

1.8.1 添付文書（案）

次頁以降に添付文書（案）を示す。

貯 法：室温保存
有効期間：24箇月ルマケラス®錠120mg
LUMAKRAS® Tablets 120mg

日本標準商品分類番号

874291

承認番号

販売開始

処方箋医薬品^{注)}注) 注意—医師等の処方箋により
使用すること

1. 警告

本剤は、緊急時に十分対応できる医療施設において、がん化学療法に十分な知識・経験を持つ医師のもとで、本剤の使用が適切と判断される症例についてのみ投与すること。また、治療開始に先立ち、患者又はその家族に有効性及び危険性を十分説明し、同意を得てから投与すること。

2. 禁忌（次の患者には投与しないこと）

本剤の成分に対し過敏症の既往歴のある患者

3. 組成・性状

3.1 組成

販 売 名	ルマケラス錠120mg
有効成分	1錠中ソトラシブ120mg
添 加 剤	結晶セルロース、乳糖水和物、クロスカルメロースナトリウム、ステアリン酸マグネシウム、ポリビニルアルコール（部分けん化物）、酸化チタン、マクロゴール4000、タルク、黄色三酸化鉄

3.2 製剤の性状

性 状		黄色のフィルムコーティング錠
外形	表面	120
	裏面	AMG
	側面	
大きさ	長径	16.00mm
	短径	7.11mm
	厚さ	5.58mm
識別コード		AMG

4. 効能又は効果

がん化学療法後に増悪したKRAS G12C変異陽性の切除不能な進行・再発の非小細胞肺癌

5. 効能又は効果に関連する注意

- 5.1 十分な経験を有する病理医又は検査施設における検査により、KRAS G12C変異陽性が確認された患者に投与すること。検査にあたっては、承認された体外診断用医薬品又は医療機器を用いること。なお、承認された体外診断用医薬品又は医療機器に関する情報については、以下のウェブサイトから入手可能である：
<https://www.pmda.go.jp/review-services/drug-reviews/review-information/cd/0001.html>
- 5.2 「17. 臨床成績」の項の内容を熟知し、本剤の有効性及び安全性を十分に理解した上で、本剤以外の治療の実施についても慎重に検討し、適応患者の選択を行うこと。〔17. 1. 1参照〕
- 5.3 本剤の一次治療における有効性及び安全性は確立していない。
- 5.4 本剤の術後補助療法における有効性及び安全性は確立していない。

6. 用法及び用量

通常、成人にはソトラシブとして960mgを1日1回経口投与する。なお、患者の状態により適宜減量する。

7. 用法及び用量に関連する注意

- 7.1 他の抗悪性腫瘍剤との併用について、有効性及び安全性は確立していない。
- 7.2 本剤投与により副作用が発現した場合には、以下の基準を考慮して、休薬・減量・中止すること。240mg/日の投与量に対して忍容性が認められない場合は投与を中止すること。

本剤の減量レベル

減量レベル	投与量
通常投与量	960mg/日
1段階減量	480mg/日
2段階減量	240mg/日

副作用発現時の休薬、減量及び中止基準

副作用	程度	処置
肝機能障害 〔8. 1、 11. 1. 1参照〕	症候性のGrade 2のAST増加若しくはALT増加又はGrade 3以上のAST増加若しくはALT増加	Grade 1以下又はベースラインに回復するまで休薬し、回復後は1段階減量して投与再開できる。
	正常値上限の3倍を超えるAST増加又はALT増加、かつ正常値上限の2倍を超える総ビリルビン増加	本剤の投与を中止する。
間質性肺炎患 〔8. 2、 11. 1. 2参照〕	全Grade	本剤の投与を中止する。
上記以外の副作用	Grade 3又は4（ただし、悪心、嘔吐、下痢は適切な処置を行っても症状が継続する場合）	Grade 1以下又はベースラインに回復するまで休薬し、回復後は1段階減量して投与再開できる。

注) GradeはNCI-CTCAE version 5.0に準じる。

8. 重要な基本的注意

- 8.1 肝機能障害があらわれることがあるので、本剤の投与開始前及び投与中は定期的に肝機能検査を行い、患者の状態を十分に観察すること。〔7. 2、11. 1. 1参照〕
- 8.2 間質性肺炎患があらわれることがあるので、本剤の投与にあたっては、初期症状（発熱、咳嗽、呼吸困難等）の確認及び胸部X線検査の実施等、観察を十分に行うこと。〔7. 2、11. 1. 2参照〕

9. 特定の背景を有する患者に関する注意

9.3 肝機能障害患者

- 9.3.1 中等度以上の肝機能障害患者（AST若しくはALTが基準値上限の2.5倍以上、又は総ビリルビンが基準値上限の1.5倍以上）
本剤は主に肝臓で代謝されるため、血中濃度が上昇する可能性がある。なお、中等度以上の肝機能障害患者を対象とした臨床試験は実施していない。

9.5 妊婦

妊婦又は妊娠している可能性のある女性には、治療上の有益性が危険性を上回ると判断される場合にのみ投与すること。ウサギを用いた胚・胎児発生に関する試験において、臨床曝露量（AUC）の約2.2倍の曝露に相当する用量で、胎児の体重減少及び中手骨骨化数の減少が認められている。

9.6 授乳婦

治療上の有益性及び母乳栄養の有益性を考慮し、授乳の継続又は中止を検討すること。本剤の母乳中への移行は不明である。

9.7 小児等

小児等を対象とした臨床試験は実施していない。

10. 相互作用

本剤はCYP3Aの基質となり、CYP3Aの誘導作用及びP-gpの阻害作用を示す。また、本剤の溶解度はpHの上昇により低下する。

10.2 併用注意（併用に注意すること）

薬剤名等	臨床症状・措置方法	機序・危険因子
CYP3A誘導剤 リファンピシン、フェニトイン、カルバマゼピン等 [16.7.1参照]	本剤の有効性が減弱するおそれがあるため、併用を避けることが望ましい。	これらの薬剤がCYP3Aを誘導することにより、本剤の血中濃度が低下する可能性がある。
CYP3Aの基質となる薬剤 ミダゾラム、トリアゾラム、ロミタビド等 [16.7.2参照]	これらの薬剤の有効性が減弱するおそれがある。	本剤がCYP3Aを誘導することにより、これらの薬剤の血中濃度が低下する可能性がある。
治療域の狭いP-gpの基質となる薬剤 ジゴキシン、エベロリムス、シロリムス等 [16.7.3参照]	これらの薬剤の副作用が増強されるおそれがあるため、患者の状態を慎重に観察し、副作用の発現に十分注意すること。	本剤がP-gpを阻害することにより、これらの薬剤の血中濃度が上昇する可能性がある。
胃内pHを上昇させる薬剤 プロトンポンプ阻害剤 オメプラゾール、ラベプラゾール、ランソプラゾール等 H ₂ 受容体拮抗剤 ファモチジン、ラニチジン、シメチジン等 [16.7.4、16.7.5参照]	本剤の有効性が減弱するおそれがあるため、これらの薬剤との併用を避けることが望ましい。	これらの薬剤による胃内pHの上昇により本剤の溶解性が低下し、本剤の血中濃度が低下する可能性がある。

11. 副作用

次の副作用があらわれることがあるので、観察を十分に行い、異常が認められた場合には投与を中止するなど適切な処置を行うこと。

11.1 重大な副作用

11.1.1 肝機能障害

ALT増加（16.3%）、AST増加（16.3%）等の肝機能障害があらわれることがある。 [7.2、8.1参照]

11.1.2 間質性肺疾患

肺臓炎（1.1%）等があらわれることがある。 [7.2、8.2参照]

