | 0.301 | -1.4 | -0.20 | -0.10  | 0.10 | 0.9 |

| P (mg/dL) |                | Summary Statistics |       |       |      |       |        |      |     |
|-----------|----------------|--------------------|-------|-------|------|-------|--------|------|-----|
| Visit     | Treatment      | N                  | Mean  | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2    | ATL-962 120 mg | 504                | 0.02  | 0.370 | -1.3 | -0.20 | 0.00   | 0.30 | 1.2 |
| Week 4    | ATL-962 120 mg | 495                | 0.01  | 0.367 | -2.1 | -0.20 | 0.00   | 0.20 | 1.2 |
| Week 8    | ATL-962 120 mg | 489                | 0.01  | 0.398 | -1.7 | -0.20 | 0.00   | 0.30 | 1.3 |
| Week 12   | ATL-962 120 mg | 479                | 0.02  | 0.413 | -2.5 | -0.20 | 0.00   | 0.30 | 1.3 |
| Week 16   | ATL-962 120 mg | 472                | 0.03  | 0.421 | -1.5 | -0.20 | 0.00   | 0.30 | 1.5 |
| Week 20   | ATL-962 120 mg | 469                | -0.02 | 0.393 | -1.4 | -0.30 | 0.00   | 0.20 | 1.5 |
| Week 24   | ATL-962 120 mg | 462                | 0.01  | 0.425 | -2.0 | -0.30 | 0.00   | 0.30 | 1.5 |
| Week 28   | ATL-962 120 mg | 453                | 0.00  | 0.392 | -1.8 | -0.20 | 0.00   | 0.30 | 1.6 |
| Week 32   | ATL-962 120 mg | 456                | 0.02  | 0.411 | -1.9 | -0.20 | 0.00   | 0.30 | 1.3 |
| Week 36   | ATL-962 120 mg | 452                | -0.02 | 0.416 | -1.9 | -0.30 | 0.00   | 0.30 | 1.1 |
| Week 40   | ATL-962 120 mg | 445                | -0.01 | 0.400 | -1.4 | -0.30 | 0.00   | 0.20 | 1.1 |
| Week 44   | ATL-962 120 mg | 442                | 0.01  | 0.423 | -1.7 | -0.20 | 0.00   | 0.30 | 1.3 |
| Week 48   | ATL-962 120 mg | 439                | 0.01  | 0.399 | -1.6 | -0.20 | 0.00   | 0.30 | 1.1 |
| Week 52   | ATL-962 120 mg | 439                | 0.01  | 0.414 | -1.7 | -0.20 | 0.00   | 0.30 | 1.8 |

表 2.7.4.7-付録-32 血液生化学検査の評価時点別の変化量 (OCT-002 試験) (7/8)

| Vitamin A (ng/mL) |                | Summary Statistics |       |        |      |        |        |      |     |
|-------------------|----------------|--------------------|-------|--------|------|--------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean  | SD     | Min  | Q1     | Median | Q3   | Max |
| Week 2            | ATL-962 120 mg | 504                | -18.5 | 99.69  | -600 | -75.5  | -14.0  | 36.0 | 896 |
| Week 4            | ATL-962 120 mg | 495                | -20.9 | 92.26  | -674 | -66.0  | -17.0  | 30.0 | 451 |
| Week 8            | ATL-962 120 mg | 489                | -28.2 | 100.33 | -756 | -78.0  | -23.0  | 21.0 | 626 |
| Week 12           | ATL-962 120 mg | 479                | -27.2 | 101.59 | -645 | -80.0  | -22.0  | 22.0 | 772 |
| Week 16           | ATL-962 120 mg | 472                | -32.6 | 111.25 | -745 | -88.0  | -31.5  | 22.0 | 688 |
| Week 20           | ATL-962 120 mg | 469                | -33.4 | 102.16 | -690 | -81.0  | -33.0  | 19.0 | 604 |
| Week 24           | ATL-962 120 mg | 462                | -42.1 | 110.03 | -696 | -94.0  | -46.0  | 14.0 | 559 |
| Week 28           | ATL-962 120 mg | 452                | -48.8 | 109.93 | -659 | -107.0 | -45.0  | 9.0  | 775 |
| Week 32           | ATL-962 120 mg | 456                | -44.3 | 116.87 | -605 | -105.5 | -39.5  | 21.5 | 620 |
| Week 36           | ATL-962 120 mg | 452                | -43.0 | 109.66 | -627 | -98.0  | -33.0  | 13.0 | 690 |
| Week 40           | ATL-962 120 mg | 445                | -42.8 | 112.49 | -640 | -104.0 | -40.0  | 17.0 | 683 |
| Week 44           | ATL-962 120 mg | 442                | -25.3 | 110.07 | -568 | -81.0  | -21.5  | 37.0 | 679 |
| Week 48           | ATL-962 120 mg | 439                | -18.4 | 110.76 | -571 | -77.0  | -17.0  | 44.0 | 669 |
| Week 52           | ATL-962 120 mg | 438                | -13.7 | 110.32 | -656 | -67.0  | -11.0  | 48.0 | 832 |

| Vitamin D (pg/mL) |                | Summary Statistics |      |       |      |       |        |      |     |
|-------------------|----------------|--------------------|------|-------|------|-------|--------|------|-----|
| Visit             | Treatment      | N                  | Mean | SD    | Min  | Q1    | Median | Q3   | Max |
| Week 2            | ATL-962 120 mg | 504                | 0.2  | 25.38 | -96  | -14.0 | 1.0    | 15.0 | 82  |
| Week 4            | ATL-962 120 mg | 495                | -3.3 | 26.00 | -99  | -18.0 | -2.0   | 12.0 | 83  |
| Week 8            | ATL-962 120 mg | 489                | -2.6 | 23.53 | -94  | -15.0 | -2.0   | 10.0 | 83  |
| Week 12           | ATL-962 120 mg | 479                | -3.4 | 23.54 | -94  | -16.0 | -3.0   | 9.0  | 75  |
| Week 16           | ATL-962 120 mg | 472                | -3.4 | 22.95 | -88  | -16.0 | -1.0   | 11.0 | 65  |
| Week 20           | ATL-962 120 mg | 469                | -4.6 | 23.15 | -94  | -16.0 | -2.0   | 7.0  | 78  |
| Week 24           | ATL-962 120 mg | 462                | -3.6 | 24.75 | -87  | -17.0 | -2.0   | 11.0 | 111 |
| Week 28           | ATL-962 120 mg | 453                | -4.4 | 23.76 | -99  | -16.0 | -2.0   | 9.0  | 62  |
| Week 32           | ATL-962 120 mg | 456                | -5.1 | 24.39 | -98  | -17.0 | -3.0   | 9.5  | 73  |
| Week 36           | ATL-962 120 mg | 452                | -3.9 | 26.42 | -112 | -17.0 | -3.0   | 12.5 | 103 |
| Week 40           | ATL-962 120 mg | 445                | -5.2 | 25.89 | -99  | -17.0 | -4.0   | 11.0 | 131 |
| Week 44           | ATL-962 120 mg | 442                | -3.4 | 25.25 | -94  | -17.0 | -2.0   | 10.0 | 80  |
| Week 48           | ATL-962 120 mg | 439                | -2.0 | 24.56 | -79  | -15.0 | -1.0   | 13.0 | 97  |
| Week 52           | ATL-962 120 mg | 439                | -0.5 | 24.91 | -93  | -12.0 | 0.0    | 14.0 | 91  |

| Vitamin E (mg/dL) |                | Summary Statistics |        |        |       |        |        |        |      |
|-------------------|----------------|--------------------|--------|--------|-------|--------|--------|--------|------|
| Visit             | Treatment      | N                  | Mean   | SD     | Min   | Q1     | Median | Q3     | Max  |
| Week 2            | ATL-962 120 mg | 504                | -0.186 | 0.2576 | -1.79 | -0.300 | -0.160 | -0.035 | 0.71 |
| Week 4            | ATL-962 120 mg | 495                | -0.194 | 0.2640 | -1.85 | -0.310 | -0.170 | -0.040 | 0.93 |
| Week 8            | ATL-962 120 mg | 489                | -0.217 | 0.2818 | -2.46 | -0.350 | -0.180 | -0.060 | 0.57 |
| Week 12           | ATL-962 120 mg | 479                | -0.224 | 0.2957 | -2.78 | -0.350 | -0.210 | -0.070 | 0.65 |
| Week 16           | ATL-962 120 mg | 472                | -0.214 | 0.2802 | -2.56 | -0.340 | -0.200 | -0.060 | 0.74 |
| Week 20           | ATL-962 120 mg | 469                | -0.222 | 0.2788 | -2.39 | -0.350 | -0.210 | -0.070 | 0.92 |
| Week 24           | ATL-962 120 mg | 462                | -0.213 | 0.2737 | -2.14 | -0.320 | -0.190 | -0.060 | 0.96 |
| Week 28           | ATL-962 120 mg | 453                | -0.209 | 0.2776 | -1.75 | -0.330 | -0.200 | -0.050 | 1.52 |
| Week 32           | ATL-962 120 mg | 456                | -0.224 | 0.2601 | -2.03 | -0.350 | -0.200 | -0.070 | 0.55 |
| Week 36           | ATL-962 120 mg | 452                | -0.241 | 0.2605 | -1.93 | -0.365 | -0.220 | -0.080 | 0.50 |
| Week 40           | ATL-962 120 mg | 445                | -0.241 | 0.2737 | -2.39 | -0.360 | -0.230 | -0.080 | 0.64 |
| Week 44           | ATL-962 120 mg | 442                | -0.257 | 0.2869 | -2.01 | -0.360 | -0.230 | -0.100 | 0.75 |
| Week 48           | ATL-962 120 mg | 439                | -0.257 | 0.2724 | -2.00 | -0.370 | -0.230 | -0.110 | 0.49 |
| Week 52           | ATL-962 120 mg | 438                | -0.271 | 0.2927 | -2.03 | -0.410 | -0.250 | -0.110 | 1.03 |

表 2.7.4.7-付録-32 血液生化学検査の評価時点別の変化量 (OCT-002 試験) (8/8)

| β-Carotene (μg/dL) |                | Summary Statistics |       |        |        |       |        |      |      |
|--------------------|----------------|--------------------|-------|--------|--------|-------|--------|------|------|
| Visit              | Treatment      | N                  | Mean  | SD     | Min    | Q1    | Median | Q3   | Max  |
| Week 2             | ATL-962 120 mg | 504                | -2.31 | 10.443 | -108.9 | -4.55 | -1.20  | 1.00 | 45.7 |
| Week 4             | ATL-962 120 mg | 495                | -2.98 | 11.499 | -110.6 | -5.60 | -1.30  | 1.60 | 35.8 |
| Week 8             | ATL-962 120 mg | 489                | -3.24 | 14.231 | -120.9 | -5.60 | -1.00  | 2.10 | 56.4 |
| Week 12            | ATL-962 120 mg | 479                | -3.90 | 15.705 | -149.9 | -6.10 | -1.70  | 2.20 | 72.7 |
| Week 16            | ATL-962 120 mg | 472                | -3.59 | 16.387 | -174.1 | -5.65 | -0.95  | 1.90 | 70.0 |
| Week 20            | ATL-962 120 mg | 469                | -2.84 | 15.113 | -124.2 | -6.30 | -0.20  | 3.40 | 79.0 |
| Week 24            | ATL-962 120 mg | 462                | -2.79 | 15.847 | -123.6 | -6.10 | -0.55  | 3.40 | 78.3 |
| Week 28            | ATL-962 120 mg | 453                | -2.14 | 16.641 | -133.7 | -5.30 | -0.30  | 3.30 | 76.0 |
| Week 32            | ATL-962 120 mg | 456                | -1.24 | 16.041 | -118.4 | -4.60 | -0.20  | 4.00 | 66.1 |
| Week 36            | ATL-962 120 mg | 452                | -1.38 | 16.181 | -122.6 | -5.25 | 0.10   | 3.95 | 55.6 |
| Week 40            | ATL-962 120 mg | 445                | -1.17 | 17.390 | -155.2 | -4.20 | 0.20   | 3.70 | 67.3 |
| Week 44            | ATL-962 120 mg | 442                | -0.83 | 17.551 | -153.8 | -4.20 | 0.40   | 4.80 | 68.1 |
| Week 48            | ATL-962 120 mg | 439                | -0.70 | 16.462 | -162.2 | -4.20 | 0.40   | 4.70 | 58.0 |
| Week 52            | ATL-962 120 mg | 439                | -0.67 | 16.091 | -145.9 | -4.00 | 0.50   | 4.10 | 52.1 |

| m-AST (IU/L) |                | Summary Statistics |      |      |     |      |        |     |     |
|--------------|----------------|--------------------|------|------|-----|------|--------|-----|-----|
| Visit        | Treatment      | N                  | Mean | SD   | Min | Q1   | Median | Q3  | Max |
| Week 2       | ATL-962 120 mg | 504                | 0.3  | 2.05 | -7  | -1.0 | 0.0    | 1.0 | 18  |
| Week 4       | ATL-962 120 mg | 495                | 0.2  | 3.79 | -10 | -1.0 | 0.0    | 1.0 | 70  |
| Week 8       | ATL-962 120 mg | 489                | 0.1  | 2.17 | -9  | -1.0 | 0.0    | 1.0 | 25  |
| Week 12      | ATL-962 120 mg | 478                | 0.0  | 2.48 | -13 | -1.0 | 0.0    | 1.0 | 37  |
| Week 16      | ATL-962 120 mg | 472                | -0.1 | 1.78 | -9  | -1.0 | 0.0    | 1.0 | 7   |
| Week 20      | ATL-962 120 mg | 469                | -0.2 | 1.90 | -12 | -1.0 | 0.0    | 1.0 | 8   |
| Week 24      | ATL-962 120 mg | 462                | -0.2 | 2.30 | -14 | -1.0 | 0.0    | 1.0 | 23  |
| Week 28      | ATL-962 120 mg | 453                | -0.2 | 2.47 | -10 | -1.0 | 0.0    | 1.0 | 20  |
| Week 32      | ATL-962 120 mg | 456                | -0.1 | 1.98 | -10 | -1.0 | 0.0    | 1.0 | 10  |
| Week 36      | ATL-962 120 mg | 452                | 0.1  | 5.12 | -10 | -1.0 | 0.0    | 1.0 | 99  |
| Week 40      | ATL-962 120 mg | 445                | 0.1  | 2.10 | -10 | -1.0 | 0.0    | 1.0 | 11  |
| Week 44      | ATL-962 120 mg | 442                | 0.3  | 2.34 | -12 | -1.0 | 0.0    | 1.0 | 19  |
| Week 48      | ATL-962 120 mg | 439                | 0.6  | 2.34 | -10 | 0.0  | 1.0    | 2.0 | 15  |
| Week 52      | ATL-962 120 mg | 439                | 0.4  | 2.13 | -12 | -1.0 | 0.0    | 2.0 | 9   |

| CK18 (Segment) (U/L) |                | Summary Statistics |        |         |        |        |        |       |       |
|----------------------|----------------|--------------------|--------|---------|--------|--------|--------|-------|-------|
| Visit                | Treatment      | N                  | Mean   | SD      | Min    | Q1     | Median | Q3    | Max   |
| Week 2               | ATL-962 120 mg | 504                | -4.67  | 116.782 | -616.1 | -31.85 | 0.00   | 38.65 | 453.1 |
| Week 4               | ATL-962 120 mg | 495                | -5.17  | 118.527 | -679.5 | -43.30 | 0.00   | 40.20 | 479.7 |
| Week 8               | ATL-962 120 mg | 489                | -13.23 | 135.217 | -739.5 | -45.60 | 0.00   | 39.90 | 611.4 |
| Week 12              | ATL-962 120 mg | 479                | -17.40 | 141.821 | -823.1 | -54.20 | 0.00   | 35.10 | 695.5 |
| Week 16              | ATL-962 120 mg | 472                | -30.12 | 138.536 | -710.0 | -64.15 | -0.80  | 26.65 | 613.4 |
| Week 20              | ATL-962 120 mg | 469                | -35.19 | 156.444 | -749.5 | -79.00 | -11.40 | 24.40 | 695.5 |
| Week 24              | ATL-962 120 mg | 462                | -42.49 | 158.317 | -861.6 | -76.80 | -13.40 | 14.40 | 701.8 |
| Week 28              | ATL-962 120 mg | 453                | -36.68 | 160.438 | -712.7 | -80.20 | -15.50 | 20.30 | 831.4 |
| Week 32              | ATL-962 120 mg | 456                | -35.47 | 164.400 | -792.3 | -84.50 | -1.80  | 30.40 | 701.8 |
| Week 36              | ATL-962 120 mg | 452                | -34.65 | 163.278 | -804.4 | -88.70 | 0.00   | 40.60 | 831.4 |
| Week 40              | ATL-962 120 mg | 445                | -35.70 | 155.436 | -807.5 | -75.60 | -0.40  | 37.90 | 695.5 |
| Week 44              | ATL-962 120 mg | 442                | -25.38 | 160.985 | -805.5 | -74.80 | 0.00   | 42.90 | 701.8 |
| Week 48              | ATL-962 120 mg | 439                | -17.65 | 164.887 | -812.0 | -63.70 | 0.00   | 43.90 | 752.0 |
| Week 52              | ATL-962 120 mg | 439                | -17.24 | 170.984 | -861.6 | -69.70 | -1.40  | 44.50 | 761.1 |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験)(1/24)

|               |         |        | ATL-962 120 mg |        |       |       |
|---------------|---------|--------|----------------|--------|-------|-------|
|               |         |        | Week 0         |        |       |       |
|               |         |        | Lower          | Normal | Upper | Total |
| Total Protein | Week 2  | Lower  | 5              | 6      | -     | 11    |
|               |         | Normal | 12             | 470    | 3     | 485   |
|               |         | Upper  | -              | 4      | 4     | 8     |
|               |         | Total  | 17             | 480    | 7     | 504   |
|               | Week 4  | Lower  | 7              | 9      | -     | 16    |
|               |         | Normal | 10             | 459    | 2     | 471   |
|               |         | Upper  | -              | 4      | 4     | 8     |
|               |         | Total  | 17             | 472    | 6     | 495   |
|               | Week 8  | Lower  | 5              | 3      | -     | 8     |
|               |         | Normal | 11             | 460    | 3     | 474   |
|               |         | Upper  | -              | 4      | 3     | 7     |
|               |         | Total  | 16             | 467    | 6     | 489   |
|               | Week 12 | Lower  | 2              | 11     | -     | 13    |
|               |         | Normal | 14             | 445    | 4     | 463   |
|               |         | Upper  | -              | 1      | 2     | 3     |
|               |         | Total  | 16             | 457    | 6     | 479   |
|               | Week 16 | Lower  | 2              | 9      | -     | 11    |
|               |         | Normal | 14             | 432    | 2     | 448   |
|               |         | Upper  | -              | 9      | 4     | 13    |
|               |         | Total  | 16             | 450    | 6     | 472   |
|               | Week 20 | Lower  | 5              | 9      | -     | 14    |
|               |         | Normal | 11             | 436    | 3     | 450   |
|               |         | Upper  | -              | 2      | 3     | 5     |
|               |         | Total  | 16             | 447    | 6     | 469   |
|               | Week 24 | Lower  | 3              | 14     | -     | 17    |
|               |         | Normal | 13             | 420    | 4     | 437   |
|               |         | Upper  | -              | 6      | 2     | 8     |
|               |         | Total  | 16             | 440    | 6     | 462   |
|               | Week 28 | Lower  | 5              | 9      | -     | 14    |
|               |         | Normal | 11             | 419    | 4     | 434   |
|               |         | Upper  | -              | 3      | 2     | 5     |
|               |         | Total  | 16             | 431    | 6     | 453   |
|               | Week 32 | Lower  | 7              | 12     | -     | 19    |
|               |         | Normal | 9              | 418    | 3     | 430   |
|               |         | Upper  | -              | 4      | 3     | 7     |
|               |         | Total  | 16             | 434    | 6     | 456   |
|               | Week 36 | Lower  | 3              | 13     | -     | 16    |
|               |         | Normal | 12             | 412    | 2     | 426   |
|               |         | Upper  | -              | 6      | 4     | 10    |
|               |         | Total  | 15             | 431    | 6     | 452   |
|               | Week 40 | Lower  | 4              | 15     | -     | 19    |
|               |         | Normal | 10             | 407    | 1     | 418   |
|               |         | Upper  | -              | 3      | 5     | 8     |
|               |         | Total  | 14             | 425    | 6     | 445   |
|               | Week 44 | Lower  | 3              | 11     | -     | 14    |
|               |         | Normal | 10             | 410    | 3     | 423   |
|               |         | Upper  | -              | 2      | 3     | 5     |
|               |         | Total  | 13             | 423    | 6     | 442   |
|               | Week 48 | Lower  | 4              | 11     | -     | 15    |
|               |         | Normal | 9              | 404    | 4     | 417   |
|               |         | Upper  | -              | 5      | 2     | 7     |
|               |         | Total  | 13             | 420    | 6     | 439   |
| Week 52       | Lower   | 1      | 11             | -      | 12    |       |
|               | Normal  | 12     | 404            | 4      | 420   |       |
|               | Upper   | -      | 5              | 2      | 7     |       |
|               | Total   | 13     | 420            | 6      | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験) (2/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| Albumin | Week 2  | Lower  | 1              | 3      | -     | 4     |
|         |         | Normal | 3              | 496    | -     | 499   |
|         |         | Upper  | -              | -      | 1     | 1     |
|         |         | Total  | 4              | 499    | 1     | 504   |
|         | Week 4  | Lower  | 2              | 3      | -     | 5     |
|         |         | Normal | 2              | 487    | 1     | 490   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 490    | 1     | 495   |
|         | Week 8  | Lower  | 2              | 2      | -     | 4     |
|         |         | Normal | 2              | 482    | 1     | 485   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 484    | 1     | 489   |
|         | Week 12 | Lower  | 1              | 3      | -     | 4     |
|         |         | Normal | 3              | 471    | 1     | 475   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 474    | 1     | 479   |
|         | Week 16 | Lower  | 2              | 3      | -     | 5     |
|         |         | Normal | 2              | 464    | 1     | 467   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 467    | 1     | 472   |
|         | Week 20 | Lower  | 2              | 2      | -     | 4     |
|         |         | Normal | 2              | 462    | 1     | 465   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 464    | 1     | 469   |
|         | Week 24 | Lower  | 1              | 4      | -     | 5     |
|         |         | Normal | 3              | 453    | 1     | 457   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 457    | 1     | 462   |
|         | Week 28 | Lower  | 2              | 4      | -     | 6     |
|         |         | Normal | 2              | 444    | 1     | 447   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 448    | 1     | 453   |
|         | Week 32 | Lower  | 2              | 2      | -     | 4     |
|         |         | Normal | 2              | 449    | 1     | 452   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 451    | 1     | 456   |
|         | Week 36 | Lower  | 1              | 3      | -     | 4     |
|         |         | Normal | 3              | 443    | 1     | 447   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 4              | 447    | 1     | 452   |
|         | Week 40 | Lower  | 1              | 5      | -     | 6     |
|         |         | Normal | 3              | 435    | 1     | 439   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 440    | 1     | 445   |
|         | Week 44 | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | 3              | 435    | 1     | 439   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 437    | 1     | 442   |
|         | Week 48 | Lower  | 2              | 1      | -     | 3     |
|         |         | Normal | 2              | 433    | 1     | 436   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 4              | 434    | 1     | 439   |
| Week 52 | Lower   | 1      | 2              | -      | 3     |       |
|         | Normal  | 3      | 432            | 1      | 436   |       |
|         | Upper   | -      | -              | -      | -     |       |
|         | Total   | 4      | 434            | 1      | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験) (3/24)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| ALT (GPT) | Week 2  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 337    | 17    | 354   |
|           |         | Upper  | -              | 30     | 120   | 150   |
|           |         | Total  | -              | 367    | 137   | 504   |
|           | Week 4  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 333    | 13    | 346   |
|           |         | Upper  | -              | 27     | 122   | 149   |
|           |         | Total  | -              | 360    | 135   | 495   |
|           | Week 8  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 325    | 24    | 349   |
|           |         | Upper  | -              | 29     | 111   | 140   |
|           |         | Total  | -              | 354    | 135   | 489   |
|           | Week 12 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 319    | 25    | 344   |
|           |         | Upper  | -              | 28     | 107   | 135   |
|           |         | Total  | -              | 347    | 132   | 479   |
|           | Week 16 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 311    | 32    | 343   |
|           |         | Upper  | -              | 30     | 99    | 129   |
|           |         | Total  | -              | 341    | 131   | 472   |
|           | Week 20 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 316    | 47    | 363   |
|           |         | Upper  | -              | 23     | 83    | 106   |
|           |         | Total  | -              | 339    | 130   | 469   |
|           | Week 24 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 305    | 53    | 358   |
|           |         | Upper  | -              | 28     | 76    | 104   |
|           |         | Total  | -              | 333    | 129   | 462   |
|           | Week 28 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 302    | 57    | 359   |
|           |         | Upper  | -              | 25     | 69    | 94    |
|           |         | Total  | -              | 327    | 126   | 453   |
|           | Week 32 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 313    | 61    | 374   |
|           |         | Upper  | -              | 16     | 66    | 82    |
|           |         | Total  | -              | 329    | 127   | 456   |
|           | Week 36 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 303    | 60    | 363   |
|           |         | Upper  | -              | 23     | 66    | 89    |
|           |         | Total  | -              | 326    | 126   | 452   |
|           | Week 40 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 307    | 60    | 367   |
|           |         | Upper  | -              | 15     | 63    | 78    |
|           |         | Total  | -              | 322    | 123   | 445   |
|           | Week 44 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 294    | 64    | 358   |
|           |         | Upper  | -              | 25     | 59    | 84    |
|           |         | Total  | -              | 319    | 123   | 442   |
|           | Week 48 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 300    | 65    | 365   |
|           |         | Upper  | -              | 16     | 58    | 74    |
|           |         | Total  | -              | 316    | 123   | 439   |
| Week 52   | Lower   | -      | -              | -      | -     |       |
|           | Normal  | -      | 294            | 68     | 362   |       |
|           | Upper   | -      | 22             | 55     | 77    |       |
|           | Total   | -      | 316            | 123    | 439   |       |



表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験) (4/24)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| AST (GOT) | Week 2  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 376    | 24    | 400   |
|           |         | Upper  | -              | 34     | 70    | 104   |
|           |         | Total  | -              | 410    | 94    | 504   |
|           | Week 4  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 365    | 22    | 387   |
|           |         | Upper  | -              | 38     | 70    | 108   |
|           |         | Total  | -              | 403    | 92    | 495   |
|           | Week 8  | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 358    | 33    | 391   |
|           |         | Upper  | -              | 39     | 59    | 98    |
|           |         | Total  | -              | 397    | 92    | 489   |
|           | Week 12 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 362    | 40    | 402   |
|           |         | Upper  | -              | 26     | 51    | 77    |
|           |         | Total  | -              | 388    | 91    | 479   |
|           | Week 16 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 360    | 43    | 403   |
|           |         | Upper  | -              | 22     | 47    | 69    |
|           |         | Total  | -              | 382    | 90    | 472   |
|           | Week 20 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 364    | 49    | 413   |
|           |         | Upper  | -              | 16     | 40    | 56    |
|           |         | Total  | -              | 380    | 89    | 469   |
|           | Week 24 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 351    | 54    | 405   |
|           |         | Upper  | -              | 22     | 35    | 57    |
|           |         | Total  | -              | 373    | 89    | 462   |
|           | Week 28 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 352    | 54    | 406   |
|           |         | Upper  | -              | 14     | 33    | 47    |
|           |         | Total  | -              | 366    | 87    | 453   |
|           | Week 32 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 351    | 49    | 400   |
|           |         | Upper  | -              | 18     | 38    | 56    |
|           |         | Total  | -              | 369    | 87    | 456   |
|           | Week 36 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 345    | 53    | 398   |
|           |         | Upper  | -              | 21     | 33    | 54    |
|           |         | Total  | -              | 366    | 86    | 452   |
|           | Week 40 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 345    | 54    | 399   |
|           |         | Upper  | -              | 15     | 31    | 46    |
|           |         | Total  | -              | 360    | 85    | 445   |
|           | Week 44 | Lower  | -              | -      | -     | -     |
|           |         | Normal | -              | 339    | 50    | 389   |
|           |         | Upper  | -              | 18     | 35    | 53    |
|           |         | Total  | -              | 357    | 85    | 442   |
| Week 48   | Lower   | -      | -              | -      | -     |       |
|           | Normal  | -      | 331            | 47     | 378   |       |
|           | Upper   | -      | 23             | 38     | 61    |       |
|           | Total   | -      | 354            | 85     | 439   |       |
| Week 52   | Lower   | -      | -              | -      | -     |       |
|           | Normal  | -      | 338            | 52     | 390   |       |
|           | Upper   | -      | 16             | 33     | 49    |       |
|           | Total   | -      | 354            | 85     | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験) (5/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| γ-GTP   | Week 2  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 330    | 15    | 345   |
|         |         | Upper  | -              | 15     | 144   | 159   |
|         |         | Total  | -              | 345    | 159   | 504   |
|         | Week 4  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 323    | 15    | 338   |
|         |         | Upper  | -              | 16     | 141   | 157   |
|         |         | Total  | -              | 339    | 156   | 495   |
|         | Week 8  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 314    | 24    | 338   |
|         |         | Upper  | -              | 20     | 131   | 151   |
|         |         | Total  | -              | 334    | 155   | 489   |
|         | Week 12 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 306    | 25    | 331   |
|         |         | Upper  | -              | 21     | 127   | 148   |
|         |         | Total  | -              | 327    | 152   | 479   |
|         | Week 16 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 310    | 30    | 340   |
|         |         | Upper  | -              | 13     | 119   | 132   |
|         |         | Total  | -              | 323    | 149   | 472   |
|         | Week 20 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 304    | 37    | 341   |
|         |         | Upper  | -              | 15     | 113   | 128   |
|         |         | Total  | -              | 319    | 150   | 469   |
|         | Week 24 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 298    | 41    | 339   |
|         |         | Upper  | -              | 16     | 107   | 123   |
|         |         | Total  | -              | 314    | 148   | 462   |
|         | Week 28 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 297    | 54    | 351   |
|         |         | Upper  | -              | 9      | 93    | 102   |
|         |         | Total  | -              | 306    | 147   | 453   |
|         | Week 32 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 302    | 51    | 353   |
|         |         | Upper  | -              | 9      | 94    | 103   |
|         |         | Total  | -              | 311    | 145   | 456   |
|         | Week 36 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 295    | 55    | 350   |
|         |         | Upper  | -              | 12     | 90    | 102   |
|         |         | Total  | -              | 307    | 145   | 452   |
|         | Week 40 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 290    | 51    | 341   |
|         |         | Upper  | -              | 12     | 92    | 104   |
|         |         | Total  | -              | 302    | 143   | 445   |
|         | Week 44 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 286    | 53    | 339   |
|         |         | Upper  | -              | 13     | 90    | 103   |
|         |         | Total  | -              | 299    | 143   | 442   |
|         | Week 48 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 285    | 52    | 337   |
|         |         | Upper  | -              | 12     | 90    | 102   |
|         |         | Total  | -              | 297    | 142   | 439   |
| Week 52 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 285            | 45     | 330   |       |
|         | Upper   | -      | 12             | 97     | 109   |       |
|         | Total   | -      | 297            | 142    | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験) (6/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| ALP     | Week 2  | Lower  | 2              | 1      | -     | 3     |
|         |         | Normal | 1              | 453    | 4     | 458   |
|         |         | Upper  | -              | 13     | 30    | 43    |
|         |         | Total  | 3              | 467    | 34    | 504   |
|         | Week 4  | Lower  | 2              | 1      | -     | 3     |
|         |         | Normal | 1              | 443    | 7     | 451   |
|         |         | Upper  | -              | 15     | 26    | 41    |
|         |         | Total  | 3              | 459    | 33    | 495   |
|         | Week 8  | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 3              | 439    | 7     | 449   |
|         |         | Upper  | -              | 13     | 26    | 39    |
|         |         | Total  | 3              | 453    | 33    | 489   |
|         | Week 12 | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | 2              | 429    | 4     | 435   |
|         |         | Upper  | -              | 13     | 29    | 42    |
|         |         | Total  | 3              | 443    | 33    | 479   |
|         | Week 16 | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | 2              | 419    | 6     | 427   |
|         |         | Upper  | -              | 16     | 27    | 43    |
|         |         | Total  | 3              | 436    | 33    | 472   |
|         | Week 20 | Lower  | 2              | 1      | -     | 3     |
|         |         | Normal | 1              | 411    | 7     | 419   |
|         |         | Upper  | -              | 21     | 26    | 47    |
|         |         | Total  | 3              | 433    | 33    | 469   |
|         | Week 24 | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | 2              | 411    | 5     | 418   |
|         |         | Upper  | -              | 16     | 26    | 42    |
|         |         | Total  | 3              | 428    | 31    | 462   |
|         | Week 28 | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | 2              | 393    | 8     | 403   |
|         |         | Upper  | -              | 26     | 23    | 49    |
|         |         | Total  | 3              | 419    | 31    | 453   |
|         | Week 32 | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | 2              | 405    | 10    | 417   |
|         |         | Upper  | -              | 15     | 22    | 37    |
|         |         | Total  | 3              | 421    | 32    | 456   |
|         | Week 36 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 3              | 400    | 9     | 412   |
|         |         | Upper  | -              | 17     | 23    | 40    |
|         |         | Total  | 3              | 417    | 32    | 452   |
|         | Week 40 | Lower  | 2              | 1      | -     | 3     |
|         |         | Normal | 1              | 387    | 8     | 396   |
|         |         | Upper  | -              | 22     | 24    | 46    |
|         |         | Total  | 3              | 410    | 32    | 445   |
|         | Week 44 | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | 2              | 387    | 11    | 400   |
|         |         | Upper  | -              | 20     | 20    | 40    |
|         |         | Total  | 3              | 408    | 31    | 442   |
| Week 48 | Lower   | 1      | 1              | -      | 2     |       |
|         | Normal  | 2      | 380            | 10     | 392   |       |
|         | Upper   | -      | 24             | 21     | 45    |       |
|         | Total   | 3      | 405            | 31     | 439   |       |
| Week 52 | Lower   | 1      | 1              | -      | 2     |       |
|         | Normal  | 2      | 391            | 11     | 404   |       |
|         | Upper   | -      | 13             | 20     | 33    |       |
|         | Total   | 3      | 405            | 31     | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験)(7/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| LDH     | Week 2  | Lower  | 3              | 1      | -     | 4     |
|         |         | Normal | -              | 435    | 17    | 452   |
|         |         | Upper  | -              | 14     | 34    | 48    |
|         |         | Total  | 3              | 450    | 51    | 504   |
|         | Week 4  | Lower  | 2              | 1      | -     | 3     |
|         |         | Normal | 1              | 432    | 17    | 450   |
|         |         | Upper  | -              | 9      | 33    | 42    |
|         |         | Total  | 3              | 442    | 50    | 495   |
|         | Week 8  | Lower  | 2              | 3      | -     | 5     |
|         |         | Normal | 1              | 420    | 18    | 439   |
|         |         | Upper  | -              | 15     | 30    | 45    |
|         |         | Total  | 3              | 438    | 48    | 489   |
|         | Week 12 | Lower  | 2              | 3      | -     | 5     |
|         |         | Normal | 1              | 408    | 19    | 428   |
|         |         | Upper  | -              | 17     | 29    | 46    |
|         |         | Total  | 3              | 428    | 48    | 479   |
|         | Week 16 | Lower  | 2              | 3      | -     | 5     |
|         |         | Normal | 1              | 406    | 21    | 428   |
|         |         | Upper  | -              | 14     | 25    | 39    |
|         |         | Total  | 3              | 423    | 46    | 472   |
|         | Week 20 | Lower  | 1              | 3      | -     | 4     |
|         |         | Normal | 1              | 408    | 23    | 432   |
|         |         | Upper  | -              | 10     | 23    | 33    |
|         |         | Total  | 2              | 421    | 46    | 469   |
|         | Week 24 | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | 1              | 404    | 20    | 425   |
|         |         | Upper  | -              | 9      | 25    | 34    |
|         |         | Total  | 2              | 415    | 45    | 462   |
|         | Week 28 | Lower  | 1              | 4      | -     | 5     |
|         |         | Normal | -              | 389    | 24    | 413   |
|         |         | Upper  | -              | 14     | 21    | 35    |
|         |         | Total  | 1              | 407    | 45    | 453   |
|         | Week 32 | Lower  | 1              | 4      | -     | 5     |
|         |         | Normal | -              | 394    | 22    | 416   |
|         |         | Upper  | -              | 12     | 23    | 35    |
|         |         | Total  | 1              | 410    | 45    | 456   |
|         | Week 36 | Lower  | 1              | 3      | -     | 4     |
|         |         | Normal | -              | 395    | 26    | 421   |
|         |         | Upper  | -              | 9      | 18    | 27    |
|         |         | Total  | 1              | 407    | 44    | 452   |
|         | Week 40 | Lower  | -              | 4      | -     | 4     |
|         |         | Normal | 1              | 380    | 23    | 404   |
|         |         | Upper  | -              | 15     | 22    | 37    |
|         |         | Total  | 1              | 399    | 45    | 445   |
|         | Week 44 | Lower  | -              | 5      | -     | 5     |
|         |         | Normal | 1              | 382    | 20    | 403   |
|         |         | Upper  | -              | 10     | 24    | 34    |
|         |         | Total  | 1              | 397    | 44    | 442   |
|         | Week 48 | Lower  | -              | 5      | -     | 5     |
|         |         | Normal | 1              | 383    | 21    | 405   |
|         |         | Upper  | -              | 7      | 22    | 29    |
|         |         | Total  | 1              | 395    | 43    | 439   |
| Week 52 | Lower   | -      | 6              | -      | 6     |       |
|         | Normal  | 1      | 382            | 22     | 405   |       |
|         | Upper   | -      | 6              | 22     | 28    |       |
|         | Total   | 1      | 394            | 44     | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験) (8/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| CPK     | Week 2  | Lower  | 7              | 7      | -     | 14    |
|         |         | Normal | 10             | 403    | 20    | 433   |
|         |         | Upper  | -              | 21     | 36    | 57    |
|         |         | Total  | 17             | 431    | 56    | 504   |
|         | Week 4  | Lower  | 9              | 8      | -     | 17    |
|         |         | Normal | 8              | 396    | 20    | 424   |
|         |         | Upper  | -              | 20     | 34    | 54    |
|         |         | Total  | 17             | 424    | 54    | 495   |
|         | Week 8  | Lower  | 11             | 6      | -     | 17    |
|         |         | Normal | 6              | 385    | 24    | 415   |
|         |         | Upper  | -              | 28     | 29    | 57    |
|         |         | Total  | 17             | 419    | 53    | 489   |
|         | Week 12 | Lower  | 12             | 8      | -     | 20    |
|         |         | Normal | 5              | 377    | 21    | 403   |
|         |         | Upper  | -              | 25     | 31    | 56    |
|         |         | Total  | 17             | 410    | 52    | 479   |
|         | Week 16 | Lower  | 8              | 6      | -     | 14    |
|         |         | Normal | 9              | 379    | 25    | 413   |
|         |         | Upper  | -              | 19     | 26    | 45    |
|         |         | Total  | 17             | 404    | 51    | 472   |
|         | Week 20 | Lower  | 11             | 7      | -     | 18    |
|         |         | Normal | 6              | 368    | 25    | 399   |
|         |         | Upper  | -              | 26     | 26    | 52    |
|         |         | Total  | 17             | 401    | 51    | 469   |
|         | Week 24 | Lower  | 11             | 10     | -     | 21    |
|         |         | Normal | 5              | 368    | 18    | 391   |
|         |         | Upper  | -              | 18     | 32    | 50    |
|         |         | Total  | 16             | 396    | 50    | 462   |
|         | Week 28 | Lower  | 7              | 6      | -     | 13    |
|         |         | Normal | 9              | 360    | 28    | 397   |
|         |         | Upper  | -              | 22     | 21    | 43    |
|         |         | Total  | 16             | 388    | 49    | 453   |
|         | Week 32 | Lower  | 7              | 8      | -     | 15    |
|         |         | Normal | 10             | 359    | 26    | 395   |
|         |         | Upper  | -              | 23     | 23    | 46    |
|         |         | Total  | 17             | 390    | 49    | 456   |
|         | Week 36 | Lower  | 7              | 5      | -     | 12    |
|         |         | Normal | 10             | 356    | 21    | 387   |
|         |         | Upper  | -              | 26     | 27    | 53    |
|         |         | Total  | 17             | 387    | 48    | 452   |
|         | Week 40 | Lower  | 8              | 7      | -     | 15    |
|         |         | Normal | 9              | 352    | 26    | 387   |
|         |         | Upper  | -              | 23     | 20    | 43    |
|         |         | Total  | 17             | 382    | 46    | 445   |
|         | Week 44 | Lower  | 10             | 9      | -     | 19    |
|         |         | Normal | 6              | 348    | 19    | 373   |
|         |         | Upper  | -              | 24     | 26    | 50    |
|         |         | Total  | 16             | 381    | 45    | 442   |
| Week 48 | Lower   | 8      | 11             | -      | 19    |       |
|         | Normal  | 8      | 335            | 26     | 369   |       |
|         | Upper   | -      | 32             | 19     | 51    |       |
|         | Total   | 16     | 378            | 45     | 439   |       |
| Week 52 | Lower   | 9      | 12             | -      | 21    |       |
|         | Normal  | 7      | 349            | 20     | 376   |       |
|         | Upper   | -      | 18             | 24     | 42    |       |
|         | Total   | 16     | 379            | 44     | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験)(9/24)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| Uric Acid | Week 2  | Lower  | 8              | 4      | -     | 12    |
|           |         | Normal | 4              | 397    | 27    | 428   |
|           |         | Upper  | -              | 19     | 45    | 64    |
|           |         | Total  | 12             | 420    | 72    | 504   |
|           | Week 4  | Lower  | 9              | 3      | -     | 12    |
|           |         | Normal | 2              | 386    | 25    | 413   |
|           |         | Upper  | -              | 24     | 46    | 70    |
|           |         | Total  | 11             | 413    | 71    | 495   |
|           | Week 8  | Lower  | 7              | 5      | -     | 12    |
|           |         | Normal | 4              | 377    | 28    | 409   |
|           |         | Upper  | -              | 25     | 43    | 68    |
|           |         | Total  | 11             | 407    | 71    | 489   |
|           | Week 12 | Lower  | 6              | 5      | -     | 11    |
|           |         | Normal | 5              | 372    | 35    | 412   |
|           |         | Upper  | -              | 22     | 34    | 56    |
|           |         | Total  | 11             | 399    | 69    | 479   |
|           | Week 16 | Lower  | 8              | 10     | -     | 18    |
|           |         | Normal | 3              | 361    | 41    | 405   |
|           |         | Upper  | -              | 21     | 28    | 49    |
|           |         | Total  | 11             | 392    | 69    | 472   |
|           | Week 20 | Lower  | 8              | 8      | -     | 16    |
|           |         | Normal | 2              | 371    | 41    | 414   |
|           |         | Upper  | -              | 11     | 28    | 39    |
|           |         | Total  | 10             | 390    | 69    | 469   |
|           | Week 24 | Lower  | 5              | 6      | 1     | 12    |
|           |         | Normal | 5              | 367    | 43    | 415   |
|           |         | Upper  | -              | 10     | 25    | 35    |
|           |         | Total  | 10             | 383    | 69    | 462   |
|           | Week 28 | Lower  | 7              | 10     | -     | 17    |
|           |         | Normal | 3              | 355    | 39    | 397   |
|           |         | Upper  | -              | 12     | 27    | 39    |
|           |         | Total  | 10             | 377    | 66    | 453   |
|           | Week 32 | Lower  | 6              | 7      | -     | 13    |
|           |         | Normal | 4              | 354    | 34    | 392   |
|           |         | Upper  | -              | 18     | 33    | 51    |
|           |         | Total  | 10             | 379    | 67    | 456   |
|           | Week 36 | Lower  | 5              | 3      | -     | 8     |
|           |         | Normal | 4              | 350    | 32    | 386   |
|           |         | Upper  | -              | 22     | 36    | 58    |
|           |         | Total  | 9              | 375    | 68    | 452   |
|           | Week 40 | Lower  | 5              | 6      | -     | 11    |
|           |         | Normal | 4              | 346    | 33    | 383   |
|           |         | Upper  | -              | 16     | 35    | 51    |
|           |         | Total  | 9              | 368    | 68    | 445   |
|           | Week 44 | Lower  | 5              | 3      | -     | 8     |
|           |         | Normal | 4              | 338    | 30    | 372   |
|           |         | Upper  | -              | 24     | 38    | 62    |
|           |         | Total  | 9              | 365    | 68    | 442   |
| Week 48   | Lower   | 3      | 5              | 1      | 9     |       |
|           | Normal  | 6      | 332            | 31     | 369   |       |
|           | Upper   | -      | 25             | 36     | 61    |       |
|           | Total   | 9      | 362            | 68     | 439   |       |
| Week 52   | Lower   | 3      | 5              | 1      | 9     |       |
|           | Normal  | 6      | 333            | 24     | 363   |       |
|           | Upper   | -      | 24             | 43     | 67    |       |
|           | Total   | 9      | 362            | 68     | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (10/24)

|                 |         |        | ATL-962 120 mg |        |       |       |
|-----------------|---------|--------|----------------|--------|-------|-------|
|                 |         |        | Week 0         |        |       |       |
|                 |         |        | Lower          | Normal | Upper | Total |
| Total Bilirubin | Week 2  | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 456    | 15    | 471   |
|                 |         | Upper  | -              | 16     | 17    | 33    |
|                 |         | Total  | -              | 472    | 32    | 504   |
|                 | Week 4  | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 449    | 19    | 468   |
|                 |         | Upper  | -              | 14     | 13    | 27    |
|                 |         | Total  | -              | 463    | 32    | 495   |
|                 | Week 8  | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 447    | 10    | 457   |
|                 |         | Upper  | -              | 10     | 22    | 32    |
|                 |         | Total  | -              | 457    | 32    | 489   |
|                 | Week 12 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 435    | 18    | 453   |
|                 |         | Upper  | -              | 13     | 13    | 26    |
|                 |         | Total  | -              | 448    | 31    | 479   |
|                 | Week 16 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 431    | 17    | 448   |
|                 |         | Upper  | -              | 10     | 14    | 24    |
|                 |         | Total  | -              | 441    | 31    | 472   |
|                 | Week 20 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 432    | 13    | 445   |
|                 |         | Upper  | -              | 7      | 17    | 24    |
|                 |         | Total  | -              | 439    | 30    | 469   |
|                 | Week 24 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 423    | 14    | 437   |
|                 |         | Upper  | -              | 9      | 16    | 25    |
|                 |         | Total  | -              | 432    | 30    | 462   |
|                 | Week 28 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 413    | 13    | 426   |
|                 |         | Upper  | -              | 11     | 16    | 27    |
|                 |         | Total  | -              | 424    | 29    | 453   |
|                 | Week 32 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 416    | 14    | 430   |
|                 |         | Upper  | -              | 11     | 15    | 26    |
|                 |         | Total  | -              | 427    | 29    | 456   |
|                 | Week 36 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 409    | 12    | 421   |
|                 |         | Upper  | -              | 14     | 17    | 31    |
|                 |         | Total  | -              | 423    | 29    | 452   |
|                 | Week 40 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 403    | 11    | 414   |
|                 |         | Upper  | -              | 13     | 18    | 31    |
|                 |         | Total  | -              | 416    | 29    | 445   |
|                 | Week 44 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 400    | 12    | 412   |
|                 |         | Upper  | -              | 13     | 17    | 30    |
|                 |         | Total  | -              | 413    | 29    | 442   |
|                 | Week 48 | Lower  | -              | -      | -     | -     |
|                 |         | Normal | -              | 395    | 13    | 408   |
|                 |         | Upper  | -              | 15     | 16    | 31    |
|                 |         | Total  | -              | 410    | 29    | 439   |
| Week 52         | Lower   | -      | -              | -      | -     |       |
|                 | Normal  | -      | 400            | 12     | 412   |       |
|                 | Upper   | -      | 10             | 17     | 27    |       |
|                 | Total   | -      | 410            | 29     | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (11/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| ChE     | Week 2  | Lower  | 3              | -      | -     | 3     |
|         |         | Normal | -              | 409    | 27    | 436   |
|         |         | Upper  | -              | 7      | 58    | 65    |
|         |         | Total  | 3              | 416    | 85    | 504   |
|         | Week 4  | Lower  | 2              | -      | -     | 2     |
|         |         | Normal | 1              | 407    | 28    | 436   |
|         |         | Upper  | -              | 3      | 54    | 57    |
|         |         | Total  | 3              | 410    | 82    | 495   |
|         | Week 8  | Lower  | 1              | -      | -     | 1     |
|         |         | Normal | 2              | 396    | 29    | 427   |
|         |         | Upper  | -              | 9      | 52    | 61    |
|         |         | Total  | 3              | 405    | 81    | 489   |
|         | Week 12 | Lower  | 2              | -      | -     | 2     |
|         |         | Normal | 1              | 389    | 28    | 418   |
|         |         | Upper  | -              | 7      | 52    | 59    |
|         |         | Total  | 3              | 396    | 80    | 479   |
|         | Week 16 | Lower  | 2              | -      | -     | 2     |
|         |         | Normal | 1              | 374    | 28    | 403   |
|         |         | Upper  | -              | 16     | 51    | 67    |
|         |         | Total  | 3              | 390    | 79    | 472   |
|         | Week 20 | Lower  | 2              | -      | -     | 2     |
|         |         | Normal | 1              | 378    | 25    | 404   |
|         |         | Upper  | -              | 10     | 53    | 63    |
|         |         | Total  | 3              | 388    | 78    | 469   |
|         | Week 24 | Lower  | 2              | -      | -     | 2     |
|         |         | Normal | 1              | 373    | 21    | 395   |
|         |         | Upper  | -              | 12     | 53    | 65    |
|         |         | Total  | 3              | 385    | 74    | 462   |
|         | Week 28 | Lower  | 2              | 1      | -     | 3     |
|         |         | Normal | 1              | 368    | 26    | 395   |
|         |         | Upper  | -              | 7      | 48    | 55    |
|         |         | Total  | 3              | 376    | 74    | 453   |
|         | Week 32 | Lower  | 2              | -      | -     | 2     |
|         |         | Normal | 1              | 369    | 27    | 397   |
|         |         | Upper  | -              | 9      | 48    | 57    |
|         |         | Total  | 3              | 378    | 75    | 456   |
|         | Week 36 | Lower  | 3              | 2      | -     | 5     |
|         |         | Normal | -              | 364    | 28    | 392   |
|         |         | Upper  | -              | 8      | 47    | 55    |
|         |         | Total  | 3              | 374    | 75    | 452   |
|         | Week 40 | Lower  | 3              | 2      | -     | 5     |
|         |         | Normal | -              | 357    | 22    | 379   |
|         |         | Upper  | -              | 10     | 51    | 61    |
|         |         | Total  | 3              | 369    | 73    | 445   |
|         | Week 44 | Lower  | 3              | 2      | -     | 5     |
|         |         | Normal | -              | 353    | 28    | 381   |
|         |         | Upper  | -              | 10     | 46    | 56    |
|         |         | Total  | 3              | 365    | 74    | 442   |
|         | Week 48 | Lower  | 2              | 2      | -     | 4     |
|         |         | Normal | 1              | 352    | 29    | 382   |
|         |         | Upper  | -              | 9      | 44    | 53    |
|         |         | Total  | 3              | 363    | 73    | 439   |
| Week 52 | Lower   | 3      | 2              | -      | 5     |       |
|         | Normal  | -      | 351            | 28     | 379   |       |
|         | Upper   | -      | 9              | 46     | 55    |       |
|         | Total   | 3      | 362            | 74     | 439   |       |