11.2 その他の副作用

	5%以上	1%～5%未満	1%未満
感染症および寄生虫症			蜂巣炎、憩室炎、口腔カンジダ症、口腔真菌感染、膿疱性皮疹、上気道感染
良性、悪性および詳細不明の新生物（嚢胞およびポリープを含む）			血管筋脂肪腫
血液およびリンパ系障害		貧血、リンパ球減少、白血球減少、好中球減少、血小板減少	赤血球減少
免疫系障害 代謝および栄養障害		食欲減退、低カリウム血症、脱水、脂質異常症、高トリグリセリド血症、低マグネシウム血症、低リン酸血症	薬物過敏症 低血糖、低ナトリウム血症

	5%以上	1%～5%未満	1%未満
精神障害			錯乱状態、不眠症
神経系障害		頭痛、味覚不全	浮動性めまい、知覚過敏、過眠症、神経系障害、末梢性ニューロパチー
眼障害			霧視、視力障害、眼球乾燥症
心臓障害			洞性徐脈
血管障害			高血圧、低血圧、末梢冷感
呼吸器、胸郭および縦隔障害		呼吸困難	咳嗽、鼻閉、湿性咳嗽
胃腸障害	下痢（27.9%）、悪心（16.3%）、嘔吐、腹痛	口内乾燥、腹部膨満、便秘、消化不良、胃食道逆流性疾患、口内炎	呼気臭、鼓腸、口腔粘膜水疱形成
肝胆道系障害			肝炎
皮膚および皮下組織障害		斑状丘疹状皮疹、皮膚乾燥、そう痒症、脱毛症、発疹	皮膚炎、さ瘡様皮膚炎、多汗症、寝汗、光線過敏性反応、紫斑、乾皮症
筋骨格系および結合組織障害		筋肉痛、関節痛	関節炎、筋力低下、変形性関節症
腎および尿路障害			白血球尿、頻尿、蛋白尿
生殖系および乳房障害			女性化乳房
一般・全身障害および投与部位の状態	疲労（11.1%）	末梢性浮腫、無力症、倦怠感、発熱	びくびく感、限局性浮腫、粘膜の炎症、非心臓性胸痛、浮腫
臨床検査		体重減少、血中コレステロール増加、血中クレアチニン増加	AST減少、血中コレステロール減少、血中クレアチニンホスホキナーゼ増加、コレチゾール減少、心電図QT延長、胃内pH低下、リパーゼ増加

14. 適用上の注意

14.1 薬剤交付時の注意

PTP包装の薬剤はPTPシートから取り出して服用するよう指導すること。PTPシートの誤飲により、硬い鋭角部が食道粘膜へ刺入し、更には穿孔をおこして縦隔洞炎等の重篤な合併症を併発することがある。

14.2 薬剤投与時の注意

粉砕や分割して使用しないこと。

16. 薬物動態

16.1 血中濃度

16.1.1 単回及び反復投与

進行固形癌患者に、本剤180、360、720又は960mgを空腹時に1日1回反復経口投与^{注)}したときのソトラシブの薬物動態パラメータ及び血漿中濃度推移を以下に示す。検討された用量範囲において、ソトラシブのC_{max}及びAUCは概ね類似した。反復投与した際、ソトラシブの蓄積は認められなかった（外国人データ）。¹⁾

注)：本剤の承認用法・用量は960mgを1日1回経口投与である。

表1 進行固形癌患者に本剤180、360、720又は960mgを1日1回経口投与^{注1)}したときの1日目及び8日目の薬物動態パラメータ〔幾何平均値 (CV%)〕

用量 (mg)	投与 (n)	C _{max} (ng/mL)	T _{max} ^{注2)} (hr)	AUC _{0-24h} (ng·hr/mL)	T _{1/2} ^{注3)} (hr)
180	1日目 (6)	6,880 (51%)	1.0 (0.50 - 2.0)	43,600 (57%)	5.71 ^{注4)} (0.815)
	8日目 (6)	6,440 (67%)	0.73 (0.50 - 1.2)	31,700 (89%)	5.13 ^{注4)} (1.99)
360	1日目 (26)	6,190 (64%)	1.1 (0.57 - 6.2)	58,400 ^{注5)} (63%)	6.45 ^{注6)} (1.80)
	8日目 (24)	6,310 (43%)	1.0 (0.50 - 4.0)	38,900 ^{注5)} (49%)	5.53 ^{注7)} (1.84)
720	1日目 (11)	7,570 (59%)	1.2 (0.50 - 4.1)	84,000 ^{注8)} (57%)	6.45 ^{注9)} (1.95)
	8日目 (11)	5,450 (50%)	1.1 (0.53 - 4.0)	42,100 (49%)	4.75 ^{注10)} (1.16)
960	1日目 (24)	8,400 (59%)	1.5 (0.25 - 4.8)	67,700 ^{注5)} (77%)	5.49 ^{注6)} (2.14)
	8日目 (24)	5,390 (65%)	1.1 (0.22 - 6.5)	32,400 ^{注11)} (75%)	5.07 ^{注12)} (1.08)

注1)：本剤の承認用法・用量は960mgを1日1回経口投与である、注2)：中央値 (範囲)、注3)：平均値 (標準偏差)、注4)：n = 5、注5)：n = 22、注6)：n = 17、注7)：n = 21、注8)：n = 9、注9)：n = 8、注10)：n = 7、注11)：n = 18、注12)：n = 15

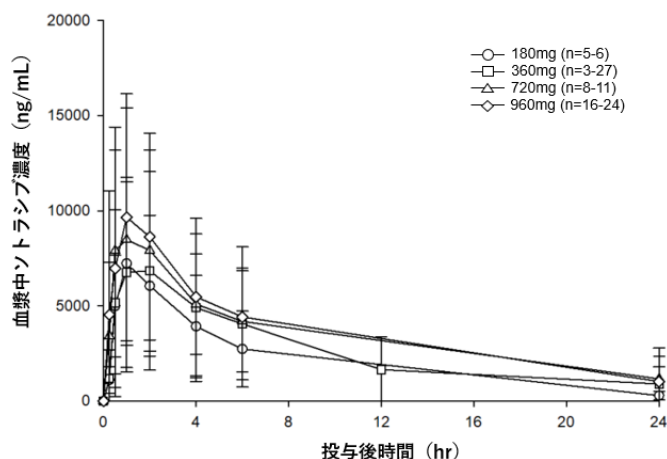


図1 進行固形癌患者に本剤180、360、720又は960mgを1日1回経口投与^{注)}したときの1日目の血漿中濃度推移 (平均値±標準偏差)

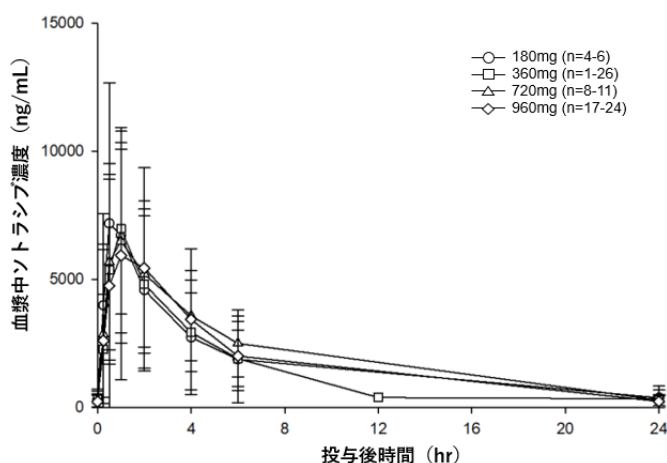


図2 進行固形癌患者に本剤180、360、720又は960mgを1日1回経口投与^{注)}したときの8日目の血漿中濃度推移 (平均値±標準偏差)