表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (12/24)

|            |         |        | ATL-962 120 mg |        |       |       |
|------------|---------|--------|----------------|--------|-------|-------|
|            |         |        | Week 0         |        |       |       |
|            |         |        | Lower          | Normal | Upper | Total |
| Creatinine | Week 2  | Lower  | 21             | 11     | -     | 32    |
|            |         | Normal | 20             | 398    | 9     | 427   |
|            |         | Upper  | -              | 11     | 34    | 45    |
|            |         | Total  | 41             | 420    | 43    | 504   |
|            | Week 4  | Lower  | 26             | 11     | -     | 37    |
|            |         | Normal | 13             | 396    | 9     | 418   |
|            |         | Upper  | 1              | 6      | 33    | 40    |
|            |         | Total  | 40             | 413    | 42    | 495   |
|            | Week 8  | Lower  | 25             | 20     | -     | 45    |
|            |         | Normal | 15             | 379    | 15    | 409   |
|            |         | Upper  | -              | 8      | 27    | 35    |
|            |         | Total  | 40             | 407    | 42    | 489   |
|            | Week 12 | Lower  | 26             | 27     | -     | 53    |
|            |         | Normal | 14             | 365    | 15    | 394   |
|            |         | Upper  | -              | 6      | 26    | 32    |
|            |         | Total  | 40             | 398    | 41    | 479   |
|            | Week 16 | Lower  | 26             | 34     | -     | 60    |
|            |         | Normal | 12             | 354    | 14    | 380   |
|            |         | Upper  | -              | 6      | 26    | 32    |
|            |         | Total  | 38             | 394    | 40    | 472   |
|            | Week 20 | Lower  | 28             | 24     | -     | 52    |
|            |         | Normal | 10             | 361    | 13    | 384   |
|            |         | Upper  | -              | 6      | 27    | 33    |
|            |         | Total  | 38             | 391    | 40    | 469   |
|            | Week 24 | Lower  | 25             | 25     | -     | 50    |
|            |         | Normal | 11             | 356    | 7     | 374   |
|            |         | Upper  | 1              | 5      | 32    | 38    |
|            |         | Total  | 37             | 386    | 39    | 462   |
|            | Week 28 | Lower  | 25             | 31     | -     | 56    |
|            |         | Normal | 11             | 345    | 11    | 367   |
|            |         | Upper  | -              | 3      | 27    | 30    |
|            |         | Total  | 36             | 379    | 38    | 453   |
|            | Week 32 | Lower  | 19             | 18     | -     | 37    |
|            |         | Normal | 17             | 355    | 7     | 379   |
|            |         | Upper  | -              | 9      | 31    | 40    |
|            |         | Total  | 36             | 382    | 38    | 456   |
|            | Week 36 | Lower  | 18             | 11     | -     | 29    |
|            |         | Normal | 17             | 355    | 16    | 388   |
|            |         | Upper  | -              | 13     | 22    | 35    |
|            |         | Total  | 35             | 379    | 38    | 452   |
|            | Week 40 | Lower  | 16             | 14     | -     | 30    |
|            |         | Normal | 19             | 350    | 12    | 381   |
|            |         | Upper  | -              | 8      | 26    | 34    |
|            |         | Total  | 35             | 372    | 38    | 445   |
|            | Week 44 | Lower  | 18             | 18     | -     | 36    |
|            |         | Normal | 16             | 340    | 11    | 367   |
|            |         | Upper  | 1              | 11     | 27    | 39    |
|            |         | Total  | 35             | 369    | 38    | 442   |
| Week 48    | Lower   | 15     | 15             | -      | 30    |       |
|            | Normal  | 19     | 342            | 12     | 373   |       |
|            | Upper   | -      | 11             | 25     | 36    |       |
|            | Total   | 34     | 368            | 37     | 439   |       |
| Week 52    | Lower   | 16     | 18             | -      | 34    |       |
|            | Normal  | 19     | 336            | 10     | 365   |       |
|            | Upper   | -      | 14             | 26     | 40    |       |
|            | Total   | 35     | 368            | 36     | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験) (13/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| BUN     | Week 2  | Lower  | -              | 4      | -     | 4     |
|         |         | Normal | 3              | 479    | 4     | 486   |
|         |         | Upper  | -              | 5      | 9     | 14    |
|         |         | Total  | 3              | 488    | 13    | 504   |
|         | Week 4  | Lower  | -              | 3      | -     | 3     |
|         |         | Normal | 3              | 471    | 6     | 480   |
|         |         | Upper  | -              | 6      | 6     | 12    |
|         |         | Total  | 3              | 480    | 12    | 495   |
|         | Week 8  | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 3              | 466    | 6     | 475   |
|         |         | Upper  | -              | 7      | 6     | 13    |
|         |         | Total  | 3              | 474    | 12    | 489   |
|         | Week 12 | Lower  | 1              | 6      | -     | 7     |
|         |         | Normal | 2              | 449    | 8     | 459   |
|         |         | Upper  | -              | 9      | 4     | 13    |
|         |         | Total  | 3              | 464    | 12    | 479   |
|         | Week 16 | Lower  | -              | 3      | -     | 3     |
|         |         | Normal | 3              | 448    | 8     | 459   |
|         |         | Upper  | -              | 6      | 4     | 10    |
|         |         | Total  | 3              | 457    | 12    | 472   |
|         | Week 20 | Lower  | -              | 3      | -     | 3     |
|         |         | Normal | 3              | 444    | 5     | 452   |
|         |         | Upper  | -              | 8      | 6     | 14    |
|         |         | Total  | 3              | 455    | 11    | 469   |
|         | Week 24 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 3              | 438    | 5     | 446   |
|         |         | Upper  | -              | 11     | 5     | 16    |
|         |         | Total  | 3              | 449    | 10    | 462   |
|         | Week 28 | Lower  | -              | 6      | -     | 6     |
|         |         | Normal | 3              | 428    | 4     | 435   |
|         |         | Upper  | -              | 6      | 6     | 12    |
|         |         | Total  | 3              | 440    | 10    | 453   |
|         | Week 32 | Lower  | 1              | 4      | -     | 5     |
|         |         | Normal | 2              | 426    | 6     | 434   |
|         |         | Upper  | -              | 13     | 4     | 17    |
|         |         | Total  | 3              | 443    | 10    | 456   |
|         | Week 36 | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | 2              | 425    | 6     | 433   |
|         |         | Upper  | -              | 13     | 4     | 17    |
|         |         | Total  | 3              | 439    | 10    | 452   |
|         | Week 40 | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | 2              | 420    | 4     | 426   |
|         |         | Upper  | -              | 10     | 6     | 16    |
|         |         | Total  | 3              | 432    | 10    | 445   |
|         | Week 44 | Lower  | -              | 5      | -     | 5     |
|         |         | Normal | 3              | 409    | 5     | 417   |
|         |         | Upper  | -              | 15     | 5     | 20    |
|         |         | Total  | 3              | 429    | 10    | 442   |
| Week 48 | Lower   | 1      | 2              | -      | 3     |       |
|         | Normal  | 2      | 418            | 6      | 426   |       |
|         | Upper   | -      | 6              | 4      | 10    |       |
|         | Total   | 3      | 426            | 10     | 439   |       |
| Week 52 | Lower   | -      | 2              | -      | 2     |       |
|         | Normal  | 3      | 415            | 3      | 421   |       |
|         | Upper   | -      | 10             | 6      | 16    |       |
|         | Total   | 3      | 427            | 9      | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (14/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| Na      | Week 2  | Lower  | 10             | 16     | -     | 26    |
|         |         | Normal | 11             | 467    | -     | 478   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 21             | 483    | -     | 504   |
|         | Week 4  | Lower  | 12             | 7      | -     | 19    |
|         |         | Normal | 9              | 467    | -     | 476   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 21             | 474    | -     | 495   |
|         | Week 8  | Lower  | 11             | 7      | -     | 18    |
|         |         | Normal | 9              | 462    | -     | 471   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 20             | 469    | -     | 489   |
|         | Week 12 | Lower  | 11             | 13     | -     | 24    |
|         |         | Normal | 9              | 446    | -     | 455   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 20             | 459    | -     | 479   |
|         | Week 16 | Lower  | 8              | 18     | -     | 26    |
|         |         | Normal | 11             | 435    | -     | 446   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 19             | 453    | -     | 472   |
|         | Week 20 | Lower  | 8              | 11     | -     | 19    |
|         |         | Normal | 11             | 439    | -     | 450   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 19             | 450    | -     | 469   |
|         | Week 24 | Lower  | 6              | 9      | -     | 15    |
|         |         | Normal | 12             | 435    | -     | 447   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 18             | 444    | -     | 462   |
|         | Week 28 | Lower  | 7              | 7      | -     | 14    |
|         |         | Normal | 11             | 428    | -     | 439   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 18             | 435    | -     | 453   |
|         | Week 32 | Lower  | 8              | 11     | -     | 19    |
|         |         | Normal | 10             | 427    | -     | 437   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 18             | 438    | -     | 456   |
|         | Week 36 | Lower  | 7              | 7      | -     | 14    |
|         |         | Normal | 10             | 428    | -     | 438   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 17             | 435    | -     | 452   |
|         | Week 40 | Lower  | 4              | 7      | -     | 11    |
|         |         | Normal | 13             | 421    | -     | 434   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 17             | 428    | -     | 445   |
|         | Week 44 | Lower  | 3              | 7      | -     | 10    |
|         |         | Normal | 14             | 418    | -     | 432   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 17             | 425    | -     | 442   |
|         | Week 48 | Lower  | 7              | 7      | -     | 14    |
|         |         | Normal | 10             | 415    | -     | 425   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 17             | 422    | -     | 439   |
| Week 52 | Lower   | 4      | 3              | -      | 7     |       |
|         | Normal  | 13     | 419            | -      | 432   |       |
|         | Upper   | -      | -              | -      | -     |       |
|         | Total   | 17     | 422            | -      | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (15/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| K       | Week 2  | Lower  | 1              | 4      | -     | 5     |
|         |         | Normal | 5              | 489    | 4     | 498   |
|         |         | Upper  | -              | -      | 1     | 1     |
|         |         | Total  | 6              | 493    | 5     | 504   |
|         | Week 4  | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | 5              | 478    | 3     | 486   |
|         |         | Upper  | -              | 4      | 2     | 6     |
|         |         | Total  | 6              | 484    | 5     | 495   |
|         | Week 8  | Lower  | 1              | 4      | -     | 5     |
|         |         | Normal | 5              | 472    | 1     | 478   |
|         |         | Upper  | -              | 3      | 3     | 6     |
|         |         | Total  | 6              | 479    | 4     | 489   |
|         | Week 12 | Lower  | 2              | 1      | -     | 3     |
|         |         | Normal | 4              | 462    | 3     | 469   |
|         |         | Upper  | -              | 6      | 1     | 7     |
|         |         | Total  | 6              | 469    | 4     | 479   |
|         | Week 16 | Lower  | -              | 5      | -     | 5     |
|         |         | Normal | 5              | 456    | 3     | 464   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 5              | 463    | 4     | 472   |
|         | Week 20 | Lower  | 1              | 3      | -     | 4     |
|         |         | Normal | 5              | 453    | 1     | 459   |
|         |         | Upper  | -              | 3      | 3     | 6     |
|         |         | Total  | 6              | 459    | 4     | 469   |
|         | Week 24 | Lower  | 1              | 5      | -     | 6     |
|         |         | Normal | 5              | 443    | 3     | 451   |
|         |         | Upper  | -              | 5      | -     | 5     |
|         |         | Total  | 6              | 453    | 3     | 462   |
|         | Week 28 | Lower  | 2              | 7      | -     | 9     |
|         |         | Normal | 4              | 434    | 2     | 440   |
|         |         | Upper  | -              | 3      | 1     | 4     |
|         |         | Total  | 6              | 444    | 3     | 453   |
|         | Week 32 | Lower  | -              | 2      | -     | 2     |
|         |         | Normal | 6              | 440    | 2     | 448   |
|         |         | Upper  | -              | 5      | 1     | 6     |
|         |         | Total  | 6              | 447    | 3     | 456   |
|         | Week 36 | Lower  | 1              | 3      | -     | 4     |
|         |         | Normal | 5              | 437    | 3     | 445   |
|         |         | Upper  | -              | 3      | -     | 3     |
|         |         | Total  | 6              | 443    | 3     | 452   |
|         | Week 40 | Lower  | 1              | 3      | -     | 4     |
|         |         | Normal | 5              | 427    | 2     | 434   |
|         |         | Upper  | -              | 6      | 1     | 7     |
|         |         | Total  | 6              | 436    | 3     | 445   |
|         | Week 44 | Lower  | 1              | 5      | -     | 6     |
|         |         | Normal | 5              | 426    | 2     | 433   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 6              | 433    | 3     | 442   |
|         | Week 48 | Lower  | 2              | 4      | -     | 6     |
|         |         | Normal | 4              | 425    | 2     | 431   |
|         |         | Upper  | -              | 2      | -     | 2     |
|         |         | Total  | 6              | 431    | 2     | 439   |
| Week 52 | Lower   | 2      | 3              | -      | 5     |       |
|         | Normal  | 4      | 428            | 1      | 433   |       |
|         | Upper   | -      | -              | 1      | 1     |       |
|         | Total   | 6      | 431            | 2      | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (16/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| CI      | Week 2  | Lower  | 4              | 10     | -     | 14    |
|         |         | Normal | 8              | 475    | -     | 483   |
|         |         | Upper  | -              | 6      | 1     | 7     |
|         |         | Total  | 12             | 491    | 1     | 504   |
|         | Week 4  | Lower  | 4              | 12     | -     | 16    |
|         |         | Normal | 8              | 469    | -     | 477   |
|         |         | Upper  | -              | 1      | 1     | 2     |
|         |         | Total  | 12             | 482    | 1     | 495   |
|         | Week 8  | Lower  | 3              | 12     | -     | 15    |
|         |         | Normal | 9              | 462    | -     | 471   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 12             | 476    | 1     | 489   |
|         | Week 12 | Lower  | 2              | 11     | -     | 13    |
|         |         | Normal | 10             | 452    | -     | 462   |
|         |         | Upper  | -              | 3      | 1     | 4     |
|         |         | Total  | 12             | 466    | 1     | 479   |
|         | Week 16 | Lower  | 2              | 15     | -     | 17    |
|         |         | Normal | 10             | 443    | 1     | 454   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 12             | 459    | 1     | 472   |
|         | Week 20 | Lower  | 3              | 10     | -     | 13    |
|         |         | Normal | 9              | 442    | 1     | 452   |
|         |         | Upper  | -              | 4      | -     | 4     |
|         |         | Total  | 12             | 456    | 1     | 469   |
|         | Week 24 | Lower  | 2              | 11     | -     | 13    |
|         |         | Normal | 10             | 439    | -     | 449   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 12             | 450    | -     | 462   |
|         | Week 28 | Lower  | 2              | 8      | -     | 10    |
|         |         | Normal | 10             | 433    | -     | 443   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 12             | 441    | -     | 453   |
|         | Week 32 | Lower  | 1              | 7      | -     | 8     |
|         |         | Normal | 11             | 436    | -     | 447   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 12             | 444    | -     | 456   |
|         | Week 36 | Lower  | 4              | 5      | -     | 9     |
|         |         | Normal | 8              | 433    | -     | 441   |
|         |         | Upper  | -              | 2      | -     | 2     |
|         |         | Total  | 12             | 440    | -     | 452   |
|         | Week 40 | Lower  | 2              | 4      | -     | 6     |
|         |         | Normal | 9              | 424    | -     | 433   |
|         |         | Upper  | -              | 6      | -     | 6     |
|         |         | Total  | 11             | 434    | -     | 445   |
|         | Week 44 | Lower  | 1              | 5      | -     | 6     |
|         |         | Normal | 10             | 425    | -     | 435   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 11             | 431    | -     | 442   |
| Week 48 | Lower   | -      | 7              | -      | 7     |       |
|         | Normal  | 11     | 420            | -      | 431   |       |
|         | Upper   | -      | 1              | -      | 1     |       |
|         | Total   | 11     | 428            | -      | 439   |       |
| Week 52 | Lower   | 3      | 5              | -      | 8     |       |
|         | Normal  | 8      | 422            | -      | 430   |       |
|         | Upper   | -      | 1              | -      | 1     |       |
|         | Total   | 11     | 428            | -      | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (17/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| Ca      | Week 2  | Lower  | 1              | 1      | -     | 2     |
|         |         | Normal | -              | 499    | 3     | 502   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 1              | 500    | 3     | 504   |
|         | Week 4  | Lower  | 1              | 2      | -     | 3     |
|         |         | Normal | -              | 489    | 3     | 492   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | 1              | 491    | 3     | 495   |
|         | Week 8  | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 1              | 483    | 2     | 486   |
|         |         | Upper  | -              | 1      | 1     | 2     |
|         |         | Total  | 1              | 485    | 3     | 489   |
|         | Week 12 | Lower  | -              | 2      | -     | 2     |
|         |         | Normal | 1              | 473    | 2     | 476   |
|         |         | Upper  | -              | -      | 1     | 1     |
|         |         | Total  | 1              | 475    | 3     | 479   |
|         | Week 16 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 1              | 467    | 2     | 470   |
|         |         | Upper  | -              | -      | 1     | 1     |
|         |         | Total  | 1              | 468    | 3     | 472   |
|         | Week 20 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 1              | 464    | 2     | 467   |
|         |         | Upper  | -              | 1      | 1     | 2     |
|         |         | Total  | 1              | 465    | 3     | 469   |
|         | Week 24 | Lower  | -              | -      | -     | -     |
|         |         | Normal | 1              | 457    | 3     | 461   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 1              | 458    | 3     | 462   |
|         | Week 28 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 1              | 448    | 2     | 451   |
|         |         | Upper  | -              | -      | 1     | 1     |
|         |         | Total  | 1              | 449    | 3     | 453   |
|         | Week 32 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | 1              | 451    | 2     | 454   |
|         |         | Upper  | -              | -      | 1     | 1     |
|         |         | Total  | 1              | 452    | 3     | 456   |
|         | Week 36 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | -              | 448    | 3     | 451   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 449    | 3     | 452   |
|         | Week 40 | Lower  | -              | 2      | -     | 2     |
|         |         | Normal | -              | 439    | 3     | 442   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | -              | 442    | 3     | 445   |
|         | Week 44 | Lower  | -              | 1      | -     | 1     |
|         |         | Normal | -              | 437    | 2     | 439   |
|         |         | Upper  | -              | 1      | 1     | 2     |
|         |         | Total  | -              | 439    | 3     | 442   |
| Week 48 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 435            | 2      | 437   |       |
|         | Upper   | -      | 1              | 1      | 2     |       |
|         | Total   | -      | 436            | 3      | 439   |       |
| Week 52 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 435            | 3      | 438   |       |
|         | Upper   | -      | 1              | -      | 1     |       |
|         | Total   | -      | 436            | 3      | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (18/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| P       | Week 2  | Lower  | 2              | 4      | -     | 6     |
|         |         | Normal | 5              | 487    | 2     | 494   |
|         |         | Upper  | -              | 3      | 1     | 4     |
|         |         | Total  | 7              | 494    | 3     | 504   |
|         | Week 4  | Lower  | 2              | 6      | -     | 8     |
|         |         | Normal | 5              | 477    | 2     | 484   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 7              | 485    | 3     | 495   |
|         | Week 8  | Lower  | 1              | 8      | -     | 9     |
|         |         | Normal | 6              | 470    | 2     | 478   |
|         |         | Upper  | -              | 1      | 1     | 2     |
|         |         | Total  | 7              | 479    | 3     | 489   |
|         | Week 12 | Lower  | 1              | 6      | -     | 7     |
|         |         | Normal | 6              | 460    | 3     | 469   |
|         |         | Upper  | -              | 3      | -     | 3     |
|         |         | Total  | 7              | 469    | 3     | 479   |
|         | Week 16 | Lower  | 1              | 9      | -     | 10    |
|         |         | Normal | 6              | 448    | 3     | 457   |
|         |         | Upper  | -              | 5      | -     | 5     |
|         |         | Total  | 7              | 462    | 3     | 472   |
|         | Week 20 | Lower  | 2              | 8      | -     | 10    |
|         |         | Normal | 5              | 449    | 2     | 456   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 7              | 459    | 3     | 469   |
|         | Week 24 | Lower  | 1              | 11     | -     | 12    |
|         |         | Normal | 5              | 440    | 2     | 447   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 6              | 453    | 3     | 462   |
|         | Week 28 | Lower  | 2              | 6      | -     | 8     |
|         |         | Normal | 4              | 433    | 2     | 439   |
|         |         | Upper  | -              | 5      | 1     | 6     |
|         |         | Total  | 6              | 444    | 3     | 453   |
|         | Week 32 | Lower  | 2              | 8      | -     | 10    |
|         |         | Normal | 4              | 435    | 2     | 441   |
|         |         | Upper  | -              | 4      | 1     | 5     |
|         |         | Total  | 6              | 447    | 3     | 456   |
|         | Week 36 | Lower  | 1              | 8      | -     | 9     |
|         |         | Normal | 4              | 434    | 2     | 440   |
|         |         | Upper  | -              | 2      | 1     | 3     |
|         |         | Total  | 5              | 444    | 3     | 452   |
|         | Week 40 | Lower  | -              | 7      | -     | 7     |
|         |         | Normal | 5              | 429    | 3     | 437   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | 5              | 437    | 3     | 445   |
|         | Week 44 | Lower  | 1              | 6      | -     | 7     |
|         |         | Normal | 4              | 423    | 2     | 429   |
|         |         | Upper  | -              | 5      | 1     | 6     |
|         |         | Total  | 5              | 434    | 3     | 442   |
|         | Week 48 | Lower  | 3              | 5      | -     | 8     |
|         |         | Normal | 2              | 422    | 2     | 426   |
|         |         | Upper  | -              | 4      | 1     | 5     |
|         |         | Total  | 5              | 431    | 3     | 439   |
| Week 52 | Lower   | 2      | 11             | -      | 13    |       |
|         | Normal  | 3      | 415            | 2      | 420   |       |
|         | Upper   | -      | 5              | 1      | 6     |       |
|         | Total   | 5      | 431            | 3      | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表(OCT-002 試験) (19/24)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| Vitamin A | Week 2  | Lower  | 60             | 35     | -     | 95    |
|           |         | Normal | 23             | 372    | 5     | 400   |
|           |         | Upper  | -              | 5      | 4     | 9     |
|           |         | Total  | 83             | 412    | 9     | 504   |
|           | Week 4  | Lower  | 61             | 34     | -     | 95    |
|           |         | Normal | 20             | 370    | 5     | 395   |
|           |         | Upper  | -              | 1      | 4     | 5     |
|           |         | Total  | 81             | 405    | 9     | 495   |
|           | Week 8  | Lower  | 70             | 36     | -     | 106   |
|           |         | Normal | 11             | 360    | 6     | 377   |
|           |         | Upper  | -              | 3      | 3     | 6     |
|           |         | Total  | 81             | 399    | 9     | 489   |
|           | Week 12 | Lower  | 62             | 37     | -     | 99    |
|           |         | Normal | 18             | 352    | 4     | 374   |
|           |         | Upper  | 1              | 1      | 4     | 6     |
|           |         | Total  | 81             | 390    | 8     | 479   |
|           | Week 16 | Lower  | 64             | 51     | 1     | 116   |
|           |         | Normal | 15             | 331    | 4     | 350   |
|           |         | Upper  | -              | 3      | 3     | 6     |
|           |         | Total  | 79             | 385    | 8     | 472   |
|           | Week 20 | Lower  | 61             | 46     | -     | 107   |
|           |         | Normal | 17             | 334    | 4     | 355   |
|           |         | Upper  | -              | 3      | 4     | 7     |
|           |         | Total  | 78             | 383    | 8     | 469   |
|           | Week 24 | Lower  | 62             | 53     | -     | 115   |
|           |         | Normal | 15             | 320    | 4     | 339   |
|           |         | Upper  | -              | 4      | 4     | 8     |
|           |         | Total  | 77             | 377    | 8     | 462   |
|           | Week 28 | Lower  | 61             | 59     | -     | 120   |
|           |         | Normal | 14             | 308    | 4     | 326   |
|           |         | Upper  | 1              | 1      | 4     | 6     |
|           |         | Total  | 76             | 368    | 8     | 452   |
|           | Week 32 | Lower  | 58             | 50     | -     | 108   |
|           |         | Normal | 18             | 320    | 5     | 343   |
|           |         | Upper  | -              | 2      | 3     | 5     |
|           |         | Total  | 76             | 372    | 8     | 456   |
|           | Week 36 | Lower  | 60             | 49     | -     | 109   |
|           |         | Normal | 14             | 320    | 4     | 338   |
|           |         | Upper  | -              | 1      | 4     | 5     |
|           |         | Total  | 74             | 370    | 8     | 452   |
|           | Week 40 | Lower  | 51             | 60     | -     | 111   |
|           |         | Normal | 22             | 301    | 4     | 327   |
|           |         | Upper  | -              | 3      | 4     | 7     |
|           |         | Total  | 73             | 364    | 8     | 445   |
|           | Week 44 | Lower  | 53             | 39     | -     | 92    |
|           |         | Normal | 18             | 320    | 3     | 341   |
|           |         | Upper  | -              | 4      | 5     | 9     |
|           |         | Total  | 71             | 363    | 8     | 442   |
| Week 48   | Lower   | 51     | 36             | -      | 87    |       |
|           | Normal  | 20     | 320            | 3      | 343   |       |
|           | Upper   | -      | 4              | 5      | 9     |       |
|           | Total   | 71     | 360            | 8      | 439   |       |
| Week 52   | Lower   | 49     | 38             | -      | 87    |       |
|           | Normal  | 21     | 319            | 3      | 343   |       |
|           | Upper   | 1      | 2              | 5      | 8     |       |
|           | Total   | 71     | 359            | 8      | 438   |       |



表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (20/24)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| Vitamin D | Week 2  | Lower  | 2              | 6      | -     | 8     |
|           |         | Normal | 10             | 258    | 72    | 340   |
|           |         | Upper  | -              | 87     | 69    | 156   |
|           |         | Total  | 12             | 351    | 141   | 504   |
|           | Week 4  | Lower  | 4              | 11     | 1     | 16    |
|           |         | Normal | 8              | 252    | 78    | 338   |
|           |         | Upper  | -              | 81     | 60    | 141   |
|           |         | Total  | 12             | 344    | 139   | 495   |
|           | Week 8  | Lower  | 2              | 9      | 1     | 12    |
|           |         | Normal | 10             | 277    | 70    | 357   |
|           |         | Upper  | -              | 52     | 68    | 120   |
|           |         | Total  | 12             | 338    | 139   | 489   |
|           | Week 12 | Lower  | 4              | 8      | -     | 12    |
|           |         | Normal | 8              | 264    | 67    | 339   |
|           |         | Upper  | -              | 59     | 69    | 128   |
|           |         | Total  | 12             | 331    | 136   | 479   |
|           | Week 16 | Lower  | 2              | 5      | 1     | 8     |
|           |         | Normal | 9              | 262    | 70    | 341   |
|           |         | Upper  | -              | 59     | 64    | 123   |
|           |         | Total  | 11             | 326    | 135   | 472   |
|           | Week 20 | Lower  | 3              | 7      | -     | 10    |
|           |         | Normal | 8              | 268    | 63    | 339   |
|           |         | Upper  | -              | 49     | 71    | 120   |
|           |         | Total  | 11             | 324    | 134   | 469   |
|           | Week 24 | Lower  | 1              | 5      | -     | 6     |
|           |         | Normal | 9              | 251    | 76    | 336   |
|           |         | Upper  | -              | 64     | 56    | 120   |
|           |         | Total  | 10             | 320    | 132   | 462   |
|           | Week 28 | Lower  | 3              | 6      | -     | 9     |
|           |         | Normal | 6              | 255    | 74    | 335   |
|           |         | Upper  | 1              | 51     | 57    | 109   |
|           |         | Total  | 10             | 312    | 131   | 453   |
|           | Week 32 | Lower  | 1              | 11     | -     | 12    |
|           |         | Normal | 8              | 257    | 77    | 342   |
|           |         | Upper  | 1              | 47     | 54    | 102   |
|           |         | Total  | 10             | 315    | 131   | 456   |
|           | Week 36 | Lower  | 1              | 9      | -     | 10    |
|           |         | Normal | 8              | 236    | 71    | 315   |
|           |         | Upper  | -              | 68     | 59    | 127   |
|           |         | Total  | 9              | 313    | 130   | 452   |
|           | Week 40 | Lower  | 1              | 10     | -     | 11    |
|           |         | Normal | 8              | 246    | 71    | 325   |
|           |         | Upper  | -              | 53     | 56    | 109   |
|           |         | Total  | 9              | 309    | 127   | 445   |
|           | Week 44 | Lower  | 2              | 7      | -     | 9     |
|           |         | Normal | 7              | 229    | 68    | 304   |
|           |         | Upper  | -              | 71     | 58    | 129   |
|           |         | Total  | 9              | 307    | 126   | 442   |
| Week 48   | Lower   | 3      | 9              | 2      | 14    |       |
|           | Normal  | 6      | 229            | 63     | 298   |       |
|           | Upper   | -      | 66             | 61     | 127   |       |
|           | Total   | 9      | 304            | 126    | 439   |       |
| Week 52   | Lower   | 1      | 5              | -      | 6     |       |
|           | Normal  | 7      | 220            | 62     | 289   |       |
|           | Upper   | -      | 80             | 64     | 144   |       |
|           | Total   | 8      | 305            | 126    | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (21/24)

|           |         |        | ATL-962 120 mg |        |       |       |
|-----------|---------|--------|----------------|--------|-------|-------|
|           |         |        | Week 0         |        |       |       |
|           |         |        | Lower          | Normal | Upper | Total |
| Vitamin E | Week 2  | Lower  | 7              | 14     | -     | 21    |
|           |         | Normal | 1              | 294    | 105   | 400   |
|           |         | Upper  | -              | 17     | 66    | 83    |
|           |         | Total  | 8              | 325    | 171   | 504   |
|           | Week 4  | Lower  | 6              | 20     | -     | 26    |
|           |         | Normal | 2              | 288    | 101   | 391   |
|           |         | Upper  | -              | 9      | 69    | 78    |
|           |         | Total  | 8              | 317    | 170   | 495   |
|           | Week 8  | Lower  | 8              | 29     | -     | 37    |
|           |         | Normal | -              | 271    | 104   | 375   |
|           |         | Upper  | -              | 13     | 64    | 77    |
|           |         | Total  | 8              | 313    | 168   | 489   |
|           | Week 12 | Lower  | 5              | 28     | -     | 33    |
|           |         | Normal | 3              | 270    | 109   | 382   |
|           |         | Upper  | -              | 7      | 57    | 64    |
|           |         | Total  | 8              | 305    | 166   | 479   |
|           | Week 16 | Lower  | 5              | 20     | -     | 25    |
|           |         | Normal | 3              | 265    | 100   | 368   |
|           |         | Upper  | -              | 15     | 64    | 79    |
|           |         | Total  | 8              | 300    | 164   | 472   |
|           | Week 20 | Lower  | 7              | 26     | 1     | 34    |
|           |         | Normal | 1              | 264    | 104   | 369   |
|           |         | Upper  | -              | 8      | 58    | 66    |
|           |         | Total  | 8              | 298    | 163   | 469   |
|           | Week 24 | Lower  | 8              | 19     | 1     | 28    |
|           |         | Normal | -              | 267    | 104   | 371   |
|           |         | Upper  | -              | 9      | 54    | 63    |
|           |         | Total  | 8              | 295    | 159   | 462   |
|           | Week 28 | Lower  | 7              | 15     | 1     | 23    |
|           |         | Normal | 1              | 259    | 99    | 359   |
|           |         | Upper  | -              | 14     | 57    | 71    |
|           |         | Total  | 8              | 288    | 157   | 453   |
|           | Week 32 | Lower  | 7              | 22     | 2     | 31    |
|           |         | Normal | 1              | 261    | 100   | 362   |
|           |         | Upper  | -              | 8      | 55    | 63    |
|           |         | Total  | 8              | 291    | 157   | 456   |
|           | Week 36 | Lower  | 6              | 25     | 1     | 32    |
|           |         | Normal | 1              | 262    | 110   | 373   |
|           |         | Upper  | -              | 1      | 46    | 47    |
|           |         | Total  | 7              | 288    | 157   | 452   |
|           | Week 40 | Lower  | 6              | 33     | 1     | 40    |
|           |         | Normal | 1              | 246    | 103   | 350   |
|           |         | Upper  | -              | 5      | 50    | 55    |
|           |         | Total  | 7              | 284    | 154   | 445   |
|           | Week 44 | Lower  | 5              | 26     | 1     | 32    |
|           |         | Normal | 2              | 250    | 115   | 367   |
|           |         | Upper  | -              | 5      | 38    | 43    |
|           |         | Total  | 7              | 281    | 154   | 442   |
| Week 48   | Lower   | 7      | 33             | -      | 40    |       |
|           | Normal  | -      | 239            | 106    | 345   |       |
|           | Upper   | -      | 8              | 46     | 54    |       |
|           | Total   | 7      | 280            | 152    | 439   |       |
| Week 52   | Lower   | 6      | 34             | 2      | 42    |       |
|           | Normal  | 1      | 241            | 111    | 353   |       |
|           | Upper   | -      | 4              | 39     | 43    |       |
|           | Total   | 7      | 279            | 152    | 438   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (22/24)

|            |         |        | ATL-962 120 mg |        |       |       |
|------------|---------|--------|----------------|--------|-------|-------|
|            |         |        | Week 0         |        |       |       |
|            |         |        | Lower          | Normal | Upper | Total |
| β-Carotene | Week 2  | Lower  | 167            | 53     | -     | 220   |
|            |         | Normal | 21             | 247    | 5     | 273   |
|            |         | Upper  | -              | 2      | 9     | 11    |
|            |         | Total  | 188            | 302    | 14    | 504   |
|            | Week 4  | Lower  | 151            | 67     | -     | 218   |
|            |         | Normal | 33             | 229    | 7     | 269   |
|            |         | Upper  | -              | 1      | 7     | 8     |
|            |         | Total  | 184            | 297    | 14    | 495   |
|            | Week 8  | Lower  | 150            | 58     | -     | 208   |
|            |         | Normal | 34             | 231    | 7     | 272   |
|            |         | Upper  | -              | 3      | 6     | 9     |
|            |         | Total  | 184            | 292    | 13    | 489   |
|            | Week 12 | Lower  | 152            | 60     | -     | 212   |
|            |         | Normal | 32             | 220    | 8     | 260   |
|            |         | Upper  | -              | 2      | 5     | 7     |
|            |         | Total  | 184            | 282    | 13    | 479   |
|            | Week 16 | Lower  | 140            | 64     | -     | 204   |
|            |         | Normal | 40             | 212    | 8     | 260   |
|            |         | Upper  | -              | 3      | 5     | 8     |
|            |         | Total  | 180            | 279    | 13    | 472   |
|            | Week 20 | Lower  | 140            | 59     | -     | 199   |
|            |         | Normal | 39             | 215    | 9     | 263   |
|            |         | Upper  | -              | 3      | 4     | 7     |
|            |         | Total  | 179            | 277    | 13    | 469   |
|            | Week 24 | Lower  | 133            | 61     | -     | 194   |
|            |         | Normal | 41             | 211    | 9     | 261   |
|            |         | Upper  | -              | 3      | 4     | 7     |
|            |         | Total  | 174            | 275    | 13    | 462   |
|            | Week 28 | Lower  | 134            | 47     | -     | 181   |
|            |         | Normal | 41             | 213    | 10    | 264   |
|            |         | Upper  | -              | 5      | 3     | 8     |
|            |         | Total  | 175            | 265    | 13    | 453   |
|            | Week 32 | Lower  | 123            | 43     | -     | 166   |
|            |         | Normal | 52             | 220    | 8     | 280   |
|            |         | Upper  | -              | 5      | 5     | 10    |
|            |         | Total  | 175            | 268    | 13    | 456   |
|            | Week 36 | Lower  | 132            | 42     | -     | 174   |
|            |         | Normal | 41             | 218    | 8     | 267   |
|            |         | Upper  | -              | 6      | 5     | 11    |
|            |         | Total  | 173            | 266    | 13    | 452   |
|            | Week 40 | Lower  | 127            | 37     | -     | 164   |
|            |         | Normal | 45             | 215    | 8     | 268   |
|            |         | Upper  | -              | 8      | 5     | 13    |
|            |         | Total  | 172            | 260    | 13    | 445   |
|            | Week 44 | Lower  | 125            | 35     | -     | 160   |
|            |         | Normal | 44             | 219    | 8     | 271   |
|            |         | Upper  | -              | 6      | 5     | 11    |
|            |         | Total  | 169            | 260    | 13    | 442   |
| Week 48    | Lower   | 121    | 24             | -      | 145   |       |
|            | Normal  | 47     | 229            | 6      | 282   |       |
|            | Upper   | -      | 5              | 7      | 12    |       |
|            | Total   | 168    | 258            | 13     | 439   |       |
| Week 52    | Lower   | 123    | 35             | -      | 158   |       |
|            | Normal  | 46     | 217            | 6      | 269   |       |
|            | Upper   | -      | 5              | 7      | 12    |       |
|            | Total   | 169    | 257            | 13     | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (23/24)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| m-AST   | Week 2  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 468    | 11    | 479   |
|         |         | Upper  | -              | 13     | 12    | 25    |
|         |         | Total  | -              | 481    | 23    | 504   |
|         | Week 4  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 463    | 13    | 476   |
|         |         | Upper  | -              | 10     | 9     | 19    |
|         |         | Total  | -              | 473    | 22    | 495   |
|         | Week 8  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 456    | 15    | 471   |
|         |         | Upper  | -              | 12     | 6     | 18    |
|         |         | Total  | -              | 468    | 21    | 489   |
|         | Week 12 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 449    | 13    | 462   |
|         |         | Upper  | -              | 8      | 8     | 16    |
|         |         | Total  | -              | 457    | 21    | 478   |
|         | Week 16 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 443    | 16    | 459   |
|         |         | Upper  | -              | 8      | 5     | 13    |
|         |         | Total  | -              | 451    | 21    | 472   |
|         | Week 20 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 437    | 15    | 452   |
|         |         | Upper  | -              | 11     | 6     | 17    |
|         |         | Total  | -              | 448    | 21    | 469   |
|         | Week 24 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 436    | 18    | 454   |
|         |         | Upper  | -              | 5      | 3     | 8     |
|         |         | Total  | -              | 441    | 21    | 462   |
|         | Week 28 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 421    | 14    | 435   |
|         |         | Upper  | -              | 12     | 6     | 18    |
|         |         | Total  | -              | 433    | 20    | 453   |
|         | Week 32 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 428    | 13    | 441   |
|         |         | Upper  | -              | 8      | 7     | 15    |
|         |         | Total  | -              | 436    | 20    | 456   |
|         | Week 36 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 423    | 17    | 440   |
|         |         | Upper  | -              | 9      | 3     | 12    |
|         |         | Total  | -              | 432    | 20    | 452   |
|         | Week 40 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 413    | 16    | 429   |
|         |         | Upper  | -              | 12     | 4     | 16    |
|         |         | Total  | -              | 425    | 20    | 445   |
|         | Week 44 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 409    | 15    | 424   |
|         |         | Upper  | -              | 13     | 5     | 18    |
|         |         | Total  | -              | 422    | 20    | 442   |
| Week 48 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 403            | 12     | 415   |       |
|         | Upper   | -      | 16             | 8      | 24    |       |
|         | Total   | -      | 419            | 20     | 439   |       |
| Week 52 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 404            | 15     | 419   |       |
|         | Upper   | -      | 15             | 5      | 20    |       |
|         | Total   | -      | 419            | 20     | 439   |       |

表 2.7.4.7-付録-33 血液生化学検査の評価時点別正異判定クロス表 (OCT-002 試験) (24/24)

|                |         |        | ATL-962 120 mg |        |       |       |
|----------------|---------|--------|----------------|--------|-------|-------|
|                |         |        | Week 0         |        |       |       |
|                |         |        | Lower          | Normal | Upper | Total |
| CK18 (Segment) | Week 2  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 395    | 31    | 426   |
|                |         | Upper  | -              | 25     | 53    | 78    |
|                |         | Total  | -              | 420    | 84    | 504   |
|                | Week 4  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 382    | 29    | 411   |
|                |         | Upper  | -              | 31     | 53    | 84    |
|                |         | Total  | -              | 413    | 82    | 495   |
|                | Week 8  | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 382    | 32    | 414   |
|                |         | Upper  | -              | 25     | 50    | 75    |
|                |         | Total  | -              | 407    | 82    | 489   |
|                | Week 12 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 374    | 35    | 409   |
|                |         | Upper  | -              | 24     | 46    | 70    |
|                |         | Total  | -              | 398    | 81    | 479   |
|                | Week 16 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 368    | 43    | 411   |
|                |         | Upper  | -              | 24     | 37    | 61    |
|                |         | Total  | -              | 392    | 80    | 472   |
|                | Week 20 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 370    | 42    | 412   |
|                |         | Upper  | -              | 20     | 37    | 57    |
|                |         | Total  | -              | 390    | 79    | 469   |
|                | Week 24 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 362    | 48    | 410   |
|                |         | Upper  | -              | 21     | 31    | 52    |
|                |         | Total  | -              | 383    | 79    | 462   |
|                | Week 28 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 350    | 49    | 399   |
|                |         | Upper  | -              | 26     | 28    | 54    |
|                |         | Total  | -              | 376    | 77    | 453   |
|                | Week 32 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 359    | 50    | 409   |
|                |         | Upper  | -              | 21     | 26    | 47    |
|                |         | Total  | -              | 380    | 76    | 456   |
|                | Week 36 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 354    | 47    | 401   |
|                |         | Upper  | -              | 22     | 29    | 51    |
|                |         | Total  | -              | 376    | 76    | 452   |
|                | Week 40 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 350    | 48    | 398   |
|                |         | Upper  | -              | 19     | 28    | 47    |
|                |         | Total  | -              | 369    | 76    | 445   |
|                | Week 44 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 343    | 44    | 387   |
|                |         | Upper  | -              | 23     | 32    | 55    |
|                |         | Total  | -              | 366    | 76    | 442   |
|                | Week 48 | Lower  | -              | -      | -     | -     |
|                |         | Normal | -              | 341    | 41    | 382   |
|                |         | Upper  | -              | 22     | 35    | 57    |
|                |         | Total  | -              | 363    | 76    | 439   |
| Week 52        | Lower   | -      | -              | -      | -     |       |
|                | Normal  | -      | 337            | 40     | 377   |       |
|                | Upper   | -      | 26             | 36     | 62    |       |
|                | Total   | -      | 363            | 76     | 439   |       |

表 2.7.4.7-付録-34 尿検査の評価時点別の変化量（OCT-002 試験）

| Specific Gravity |                | Summary Statistics |         |         |        |         |         |        |       |
|------------------|----------------|--------------------|---------|---------|--------|---------|---------|--------|-------|
| Visit            | Treatment      | N                  | Mean    | SD      | Min    | Q1      | Median  | Q3     | Max   |
| Week 2           | ATL-962 120 mg | 504                | -0.0006 | 0.00555 | -0.024 | -0.0040 | -0.0010 | 0.0030 | 0.020 |
| Week 4           | ATL-962 120 mg | 495                | -0.0006 | 0.00580 | -0.021 | -0.0040 | 0.0000  | 0.0030 | 0.024 |
| Week 8           | ATL-962 120 mg | 489                | -0.0010 | 0.00599 | -0.020 | -0.0050 | -0.0010 | 0.0020 | 0.024 |
| Week 12          | ATL-962 120 mg | 479                | -0.0014 | 0.00610 | -0.022 | -0.0050 | -0.0010 | 0.0020 | 0.023 |
| Week 16          | ATL-962 120 mg | 472                | -0.0012 | 0.00589 | -0.022 | -0.0050 | -0.0010 | 0.0020 | 0.015 |
| Week 20          | ATL-962 120 mg | 469                | -0.0014 | 0.00610 | -0.020 | -0.0050 | -0.0020 | 0.0030 | 0.020 |
| Week 24          | ATL-962 120 mg | 462                | -0.0010 | 0.00590 | -0.023 | -0.0050 | -0.0010 | 0.0020 | 0.017 |
| Week 28          | ATL-962 120 mg | 453                | -0.0010 | 0.00593 | -0.020 | -0.0050 | -0.0010 | 0.0030 | 0.019 |
| Week 32          | ATL-962 120 mg | 456                | -0.0011 | 0.00605 | -0.021 | -0.0050 | -0.0010 | 0.0030 | 0.018 |
| Week 36          | ATL-962 120 mg | 452                | -0.0009 | 0.00599 | -0.018 | -0.0040 | -0.0010 | 0.0030 | 0.020 |
| Week 40          | ATL-962 120 mg | 445                | -0.0008 | 0.00619 | -0.020 | -0.0050 | -0.0010 | 0.0030 | 0.019 |
| Week 44          | ATL-962 120 mg | 442                | -0.0002 | 0.00657 | -0.025 | -0.0040 | 0.0000  | 0.0040 | 0.026 |
| Week 48          | ATL-962 120 mg | 439                | -0.0005 | 0.00592 | -0.019 | -0.0040 | 0.0000  | 0.0030 | 0.020 |
| Week 52          | ATL-962 120 mg | 439                | -0.0008 | 0.00591 | -0.019 | -0.0040 | 0.0000  | 0.0030 | 0.017 |