注)：本剤の承認用法・用量は960mgを1日1回経口投与である。

日本人固形癌患者に、本剤960mgを空腹時に1日1回反復経口投与したときのソトラシブの薬物動態パラメータを以下に示す。²⁾

表2 日本人患者に本剤960mgを1日1回経口投与したときの1日目及び8日目の薬物動態パラメータ〔幾何平均値 (CV%)〕

投与 (n)	C _{max} (ng/mL)	T _{max} ^{注1)} (hr)	AUC _{0-24h} ^{注2)} (ng·hr/mL)	T _{1/2} ^{注2), 3)} (hr)
1日目 (7)	5,190 (50%)	2.0 (0.53 - 6.1)	47,800 (38%)	4.59 (1.50)
8日目 (8)	4,150 (40%)	1.9 (0.90 - 5.9)	25,400 (23%)	3.95 (0.832)

注1)：中央値 (範囲)、注2)：n = 5、注3)：平均値 (標準偏差)

16.2 吸収

16.2.1 食事の影響

健康被験者 (14例) に本剤360mgを単回経口投与^{注)}したとき、空腹時投与に対する高脂肪食後投与におけるソトラシブのC_{max}及びAUC_{last}の最小二乗幾何平均値の比は、それぞれ1.03及び1.38であった (外国人データ)。³⁾

注)：本剤の承認用法・用量は960mgを1日1回経口投与である。

16.3 分布

非小細胞肺癌患者 (21例) に本剤960mgを1日1回反復経口投与したときのソトラシブの定常状態における見かけの分布容積の幾何平均値は、211Lであった (外国人データ)。¹⁾

ソトラシブのヒト血漿タンパク結合率は約89%であった (*in vitro*)。⁴⁾ソトラシブのヒト血液/血漿中濃度比は約0.7であった (*in vitro*)。⁵⁾

16.4 代謝

ソトラシブの代謝に関与する主な代謝酵素はCYP3Aである (*in vitro*)。⁶⁾

健康被験者 (8例) に¹⁴C-標識ソトラシブ720mgを単回経口投与^{注)}したとき、投与192時間後までの血漿中には、主に未変化体、M10 (システイン付加体) 及びM24 (ピペラジンアクリルアミドが脱離した酸化体) が認められ、血漿中総放射能のそれぞれ17.1%、26.8%及び7.81%を占めた (外国人データ)。⁷⁾

注)：本剤の承認用法・用量は960mgを1日1回経口投与である。

16.5 排泄

健康被験者 (8例) に¹⁴C-標識ソトラシブ720mgを単回経口投与^{注)}したとき、投与312時間後までに、投与された全放射能の74.4%が糞中に、5.81%が尿中に排泄された。また、投与192時間後までに、投与された全放射能の53.0%が糞中に、1.39%が尿中に、それぞれ未変化体として排泄された (外国人データ)。⁷⁾

注)：本剤の承認用法・用量は960mgを1日1回経口投与である。

16.7 薬物相互作用

16.7.1 リファンピシン

健康被験者 (14例) にリファンピシン (強いCYP3A誘導剤) 600mgを1日1回15日間反復経口投与し、本剤960mgを単回経口投与したとき、本剤単独投与時に対するリファンピシン併用投与時のソトラシブのC_{max}及びAUC_{inf}の最小二乗幾何平均値の比は、それぞれ0.647及び0.487であった (外国人データ)。⁸⁾ [10.2参照]

16.7.2 ミダゾラム

非小細胞肺癌患者 (7例) に本剤960mgを1日1回15日間反復経口投与し、ミダゾラム (CYP3A基質) 2mgを単回経口投与したとき、ミダゾラム単独投与時に対する本剤併用投与時のミダゾラムのC_{max}及びAUC_{0h}の最小二乗幾何平均値の比は、それぞれ0.52及び0.47であった (外国人データ)。⁹⁾ [10.2参照]

16.7.3 ジゴキシン

健康被験者 (14例) に本剤960mg及びジゴキシン (P-gp基質) 0.5mgを単回経口投与したとき、ジゴキシン単独投与時に対する本剤併用投与時のジゴキシンのC_{max}及びAUC_{inf}の最小二乗幾何平均値の比は、それぞれ1.91及び1.21であった (外国人データ)。¹⁰⁾ [10.2参照]

16.7.4 オメプラゾール

(1) 健康被験者 (14例) にオメプラゾール (プロトンポンプ阻害

剤) 40mgを1日1回6日間反復経口投与し、本剤960mgを空腹時に単回経口投与したとき、本剤単独投与時に対するオメプラゾール併用投与時のソトラシブの C_{max} 及び AUC_{last} の最小二乗幾何平均値の比は、それぞれ0.431及び0.576であった(外国人データ)。¹¹⁾ [10.2参照]

- (2) 健康被験者(13例)にオメプラゾール(プロトンポンプ阻害剤) 40mgを1日1回6日間反復経口投与し、本剤960mgを中脂肪食後に単回経口投与したとき、本剤単独投与時に対するオメプラゾール併用投与時のソトラシブの C_{max} 及び AUC_{last} の最小二乗幾何平均値の比は、それぞれ0.349及び0.414であった(外国人データ)。¹²⁾ [10.2参照]

16.7.5 ファモチジン

健康被験者(14例)に本剤960mgを中脂肪食後に単回投与、並びにファモチジン(H_2 受容体拮抗剤) 40mgを本剤投与10時間前及び2時間後に経口投与したとき、本剤単独投与時に対するファモチジン併用投与時のソトラシブの C_{max} 及び AUC_{last} の最小二乗幾何平均値の比は、それぞれ0.654及び0.619であった(外国人データ)。¹²⁾ [10.2参照]

16.7.6 その他

- (1) 健康被験者(14例)にイトラコナゾール(強いCYP3A阻害剤) 200mgを1日2回経口投与後に4日間1日1回反復経口投与し、本剤360mgを単回経口投与^{注1)}したとき、本剤単独投与時に対するイトラコナゾール併用投与時のソトラシブの C_{max} 及び AUC_{inf} の最小二乗幾何平均値の比は、それぞれ1.04及び1.26であった(外国人データ)。¹³⁾

注)：本剤の承認用法・用量は960mgを1日1回経口投与である。

- (2) 健康被験者(13例)に本剤960mg及びメトホルミン(MATE1及びMATE2-K基質) 850mgを単回経口投与したとき、メトホルミン単独投与時に対する本剤併用投与時のメトホルミンの C_{max} 及び AUC_{inf} の最小二乗幾何平均値の比は、それぞれ0.996及び0.985であった(外国人データ)。¹⁴⁾
- (3) ソトラシブはBCRPを阻害した(*in vitro*)。¹⁵⁾

17. 臨床成績

17.1 有効性及び安全性に関する試験

17.1.1 国際共同第Ⅰ/Ⅱ相試験(20170543試験、CodeBreak 100試験)

がん化学療法後に増悪したKRAS G12C変異陽性の切除不能な進行・再発の固形癌患者を対象とした国際共同第Ⅰ/Ⅱ相試験の第Ⅱ相パートにおいて、非小細胞肺癌患者126例(日本人11例を含む)に本剤960mgを1日1回経口投与した。

主要評価項目であるRECIST ver. 1.1に基づく独立中央画像判定機関による奏効率(データカットオフ日：2020年9月1日)は、有効性評価対象^{注1)} 123例(日本人10例を含む)で37.4%(95%信頼区間：28.8, 46.6)であった。¹⁶⁾

安全性評価対象^{注2)} 190例(日本人13例を含む)中128例(67.4%)に副作用が認められ、主な副作用(発現率5%以上)は、下痢(27.9%)、悪心、ALT増加、AST増加(各16.3%)、疲労(11.1%)、血中アルカリホスファターゼ増加(7.9%)、嘔吐(7.4%)及び腹痛(5.3%)であった。[5.2参照]

注1) 中央判定によりベースライン時に測定可能病変を有しないと判定された3例(日本人1例を含む)が除外された。

注2) 第Ⅰ相及び第Ⅱ相パートにおいて本剤960mgを1回以上投与された非小細胞肺癌患者

18. 薬効薬理

18.1 作用機序

ソトラシブは、KRAS G12C変異を有するKRASに対する阻害作用を有する低分子化合物である。ソトラシブは、G12C変異を有するKRASに結合することで、KRASの活性化を阻害し、下流のシグナル伝達を阻害することにより、腫瘍増殖抑制作用を示すと考えられている。