表 2.7.4.7-付録-35 尿検査の評価時点別クロス表 (OCT-002 試験) (1/6)

|                  |         |        | ATL-962 120 mg |        |       |       |
|------------------|---------|--------|----------------|--------|-------|-------|
|                  |         |        | Week 0         |        |       |       |
|                  |         |        | Lower          | Normal | Upper | Total |
| Specific Gravity | Week 2  | Lower  | -              | 2      | -     | 2     |
|                  |         | Normal | -              | 497    | 1     | 498   |
|                  |         | Upper  | -              | 4      | -     | 4     |
|                  |         | Total  | -              | 503    | 1     | 504   |
|                  | Week 4  | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 489    | 1     | 490   |
|                  |         | Upper  | -              | 5      | -     | 5     |
|                  |         | Total  | -              | 494    | 1     | 495   |
|                  | Week 8  | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 483    | 1     | 484   |
|                  |         | Upper  | -              | 5      | -     | 5     |
|                  |         | Total  | -              | 488    | 1     | 489   |
|                  | Week 12 | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 475    | 1     | 476   |
|                  |         | Upper  | -              | 3      | -     | 3     |
|                  |         | Total  | -              | 478    | 1     | 479   |
|                  | Week 16 | Lower  | -              | 1      | -     | 1     |
|                  |         | Normal | -              | 467    | 1     | 468   |
|                  |         | Upper  | -              | 3      | -     | 3     |
|                  |         | Total  | -              | 471    | 1     | 472   |
|                  | Week 20 | Lower  | -              | 1      | -     | 1     |
|                  |         | Normal | -              | 466    | 1     | 467   |
|                  |         | Upper  | -              | 1      | -     | 1     |
|                  |         | Total  | -              | 468    | 1     | 469   |
|                  | Week 24 | Lower  | -              | 1      | -     | 1     |
|                  |         | Normal | -              | 459    | 1     | 460   |
|                  |         | Upper  | -              | 1      | -     | 1     |
|                  |         | Total  | -              | 461    | 1     | 462   |
|                  | Week 28 | Lower  | -              | 1      | -     | 1     |
|                  |         | Normal | -              | 449    | 1     | 450   |
|                  |         | Upper  | -              | 2      | -     | 2     |
|                  |         | Total  | -              | 452    | 1     | 453   |
|                  | Week 32 | Lower  | -              | 1      | -     | 1     |
|                  |         | Normal | -              | 454    | 1     | 455   |
|                  |         | Upper  | -              | -      | -     | -     |
|                  |         | Total  | -              | 455    | 1     | 456   |
|                  | Week 36 | Lower  | -              | 1      | -     | 1     |
|                  |         | Normal | -              | 447    | 1     | 448   |
|                  |         | Upper  | -              | 3      | -     | 3     |
|                  |         | Total  | -              | 451    | 1     | 452   |
|                  | Week 40 | Lower  | -              | 1      | -     | 1     |
|                  |         | Normal | -              | 441    | 1     | 442   |
|                  |         | Upper  | -              | 2      | -     | 2     |
|                  |         | Total  | -              | 444    | 1     | 445   |
|                  | Week 44 | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 438    | 1     | 439   |
|                  |         | Upper  | -              | 3      | -     | 3     |
|                  |         | Total  | -              | 441    | 1     | 442   |
|                  | Week 48 | Lower  | -              | -      | -     | -     |
|                  |         | Normal | -              | 434    | 1     | 435   |
|                  |         | Upper  | -              | 4      | -     | 4     |
|                  |         | Total  | -              | 438    | 1     | 439   |
| Week 52          | Lower   | -      | -              | -      | -     |       |
|                  | Normal  | -      | 435            | 1      | 436   |       |
|                  | Upper   | -      | 3              | -      | 3     |       |
|                  | Total   | -      | 438            | 1      | 439   |       |

表 2.7.4.7-付録-35 尿検査の評価時点別クロス表 (OCT-002 試験) (2/6)

|         |         |        | ATL-962 120 mg |        |       |       |
|---------|---------|--------|----------------|--------|-------|-------|
|         |         |        | Week 0         |        |       |       |
|         |         |        | Lower          | Normal | Upper | Total |
| pH      | Week 2  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 503    | -     | 503   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | -              | 504    | -     | 504   |
|         | Week 4  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 495    | -     | 495   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 495    | -     | 495   |
|         | Week 8  | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 489    | -     | 489   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 489    | -     | 489   |
|         | Week 12 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 479    | -     | 479   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 479    | -     | 479   |
|         | Week 16 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 472    | -     | 472   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 472    | -     | 472   |
|         | Week 20 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 469    | -     | 469   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 469    | -     | 469   |
|         | Week 24 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 461    | -     | 461   |
|         |         | Upper  | -              | 1      | -     | 1     |
|         |         | Total  | -              | 462    | -     | 462   |
|         | Week 28 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 453    | -     | 453   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 453    | -     | 453   |
|         | Week 32 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 456    | -     | 456   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 456    | -     | 456   |
|         | Week 36 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 452    | -     | 452   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 452    | -     | 452   |
|         | Week 40 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 445    | -     | 445   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 445    | -     | 445   |
|         | Week 44 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 442    | -     | 442   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 442    | -     | 442   |
|         | Week 48 | Lower  | -              | -      | -     | -     |
|         |         | Normal | -              | 439    | -     | 439   |
|         |         | Upper  | -              | -      | -     | -     |
|         |         | Total  | -              | 439    | -     | 439   |
| Week 52 | Lower   | -      | -              | -      | -     |       |
|         | Normal  | -      | 439            | -      | 439   |       |
|         | Upper   | -      | -              | -      | -     |       |
|         | Total   | -      | 439            | -      | 439   |       |



表 2.7.4.7-付録-35 尿検査の評価時点別クロス表 (OCT-002 試験) (3/6)

Glucose

| Frequency |                | Week 0  |          |       |    |     |
|-----------|----------------|---------|----------|-------|----|-----|
|           |                | Normal  | Abnormal | Total |    |     |
| Treatment | ATL-962 120 mg | Week 2  | Normal   | 405   | 39 | 444 |
|           |                |         | Abnormal | 9     | 51 | 60  |
|           |                |         | Total    | 414   | 90 | 504 |
|           |                | Week 4  | Normal   | 399   | 42 | 441 |
|           |                |         | Abnormal | 7     | 47 | 54  |
|           |                |         | Total    | 406   | 89 | 495 |
|           |                | Week 8  | Normal   | 386   | 43 | 429 |
|           |                |         | Abnormal | 15    | 45 | 60  |
|           |                |         | Total    | 401   | 88 | 489 |
|           |                | Week 12 | Normal   | 377   | 43 | 420 |
|           |                |         | Abnormal | 16    | 43 | 59  |
|           |                |         | Total    | 393   | 86 | 479 |
|           |                | Week 16 | Normal   | 370   | 39 | 409 |
|           |                |         | Abnormal | 18    | 45 | 63  |
|           |                |         | Total    | 388   | 84 | 472 |
|           |                | Week 20 | Normal   | 366   | 37 | 403 |
|           |                |         | Abnormal | 20    | 46 | 66  |
|           |                |         | Total    | 386   | 83 | 469 |
|           |                | Week 24 | Normal   | 355   | 40 | 395 |
|           |                |         | Abnormal | 26    | 41 | 67  |
|           |                |         | Total    | 381   | 81 | 462 |
|           |                | Week 28 | Normal   | 353   | 34 | 387 |
|           |                |         | Abnormal | 20    | 46 | 66  |
|           |                |         | Total    | 373   | 80 | 453 |
|           |                | Week 32 | Normal   | 354   | 35 | 389 |
|           |                |         | Abnormal | 21    | 46 | 67  |
|           |                |         | Total    | 375   | 81 | 456 |
|           |                | Week 36 | Normal   | 355   | 40 | 395 |
|           |                |         | Abnormal | 17    | 40 | 57  |
|           |                |         | Total    | 372   | 80 | 452 |
|           |                | Week 40 | Normal   | 341   | 37 | 378 |
|           |                |         | Abnormal | 26    | 41 | 67  |
|           |                |         | Total    | 367   | 78 | 445 |
|           |                | Week 44 | Normal   | 343   | 36 | 379 |
|           |                |         | Abnormal | 21    | 42 | 63  |
|           |                |         | Total    | 364   | 78 | 442 |
|           |                | Week 48 | Normal   | 343   | 43 | 386 |
|           |                |         | Abnormal | 18    | 35 | 53  |
|           |                |         | Total    | 361   | 78 | 439 |
|           |                | Week 52 | Normal   | 337   | 38 | 375 |
|           |                |         | Abnormal | 24    | 40 | 64  |
|           |                |         | Total    | 361   | 78 | 439 |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-35 尿検査の評価時点別クロス表 (OCT-002 試験) (4/6)

| Protein   |                | Frequency |          |     | Week 0 |          |       |
|-----------|----------------|-----------|----------|-----|--------|----------|-------|
|           |                |           |          |     | Normal | Abnormal | Total |
| Treatment | ATL-962 120 mg | Week 2    | Normal   | 334 | 52     | 386      |       |
|           |                |           | Abnormal | 30  | 88     | 118      |       |
|           |                |           | Total    | 364 | 140    | 504      |       |
|           |                | Week 4    | Normal   | 327 | 41     | 368      |       |
|           |                |           | Abnormal | 30  | 97     | 127      |       |
|           |                |           | Total    | 357 | 138    | 495      |       |
|           |                | Week 8    | Normal   | 314 | 52     | 366      |       |
|           |                |           | Abnormal | 37  | 86     | 123      |       |
|           |                |           | Total    | 351 | 138    | 489      |       |
|           |                | Week 12   | Normal   | 311 | 43     | 354      |       |
|           |                |           | Abnormal | 33  | 92     | 125      |       |
|           |                |           | Total    | 344 | 135    | 479      |       |
|           |                | Week 16   | Normal   | 301 | 50     | 351      |       |
|           |                |           | Abnormal | 38  | 83     | 121      |       |
|           |                |           | Total    | 339 | 133    | 472      |       |
|           |                | Week 20   | Normal   | 309 | 40     | 349      |       |
|           |                |           | Abnormal | 30  | 90     | 120      |       |
|           |                |           | Total    | 339 | 130    | 469      |       |
|           |                | Week 24   | Normal   | 305 | 41     | 346      |       |
|           |                |           | Abnormal | 31  | 85     | 116      |       |
|           |                |           | Total    | 336 | 126    | 462      |       |
|           |                | Week 28   | Normal   | 295 | 39     | 334      |       |
|           |                |           | Abnormal | 35  | 84     | 119      |       |
|           |                |           | Total    | 330 | 123    | 453      |       |
|           |                | Week 32   | Normal   | 298 | 46     | 344      |       |
|           |                |           | Abnormal | 35  | 77     | 112      |       |
|           |                |           | Total    | 333 | 123    | 456      |       |
|           |                | Week 36   | Normal   | 287 | 39     | 326      |       |
|           |                |           | Abnormal | 41  | 85     | 126      |       |
|           |                |           | Total    | 328 | 124    | 452      |       |
|           |                | Week 40   | Normal   | 290 | 44     | 334      |       |
|           |                |           | Abnormal | 34  | 77     | 111      |       |
|           |                |           | Total    | 324 | 121    | 445      |       |
|           |                | Week 44   | Normal   | 279 | 43     | 322      |       |
|           |                |           | Abnormal | 43  | 77     | 120      |       |
|           |                |           | Total    | 322 | 120    | 442      |       |
|           |                | Week 48   | Normal   | 281 | 44     | 325      |       |
|           |                |           | Abnormal | 39  | 75     | 114      |       |
|           |                |           | Total    | 320 | 119    | 439      |       |
|           |                | Week 52   | Normal   | 284 | 40     | 324      |       |
|           |                |           | Abnormal | 36  | 79     | 115      |       |
|           |                |           | Total    | 320 | 119    | 439      |       |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-35 尿検査の評価時点別クロス表 (OCT-002 試験) (5/6)

Ketones

| Frequency |                | Week 0  |          |       |    |     |
|-----------|----------------|---------|----------|-------|----|-----|
|           |                | Normal  | Abnormal | Total |    |     |
| Treatment | ATL-962 120 mg | Week 2  | Normal   | 444   | 28 | 472 |
|           |                |         | Abnormal | 20    | 12 | 32  |
|           |                |         | Total    | 464   | 40 | 504 |
|           |                | Week 4  | Normal   | 437   | 27 | 464 |
|           |                |         | Abnormal | 18    | 13 | 31  |
|           |                |         | Total    | 455   | 40 | 495 |
|           |                | Week 8  | Normal   | 430   | 30 | 460 |
|           |                |         | Abnormal | 19    | 10 | 29  |
|           |                |         | Total    | 449   | 40 | 489 |
|           |                | Week 12 | Normal   | 424   | 30 | 454 |
|           |                |         | Abnormal | 15    | 10 | 25  |
|           |                |         | Total    | 439   | 40 | 479 |
|           |                | Week 16 | Normal   | 411   | 28 | 439 |
|           |                |         | Abnormal | 21    | 12 | 33  |
|           |                |         | Total    | 432   | 40 | 472 |
|           |                | Week 20 | Normal   | 417   | 30 | 447 |
|           |                |         | Abnormal | 14    | 8  | 22  |
|           |                |         | Total    | 431   | 38 | 469 |
|           |                | Week 24 | Normal   | 408   | 26 | 434 |
|           |                |         | Abnormal | 17    | 11 | 28  |
|           |                |         | Total    | 425   | 37 | 462 |
|           |                | Week 28 | Normal   | 400   | 27 | 427 |
|           |                |         | Abnormal | 17    | 9  | 26  |
|           |                |         | Total    | 417   | 36 | 453 |
|           |                | Week 32 | Normal   | 397   | 26 | 423 |
|           |                |         | Abnormal | 22    | 11 | 33  |
|           |                |         | Total    | 419   | 37 | 456 |
|           |                | Week 36 | Normal   | 393   | 24 | 417 |
|           |                |         | Abnormal | 22    | 13 | 35  |
|           |                |         | Total    | 415   | 37 | 452 |
|           |                | Week 40 | Normal   | 382   | 26 | 408 |
|           |                |         | Abnormal | 26    | 11 | 37  |
|           |                |         | Total    | 408   | 37 | 445 |
|           |                | Week 44 | Normal   | 380   | 24 | 404 |
|           |                |         | Abnormal | 25    | 13 | 38  |
|           |                |         | Total    | 405   | 37 | 442 |
|           |                | Week 48 | Normal   | 371   | 23 | 394 |
|           |                |         | Abnormal | 31    | 14 | 45  |
|           |                |         | Total    | 402   | 37 | 439 |
|           |                | Week 52 | Normal   | 376   | 21 | 397 |
|           |                |         | Abnormal | 26    | 16 | 42  |
|           |                |         | Total    | 402   | 37 | 439 |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-35 尿検査の評価時点別クロス表 (OCT-002 試験) (6/6)

Occult Blood

| Frequency |                | Week 0  |          |       |    |     |
|-----------|----------------|---------|----------|-------|----|-----|
|           |                | Normal  | Abnormal | Total |    |     |
| Treatment | ATL-962 120 mg | Week 2  | Normal   | 422   | 19 | 441 |
|           |                |         | Abnormal | 21    | 42 | 63  |
|           |                |         | Total    | 443   | 61 | 504 |
|           |                | Week 4  | Normal   | 412   | 19 | 431 |
|           |                |         | Abnormal | 24    | 40 | 64  |
|           |                |         | Total    | 436   | 59 | 495 |
|           |                | Week 8  | Normal   | 409   | 22 | 431 |
|           |                |         | Abnormal | 21    | 37 | 58  |
|           |                |         | Total    | 430   | 59 | 489 |
|           |                | Week 12 | Normal   | 398   | 18 | 416 |
|           |                |         | Abnormal | 22    | 41 | 63  |
|           |                |         | Total    | 420   | 59 | 479 |
|           |                | Week 16 | Normal   | 382   | 19 | 401 |
|           |                |         | Abnormal | 32    | 39 | 71  |
|           |                |         | Total    | 414   | 58 | 472 |
|           |                | Week 20 | Normal   | 382   | 15 | 397 |
|           |                |         | Abnormal | 28    | 44 | 72  |
|           |                |         | Total    | 410   | 59 | 469 |
|           |                | Week 24 | Normal   | 372   | 26 | 398 |
|           |                |         | Abnormal | 32    | 32 | 64  |
|           |                |         | Total    | 404   | 58 | 462 |
|           |                | Week 28 | Normal   | 361   | 21 | 382 |
|           |                |         | Abnormal | 34    | 37 | 71  |
|           |                |         | Total    | 395   | 58 | 453 |
|           |                | Week 32 | Normal   | 368   | 24 | 392 |
|           |                |         | Abnormal | 30    | 34 | 64  |
|           |                |         | Total    | 398   | 58 | 456 |
|           |                | Week 36 | Normal   | 365   | 20 | 385 |
|           |                |         | Abnormal | 30    | 37 | 67  |
|           |                |         | Total    | 395   | 57 | 452 |
|           |                | Week 40 | Normal   | 359   | 23 | 382 |
|           |                |         | Abnormal | 29    | 34 | 63  |
|           |                |         | Total    | 388   | 57 | 445 |
|           |                | Week 44 | Normal   | 356   | 24 | 380 |
|           |                |         | Abnormal | 31    | 31 | 62  |
|           |                |         | Total    | 387   | 55 | 442 |
|           |                | Week 48 | Normal   | 359   | 22 | 381 |
|           |                |         | Abnormal | 26    | 32 | 58  |
|           |                |         | Total    | 385   | 54 | 439 |
|           |                | Week 52 | Normal   | 354   | 20 | 374 |
|           |                |         | Abnormal | 31    | 34 | 65  |
|           |                |         | Total    | 385   | 54 | 439 |

"Abnormal" is used for this summary table in place of "Upper".

表 2.7.4.7-付録-36 バイタルサイン（収縮期血圧）の評価時点別の変化量（CCT-001 試験）

| Variable / Visit   | Treatment | Summary Statistics |      |       |       |       |        |      |     |       | 95% CI |         | One-Sample t-test |        |
|--|-----------|--------------------|------|-------|-------|-------|--------|------|-----|-------|--------|---------|-------------------|--------|
|  |           | N                  | Mean | SD    | Min   | Q1    | Median | Q3   | Max | Lower | Upper  | t-value | p-value           |        |
| Change from Baseline<br>in Systolic Blood Pressure<br>(mmHg) | Week 2    | Placebo            | 98   | 0.6   | 13.58 | -46   | -8.0   | 0.0  | 8.0 | 34    | -2.15  | 3.29    | 0.4165            | 0.6780 |
|  |           | 15 mg              | 89   | -1.8  | 12.76 | -33   | -8.0   | -1.0 | 5.0 | 33    | -4.52  | 0.86    | -1.3536           | 0.1793 |
|  |           | 30 mg              | 92   | -0.8  | 13.32 | -39   | -8.0   | 0.0  | 7.0 | 34    | -3.57  | 1.94    | -0.5869           | 0.5587 |
|  |           | 60 mg              | 93   | -3.3  | 16.15 | -42   | -12.0  | -3.0 | 8.0 | 52    | -6.61  | 0.05    | -1.9587           | 0.0532 |
|  |           | 120 mg             | 89   | -1.8  | 12.89 | -33   | -9.0   | -2.0 | 6.0 | 28    | -4.51  | 0.92    | -1.3160           | 0.1916 |
|  | Week 4    | Placebo            | 96   | -0.7  | 13.89 | -44   | -7.0   | 0.0  | 6.5 | 39    | -3.54  | 2.09    | -0.5143           | 0.6082 |
|  |           | 15 mg              | 89   | -1.9  | 12.26 | -32   | -9.0   | -2.0 | 5.0 | 28    | -4.47  | 0.69    | -1.4531           | 0.1498 |
|  |           | 30 mg              | 91   | -2.5  | 12.76 | -42   | -10.0  | -1.0 | 6.0 | 25    | -5.18  | 0.13    | -1.8897           | 0.0620 |
|  |           | 60 mg              | 90   | -3.0  | 14.29 | -32   | -14.0  | -3.5 | 7.0 | 32    | -6.03  | -0.04   | -2.0138           | 0.0470 |
|  |           | 120 mg             | 88   | -2.0  | 13.10 | -40   | -9.5   | -1.0 | 5.0 | 26    | -4.79  | 0.77    | -1.4398           | 0.1535 |
|  | Week 8    | Placebo            | 95   | -2.0  | 14.67 | -50   | -10.0  | -1.0 | 4.0 | 54    | -4.95  | 1.03    | -1.3005           | 0.1966 |
|  |           | 15 mg              | 87   | -1.2  | 11.63 | -30   | -8.0   | -3.0 | 5.0 | 40    | -3.66  | 1.30    | -0.9493           | 0.3451 |
|  |           | 30 mg              | 89   | -2.8  | 12.22 | -34   | -10.0  | -2.0 | 6.0 | 28    | -5.42  | -0.27   | -2.1945           | 0.0308 |
|  |           | 60 mg              | 86   | -3.8  | 13.77 | -36   | -13.0  | -3.5 | 6.0 | 34    | -6.71  | -0.80   | -2.5287           | 0.0133 |
|  |           | 120 mg             | 85   | -2.9  | 13.65 | -36   | -12.0  | -2.0 | 6.0 | 33    | -5.81  | 0.07    | -1.9388           | 0.0559 |
|  | Week 12   | Placebo            | 95   | -1.5  | 14.77 | -38   | -9.0   | 0.0  | 5.0 | 42    | -4.49  | 1.53    | -0.9792           | 0.3300 |
|  |           | 15 mg              | 86   | -2.3  | 15.72 | -36   | -10.0  | 0.5  | 7.0 | 65    | -5.72  | 1.02    | -1.3852           | 0.1696 |
|  |           | 30 mg              | 86   | -0.7  | 13.50 | -36   | -10.0  | 0.5  | 9.0 | 27    | -3.63  | 2.16    | -0.5031           | 0.6162 |
|  |           | 60 mg              | 82   | -4.9  | 15.60 | -38   | -16.0  | -4.0 | 4.0 | 38    | -8.28  | -1.43   | -2.8170           | 0.0061 |
|  |           | 120 mg             | 83   | -0.4  | 13.75 | -28   | -10.0  | -2.0 | 8.0 | 44    | -3.45  | 2.56    | -0.2953           | 0.7685 |
| Week 16  | Placebo   | 95                 | -1.0 | 13.96 | -34   | -10.0 | 0.0    | 8.0  | 52  | -3.86 | 1.82   | -0.7131 | 0.4775            |        |
|  | 15 mg     | 84                 | -3.4 | 15.48 | -54   | -11.0 | -2.0   | 6.0  | 47  | -6.73 | -0.01  | -1.9945 | 0.0494            |        |
|  | 30 mg     | 85                 | -1.2 | 12.80 | -33   | -11.0 | -2.0   | 9.0  | 24  | -3.91 | 1.61   | -0.8302 | 0.4088            |        |
|  | 60 mg     | 80                 | -5.2 | 15.92 | -39   | -17.5 | -5.0   | 4.0  | 34  | -8.78 | -1.69  | -2.9427 | 0.0043            |        |
|  | 120 mg    | 82                 | -1.3 | 11.84 | -37   | -10.0 | 0.0    | 6.0  | 38  | -3.92 | 1.28   | -1.0075 | 0.3167            |        |
| Week 20  | Placebo   | 95                 | -1.6 | 15.14 | -46   | -11.0 | -2.0   | 8.0  | 50  | -4.73 | 1.44   | -1.0570 | 0.2932            |        |
|  | 15 mg     | 84                 | -2.8 | 13.96 | -33   | -12.0 | -3.5   | 5.0  | 41  | -5.88 | 0.19   | -1.8674 | 0.0654            |        |
|  | 30 mg     | 85                 | -1.7 | 16.06 | -46   | -10.0 | -1.0   | 8.0  | 54  | -5.21 | 1.72   | -0.9995 | 0.3204            |        |
|  | 60 mg     | 79                 | -4.0 | 15.24 | -33   | -16.0 | -6.0   | 6.0  | 36  | -7.45 | -0.62  | -2.3553 | 0.0210            |        |
|  | 120 mg    | 80                 | -1.8 | 12.97 | -41   | -8.5  | -2.0   | 5.0  | 40  | -4.72 | 1.05   | -1.2674 | 0.2087            |        |
| Week 24  | Placebo   | 94                 | -2.0 | 13.39 | -42   | -10.0 | -1.0   | 6.0  | 37  | -4.72 | 0.76   | -1.4327 | 0.1553            |        |
|  | 15 mg     | 84                 | -2.1 | 13.25 | -32   | -10.5 | -4.0   | 9.5  | 51  | -4.98 | 0.77   | -1.4576 | 0.1487            |        |
|  | 30 mg     | 84                 | -1.8 | 12.86 | -30   | -10.0 | -2.5   | 6.5  | 43  | -4.61 | 0.97   | -1.2985 | 0.1977            |        |
|  | 60 mg     | 78                 | -6.2 | 13.48 | -40   | -16.0 | -6.0   | 4.0  | 23  | -9.19 | -3.11  | -4.0306 | 0.0001            |        |
|  | 120 mg    | 79                 | -0.9 | 13.53 | -32   | -11.0 | 0.0    | 9.0  | 33  | -3.97 | 2.09   | -0.6152 | 0.5402            |        |
| Week 28  | Placebo   | 94                 | -0.2 | 14.61 | -43   | -9.0  | -0.5   | 9.0  | 59  | -3.20 | 2.78   | -0.1412 | 0.8880            |        |
|  | 15 mg     | 83                 | -2.6 | 15.13 | -38   | -12.0 | -3.0   | 6.0  | 51  | -5.92 | 0.69   | -1.5745 | 0.1192            |        |
|  | 30 mg     | 84                 | -0.8 | 13.64 | -28   | -10.0 | 0.0    | 8.0  | 40  | -3.73 | 2.19   | -0.5200 | 0.6044            |        |
|  | 60 mg     | 77                 | -4.2 | 13.82 | -36   | -14.0 | -4.0   | 6.0  | 27  | -7.33 | -1.06  | -2.6632 | 0.0094            |        |
|  | 120 mg    | 78                 | -1.2 | 13.71 | -34   | -9.0  | -2.0   | 5.0  | 45  | -4.32 | 1.86   | -0.7927 | 0.4304            |        |
| Follow-up  | Placebo   | 98                 | -0.2 | 14.93 | -43   | -9.0  | 0.0    | 9.0  | 59  | -3.22 | 2.77   | -0.1488 | 0.8820            |        |
|  | 15 mg     | 86                 | -2.2 | 15.22 | -38   | -11.0 | -2.5   | 6.0  | 51  | -5.49 | 1.03   | -1.3607 | 0.1772            |        |
|  | 30 mg     | 92                 | -1.2 | 14.09 | -35   | -11.0 | 0.0    | 8.0  | 40  | -4.13 | 1.70   | -0.8289 | 0.4093            |        |
|  | 60 mg     | 87                 | -4.2 | 13.53 | -36   | -14.0 | -2.0   | 6.0  | 27  | -7.04 | -1.28  | -2.8681 | 0.0052            |        |
|  | 120 mg    | 88                 | -1.0 | 13.57 | -34   | -9.5  | -1.5   | 5.5  | 45  | -3.92 | 1.83   | -0.7228 | 0.4717            |        |

表 2.7.4.7-付録-37 バイタルサイン（拡張期血圧）の評価時点別の変化量（CCT-001 試験）

| Variable / Visit   | Treatment | Summary Statistics |      |      |       |     |        |      |     |       | 95% CI |         | One-Sample t-test |        |
|--|-----------|--------------------|------|------|-------|-----|--------|------|-----|-------|--------|---------|-------------------|--------|
|  |           | N                  | Mean | SD   | Min   | Q1  | Median | Q3   | Max | Lower | Upper  | t-value | p-value           |        |
| Change from Baseline<br>in Diastolic Blood<br>Pressure<br>(mmHg) | Week 2    | Placebo            | 98   | 0.0  | 8.25  | -19 | -4.0   | 0.0  | 4.0 | 33    | -1.62  | 1.69    | 0.0367            | 0.9708 |
|  |           | 15 mg              | 89   | -1.3 | 9.02  | -29 | -8.0   | -2.0 | 4.0 | 24    | -3.16  | 0.64    | -1.3161           | 0.1916 |
|  |           | 30 mg              | 92   | 0.0  | 9.17  | -30 | -5.5   | 0.0  | 5.5 | 30    | -1.88  | 1.92    | 0.0228            | 0.9819 |
|  |           | 60 mg              | 93   | -0.2 | 9.43  | -30 | -6.0   | -1.0 | 6.0 | 28    | -2.16  | 1.73    | -0.2200           | 0.8263 |
|  |           | 120 mg             | 89   | -0.8 | 8.92  | -22 | -6.0   | -1.0 | 4.0 | 20    | -2.64  | 1.11    | -0.8085           | 0.4210 |
|  | Week 4    | Placebo            | 96   | -0.9 | 8.40  | -28 | -5.0   | 0.0  | 4.5 | 28    | -2.63  | 0.78    | -1.0808           | 0.2825 |
|  |           | 15 mg              | 89   | -1.5 | 8.53  | -24 | -6.0   | -2.0 | 2.0 | 24    | -3.32  | 0.27    | -1.6901           | 0.0945 |
|  |           | 30 mg              | 91   | -0.3 | 7.40  | -24 | -4.0   | 0.0  | 5.0 | 17    | -1.80  | 1.28    | -0.3400           | 0.7346 |
|  |           | 60 mg              | 90   | -1.1 | 9.08  | -23 | -6.0   | -1.0 | 4.0 | 28    | -2.99  | 0.81    | -1.1371           | 0.2586 |
|  |           | 120 mg             | 88   | -2.0 | 8.48  | -26 | -8.0   | -3.0 | 4.0 | 14    | -3.77  | -0.18   | -2.1884           | 0.0313 |
|  | Week 8    | Placebo            | 95   | -3.0 | 7.86  | -28 | -7.0   | -3.0 | 2.0 | 16    | -4.63  | -1.43   | -3.7594           | 0.0003 |
|  |           | 15 mg              | 87   | -1.7 | 9.00  | -23 | -8.0   | -2.0 | 4.0 | 26    | -3.60  | 0.24    | -1.7388           | 0.0856 |
|  |           | 30 mg              | 89   | -1.6 | 9.09  | -25 | -7.0   | -1.0 | 4.0 | 24    | -3.49  | 0.34    | -1.6323           | 0.1062 |
|  |           | 60 mg              | 86   | -0.4 | 8.69  | -22 | -6.0   | 0.0  | 6.0 | 20    | -2.26  | 1.47    | -0.4220           | 0.6741 |
|  |           | 120 mg             | 85   | -1.0 | 9.24  | -30 | -6.0   | -2.0 | 5.0 | 20    | -2.97  | 1.02    | -0.9742           | 0.3328 |
|  | Week 12   | Placebo            | 95   | -1.5 | 8.48  | -25 | -7.0   | -2.0 | 5.0 | 22    | -3.19  | 0.26    | -1.6812           | 0.0960 |
|  |           | 15 mg              | 86   | -1.6 | 9.28  | -28 | -6.0   | -1.0 | 4.0 | 26    | -3.57  | 0.41    | -1.5796           | 0.1179 |
|  |           | 30 mg              | 86   | -0.4 | 8.99  | -20 | -6.0   | 0.0  | 6.0 | 24    | -2.33  | 1.52    | -0.4198           | 0.6757 |
|  |           | 60 mg              | 82   | -2.3 | 8.26  | -23 | -8.0   | -3.0 | 4.0 | 16    | -4.16  | -0.53   | -2.5660           | 0.0121 |
|  |           | 120 mg             | 83   | -0.7 | 10.56 | -27 | -7.0   | 0.0  | 6.0 | 27    | -3.00  | 1.61    | -0.6030           | 0.5482 |
|  | Week 16   | Placebo            | 95   | -1.9 | 7.70  | -24 | -8.0   | -1.0 | 2.0 | 15    | -3.47  | -0.34   | -2.4115           | 0.0178 |
|  |           | 15 mg              | 84   | -3.0 | 12.48 | -42 | -9.5   | -2.5 | 4.0 | 24    | -5.72  | -0.30   | -2.2123           | 0.0297 |
|  |           | 30 mg              | 85   | -1.8 | 8.72  | -22 | -7.0   | -2.0 | 2.0 | 23    | -3.70  | 0.06    | -1.9287           | 0.0571 |
|  |           | 60 mg              | 80   | -2.4 | 10.84 | -32 | -9.0   | -2.0 | 4.0 | 28    | -4.81  | 0.01    | -1.9803           | 0.0512 |
|  |           | 120 mg             | 82   | -1.4 | 9.70  | -26 | -8.0   | 0.0  | 4.0 | 20    | -3.52  | 0.74    | -1.2980           | 0.1980 |
|  | Week 20   | Placebo            | 95   | -1.6 | 8.44  | -28 | -6.0   | -1.0 | 4.0 | 18    | -3.28  | 0.16    | -1.7997           | 0.0751 |
|  |           | 15 mg              | 84   | -1.4 | 9.38  | -33 | -6.0   | -1.0 | 2.5 | 30    | -3.44  | 0.63    | -1.3719           | 0.1738 |
|  |           | 30 mg              | 85   | -0.3 | 9.37  | -20 | -6.0   | -1.0 | 6.0 | 26    | -2.29  | 1.75    | -0.2662           | 0.7907 |
|  |           | 60 mg              | 79   | -2.8 | 8.68  | -21 | -8.0   | -4.0 | 2.0 | 27    | -4.78  | -0.89   | -2.9049           | 0.0048 |
|  |           | 120 mg             | 80   | -2.4 | 10.52 | -30 | -8.5   | -2.0 | 5.0 | 25    | -4.75  | -0.07   | -2.0518           | 0.0435 |
|  | Week 24   | Placebo            | 94   | -2.0 | 8.80  | -26 | -8.0   | -2.0 | 4.0 | 29    | -3.85  | -0.24   | -2.2501           | 0.0268 |
|  |           | 15 mg              | 84   | -1.9 | 10.42 | -35 | -9.0   | -2.0 | 4.0 | 30    | -4.17  | 0.36    | -1.6754           | 0.0976 |
|  |           | 30 mg              | 84   | -2.7 | 9.56  | -28 | -8.0   | -2.0 | 2.5 | 25    | -4.73  | -0.58   | -2.5462           | 0.0127 |
|  |           | 60 mg              | 78   | -3.0 | 9.15  | -20 | -9.0   | -4.5 | 4.0 | 28    | -5.06  | -0.94   | -2.8949           | 0.0049 |
|  |           | 120 mg             | 79   | -1.8 | 10.83 | -34 | -10.0  | -2.0 | 6.0 | 29    | -4.26  | 0.59    | -1.5070           | 0.1359 |
|  | Week 28   | Placebo            | 94   | -1.1 | 9.61  | -25 | -7.0   | -1.0 | 4.0 | 21    | -3.11  | 0.83    | -1.1480           | 0.2539 |
|  |           | 15 mg              | 83   | -1.7 | 11.37 | -33 | -8.0   | -3.0 | 4.0 | 40    | -4.18  | 0.78    | -1.3614           | 0.1771 |
|  |           | 30 mg              | 84   | -1.2 | 9.50  | -21 | -8.0   | -1.5 | 4.5 | 25    | -3.26  | 0.86    | -1.1601           | 0.2493 |
|  |           | 60 mg              | 77   | -2.1 | 9.41  | -19 | -7.0   | -4.0 | 4.0 | 24    | -4.24  | 0.03    | -1.9626           | 0.0534 |
|  |           | 120 mg             | 78   | -2.3 | 10.01 | -34 | -8.0   | -3.0 | 2.0 | 20    | -4.60  | -0.09   | -2.0702           | 0.0418 |
|  | Follow-up | Placebo            | 98   | -1.1 | 9.53  | -25 | -7.0   | -1.0 | 4.0 | 21    | -2.99  | 0.83    | -1.1232           | 0.2641 |
|  |           | 15 mg              | 86   | -1.3 | 11.38 | -33 | -8.0   | -2.5 | 5.0 | 40    | -3.79  | 1.09    | -1.0989           | 0.2749 |
|  |           | 30 mg              | 92   | -1.5 | 9.30  | -21 | -8.0   | -2.0 | 4.0 | 25    | -3.43  | 0.43    | -1.5464           | 0.1255 |
|  |           | 60 mg              | 87   | -1.6 | 9.16  | -19 | -7.0   | -2.0 | 4.0 | 24    | -3.56  | 0.34    | -1.6387           | 0.1049 |
|  |           | 120 mg             | 88   | -2.4 | 9.60  | -34 | -8.0   | -3.0 | 2.0 | 20    | -4.41  | -0.34   | -2.3213           | 0.0226 |

表 2.7.4.7-付録-38 バイタルサイン（脈拍数）の評価時点別の変化量（CCT-001 試験）

| Item                      | Treatment | Summary Statistics |      |                      |      |       |     |        |      |     |         | One-sample t-test |        |
|---------------------------|-----------|--------------------|------|----------------------|------|-------|-----|--------|------|-----|---------|-------------------|--------|
|                           |           | Baseline           |      | Change from Baseline |      |       |     |        |      |     | t-value | p-value           |        |
|                           |           | Mean               | N    | Mean                 | SD   | Min   | Q1  | Median | Q3   | Max |         |                   |        |
| Pulse Rate<br>(beats/min) | Week 2    | Placebo            | 75.8 | 98                   | -1.7 | 7.70  | -20 | -7.0   | -1.0 | 3.0 | 25      | -2.1262           | 0.0360 |
|                           |           | 15 mg              | 76.8 | 89                   | -0.0 | 9.49  | -30 | -5.0   | 0.0  | 6.0 | 23      | -0.0335           | 0.9733 |
|                           |           | 30 mg              | 76.2 | 92                   | -0.7 | 8.37  | -20 | -5.5   | -1.0 | 2.0 | 25      | -0.7972           | 0.4274 |
|                           |           | 60 mg              | 74.1 | 93                   | 1.7  | 7.07  | -15 | -2.0   | 2.0  | 6.0 | 19      | 2.3029            | 0.0235 |
|                           |           | 120 mg             | 74.1 | 89                   | -0.0 | 8.38  | -22 | -5.0   | 0.0  | 5.0 | 20      | -0.0253           | 0.9799 |
|                           | Week 4    | Placebo            | 75.9 | 96                   | -1.5 | 8.01  | -32 | -6.0   | 0.0  | 4.0 | 14      | -1.8487           | 0.0676 |
|                           |           | 15 mg              | 76.8 | 89                   | -0.4 | 9.58  | -38 | -6.0   | 1.0  | 6.0 | 18      | -0.4203           | 0.6753 |
|                           |           | 30 mg              | 76.0 | 91                   | -0.9 | 8.26  | -31 | -6.0   | -1.0 | 4.0 | 19      | -1.0666           | 0.2890 |
|                           |           | 60 mg              | 74.6 | 90                   | 0.2  | 7.48  | -26 | -4.0   | 0.0  | 6.0 | 18      | 0.2959            | 0.7680 |
|                           |           | 120 mg             | 74.2 | 88                   | -0.8 | 8.05  | -23 | -5.5   | 0.0  | 4.5 | 20      | -0.9140           | 0.3633 |
|                           | Week 8    | Placebo            | 75.9 | 95                   | -1.3 | 7.42  | -26 | -6.0   | 0.0  | 4.0 | 12      | -1.7426           | 0.0847 |
|                           |           | 15 mg              | 76.6 | 87                   | -0.5 | 9.26  | -30 | -5.0   | -1.0 | 5.0 | 17      | -0.5442           | 0.5877 |
|                           |           | 30 mg              | 76.4 | 89                   | -0.9 | 8.66  | -18 | -6.0   | -1.0 | 4.0 | 31      | -0.9543           | 0.3425 |
|                           |           | 60 mg              | 74.4 | 86                   | 2.7  | 8.95  | -22 | -1.0   | 4.0  | 8.0 | 30      | 2.8322            | 0.0058 |
|                           |           | 120 mg             | 74.2 | 85                   | 0.2  | 10.23 | -23 | -6.0   | 0.0  | 5.0 | 39      | 0.1484            | 0.8824 |
|                           | Week 12   | Placebo            | 75.9 | 95                   | -2.1 | 9.94  | -32 | -6.0   | -2.0 | 4.0 | 26      | -2.0130           | 0.0470 |
|                           |           | 15 mg              | 76.3 | 86                   | -2.4 | 8.70  | -23 | -7.0   | -2.5 | 2.0 | 30      | -2.5415           | 0.0129 |
|                           |           | 30 mg              | 76.4 | 86                   | -2.2 | 8.39  | -21 | -7.0   | -2.5 | 3.0 | 21      | -2.4418           | 0.0167 |
|                           |           | 60 mg              | 74.8 | 82                   | 0.3  | 9.47  | -32 | -5.0   | 0.0  | 5.0 | 34      | 0.2450            | 0.8071 |
|                           |           | 120 mg             | 73.8 | 83                   | -1.7 | 9.47  | -30 | -7.0   | -2.0 | 2.0 | 35      | -1.5880           | 0.1161 |
|                           | Week 16   | Placebo            | 75.9 | 95                   | -0.9 | 8.95  | -27 | -5.0   | 0.0  | 4.0 | 21      | -1.0086           | 0.3158 |
|                           |           | 15 mg              | 76.2 | 84                   | -2.8 | 10.17 | -33 | -9.5   | -1.5 | 5.0 | 17      | -2.5647           | 0.0121 |
|                           |           | 30 mg              | 76.4 | 85                   | -2.1 | 7.97  | -20 | -8.0   | -1.0 | 4.0 | 15      | -2.4351           | 0.0170 |
|                           |           | 60 mg              | 74.6 | 80                   | -0.7 | 8.22  | -32 | -5.0   | -1.0 | 5.5 | 14      | -0.7890           | 0.4325 |
|                           |           | 120 mg             | 73.9 | 82                   | 0.7  | 9.97  | -25 | -6.0   | 0.0  | 6.0 | 30      | 0.6425            | 0.5223 |
|                           | Week 20   | Placebo            | 75.9 | 95                   | -0.8 | 9.82  | -32 | -6.0   | 0.0  | 5.0 | 18      | -0.7629           | 0.4474 |
|                           |           | 15 mg              | 76.2 | 84                   | -1.3 | 10.74 | -36 | -8.0   | -1.5 | 4.0 | 24      | -1.1279           | 0.2626 |
|                           |           | 30 mg              | 76.4 | 85                   | -0.6 | 7.80  | -21 | -5.0   | -2.0 | 5.0 | 22      | -0.7093           | 0.4801 |
|                           |           | 60 mg              | 74.7 | 79                   | -1.0 | 8.21  | -30 | -6.0   | -1.0 | 4.0 | 19      | -1.0410           | 0.3011 |
|                           |           | 120 mg             | 74.3 | 80                   | 0.1  | 9.62  | -20 | -5.0   | 0.0  | 2.5 | 28      | 0.0465            | 0.9630 |
|                           | Week 24   | Placebo            | 75.9 | 94                   | -1.2 | 8.78  | -26 | -6.0   | 0.0  | 4.0 | 26      | -1.3392           | 0.1838 |
|                           |           | 15 mg              | 76.2 | 84                   | -1.7 | 9.13  | -36 | -8.0   | -1.0 | 5.0 | 22      | -1.6611           | 0.1005 |
|                           |           | 30 mg              | 76.3 | 84                   | -1.9 | 8.13  | -19 | -7.0   | -2.5 | 3.0 | 23      | -2.1207           | 0.0369 |
|                           |           | 60 mg              | 74.8 | 78                   | -0.4 | 7.03  | -16 | -5.0   | 0.0  | 4.0 | 15      | -0.4995           | 0.6189 |
|                           |           | 120 mg             | 74.1 | 79                   | -0.1 | 10.37 | -23 | -7.0   | -1.0 | 5.0 | 34      | -0.0868           | 0.9311 |
|                           | Week 28   | Placebo            | 75.9 | 94                   | -0.7 | 9.18  | -26 | -6.0   | -1.0 | 5.0 | 22      | -0.7638           | 0.4469 |
|                           |           | 15 mg              | 76.4 | 83                   | -2.1 | 10.43 | -38 | -9.0   | -1.0 | 5.0 | 27      | -1.8315           | 0.0707 |
|                           |           | 30 mg              | 76.3 | 84                   | -2.0 | 7.97  | -16 | -8.0   | -2.0 | 3.0 | 20      | -2.3561           | 0.0208 |
|                           |           | 60 mg              | 74.8 | 77                   | -1.6 | 6.78  | -18 | -4.0   | 0.0  | 2.0 | 12      | -2.0172           | 0.0472 |
|                           |           | 120 mg             | 73.9 | 78                   | 1.5  | 10.77 | -21 | -5.0   | 0.0  | 6.0 | 42      | 1.2088            | 0.2305 |
|                           | Follow-up | Placebo            | 75.8 | 98                   | -0.7 | 9.20  | -26 | -6.0   | -1.0 | 5.0 | 22      | -0.7684           | 0.4441 |
|                           |           | 15 mg              | 77.0 | 86                   | -2.2 | 10.42 | -38 | -9.0   | -1.0 | 5.0 | 27      | -1.9246           | 0.0576 |
|                           |           | 30 mg              | 76.1 | 92                   | -1.9 | 7.87  | -16 | -7.0   | -2.0 | 3.0 | 20      | -2.3317           | 0.0219 |
|                           |           | 60 mg              | 74.9 | 87                   | -1.1 | 7.15  | -18 | -4.0   | 0.0  | 3.0 | 20      | -1.4101           | 0.1621 |
|                           |           | 120 mg             | 74.0 | 88                   | 1.2  | 10.71 | -22 | -5.0   | 0.0  | 6.0 | 42      | 1.0552            | 0.2943 |