18.2 抗腫瘍効果

ソトラシブは、KRAS G12C変異を有するヒト非小細胞肺癌由来NCI-H358細胞株を皮下移植したヌードマウスにおいて、腫瘍増殖

抑制作用を示した。¹⁷⁾

19. 有効成分に関する理化学的知見

一般的名称：ソトラシブ(Sotorasib)

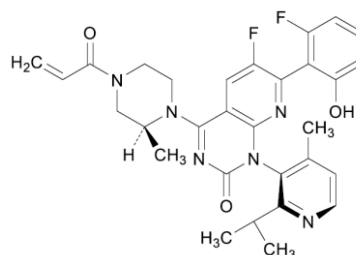
化学名：(1*M*)-6-Fluoro-7-(2-fluoro-6-hydroxyphenyl)-1-[4-methyl-2-(propan-2-yl)pyridin-3-yl]-4-[(2*S*)-2-methyl-4-(prop-2-en-1-yl)piperazin-1-yl]pyrido[2,3-*d*]pyrimidin-2(1*H*)-one

分子式： $C_{30}H_{30}F_2N_6O_3$

分子量：560.59

性状：白色～薄い褐色の粉末である。水にほとんど溶けず、エタノールにやや溶けにくい。

化学構造式：



分配係数：Log D = 2.44(n-オクタノール/pH7.4 リン酸塩緩衝液)

22. 包装

56錠 [8錠(PTP)×7]

23. 主要文献

- 社内資料：20170543試験 患者における薬物動態(X年X月X日承認、CTD 2.7.2.2.1.1)
- 社内資料：20170543試験 日本人患者における薬物動態(X年X月X日承認、CTD 2.7.2.5.1)
- 社内資料：20190316試験 食事の影響を検討した試験(X年X月X日承認、CTD 2.7.6.1)
- 社内資料：150530試験 ソトラシブのヒト血漿タンパク結合率(X年X月X日承認、CTD 2.6.4.4)
- 社内資料：150532試験 ソトラシブのヒト血液/血漿中濃度比(X年X月X日承認、CTD 2.6.4.1)
- 社内資料：150537試験 ソトラシブのヒトにおける代謝(X年X月X日承認、CTD 2.6.4.5.2)
- 社内資料：20190321試験 マスバランス試験(X年X月X日承認、CTD 2.7.2.2.1.2)
- 社内資料：20190319試験 薬物相互作用試験(X年X月X日承認、CTD 2.7.6.7)
- 社内資料：20170543試験 薬物相互作用サブスタディ(X年X月X日承認、CTD 2.7.6.10)
- 社内資料：20190315試験 薬物相互作用試験(X年X月X日承認、CTD 2.7.6.4)
- 社内資料：20190320試験 薬物相互作用試験(X年X月X日承認、CTD 2.7.6.8)
- 社内資料：20200199試験 薬物相互作用試験(X年X月X日承認、CTD 2.7.6.9)
- 社内資料：20190318試験 薬物相互作用試験(X年X月X日承認、CTD 2.7.6.6)
- 社内資料：20190317試験 薬物相互作用試験(X年X月X日承認、CTD 2.7.6.5)
- 社内資料：150539試験 薬物トランスポーターに対するソトラシブの阻害作用試験(X年X月X日承認、CTD 2.6.4.7)
- 社内資料：20170543試験 国際共同第Ⅰ/Ⅱ相試験(X年X月X日承認、CTD 2.7.6.11)
- Canon J, et al.:Nature. 2019;575:217-223.

24. 文献請求先及び問い合わせ先

アムジェン株式会社

メディカルインフォメーションセンター

〒107-6239 東京都港区赤坂九丁目7番1号

0120-790-549

26. 製造販売業者等

26.1 製造販売

アムジェン株式会社

東京都港区赤坂九丁目7番1号

目次

1.	効能又は効果、効能又は効果に関連する注意.....	3
2.	効能又は効果の設定根拠.....	3
3.	効能又は効果に関連する注意の設定根拠.....	4
4.	用法及び用量、用法及び用量に関連する注意.....	4
5.	用法及び用量の設定根拠.....	5
6.	用法及び用量に関連する注意の設定根拠.....	6

略語一覧

略語又は用語	定義／説明
DLT	dose-limiting toxicity 用量制限毒性
ERK	extracellular signal-regulated kinase 細胞外シグナル調節キナーゼ
GDP	guanosine diphosphate グアノシン二リン酸
KRAS	Kirsten rat sarcoma viral oncogene homolog
KRAS G12C	KRAS protein with a G12C amino acid substitution 12番目のアミノ酸グリシン（G）がシステイン（C）に置換されたKRASタンパク
MTD	maximum tolerated dose 最大耐量
NSCLC	non-small cell lung cancer 非小細胞肺癌
PK	pharmacokinetics 薬物動態
PR	partial response 部分奏効
QD	once daily 1日1回
RAF	RAF proto-oncogene serine/threonine-protein kinase RAFプロトオンコジーンセリン／トレオニンプロテインキナーゼ
RP2D	recommended phase 2 dose 第II相部分における推奨用量

1. 効能又は効果、効能又は効果に関連する注意

効能又は効果（案）

がん化学療法後に増悪したKRAS G12C変異陽性の切除不能な進行・再発の非小細胞肺癌

<効能又は効果に関連する注意（案）>

5.1 十分な経験を有する病理医又は検査施設における検査により、KRAS G12C変異陽性が確認された患者に投与すること。検査にあたっては、承認された体外診断用医薬品又は医療機器を用いること。なお、承認された体外診断用医薬品又は医療機器に関する情報については、以下のウェブサイトから入手可能である：

<https://www.pmda.go.jp/review-services/drug-reviews/review-information/cd/0001.html>

5.2 「17.臨床成績」の項の内容を熟知し、本剤の有効性及び安全性を十分に理解した上で、本剤以外の治療の実施についても慎重に検討し、適応患者の選択を行うこと。

5.3 本剤の一次治療における有効性及び安全性は確立していない。

5.4 本剤の術後補助療法における有効性及び安全性は確立していない。

2. 効能又は効果の設定根拠

sotorasibは、KRAS G12C変異タンパクに特異的に結合し、不可逆的に阻害する低分子化合物である。sotorasibは、システインに変異した12番目のアミノ酸に隣接するKRAS G12CのP2ポケットと呼ばれる溝構造及びアミノ酸位置95のヒスチジンにおける溝構造に結合する。sotorasibは、チオール反応性部分を有し、システイン残基を共有結合により修飾することでKRAS G12Cをグアノシン二リン酸（GDP）が結合した不活性型の高次構造のまま保持する。これにより、Kirsten rat sarcoma viral oncogene homolog（KRAS）とRAFプロトオンコジーンセリン／トレオニンプロテインキナーゼ（RAF）等のエフェクターとの相互作用が阻害され、細胞外シグナル調節キナーゼ（ERK）のリン酸化を含め、下流のシグナル伝達が阻害される。sotorasibを用いた薬理試験でも、in vivoでこれらの作用が確認されており、KRAS G12C変異を有する腫瘍に対する選択的な細胞増殖阻害及び腫瘍退縮が認められている。sotorasibが結合する他の野生型又は変異型のタンパクや受容体は特定されておらず、KRAS G12C変異のない細胞ではsotorasibの影響は認められていない。In vitroでの選択性アッセイ及びシステインプロテオームプロファイリングの結果から、sotorasibはKRAS G12Cに対して高い選択性を有することが示唆された（モジュール2.4）。