表 2.7.4.7-付録-39 バイタルサイン（収縮期血圧）の評価時点別の変化量（CCT-002 試験）

| Systolic Blood Pressure<br>(mmHg) | Treatment      | Summary Statistics |      |       |     |       |        |      |     | 95% CI |       | One-Sample t-test |         |
|-----------------------------------|----------------|--------------------|------|-------|-----|-------|--------|------|-----|--------|-------|-------------------|---------|
|                                   |                | N                  | Mean | SD    | Min | Q1    | Median | Q3   | Max | Lower  | Upper | t-value           | p-value |
| Visit                             |                |                    |      |       |     |       |        |      |     |        |       |                   |         |
| Week 2                            | Placebo        | 65                 | 1.5  | 11.14 | -30 | -5.0  | 1.0    | 10.0 | 28  | -1.25  | 4.27  | 1.0907            | 0.2795  |
|                                   | ATL-962 120 mg | 140                | -0.9 | 12.22 | -46 | -8.0  | 0.0    | 6.0  | 36  | -2.91  | 1.18  | -0.8371           | 0.4040  |
| Week 4                            | Placebo        | 63                 | 4.4  | 12.07 | -30 | -2.0  | 5.0    | 10.0 | 41  | 1.36   | 7.44  | 2.8916            | 0.0053  |
|                                   | ATL-962 120 mg | 139                | -2.3 | 12.37 | -38 | -8.0  | -2.0   | 5.0  | 37  | -4.38  | -0.23 | -2.2003           | 0.0294  |
| Week 8                            | Placebo        | 62                 | 0.9  | 11.82 | -31 | -7.0  | 0.0    | 10.0 | 20  | -2.10  | 3.91  | 0.6017            | 0.5496  |
|                                   | ATL-962 120 mg | 137                | -1.6 | 12.56 | -48 | -9.0  | 0.0    | 6.0  | 31  | -3.69  | 0.55  | -1.4625           | 0.1459  |
| Week 12                           | Placebo        | 59                 | 0.6  | 12.66 | -23 | -8.0  | -2.0   | 9.0  | 36  | -2.72  | 3.87  | 0.3497            | 0.7278  |
|                                   | ATL-962 120 mg | 137                | -4.4 | 11.91 | -54 | -10.0 | -3.0   | 4.0  | 27  | -6.44  | -2.41 | -4.3482           | <0.0001 |
| Week 16                           | Placebo        | 59                 | 0.7  | 12.87 | -30 | -7.0  | 2.0    | 10.0 | 38  | -2.63  | 4.08  | 0.4350            | 0.6652  |
|                                   | ATL-962 120 mg | 134                | -2.0 | 11.18 | -39 | -10.0 | -1.0   | 4.0  | 30  | -3.93  | -0.11 | -2.0933           | 0.0382  |
| Week 20                           | Placebo        | 59                 | -0.5 | 12.69 | -38 | -10.0 | 0.0    | 8.0  | 38  | -3.82  | 2.80  | -0.3077           | 0.7594  |
|                                   | ATL-962 120 mg | 133                | -5.9 | 12.96 | -49 | -13.0 | -6.0   | 1.0  | 32  | -8.08  | -3.63 | -5.2121           | <0.0001 |
| Week 24                           | Placebo        | 59                 | -0.2 | 14.61 | -44 | -10.0 | 2.0    | 9.0  | 34  | -3.98  | 3.64  | -0.0891           | 0.9293  |
|                                   | ATL-962 120 mg | 130                | -4.2 | 13.88 | -41 | -13.0 | -2.0   | 4.0  | 33  | -6.65  | -1.84 | -3.4882           | 0.0007  |
| Week 28                           | Placebo        | 59                 | -1.2 | 12.06 | -26 | -10.0 | -1.0   | 7.0  | 31  | -4.35  | 1.94  | -0.7666           | 0.4465  |
|                                   | ATL-962 120 mg | 127                | -3.9 | 12.44 | -45 | -12.0 | -3.0   | 4.0  | 20  | -6.13  | -1.76 | -3.5726           | 0.0005  |
| Week 32                           | Placebo        | 58                 | -0.9 | 13.38 | -35 | -8.0  | 0.0    | 8.0  | 30  | -4.38  | 2.66  | -0.4907           | 0.6255  |
|                                   | ATL-962 120 mg | 127                | -3.5 | 13.00 | -34 | -12.0 | -3.0   | 2.0  | 36  | -5.80  | -1.24 | -3.0519           | 0.0028  |
| Week 36                           | Placebo        | 58                 | 0.1  | 13.60 | -30 | -10.0 | 2.0    | 8.0  | 34  | -3.47  | 3.68  | 0.0579            | 0.9540  |
|                                   | ATL-962 120 mg | 124                | -3.1 | 13.39 | -46 | -10.5 | -2.0   | 5.0  | 29  | -5.49  | -0.73 | -2.5882           | 0.0108  |
| Week 40                           | Placebo        | 58                 | 1.3  | 12.36 | -31 | -7.0  | 1.5    | 10.0 | 26  | -1.97  | 4.53  | 0.7861            | 0.4351  |
|                                   | ATL-962 120 mg | 122                | -3.1 | 12.59 | -41 | -10.0 | -2.0   | 4.0  | 35  | -5.31  | -0.80 | -2.6826           | 0.0083  |
| Week 44                           | Placebo        | 58                 | 1.1  | 14.21 | -40 | -10.0 | 2.5    | 10.0 | 37  | -2.68  | 4.79  | 0.5638            | 0.5751  |
|                                   | ATL-962 120 mg | 119                | -2.1 | 13.06 | -32 | -10.0 | -2.0   | 6.0  | 32  | -4.51  | 0.24  | -1.7824           | 0.0772  |
| Week 48                           | Placebo        | 58                 | 2.3  | 13.15 | -29 | -6.0  | 4.0    | 12.0 | 30  | -1.11  | 5.80  | 1.3582            | 0.1797  |
|                                   | ATL-962 120 mg | 119                | -3.0 | 12.82 | -43 | -10.0 | -3.0   | 6.0  | 22  | -5.30  | -0.65 | -2.5317           | 0.0127  |
| Week 52                           | Placebo        | 58                 | 2.2  | 14.25 | -30 | -6.0  | 2.5    | 7.0  | 36  | -1.54  | 5.95  | 1.1792            | 0.2432  |
|                                   | ATL-962 120 mg | 118                | -3.4 | 13.91 | -41 | -12.0 | -3.0   | 4.0  | 44  | -5.93  | -0.86 | -2.6548           | 0.0090  |
| Week 52(LOCF)                     | Placebo        | 65                 | 2.3  | 13.57 | -30 | -4.0  | 2.0    | 7.0  | 36  | -1.07  | 5.65  | 1.3620            | 0.1780  |
|                                   | ATL-962 120 mg | 141                | -3.7 | 13.87 | -41 | -12.0 | -3.0   | 4.0  | 44  | -5.97  | -1.35 | -3.1324           | 0.0021  |



表 2.7.4.7-付録-40 バイタルサイン（拡張期血圧）の評価時点別の変化量（CCT-002 試験）

| Diastolic Blood Pressure<br>(mmHg) | Treatment      | Summary Statistics |      |       |     |       |        |     |     | 95% CI |       | One-Sample t-test |         |
|------------------------------------|----------------|--------------------|------|-------|-----|-------|--------|-----|-----|--------|-------|-------------------|---------|
|                                    |                | N                  | Mean | SD    | Min | Q1    | Median | Q3  | Max | Lower  | Upper | t-value           | p-value |
| Visit                              |                |                    |      |       |     |       |        |     |     |        |       |                   |         |
| Week 2                             | Placebo        | 65                 | 1.5  | 9.86  | -28 | -4.0  | 0.0    | 8.0 | 22  | -0.94  | 3.95  | 1.2324            | 0.2223  |
|                                    | ATL-962 120 mg | 140                | -0.8 | 8.26  | -26 | -5.5  | 0.0    | 4.0 | 23  | -2.20  | 0.56  | -1.1766           | 0.2414  |
| Week 4                             | Placebo        | 63                 | -0.1 | 10.09 | -29 | -6.0  | 0.0    | 6.0 | 31  | -2.62  | 2.46  | -0.0624           | 0.9504  |
|                                    | ATL-962 120 mg | 139                | -0.9 | 8.00  | -24 | -5.0  | -1.0   | 4.0 | 20  | -2.29  | 0.39  | -1.3993           | 0.1640  |
| Week 8                             | Placebo        | 62                 | -0.9 | 8.91  | -29 | -4.0  | 0.0    | 3.0 | 32  | -3.13  | 1.39  | -0.7697           | 0.4445  |
|                                    | ATL-962 120 mg | 137                | -0.9 | 8.14  | -30 | -5.0  | 0.0    | 4.0 | 21  | -2.28  | 0.47  | -1.3013           | 0.1954  |
| Week 12                            | Placebo        | 59                 | -1.1 | 10.20 | -19 | -8.0  | -2.0   | 5.0 | 32  | -3.76  | 1.56  | -0.8300           | 0.4100  |
|                                    | ATL-962 120 mg | 137                | -2.3 | 8.68  | -24 | -7.0  | -3.0   | 2.0 | 34  | -3.75  | -0.82 | -3.0809           | 0.0025  |
| Week 16                            | Placebo        | 59                 | -0.6 | 10.37 | -23 | -6.0  | 0.0    | 5.0 | 27  | -3.30  | 2.11  | -0.4394           | 0.6620  |
|                                    | ATL-962 120 mg | 134                | -2.0 | 8.05  | -23 | -8.0  | -1.5   | 2.0 | 21  | -3.38  | -0.62 | -2.8744           | 0.0047  |
| Week 20                            | Placebo        | 59                 | -2.5 | 9.53  | -28 | -8.0  | -2.0   | 2.0 | 27  | -4.99  | -0.02 | -2.0216           | 0.0478  |
|                                    | ATL-962 120 mg | 133                | -3.5 | 8.56  | -26 | -10.0 | -2.0   | 1.0 | 20  | -5.00  | -2.07 | -4.7608           | <0.0001 |
| Week 24                            | Placebo        | 59                 | -0.7 | 9.45  | -27 | -8.0  | -1.0   | 6.0 | 21  | -3.18  | 1.75  | -0.5785           | 0.5652  |
|                                    | ATL-962 120 mg | 130                | -2.9 | 9.49  | -30 | -9.0  | -2.0   | 3.0 | 20  | -4.51  | -1.22 | -3.4389           | 0.0008  |
| Week 28                            | Placebo        | 59                 | -2.4 | 9.18  | -23 | -8.0  | -3.0   | 2.0 | 23  | -4.75  | 0.04  | -1.9713           | 0.0535  |
|                                    | ATL-962 120 mg | 127                | -2.6 | 9.32  | -22 | -10.0 | -2.0   | 4.0 | 20  | -4.28  | -1.01 | -3.1999           | 0.0017  |
| Week 32                            | Placebo        | 58                 | -1.4 | 9.75  | -33 | -6.0  | -2.0   | 3.0 | 25  | -3.94  | 1.18  | -1.0775           | 0.2858  |
|                                    | ATL-962 120 mg | 127                | -2.3 | 9.55  | -29 | -8.0  | -2.0   | 4.0 | 23  | -3.96  | -0.61 | -2.6957           | 0.0080  |
| Week 36                            | Placebo        | 58                 | -1.3 | 10.25 | -26 | -8.0  | 0.0    | 4.0 | 26  | -3.97  | 1.42  | -0.9484           | 0.3469  |
|                                    | ATL-962 120 mg | 124                | -2.3 | 9.64  | -35 | -8.0  | -1.0   | 4.0 | 20  | -4.06  | -0.63 | -2.7095           | 0.0077  |
| Week 40                            | Placebo        | 58                 | -0.6 | 12.22 | -30 | -10.0 | 0.0    | 8.0 | 26  | -3.85  | 2.58  | -0.3974           | 0.6925  |
|                                    | ATL-962 120 mg | 122                | -2.5 | 9.74  | -30 | -8.0  | -2.0   | 4.0 | 21  | -4.26  | -0.77 | -2.8522           | 0.0051  |
| Week 44                            | Placebo        | 58                 | -1.0 | 11.64 | -29 | -10.0 | 0.0    | 9.0 | 19  | -4.08  | 2.04  | -0.6658           | 0.5082  |
|                                    | ATL-962 120 mg | 119                | -1.9 | 9.60  | -34 | -8.0  | -1.0   | 2.0 | 31  | -3.64  | -0.16 | -2.1585           | 0.0329  |
| Week 48                            | Placebo        | 58                 | -2.2 | 12.35 | -33 | -10.0 | -1.0   | 6.0 | 27  | -5.40  | 1.09  | -1.3292           | 0.1891  |
|                                    | ATL-962 120 mg | 119                | -2.9 | 9.59  | -24 | -9.0  | -3.0   | 3.0 | 27  | -4.62  | -1.13 | -3.2684           | 0.0014  |
| Week 52                            | Placebo        | 58                 | -0.6 | 10.69 | -26 | -8.0  | 0.0    | 8.0 | 22  | -3.43  | 2.19  | -0.4423           | 0.6599  |
|                                    | ATL-962 120 mg | 118                | -3.0 | 9.78  | -27 | -10.0 | -3.0   | 2.0 | 29  | -4.77  | -1.20 | -3.3149           | 0.0012  |
| Week 52(LOCF)                      | Placebo        | 65                 | -0.2 | 10.44 | -26 | -8.0  | 0.0    | 7.0 | 22  | -2.83  | 2.34  | -0.1901           | 0.8498  |
|                                    | ATL-962 120 mg | 141                | -3.0 | 10.07 | -27 | -10.0 | -3.0   | 2.0 | 29  | -4.66  | -1.31 | -3.5224           | 0.0006  |

表 2.7.4.7-付録-41 バイタルサイン（脈拍数）の評価時点別の変化量（CCT-002 試験）

| Pulse (bpm) | Treatment      | Summary Statistics |      |       |     |       |      |        |    |
|-------------|----------------|--------------------|------|-------|-----|-------|------|--------|----|
|             |                | Visit              | N    | Mean  | SD  | Min   | Q1   | Median | Q3 |
| Week 2      | Placebo        | 65                 | -2.6 | 10.80 | -30 | -10.0 | -2.0 | 1.0    | 28 |
|             | ATL-962 120 mg | 140                | 0.7  | 10.06 | -31 | -3.5  | 0.0  | 5.5    | 46 |
| Week 4      | Placebo        | 63                 | -1.8 | 10.31 | -23 | -7.0  | -1.0 | 2.0    | 30 |
|             | ATL-962 120 mg | 139                | 0.4  | 8.40  | -28 | -5.0  | 0.0  | 5.0    | 24 |
| Week 8      | Placebo        | 62                 | -2.0 | 10.92 | -28 | -7.0  | -1.5 | 2.0    | 31 |
|             | ATL-962 120 mg | 137                | 0.5  | 7.56  | -24 | -5.0  | 0.0  | 6.0    | 24 |
| Week 12     | Placebo        | 59                 | -0.5 | 10.58 | -25 | -6.0  | 0.0  | 5.0    | 34 |
|             | ATL-962 120 mg | 137                | 0.9  | 8.94  | -24 | -5.0  | 0.0  | 6.0    | 26 |
| Week 16     | Placebo        | 59                 | -1.5 | 10.44 | -31 | -7.0  | -1.0 | 4.0    | 27 |
|             | ATL-962 120 mg | 134                | -0.9 | 7.71  | -20 | -6.0  | 0.0  | 3.0    | 22 |
| Week 20     | Placebo        | 59                 | 0.4  | 11.53 | -28 | -6.0  | 0.0  | 6.0    | 32 |
|             | ATL-962 120 mg | 133                | -0.1 | 8.45  | -20 | -6.0  | 0.0  | 6.0    | 30 |
| Week 24     | Placebo        | 59                 | -0.8 | 10.06 | -22 | -7.0  | 0.0  | 6.0    | 23 |
|             | ATL-962 120 mg | 130                | -0.5 | 9.98  | -30 | -6.0  | -1.0 | 6.0    | 38 |
| Week 28     | Placebo        | 59                 | 1.8  | 14.18 | -32 | -6.0  | 1.0  | 7.0    | 62 |
|             | ATL-962 120 mg | 127                | 0.3  | 8.22  | -26 | -5.0  | 0.0  | 6.0    | 21 |
| Week 32     | Placebo        | 58                 | -0.7 | 10.25 | -24 | -6.0  | 0.0  | 5.0    | 29 |
|             | ATL-962 120 mg | 127                | -1.0 | 8.77  | -27 | -6.0  | 0.0  | 5.0    | 20 |
| Week 36     | Placebo        | 58                 | -0.8 | 10.94 | -27 | -6.0  | -1.0 | 6.0    | 30 |
|             | ATL-962 120 mg | 124                | -0.6 | 7.85  | -27 | -5.0  | 0.0  | 4.0    | 22 |
| Week 40     | Placebo        | 58                 | -0.9 | 11.47 | -32 | -6.0  | 0.0  | 4.0    | 30 |
|             | ATL-962 120 mg | 122                | 0.3  | 8.64  | -24 | -6.0  | 0.0  | 6.0    | 21 |
| Week 44     | Placebo        | 58                 | 1.2  | 11.49 | -28 | -4.0  | 0.5  | 6.0    | 44 |
|             | ATL-962 120 mg | 119                | -0.6 | 8.18  | -28 | -6.0  | 0.0  | 5.0    | 18 |
| Week 48     | Placebo        | 58                 | -0.5 | 11.16 | -22 | -7.0  | 0.0  | 6.0    | 38 |
|             | ATL-962 120 mg | 119                | -0.4 | 9.54  | -30 | -5.0  | 0.0  | 6.0    | 25 |
| Week 52     | Placebo        | 58                 | -2.0 | 11.81 | -35 | -8.0  | 0.0  | 3.0    | 37 |
|             | ATL-962 120 mg | 118                | -0.3 | 8.88  | -24 | -6.0  | 0.0  | 5.0    | 34 |

表 2.7.4.7-付録-42 バイタルサイン（収縮期血圧）の評価時点別の変化量（OCT-001 試験）

| Systolic Blood Pressure (mmHg) | Treatment      | Summary Statistics |      |       |     |       |      |        |    | 95% CI |       | One-sample t-test |         |
|--------------------------------|----------------|--------------------|------|-------|-----|-------|------|--------|----|--------|-------|-------------------|---------|
|                                |                | Visit              | N    | Mean  | SD  | Min   | Q1   | Median | Q3 | Max    | Lower | Upper             | t-value |
| Week 2                         | ATL-962 120 mg | 127                | -1.1 | 10.94 | -36 | -8.0  | -1.0 | 6.0    | 24 | -2.99  | 0.85  | -1.1029           | 0.2722  |
|                                | ATL-962 120 mg | 126                | -2.9 | 13.47 | -36 | -12.0 | -3.0 | 4.0    | 75 | -5.28  | -0.53 | -2.4207           | 0.0169  |
| Week 8                         | ATL-962 120 mg | 126                | -2.0 | 12.75 | -43 | -10.0 | 0.0  | 6.0    | 35 | -4.28  | 0.22  | -1.7888           | 0.0761  |
|                                | ATL-962 120 mg | 124                | -1.6 | 12.76 | -32 | -10.0 | -1.0 | 7.5    | 35 | -3.88  | 0.66  | -1.4073           | 0.1619  |
| Week 16                        | ATL-962 120 mg | 124                | -0.5 | 14.23 | -33 | -10.0 | -0.5 | 7.5    | 66 | -3.03  | 2.03  | -0.3912           | 0.6963  |
|                                | ATL-962 120 mg | 123                | 1.2  | 13.00 | -32 | -8.0  | 1.0  | 9.0    | 35 | -1.14  | 3.50  | 1.0060            | 0.3164  |
| Week 24                        | ATL-962 120 mg | 122                | 1.3  | 14.08 | -45 | -8.0  | 3.0  | 10.0   | 45 | -1.22  | 3.83  | 1.0225            | 0.3086  |
|                                | ATL-962 120 mg | 127                | 1.4  | 14.10 | -45 | -8.0  | 3.0  | 11.0   | 45 | -1.10  | 3.85  | 1.1015            | 0.2728  |

表 2.7.4.7-付録-43 バイタルサイン（拡張期血圧）の評価時点別の変化量（OCT-001 試験）

| Diastolic Blood Pressure (mmHg) | Treatment      | Summary Statistics |     |      |       |     |      |        |     | 95% CI |       | One-sample t-test |         |         |
|---------------------------------|----------------|--------------------|-----|------|-------|-----|------|--------|-----|--------|-------|-------------------|---------|---------|
|                                 |                | Visit              | N   | Mean | SD    | Min | Q1   | Median | Q3  | Max    | Lower | Upper             | t-value | p-value |
|                                 | ATL-962 120 mg | Week 2             | 127 | -1.2 | 8.70  | -26 | -7.0 | 0.0    | 4.0 | 30     | -2.70 | 0.35              | -1.5198 | 0.1311  |
|                                 | ATL-962 120 mg | Week 4             | 126 | -2.2 | 9.61  | -26 | -9.0 | -2.0   | 2.0 | 41     | -3.91 | -0.52             | -2.5858 | 0.0109  |
|                                 | ATL-962 120 mg | Week 8             | 126 | -0.3 | 8.51  | -20 | -6.0 | 0.0    | 5.0 | 30     | -1.80 | 1.20              | -0.3979 | 0.6914  |
|                                 | ATL-962 120 mg | Week 12            | 124 | -0.7 | 8.96  | -22 | -6.0 | 0.0    | 3.5 | 24     | -2.29 | 0.89              | -0.8717 | 0.3851  |
|                                 | ATL-962 120 mg | Week 16            | 124 | 0.2  | 9.87  | -30 | -6.0 | 1.0    | 4.5 | 44     | -1.55 | 1.96              | 0.2365  | 0.8135  |
|                                 | ATL-962 120 mg | Week 20            | 123 | 0.3  | 9.66  | -30 | -6.0 | 0.0    | 6.0 | 29     | -1.41 | 2.04              | 0.3640  | 0.7165  |
|                                 | ATL-962 120 mg | Week 24            | 122 | 0.7  | 10.07 | -30 | -8.0 | 2.0    | 8.0 | 30     | -1.08 | 2.53              | 0.8002  | 0.4251  |
|                                 | ATL-962 120 mg | Week 24(LOCF)      | 127 | 0.9  | 10.03 | -30 | -8.0 | 2.0    | 8.0 | 30     | -0.89 | 2.64              | 0.9821  | 0.3279  |

表 2.7.4.7-付録-44 バイタルサイン（脈拍数）の評価時点別の変化量（OCT-001 試験）

| Pulse (bpm) | Treatment      | Summary Statistics |     |      |      |     |      |        |     |     |
|-------------|----------------|--------------------|-----|------|------|-----|------|--------|-----|-----|
|             |                | Visit              | N   | Mean | SD   | Min | Q1   | Median | Q3  | Max |
|             | ATL-962 120 mg | Week 2             | 127 | 0.2  | 8.66 | -52 | -4.0 | 0.0    | 5.0 | 20  |
|             | ATL-962 120 mg | Week 4             | 126 | -0.8 | 9.42 | -48 | -5.0 | 0.0    | 4.0 | 28  |
|             | ATL-962 120 mg | Week 8             | 126 | -0.8 | 9.09 | -56 | -6.0 | 0.0    | 5.0 | 18  |
|             | ATL-962 120 mg | Week 12            | 124 | -2.4 | 9.04 | -58 | -6.0 | -1.0   | 2.5 | 15  |
|             | ATL-962 120 mg | Week 16            | 124 | -0.2 | 9.26 | -56 | -3.5 | 1.0    | 5.5 | 21  |
|             | ATL-962 120 mg | Week 20            | 123 | -0.7 | 9.12 | -55 | -5.0 | -1.0   | 4.0 | 28  |
|             | ATL-962 120 mg | Week 24            | 122 | -0.1 | 9.14 | -61 | -4.0 | 0.0    | 5.0 | 27  |

表 2.7.4.7-付録-45 バイタルサイン（収縮期血圧）の評価時点別の変化量（OCT-002 試験）

| Systolic Blood Pressure (mmHg) | Treatment      | Summary Statistics |     |      |       |     |       |        |     | 95% CI |       | One-sample t-test |         |         |
|--------------------------------|----------------|--------------------|-----|------|-------|-----|-------|--------|-----|--------|-------|-------------------|---------|---------|
|                                |                | Visit              | N   | Mean | SD    | Min | Q1    | Median | Q3  | Max    | Lower | Upper             | t-value | p-value |
|                                | ATL-962 120 mg | Week 2             | 504 | -1.4 | 11.35 | -34 | -8.0  | 0.0    | 6.0 | 46     | -2.41 | -0.42             | -2.7953 | 0.0054  |
|                                | ATL-962 120 mg | Week 4             | 495 | -1.4 | 11.82 | -40 | -9.0  | -2.0   | 6.0 | 41     | -2.41 | -0.32             | -2.5716 | 0.0104  |
|                                | ATL-962 120 mg | Week 8             | 489 | -1.2 | 11.67 | -43 | -8.0  | -1.0   | 5.0 | 42     | -2.22 | -0.15             | -2.2522 | 0.0248  |
|                                | ATL-962 120 mg | Week 12            | 479 | -0.9 | 12.93 | -40 | -9.0  | -1.0   | 8.0 | 40     | -2.05 | 0.27              | -1.5086 | 0.1321  |
|                                | ATL-962 120 mg | Week 16            | 472 | -0.9 | 12.00 | -40 | -8.0  | 0.0    | 7.0 | 42     | -1.99 | 0.18              | -1.6451 | 0.1006  |
|                                | ATL-962 120 mg | Week 20            | 469 | -0.8 | 13.03 | -44 | -9.0  | 0.0    | 8.0 | 42     | -2.01 | 0.35              | -1.3785 | 0.1687  |
|                                | ATL-962 120 mg | Week 24            | 462 | -1.2 | 13.14 | -48 | -10.0 | -2.0   | 7.0 | 39     | -2.45 | -0.05             | -2.0393 | 0.0420  |
|                                | ATL-962 120 mg | Week 28            | 453 | -1.5 | 13.79 | -44 | -10.0 | 0.0    | 7.0 | 44     | -2.80 | -0.25             | -2.3517 | 0.0191  |
|                                | ATL-962 120 mg | Week 32            | 456 | -1.6 | 12.94 | -45 | -11.0 | -2.0   | 7.0 | 83     | -2.84 | -0.46             | -2.7220 | 0.0067  |
|                                | ATL-962 120 mg | Week 36            | 452 | -2.6 | 13.11 | -46 | -10.0 | -2.0   | 5.0 | 48     | -3.86 | -1.44             | -4.2936 | <0.0001 |
|                                | ATL-962 120 mg | Week 40            | 444 | -3.6 | 13.11 | -58 | -12.0 | -3.0   | 5.0 | 34     | -4.79 | -2.35             | -5.7383 | <0.0001 |
|                                | ATL-962 120 mg | Week 44            | 442 | -4.2 | 13.08 | -47 | -12.0 | -3.5   | 4.0 | 34     | -5.46 | -3.01             | -6.8064 | <0.0001 |
|                                | ATL-962 120 mg | Week 48            | 439 | -3.7 | 12.92 | -48 | -12.0 | -3.0   | 4.0 | 42     | -4.88 | -2.46             | -5.9507 | <0.0001 |
|                                | ATL-962 120 mg | Week 52            | 439 | -3.4 | 13.10 | -48 | -12.0 | -2.0   | 6.0 | 40     | -4.68 | -2.22             | -5.5168 | <0.0001 |
|                                | ATL-962 120 mg | Week 52(LOCF)      | 504 | -3.0 | 12.91 | -48 | -11.0 | -2.0   | 6.0 | 40     | -4.16 | -1.90             | -5.2617 | <0.0001 |