KRAS G12C変異を有する固形癌患者を対象とした第I/II相試験である20170543試験の第II相部分の結果、既治療のKRAS G12C変異陽性非小細胞肺癌（NSCLC）患者に対するsotorasib単独療法による臨床的意義のある有効性及び良好な安全性プロファイルが認められ、sotorasibはKRAS G12C変異陽性NSCLC患者に対して良好なリスク・ベネフィットのプロファイルを有し、有用な治療選択肢となりうると考えられた（モジュール2.5）。

以上より、効能又は効果を「がん化学療法後に増悪したKRAS G12C変異陽性の切除不能な進行・再発の非小細胞肺癌」と設定した。

3. 効能又は効果に関連する注意の設定根拠

- 5.1 十分な経験を有する病理医又は検査施設における検査により、KRAS G12C変異陽性が確認された患者に投与すること。検査にあたっては、承認された体外診断用医薬品又は医療機器を用いること。なお、承認された体外診断用医薬品又は医療機器に関する情報については、以下のウェブサイトから入手可能である：

<https://www.pmda.go.jp/review-services/drug-reviews/review-information/cd/0001.html>

本剤の作用機序及び20170543試験の結果に基づき、本剤により期待される効果を得るためには、十分な経験を有する病理医又は検査施設における検査により、KRAS G12C変異陽性を確認する必要があると考え、設定した。なお、既承認の薬剤での記載に準じた。

- 5.2 「17.臨床成績」の項の内容を熟知し、本剤の有効性及び安全性を十分に理解した上で、本剤以外の治療の実施についても慎重に検討し、適応患者の選択を行うこと。

本剤の効能又は効果は、20170543試験の結果に基づき設定したことから、本剤の適応患者の選択に際し、臨床成績の項の内容を熟知し、本剤の有効性及び安全性を十分理解する必要があると考え、注意喚起することとした。また、本剤の延命効果に関する検証的な臨床試験成績は得られていないこと等を考慮し、本剤以外の治療の実施についても慎重に検討した上で、本剤投与の可否を判断する必要がある旨を注意喚起することが適切であると判断した。

- 5.3 本剤の一次治療における有効性及び安全性は確立していない。

本剤の一次治療における有効性及び安全性は、現時点では確立していないことから設定した。

- 5.4 本剤の術後補助療法における有効性及び安全性は確立していない。

本剤の術後補助療法における有効性及び安全性は、現時点では確立していないことから設定した。

4. 用法及び用量、用法及び用量に関連する注意

用法及び用量（案）

通常、成人にはソトラシブとして960mgを1日1回経口投与する。なお、患者の状態により適宜減量する。

<用法及び用量に関連する注意（案）>

- 7.1 他の抗悪性腫瘍剤との併用について、有効性及び安全性は確立していない。

7.2 本剤投与により副作用が発現した場合には、以下の基準を考慮して、休薬・減量・中止すること。240mg/日の投与量に対して忍容性が認められない場合は投与を中止すること。

本剤の減量レベル

減量レベル	投与量
通常投与量	960mg/日
1段階減量	480mg/日
2段階減量	240mg/日

副作用発現時の休薬、減量及び中止基準

副作用	程度	処置
肝機能障害	症候性のGrade 2のAST増加若しくはALT増加 又は Grade 3以上のAST増加若しくはALT増加	Grade 1以下又はベースラインに回復するまで休薬し、回復後は1段階減量して投与再開できる。
	正常値上限の3倍を超えるAST増加又はALT増加、かつ正常値上限の2倍を超える総ビリルビン増加	本剤の投与を中止する。
間質性肺疾患	全Grade	本剤の投与を中止する。
上記以外の副作用	Grade 3又は4（ただし、悪心、嘔吐、下痢は適切な処置を行っても症状が継続する場合）	Grade 1以下又はベースラインに回復するまで休薬し、回復後は1段階減量して投与再開できる。

注）GradeはNCI-CTCAE version 5.0に準じる。

5. 用法及び用量の設定根拠

用量漸増試験である20170543試験の第I相部分のパート1aの成績に基づいて、第II相部分における推奨用量（RP2D）を選択した。本パートではsotorasib 180、360、720及び960 mgの1日1回（QD）単独投与を検討した。その結果、いずれの用量にも用量制限毒性（DLT）は認められず、検討した最高用量である960 mgにおいても最大耐量（MTD）には到達しなかった。また、本パートでの予備的な有効性データの評価において、評価可能な進行NSCLC被験者17例中7例に部分奏効（PR）が認められ、960 mgの用量では評価可能な進行NSCLC被験者7例中4例にPRが認められた。これらの結果から、960 mg QD単独投与をRP2Dと決定した。

本試験の第II相部分では、KRAS G12C変異を有する進行NSCLC被験者において、RP2Dである960 mg QD単独投与により臨床的に意味のある客観的奏効率が認められた。また、忍容かつ良好な安全性が確認された。KRAS G12C変異を有する進行NSCLC被験者における960 mg QD単独投与では有害事象による中止割合は9.5%と低かった。さらに、用量変更が行われた割合は17.9%と低く、960 mgの用量選択の妥当性を支持している。

KRAS G12C変異を有する進行固形癌被験者にsotorasibを180 mgから960 mgまでQD経口投与したときの薬物動態（PK）データでは、sotorasibの曝露量は用量比を下回る増加を示した。反復投与による薬物の蓄積は認められなかった。また、960 mgを空腹時又は食後に投与したときのsotorasibの曝露量は、空腹時投与と食後投与で同様であった。母集団PK解析により、性別、人種、病勢、体重、年齢、軽度腎機能障害又は軽度肝機能障害の影響を検討した結果、sotorasibのPKに臨床的に意味のある差は認められず、これらの共変量に基づく用量調整は必要ないことが示唆された。

以上より、sotorasibの安全性及び有効性を含むエビデンスの包括的評価に基づき、KRAS G12C変異を有する進行NSCLC患者に対するsotorasibの推奨用法・用量は、960 mg QD単独投与とすることが妥当と判断された（[モジュール2.7.3 4項](#)）。

6. 用法及び用量に関連する注意の設定根拠

7.1 他の抗悪性腫瘍剤との併用について、有効性及び安全性は確立していない。

他の抗悪性腫瘍剤との併用療法における本剤の有効性及び安全性は、現時点では確立していないことから設定した。

7.2 本剤投与により副作用が発現した場合には、以下の基準を考慮して、休薬・減量・中止すること。240mg/日の投与量に対して忍容性が認められない場合は投与を中止すること。

20170543試験の結果、本剤の休薬・減量・中止を要した副作用のうち、多く認められたのは肝酵素増加や悪心、嘔吐、下痢であり（[モジュール2.7.4 2.1.5項](#)）、当該事象が認められた場合、本剤の休薬・減量・中止を検討する必要があると考え、20170543試験での設定に基づき、肝機能障害及びそれ以外の副作用が発現した場合について、休薬・減量・中止の基準を設定した。また、間質性肺疾患は死亡を含む重大な転帰に至る可能性のある事象であり、発現時には早期の対応を行うことが肝要であると考えられることから、いずれのGradeであっても発現が確認された場合は投与中止の処置が推奨されると考え、設定した。