表 2.7.4.7-付録-46 バイタルサイン（拡張期血圧）の評価時点別の変化量（OCT-002 試験）

| Diastolic Blood Pressure (mmHg) | Treatment      | Summary Statistics |      |      |     |      |      |        |    |       | 95% CI |         | One-sample t-test |         |
|---------------------------------|----------------|--------------------|------|------|-----|------|------|--------|----|-------|--------|---------|-------------------|---------|
|                                 |                | Visit              | N    | Mean | SD  | Min  | Q1   | Median | Q3 | Max   | Lower  | Upper   | t-value           | p-value |
| Week 2                          | ATL-962 120 mg | 504                | -1.1 | 8.83 | -30 | -6.0 | 0.0  | 4.0    | 38 | -1.87 | -0.32  | -2.7901 | 0.0055            |         |
| Week 4                          | ATL-962 120 mg | 495                | -0.8 | 9.25 | -38 | -6.0 | 0.0  | 5.0    | 26 | -1.65 | -0.01  | -1.9923 | 0.0469            |         |
| Week 8                          | ATL-962 120 mg | 489                | -0.7 | 8.90 | -39 | -6.0 | 0.0  | 5.0    | 26 | -1.45 | 0.13   | -1.6464 | 0.1003            |         |
| Week 12                         | ATL-962 120 mg | 479                | -1.1 | 9.13 | -40 | -8.0 | 0.0  | 4.0    | 23 | -1.87 | -0.23  | -2.5226 | 0.0120            |         |
| Week 16                         | ATL-962 120 mg | 472                | -0.8 | 8.47 | -28 | -6.0 | 0.0  | 4.0    | 26 | -1.61 | -0.08  | -2.1675 | 0.0307            |         |
| Week 20                         | ATL-962 120 mg | 469                | -1.0 | 9.22 | -44 | -8.0 | 0.0  | 4.0    | 26 | -1.86 | -0.19  | -2.3998 | 0.0168            |         |
| Week 24                         | ATL-962 120 mg | 462                | -1.2 | 9.45 | -30 | -8.0 | 0.0  | 5.0    | 26 | -2.03 | -0.30  | -2.6486 | 0.0084            |         |
| Week 28                         | ATL-962 120 mg | 453                | -1.6 | 9.45 | -30 | -7.0 | -1.0 | 5.0    | 24 | -2.48 | -0.74  | -3.6288 | 0.0003            |         |
| Week 32                         | ATL-962 120 mg | 456                | -2.2 | 9.25 | -33 | -8.5 | -2.0 | 4.0    | 26 | -3.06 | -1.36  | -5.1045 | <0.0001           |         |
| Week 36                         | ATL-962 120 mg | 452                | -2.5 | 9.81 | -34 | -9.0 | -2.0 | 4.0    | 24 | -3.43 | -1.61  | -5.4628 | <0.0001           |         |
| Week 40                         | ATL-962 120 mg | 444                | -3.1 | 9.93 | -29 | -9.0 | -3.0 | 2.0    | 29 | -4.04 | -2.18  | -6.6005 | <0.0001           |         |
| Week 44                         | ATL-962 120 mg | 442                | -3.3 | 9.65 | -37 | -9.0 | -2.5 | 2.0    | 30 | -4.25 | -2.45  | -7.2948 | <0.0001           |         |
| Week 48                         | ATL-962 120 mg | 439                | -3.2 | 9.77 | -34 | -9.0 | -2.0 | 2.0    | 24 | -4.13 | -2.30  | -6.8881 | <0.0001           |         |
| Week 52                         | ATL-962 120 mg | 439                | -2.9 | 9.47 | -32 | -9.0 | -2.0 | 4.0    | 28 | -3.82 | -2.04  | -6.4829 | <0.0001           |         |
| Week 52(LOCF)                   | ATL-962 120 mg | 504                | -2.7 | 9.25 | -32 | -8.0 | -2.0 | 3.5    | 28 | -3.47 | -1.85  | -6.4453 | <0.0001           |         |

表 2.7.4.7-付録-47 バイタルサイン（脈拍数）の評価時点別の変化量（OCT-002 試験）

| Pulse (bpm) | Treatment      | Summary Statistics |      |      |     |      |      |        |    |     |
|-------------|----------------|--------------------|------|------|-----|------|------|--------|----|-----|
|             |                | Visit              | N    | Mean | SD  | Min  | Q1   | Median | Q3 | Max |
| Week 2      | ATL-962 120 mg | 504                | -0.5 | 8.09 | -26 | -5.0 | 0.0  | 4.0    | 33 |     |
| Week 4      | ATL-962 120 mg | 495                | -0.9 | 7.91 | -30 | -6.0 | 0.0  | 4.0    | 31 |     |
| Week 8      | ATL-962 120 mg | 489                | -0.9 | 8.05 | -32 | -6.0 | 0.0  | 4.0    | 29 |     |
| Week 12     | ATL-962 120 mg | 479                | -1.6 | 8.55 | -31 | -6.0 | -1.0 | 3.0    | 39 |     |
| Week 16     | ATL-962 120 mg | 472                | -1.4 | 8.17 | -26 | -6.0 | -1.0 | 4.0    | 27 |     |
| Week 20     | ATL-962 120 mg | 469                | -1.7 | 8.71 | -36 | -7.0 | -1.0 | 3.0    | 39 |     |
| Week 24     | ATL-962 120 mg | 462                | -2.0 | 8.51 | -35 | -6.0 | -2.0 | 3.0    | 25 |     |
| Week 28     | ATL-962 120 mg | 453                | -1.0 | 8.84 | -26 | -6.0 | -1.0 | 4.0    | 37 |     |
| Week 32     | ATL-962 120 mg | 456                | -1.5 | 8.96 | -38 | -6.0 | 0.0  | 4.0    | 24 |     |
| Week 36     | ATL-962 120 mg | 452                | -2.1 | 8.44 | -34 | -6.0 | -2.0 | 2.0    | 24 |     |
| Week 40     | ATL-962 120 mg | 444                | -1.5 | 8.99 | -34 | -7.0 | -1.0 | 4.0    | 40 |     |
| Week 44     | ATL-962 120 mg | 442                | -1.7 | 9.50 | -38 | -6.0 | -2.0 | 3.0    | 58 |     |
| Week 48     | ATL-962 120 mg | 439                | -2.2 | 8.66 | -43 | -7.0 | -2.0 | 2.0    | 30 |     |
| Week 52     | ATL-962 120 mg | 439                | -2.1 | 9.46 | -46 | -7.0 | -1.0 | 4.0    | 45 |     |

表 2.7.4.7-付録-48 心電図判定の評価時点別クロス表 (CCT-001 試験) (1/3)

| Frequency |         |           |                                      | Week 0               |                                      |                                  |       |
|-----------|---------|-----------|--------------------------------------|----------------------|--------------------------------------|----------------------------------|-------|
|           |         |           |                                      | Within Normal Limits | Abnormal, Not Clinically Significant | Abnormal, Clinically Significant | Total |
| Treatment | Placebo | Week 12   | Within Normal Limits                 | 64                   | 5                                    | -                                | 69    |
|           |         |           | Abnormal, Not Clinically Significant | 9                    | 19                                   | -                                | 28    |
|           |         |           | Abnormal, Clinically Significant     | -                    | -                                    | 1                                | 1     |
|           |         |           | Total                                | 73                   | 24                                   | 1                                | 98    |
|           |         | Week 24   | Within Normal Limits                 | 65                   | 5                                    | -                                | 70    |
|           |         |           | Abnormal, Not Clinically Significant | 5                    | 19                                   | -                                | 24    |
|           |         |           | Abnormal, Clinically Significant     | -                    | -                                    | 1                                | 1     |
|           |         |           | Total                                | 70                   | 24                                   | 1                                | 95    |
|           |         | Week 28   | Within Normal Limits                 | 62                   | 6                                    | -                                | 68    |
|           |         |           | Abnormal, Not Clinically Significant | 7                    | 18                                   | -                                | 25    |
|           |         |           | Abnormal, Clinically Significant     | -                    | -                                    | 1                                | 1     |
|           |         |           | Total                                | 69                   | 24                                   | 1                                | 94    |
|           |         | Follow-up | Within Normal Limits                 | 66                   | 6                                    | -                                | 72    |
|           |         |           | Abnormal, Not Clinically Significant | 7                    | 18                                   | -                                | 25    |
|           |         |           | Abnormal, Clinically Significant     | -                    | -                                    | 1                                | 1     |
|           |         |           | Total                                | 73                   | 24                                   | 1                                | 98    |

| Frequency |       |           |                                      | Week 0               |                                      |                                  |       |
|-----------|-------|-----------|--------------------------------------|----------------------|--------------------------------------|----------------------------------|-------|
|           |       |           |                                      | Within Normal Limits | Abnormal, Not Clinically Significant | Abnormal, Clinically Significant | Total |
| Treatment | 15 mg | Week 12   | Within Normal Limits                 | 59                   | 10                                   | -                                | 69    |
|           |       |           | Abnormal, Not Clinically Significant | 5                    | 12                                   | -                                | 17    |
|           |       |           | Abnormal, Clinically Significant     | 2                    | -                                    | 1                                | 3     |
|           |       |           | Total                                | 66                   | 22                                   | 1                                | 89    |
|           |       | Week 24   | Within Normal Limits                 | 58                   | 8                                    | -                                | 66    |
|           |       |           | Abnormal, Not Clinically Significant | 4                    | 13                                   | -                                | 17    |
|           |       |           | Abnormal, Clinically Significant     | 1                    | -                                    | -                                | 1     |
|           |       |           | Total                                | 63                   | 21                                   | -                                | 84    |
|           |       | Week 28   | Within Normal Limits                 | 57                   | 6                                    | -                                | 63    |
|           |       |           | Abnormal, Not Clinically Significant | 5                    | 15                                   | -                                | 20    |
|           |       |           | Abnormal, Clinically Significant     | -                    | -                                    | -                                | -     |
|           |       |           | Total                                | 62                   | 21                                   | -                                | 83    |
|           |       | Follow-up | Within Normal Limits                 | 60                   | 6                                    | -                                | 66    |
|           |       |           | Abnormal, Not Clinically Significant | 4                    | 16                                   | -                                | 20    |
|           |       |           | Abnormal, Clinically Significant     | -                    | -                                    | -                                | -     |
|           |       |           | Total                                | 64                   | 22                                   | -                                | 86    |

表 2.7.4.7-付録-48 心電図判定の評価時点別クロス表 (CCT-001 試験) (2/3)

| Frequency |           |                                      |                                      | Week 0               |                                      |                                  |       |
|-----------|-----------|--------------------------------------|--------------------------------------|----------------------|--------------------------------------|----------------------------------|-------|
|           |           |                                      |                                      | Within Normal Limits | Abnormal, Not Clinically Significant | Abnormal, Clinically Significant | Total |
| Treatment | 30 mg     | Week 12                              | Within Normal Limits                 | 60                   | 4                                    | -                                | 64    |
|           |           |                                      | Abnormal, Not Clinically Significant | 8                    | 18                                   | 1                                | 27    |
|           |           |                                      | Abnormal, Clinically Significant     | -                    | -                                    | 1                                | 1     |
|           |           |                                      | Total                                | 68                   | 22                                   | 2                                | 92    |
|           |           | Week 24                              | Within Normal Limits                 | 58                   | 4                                    | -                                | 62    |
|           |           |                                      | Abnormal, Not Clinically Significant | 6                    | 15                                   | 1                                | 22    |
|           |           |                                      | Abnormal, Clinically Significant     | -                    | -                                    | 1                                | 1     |
|           |           |                                      | Total                                | 64                   | 19                                   | 2                                | 85    |
|           |           | Week 28                              | Within Normal Limits                 | 54                   | 4                                    | -                                | 58    |
|           |           |                                      | Abnormal, Not Clinically Significant | 9                    | 15                                   | 1                                | 25    |
|           |           |                                      | Abnormal, Clinically Significant     | -                    | -                                    | 1                                | 1     |
|           |           |                                      | Total                                | 63                   | 19                                   | 2                                | 84    |
|           | Follow-up | Within Normal Limits                 | 58                                   | 5                    | -                                    | 63                               |       |
|           |           | Abnormal, Not Clinically Significant | 9                                    | 18                   | 1                                    | 28                               |       |
|           |           | Abnormal, Clinically Significant     | -                                    | -                    | 1                                    | 1                                |       |
|           |           | Total                                | 67                                   | 23                   | 2                                    | 92                               |       |

| Frequency |           |                                      |                                      | Week 0               |                                      |                                  |       |
|-----------|-----------|--------------------------------------|--------------------------------------|----------------------|--------------------------------------|----------------------------------|-------|
|           |           |                                      |                                      | Within Normal Limits | Abnormal, Not Clinically Significant | Abnormal, Clinically Significant | Total |
| Treatment | 60 mg     | Week 12                              | Within Normal Limits                 | 58                   | 7                                    | -                                | 65    |
|           |           |                                      | Abnormal, Not Clinically Significant | 11                   | 15                                   | -                                | 26    |
|           |           |                                      | Abnormal, Clinically Significant     | -                    | -                                    | -                                | -     |
|           |           |                                      | Total                                | 69                   | 22                                   | -                                | 91    |
|           |           | Week 24                              | Within Normal Limits                 | 57                   | 6                                    | -                                | 63    |
|           |           |                                      | Abnormal, Not Clinically Significant | 5                    | 11                                   | -                                | 16    |
|           |           |                                      | Abnormal, Clinically Significant     | -                    | -                                    | -                                | -     |
|           |           |                                      | Total                                | 62                   | 17                                   | -                                | 79    |
|           |           | Week 28                              | Within Normal Limits                 | 56                   | 7                                    | -                                | 63    |
|           |           |                                      | Abnormal, Not Clinically Significant | 3                    | 10                                   | -                                | 13    |
|           |           |                                      | Abnormal, Clinically Significant     | -                    | -                                    | -                                | -     |
|           |           |                                      | Total                                | 59                   | 17                                   | -                                | 76    |
|           | Follow-up | Within Normal Limits                 | 62                                   | 7                    | -                                    | 69                               |       |
|           |           | Abnormal, Not Clinically Significant | 3                                    | 14                   | -                                    | 17                               |       |
|           |           | Abnormal, Clinically Significant     | -                                    | -                    | -                                    | -                                |       |
|           |           | Total                                | 65                                   | 21                   | -                                    | 86                               |       |

表 2.7.4.7-付録-48 心電図判定の評価時点別クロス表 (CCT-001 試験) (3/3)

| Frequency |           |                                      | Week 0                               |                                      |                                  |       |    |
|-----------|-----------|--------------------------------------|--------------------------------------|--------------------------------------|----------------------------------|-------|----|
|           |           |                                      | Within Normal Limits                 | Abnormal, Not Clinically Significant | Abnormal, Clinically Significant | Total |    |
| Treatment | 120 mg    | Week 12                              | Within Normal Limits                 | 63                                   | 3                                | -     | 66 |
|           |           |                                      | Abnormal, Not Clinically Significant | 9                                    | 14                               | -     | 23 |
|           |           |                                      | Abnormal, Clinically Significant     | -                                    | -                                | -     | -  |
|           |           |                                      | Total                                | 72                                   | 17                               | -     | 89 |
|           |           | Week 24                              | Within Normal Limits                 | 58                                   | 5                                | -     | 63 |
|           |           |                                      | Abnormal, Not Clinically Significant | 8                                    | 9                                | -     | 17 |
|           |           |                                      | Abnormal, Clinically Significant     | -                                    | -                                | -     | -  |
|           |           |                                      | Total                                | 66                                   | 14                               | -     | 80 |
|           |           | Week 28                              | Within Normal Limits                 | 56                                   | 5                                | -     | 61 |
|           |           |                                      | Abnormal, Not Clinically Significant | 9                                    | 8                                | -     | 17 |
|           |           |                                      | Abnormal, Clinically Significant     | -                                    | -                                | -     | -  |
|           |           |                                      | Total                                | 65                                   | 13                               | -     | 78 |
|           | Follow-up | Within Normal Limits                 | 61                                   | 7                                    | -                                | 68    |    |
|           |           | Abnormal, Not Clinically Significant | 11                                   | 9                                    | -                                | 20    |    |
|           |           | Abnormal, Clinically Significant     | -                                    | -                                    | -                                | -     |    |
|           |           | Total                                | 72                                   | 16                                   | -                                | 88    |    |

表 2.7.4.7-付録-49 心電図判定の評価時点別クロス表 (CCT-002 試験)

| Frequency |                |                                      | Week 0                               |                                      |                                  |       |     |     |
|-----------|----------------|--------------------------------------|--------------------------------------|--------------------------------------|----------------------------------|-------|-----|-----|
|           |                |                                      | Within Normal Limits                 | Abnormal, Not Clinically Significant | Abnormal, Clinically Significant | Total |     |     |
| Treatment | Placebo        | Week 12                              | Within Normal Limits                 | 43                                   | 3                                | -     | 46  |     |
|           |                |                                      | Abnormal, Not Clinically Significant | 2                                    | 16                               | -     | 18  |     |
|           |                |                                      | Abnormal, Clinically Significant     | -                                    | -                                | -     | -   |     |
|           |                |                                      |                                      | Total                                | 45                               | 19    | -   | 64  |
|           |                | Week 24                              | Within Normal Limits                 | 37                                   | 3                                | -     | 40  |     |
|           |                |                                      | Abnormal, Not Clinically Significant | 5                                    | 14                               | -     | 19  |     |
|           |                |                                      | Abnormal, Clinically Significant     | -                                    | -                                | -     | -   |     |
|           |                |                                      |                                      | Total                                | 42                               | 17    | -   | 59  |
|           |                | Week 36                              | Within Normal Limits                 | 37                                   | 3                                | -     | 40  |     |
|           |                |                                      | Abnormal, Not Clinically Significant | 4                                    | 14                               | -     | 18  |     |
|           |                |                                      | Abnormal, Clinically Significant     | -                                    | -                                | -     | -   |     |
|           |                |                                      |                                      | Total                                | 41                               | 17    | -   | 58  |
|           | Week 52        | Within Normal Limits                 | 36                                   | 4                                    | -                                | 40    |     |     |
|           |                | Abnormal, Not Clinically Significant | 5                                    | 13                                   | -                                | 18    |     |     |
|           |                | Abnormal, Clinically Significant     | -                                    | -                                    | -                                | -     |     |     |
|           |                |                                      | Total                                | 41                                   | 17                               | -     | 58  |     |
|           | ATL-962 120 mg | Week 12                              | Within Normal Limits                 | 101                                  | 10                               | -     | 111 |     |
|           |                |                                      | Abnormal, Not Clinically Significant | 3                                    | 25                               | -     | 28  |     |
|           |                |                                      | Abnormal, Clinically Significant     | -                                    | -                                | -     | -   |     |
|           |                |                                      |                                      | Total                                | 104                              | 35    | -   | 139 |
| Week 24   |                | Within Normal Limits                 | 94                                   | 11                                   | -                                | 105   |     |     |
|           |                | Abnormal, Not Clinically Significant | 6                                    | 21                                   | -                                | 27    |     |     |
|           |                | Abnormal, Clinically Significant     | -                                    | -                                    | -                                | -     |     |     |
|           |                |                                      | Total                                | 100                                  | 32                               | -     | 132 |     |
| Week 36   |                | Within Normal Limits                 | 94                                   | 8                                    | -                                | 102   |     |     |
|           |                | Abnormal, Not Clinically Significant | 4                                    | 21                                   | -                                | 25    |     |     |
|           |                | Abnormal, Clinically Significant     | -                                    | -                                    | -                                | -     |     |     |
|           |                |                                      | Total                                | 98                                   | 29                               | -     | 127 |     |
| Week 52   |                | Within Normal Limits                 | 84                                   | 9                                    | -                                | 93    |     |     |
|           |                | Abnormal, Not Clinically Significant | 7                                    | 18                                   | -                                | 25    |     |     |
|           |                | Abnormal, Clinically Significant     | -                                    | -                                    | -                                | -     |     |     |
|           |                |                                      | Total                                | 91                                   | 27                               | -     | 118 |     |

表 2.7.4.7-付録-50 心電図判定の評価時点別クロス表 (OCT-001 試験)

| Frequency |                |         | Week 0                               |                                      |                                  |       |     |
|-----------|----------------|---------|--------------------------------------|--------------------------------------|----------------------------------|-------|-----|
|           |                |         | Within Normal Limits                 | Abnormal, Not Clinically Significant | Abnormal, Clinically Significant | Total |     |
| Treatment | ATL-962 120 mg | Week 12 | Within Normal Limits                 | 94                                   | 5                                | -     | 99  |
|           |                |         | Abnormal, Not Clinically Significant | 2                                    | 24                               | 1     | 27  |
|           |                |         | Abnormal, Clinically Significant     | 1                                    | -                                | -     | 1   |
|           |                |         | Total                                | 97                                   | 29                               | 1     | 127 |
|           |                | Week 24 | Within Normal Limits                 | 90                                   | 10                               | -     | 100 |
|           |                |         | Abnormal, Not Clinically Significant | 4                                    | 17                               | 1     | 22  |
|           |                |         | Abnormal, Clinically Significant     | -                                    | 1                                | -     | 1   |
|           |                | Total   | 94                                   | 28                                   | 1                                | 123   |     |



表 2.7.4.7-付録-51 心電図判定の評価時点別クロス表 (OCT-002 試験)

| Frequency |                   |                                      | Week 0                               |                                      |                                  |       |     |
|-----------|-------------------|--------------------------------------|--------------------------------------|--------------------------------------|----------------------------------|-------|-----|
|           |                   |                                      | Within Normal Limits                 | Abnormal, Not Clinically Significant | Abnormal, Clinically Significant | Total |     |
| Treatment | ATL-962<br>120 mg | Week 12                              | Within Normal Limits                 | 316                                  | 32                               | 1     | 349 |
|           |                   |                                      | Abnormal, Not Clinically Significant | 39                                   | 95                               | 2     | 136 |
|           |                   |                                      | Abnormal, Clinically Significant     | 2                                    | -                                | 14    | 16  |
|           |                   |                                      | Total                                | 357                                  | 127                              | 17    | 501 |
|           |                   | Week 24                              | Within Normal Limits                 | 290                                  | 32                               | 1     | 323 |
|           |                   |                                      | Abnormal, Not Clinically Significant | 39                                   | 86                               | 3     | 128 |
|           |                   |                                      | Abnormal, Clinically Significant     | 2                                    | 2                                | 13    | 17  |
|           |                   |                                      | Total                                | 331                                  | 120                              | 17    | 468 |
|           |                   | Week 36                              | Within Normal Limits                 | 283                                  | 37                               | 2     | 322 |
|           |                   |                                      | Abnormal, Not Clinically Significant | 36                                   | 79                               | 3     | 118 |
|           |                   |                                      | Abnormal, Clinically Significant     | 3                                    | 1                                | 11    | 15  |
|           |                   |                                      | Total                                | 322                                  | 117                              | 16    | 455 |
|           | Week 52           | Within Normal Limits                 | 277                                  | 44                                   | 2                                | 323   |     |
|           |                   | Abnormal, Not Clinically Significant | 31                                   | 69                                   | 4                                | 104   |     |
|           |                   | Abnormal, Clinically Significant     | 3                                    | 1                                    | 10                               | 14    |     |
|           |                   | Total                                | 311                                  | 114                                  | 16                               | 441   |     |