1. 使用上の注意及びその設定根拠

使用上の注意（案）	設定根拠
1. 警告 本剤は、緊急時に十分対応できる医療施設において、がん化学療法に十分な知識・経験を持つ医師のもとで、本剤の使用が適切と判断される症例についてのみ投与すること。また、治療開始に先立ち、患者又はその家族に有効性及び危険性を十分説明し、同意を得てから投与すること。	副作用に適切に対応するため、本剤の投与はがん化学療法に関する十分な知識・経験を持つ医師のもとで、かつ副作用発現による緊急時に十分な措置ができる医療施設において、本剤の使用が適切と判断される症例についてのみ投与されることが安全対策上必要であることから設定した。また、本剤投与にあたり、患者又はその家族に有効性及び安全性を理解いただき、同意を得てから投与する必要があることから設定した。
2. 禁忌（次の患者には投与しないこと） 本剤の成分に対し過敏症の既往歴のある患者	過敏症に対する一般的な注意事項として設定した。また、本剤成分に過敏症の既往歴がある場合、本剤投与により過敏症が発現する可能性が高いことから設定した。
8. 重要な基本的注意 8.1 肝機能障害があらわれることがあるので、本剤の投与開始前及び投与中は定期的に肝機能検査を行い、患者の状態を十分に観察すること。〔7.2、11.1.1参照〕	国際共同第I／II相試験（20170543試験）における安全性評価対象190例（本剤960 mgを1回以上投与された非小細胞肺癌患者）の結果及びCore Data Sheet（CDS）に基づき設定した。
8.2 間質性肺疾患があらわれることがあるので、本剤の投与にあたっては、初期症状（発熱、咳嗽、呼吸困難等）の確認及び胸部X線検査の実施等、観察を十分に行うこと。〔7.2、11.1.2参照〕	国際共同第I／II相試験（20170543試験）における安全性評価対象190例（本剤960 mgを1回以上投与された非小細胞肺癌患者）の結果に基づき設定した。
9. 特定の背景を有する患者に関する注意 9.3 肝機能障害患者 9.3.1 中等度以上の肝機能障害患者（AST若しくはALTが基準値上限の2.5倍以上、又は総ビリルビンが基準値上限の1.5倍以上） 本剤は主に肝臓で代謝されるため、血中濃度が上昇する可能性がある。なお、中等度以上の肝機能障害患者を対象とした臨床試験は実施していない。	本剤は主に肝臓で代謝されるため、血中濃度が上昇する可能性があること、中等度以上の肝機能障害患者を対象とした臨床試験は実施していないことから設定した。
9.5 妊婦 妊婦又は妊娠している可能性のある女性には、治療上の有益性が危険性を上回ると判断される場合にのみ投与すること。ウサギを用いた胚・胎児発生に関する試験において、臨床曝露量（AUC）の約2.2倍の曝露に相当する用量で、胎児の体重減少及び中手骨骨化数の減少が認められている。	妊婦又は妊娠している可能性のある女性における安全性は検討していないこと、非臨床試験の結果及びCDSに基づき設定した。
9.6 授乳婦 治療上の有益性及び母乳栄養の有益性を考慮し、授乳の継続又は中止を検討すること。本剤の母乳中への移行は不明である。	臨床試験及び非臨床試験等のデータがなく、ヒトで哺乳中の児における影響が不明であるため、設定した。
9.7 小児等 小児等を対象とした臨床試験は実施していない。	小児等を対象とした臨床試験は実施していないことから設定した。

使用上の注意（案）				設定根拠																
10. 相互作用 本剤はCYP3Aの基質となり、CYP3Aの誘導作用及びP-gpの阻害作用を示す。また、本剤の溶解度はpHの上昇により低下する。 10.2 併用注意（併用に注意すること） <table><tr><th>薬剤名等</th><th>臨床症状・措置方法</th><th>機序・危険因子</th></tr><tr><td>CYP3A誘導剤 リファンピシン、 フェニトイン、カルバマゼピン等 [16.7.1参照]</td><td>本剤の有効性が減弱するおそれがあるため、併用を避けることが望ましい。</td><td>これらの薬剤がCYP3Aを誘導することにより、本剤の血中濃度が低下する可能性がある。</td></tr><tr><td>CYP3Aの基質となる薬剤 ミダゾラム、トリアゾラム、ロミタピド等 [16.7.2参照]</td><td>これらの薬剤の有効性が減弱するおそれがある。</td><td>本剤がCYP3Aを誘導することにより、これらの薬剤の血中濃度が低下する可能性がある。</td></tr><tr><td>治療域の狭いP-gpの基質となる薬剤 ジゴキシン、エベロリムス、シロリムス等 [16.7.3参照]</td><td>これらの薬剤の副作用が増強されるおそれがあるため、患者の状態を慎重に観察し、副作用の発現に十分注意すること。</td><td>本剤がP-gpを阻害することにより、これらの薬剤の血中濃度が上昇する可能性がある。</td></tr><tr><td>胃内pHを上昇させる薬剤 プロトンポンプ阻害剤 オメプラゾール、ラベプラゾール、ランソプラゾール等 H₂受容体拮抗剤 ファモチジン、ラニチジン、シメチジン等 [16.7.4、16.7.5参照]</td><td>本剤の有効性が減弱するおそれがあるため、これらの薬剤との併用を避けることが望ましい。</td><td>これらの薬剤による胃内pHの上昇により本剤の溶解性が低下し、本剤の血中濃度が低下する可能性がある。</td></tr></table>				薬剤名等	臨床症状・措置方法	機序・危険因子	CYP3A誘導剤 リファンピシン、 フェニトイン、カルバマゼピン等 [16.7.1参照]	本剤の有効性が減弱するおそれがあるため、併用を避けることが望ましい。	これらの薬剤がCYP3Aを誘導することにより、本剤の血中濃度が低下する可能性がある。	CYP3Aの基質となる薬剤 ミダゾラム、トリアゾラム、ロミタピド等 [16.7.2参照]	これらの薬剤の有効性が減弱するおそれがある。	本剤がCYP3Aを誘導することにより、これらの薬剤の血中濃度が低下する可能性がある。	治療域の狭いP-gpの基質となる薬剤 ジゴキシン、エベロリムス、シロリムス等 [16.7.3参照]	これらの薬剤の副作用が増強されるおそれがあるため、患者の状態を慎重に観察し、副作用の発現に十分注意すること。	本剤がP-gpを阻害することにより、これらの薬剤の血中濃度が上昇する可能性がある。	胃内pHを上昇させる薬剤 プロトンポンプ阻害剤 オメプラゾール、ラベプラゾール、ランソプラゾール等 H ₂ 受容体拮抗剤 ファモチジン、ラニチジン、シメチジン等 [16.7.4、16.7.5参照]	本剤の有効性が減弱するおそれがあるため、これらの薬剤との併用を避けることが望ましい。	これらの薬剤による胃内pHの上昇により本剤の溶解性が低下し、本剤の血中濃度が低下する可能性がある。	本剤は主に代謝酵素チトクロームP450 3A（CYP3A）で代謝され、またCYP3A誘導作用を有することが確認されている。また、本剤の併用によりP-gpの基質であるジゴキシンの血中濃度が上昇することが確認されている。さらに、オメプラゾールやファモチジン等による胃内pHの上昇により、本剤の血中濃度が低下することが確認されている。併用注意に関しては、臨床薬理試験等で相互作用が確認され、かつ有効性への影響が不明もしくは有効性を減弱させる可能性または副作用が増強される可能性のある薬剤について記載した。	
薬剤名等	臨床症状・措置方法	機序・危険因子																		
CYP3A誘導剤 リファンピシン、 フェニトイン、カルバマゼピン等 [16.7.1参照]	本剤の有効性が減弱するおそれがあるため、併用を避けることが望ましい。	これらの薬剤がCYP3Aを誘導することにより、本剤の血中濃度が低下する可能性がある。																		
CYP3Aの基質となる薬剤 ミダゾラム、トリアゾラム、ロミタピド等 [16.7.2参照]	これらの薬剤の有効性が減弱するおそれがある。	本剤がCYP3Aを誘導することにより、これらの薬剤の血中濃度が低下する可能性がある。																		
治療域の狭いP-gpの基質となる薬剤 ジゴキシン、エベロリムス、シロリムス等 [16.7.3参照]	これらの薬剤の副作用が増強されるおそれがあるため、患者の状態を慎重に観察し、副作用の発現に十分注意すること。	本剤がP-gpを阻害することにより、これらの薬剤の血中濃度が上昇する可能性がある。																		
胃内pHを上昇させる薬剤 プロトンポンプ阻害剤 オメプラゾール、ラベプラゾール、ランソプラゾール等 H ₂ 受容体拮抗剤 ファモチジン、ラニチジン、シメチジン等 [16.7.4、16.7.5参照]	本剤の有効性が減弱するおそれがあるため、これらの薬剤との併用を避けることが望ましい。	これらの薬剤による胃内pHの上昇により本剤の溶解性が低下し、本剤の血中濃度が低下する可能性がある。																		
11. 副作用 次の副作用があらわれることがあるので、観察を十分に行い、異常が認められた場合には投与を中止するなど適切な処置を行うこと。 11.1 重大な副作用 11.1.1 肝機能障害 ALT増加（16.3%）、AST増加（16.3%）等の肝機能障害があらわれることがある。〔7.2、8.1参照〕 11.1.2 間質性肺疾患 肺臓炎（1.1%）等があらわれることがある。〔7.2、8.2参照〕 11.2 その他の副作用 <table><tr><th></th><th>5%以上</th><th>1%～5%未満</th><th>1%未満</th></tr><tr><td>感染症および寄生虫症</td><td></td><td></td><td>蜂巣炎、憩室炎、口腔カンジダ症、口腔真菌感染、膿疱性皮疹、上気道感染</td></tr><tr><td>良性、悪性および詳細不明の新生物（嚢胞およびポリープを含む）</td><td></td><td></td><td>血管筋脂肪腫</td></tr><tr><td>血液およびリンパ系障害</td><td></td><td>貧血、リンパ球減少、白血球減少、</td><td>赤血球減少</td></tr></table>					5%以上	1%～5%未満	1%未満	感染症および寄生虫症			蜂巣炎、憩室炎、口腔カンジダ症、口腔真菌感染、膿疱性皮疹、上気道感染	良性、悪性および詳細不明の新生物（嚢胞およびポリープを含む）			血管筋脂肪腫	血液およびリンパ系障害		貧血、リンパ球減少、白血球減少、	赤血球減少	20170543試験の結果及び/又はCDSに基づき設定した。なお、本剤の臨床試験結果はMedDRAの基本語を用いて集計した。
	5%以上	1%～5%未満	1%未満																	
感染症および寄生虫症			蜂巣炎、憩室炎、口腔カンジダ症、口腔真菌感染、膿疱性皮疹、上気道感染																	
良性、悪性および詳細不明の新生物（嚢胞およびポリープを含む）			血管筋脂肪腫																	
血液およびリンパ系障害		貧血、リンパ球減少、白血球減少、	赤血球減少																	

使用上の注意（案）				設定根拠
		好中球減少、血小板減少		
免疫系障害			薬物過敏症	
代謝および栄養障害		食欲減退、低カリウム血症、脱水、脂質異常症、高トリグリセリド血症、低マグネシウム血症、低リン酸血症	低血糖、低ナトリウム血症	
精神障害			錯乱状態、不眠症	
神経系障害		頭痛、味覚不全	浮動性めまい、知覚過敏、過眠症、神経系障害、末梢性ニューロパチー	
眼障害			霧視、視力障害、眼球乾燥症	
	5%以上	1%～5%未満	1%未満	
心臓障害			洞性徐脈	
血管障害			高血圧、低血圧、末梢冷感	
呼吸器、胸郭および縦隔障害		呼吸困難	咳嗽、鼻閉、湿性咳嗽	
胃腸障害	下痢（27.9%）、悪心（16.3%）、嘔吐、腹痛	口内乾燥、腹部膨満、便秘、消化不良、胃食道逆流性疾患、口内炎	呼気臭、鼓腸、口腔粘膜水疱形成	
肝胆道系障害			肝炎	
皮膚および皮下組織障害		斑状丘疹状皮疹、皮膚乾燥、そう痒症、脱毛症、発疹	皮膚炎、ざ瘡様皮膚炎、多汗症、寝汗、光線過敏性反応、紫斑、乾皮症	
筋骨格系および結合組織障害		筋肉痛、関節痛	関節炎、筋力低下、変形性関節症	
腎および尿路障害			白血球尿、頻尿、蛋白尿	
生殖系および乳房障害			女性化乳房	
一般・全身障害および投与部位の状態	疲労（11.1%）	末梢性浮腫、無力症、倦怠感、発熱	びくびく感、限局性浮腫、粘膜の炎症、非心臓性胸痛、浮腫	
臨床検査		体重減少、血中コレステロール増加、血中クレアチニン増加、	AST減少、血中コルチコトロピン減少、血中クレアチンホスホキナーゼ増加、コルチゾール減少、心電図QT延長、胃内pH低下、リパーゼ増加	
14. 適用上の注意				
14.1 薬剤交付時の注意 PTP包装の薬剤はPTPシートから取り出して服用するよう指導すること。PTPシートの誤飲により、硬い鋭角部が食道粘膜へ刺入し、更には穿孔をおこして縦隔洞炎等の重篤な合併症を併発することがある。				
14.2 薬剤投与時の注意 粉砕や分割して使用しないこと。				本剤を粉砕や分割して使用したデータはないことから設定した。

1. JAN

医薬品一般的名称は、2021年4月13日付薬生薬審発0413第1号により、以下のとおり通知された。

JAN :

(日本名) ソトラシブ

(英 名) Sotorasib

化学名 :

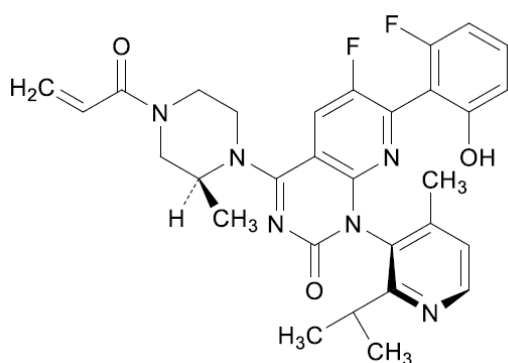
(日本名)

(1*M*)-6-フルオロ-7-(2-フルオロ-6-ヒドロキシフェニル)-1-[4-メチル-2-(プロパン-2-イル)ピリジン-3-イル]-4-[(2*S*)-2-メチル-4-(プロパ-2-エノイル)ピペラジン-1-イル]ピリド[2,3-*d*]ピリミジン-2(1*H*)-オン

(英 名)

(1*M*)-6-Fluoro-7-(2-fluoro-6-hydroxyphenyl)-1-[4-methyl-2-(propan-2-yl)pyridin-3-yl]-4-[(2*S*)-2-methyl-4-(prop-2-enoyl)piperazin-1-yl]pyrido[2,3-*d*]pyrimidin-2(1*H*)-one

化学構造式 :



2. INN

r-INN : sotorasib

WHO Drug Information, Vol. 35, No.1, 2021, Recommended INN: List 85に掲載されている。

薬生薬審発 0413 第 1 号
令和 3 年 4 月 13 日

各都道府県衛生主管部（局）長 殿

厚生労働省医薬・生活衛生局医薬品審査管理課長
（ 公 印 省 略 ）

医薬品の一般的名称について

標記については、「医薬品の一般的名称の取扱いについて（平成 18 年 3 月 31 日薬食発第 0331001 号厚生労働省医薬食品局長通知）」等により取り扱っているところです。今般、我が国における医薬品の一般的名称（以下「JAN」という。）について、新たに別添のとおり定めたので、御了知の上、貴管下関係業者に周知方よろしく御配慮願います。

（参照）

「日本医薬品一般的名称データベース」<https://jpdbs.nihs.go.jp/jan/Default.aspx>
（別添の情報のうち、JAN 以外の最新の情報は、当該データベースの情報で対応することとしています。）

(別表2) INNに収載された品目の我が国における医薬品一般的名称

(平成18年3月31日薬食審査発第0331001号厚生労働省医薬食品局審査管理課長通知に示す別表2)

登録番号 302-4-B4

JAN (日本名) : スペソリマブ (遺伝子組換え)

JAN (英 名) : Spesolimab (Genetical Recombination)

アミノ酸配列及びジスルフィド結合

L鎖

```

QIVLTQSPGT LSLSPGERAT MTCTASSSVS SSYFHWYQQK PGQAPRLWIY
                |
RTSRLASGVP DRFSGSGSGT DFTLTISRLE PEDAATYYCH QFHRSPITFG
                |
AGTKLEIKRT VAAPSVFIFP PSDEQLKSGT ASVVCLLNNF YPREAKVQWK
                |
VDNALQSGNS QESVTEQDSK DSTYSLSTL TLSKADYEKH KVIACEVTHQ
GLSSPVTKSF NRGEK
  
```

H鎖

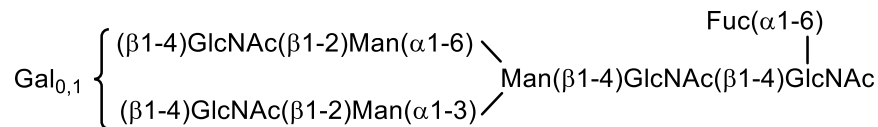
```

QVQLVQSGAE VKKPGASVKV SCKASGYSFT SSWIHWVKQA PGQGLEWMGE
                |
INPGNVRTNY NENFRNKVTM TVDTSISTAY MELSRLRSD TAVYYCTVVF
                |
YGEPIYFPYWG QGTLVTVSSA STKGPSVFPL APSSKSTSGG TAALGCLVKD
|
YFPEPVTVSW NSGALTSGVH TFPVAVLQSSG LYSLSVTV PSSSLGTQTY
|
ICNVNHNKPSN TKVDKRVKPK SCDKTHTCPP CPAPEAAGGP SVFLFPPKPK
DTLMISRTPE VTCVVDVSH EDPEVKFNWY VDGVEVHNAK TKPREEQYNS
                |
TYRVVSVLTV LHQDWLNGKE YKCKVSNKAL PAPIEKTISK AKGQPREPQV
YTLPPSREEM TKNQVSLTCL VKGFYPSDIA VEWESNGQPE NNYKTTTPVL
                |
DSDGSFFLYS KLTVDKSRWQ QGNVFSCSVM HEALHNHYTQ KSLSLSPGK
  
```

L鎖 Q1, H鎖 Q1 : 部分的ピログルタミン酸 ; H鎖 N299 : 糖鎖結合 ; H鎖 K449 : 部分的プロセシング

L鎖 C215 – H鎖 C222, H鎖 C228 – H鎖 C228, H鎖 C231 – H鎖 C231 : ジスルフィド結合

主な糖鎖の推定構造



C₆₄₈₀H₉₉₈₈N₁₇₃₆O₂₀₁₂S₄₆ (タンパク質部分, 4 本鎖)

H 鎖 C₂₂₀₁H₃₃₉₂N₅₈₂O₆₇₄S₁₇

L 鎖 C₁₀₃₉H₁₆₀₆N₂₈₆O₃₃₂S₆

スペソリマブは遺伝子組換え抗ヒトインターロイキン-36 受容体モノクローナル抗体であり, その相補性決定部はマウス抗体に由来し, その他はヒト IgG1 に由来する. スペソリマブは, チャイニーズハムスター卵巣細胞により産生される. スペソリマブは, 449 個のアミノ酸残基からなる H 鎖 (γ1 鎖) 2 本及び 215 個のアミノ酸残基からなる L 鎖 (κ 鎖) 2 本で構成される糖タンパク質 (分子量: 約 149,000) である.

Spesolimab is a recombinant anti-human interleukin-36 receptor monoclonal antibody, the complementarity-determining regions of which are derived from mouse antibody and other regions are derived from human IgG1. Spesolimab is produced in Chinese hamster ovary cells. Spesolimab is a glycoprotein (molecular weight: ca. 149,000) composed of 2 H-chains (γ1-chains) consisting of 449 amino acid residues each and 2 L-chains (κ-chains) consisting of 215 amino acid residues each.

登録番号 302-4-B6

JAN（日本名）：ガレトスマブ（遺伝子組換え）

JAN（英 名）：Garetosmab (Genetical Recombination)

アミノ酸配列及びジスルフィド結合

L鎖

```
EIVLTQSPGT LSLSPGERAT LSCRASQSVS SSYLAWYQQK PGQAPRLLIY
                                     |
GASSRATGIP DRFSGSGSGT DFTLTISRLE PEDFAVYYCQ QYGSSPWTFG
                                     |
QGTKVEIKRT VAAPSVFIFP PSDEQLKSGT ASVVCLLNNF YPREAKVQWK
                                     |
VDNALQSGNS QESVTEQDSK DSTYSLSTL TLSKADYEKH KVIACEVTHQ
                                     |
GLSSPVTKSF NRGEC
```

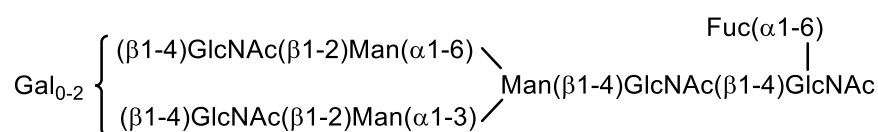
H鎖

```
QVQLQESGPG LVKPSETLSL TCTVSGGSFS SHFWSWIRQP PGKGLEWIGY
                                     |
ILYTGGTSFN PSLKSRVSMS VGTSKNQFSL KLSSVTAADT AVYYCARARS
                                     |
GITFTGIIVP GSFDIWQGT MVTVSSASTK GPSVFPLAPC SRSTSESTAA
                                     |
LGCLVKDYFP EPVTVSWNSG ALTSGVHTFP AVLQSSGLYS LSSVVTVPSS
|
SLGTKTYTCN VDHKPSNTKV DKRVESKYGP PCPPCPAPEF LGGPSVFLFP
|
PKPKDTLMIS RTPEVTCVVV DVSQEDPEVQ FNWYVDGVEV HNAKTKPREE
|
QFNSTYRVVS VLTVLHQDWL NGKEYKCKVS NKGLPSSIEK TISKAKGQPR
|
EPQVYTLPPS QEEMTKNQVS LTCLVKGFYP SDIAVEWESN GQPENNYKTT
|
PPVLDSDGSF FLYSRLTVDK SRWQEGNVFS CSVMHEALHN HYTQKSLSL
|
LGK
```

H鎖 Q1：部分的ピログルタミン酸；H鎖 N303：糖鎖結合；H鎖 K453：部分的プロセシング

L鎖 C215－H鎖 C140, H鎖 C232－H鎖 C232, H鎖 C235－H鎖 C235：ジスルフィド結合

主な糖鎖の推定構造



C₆₄₇₂H₁₀₀₀₄N₁₇₁₂O₂₀₂₄S₄₂ (タンパク質部分, 4 本鎖)

H 鎖 C₂₂₀₅H₃₄₀₉N₅₇₇O₆₇₇S₁₆

L 鎖 C₁₀₃₁H₁₅₉₇N₂₇₉O₃₃₅S₅

ガレトスマブはヒトアクチビン A, アクチビン AB 及びアクチビン AC に対する遺伝子組換えモノクローナル抗体であり, ヒト IgG4 に由来する. H 鎖の 234 番目のアミノ酸残基は Pro に置換されている. ガレトスマブはチャイニーズハムスター卵巣細胞により産生される. ガレトスマブは, 453 個のアミノ酸残基からなる H 鎖 (γ4 鎖) 2 本及び 215 個のアミノ酸残基からなる L 鎖 (κ 鎖) 2 本で構成される糖タンパク質 (分子量: 約 148,000) である.

Garetosmab is a recombinant monoclonal antibody against human activin A, activin AB and activin AC derived from human IgG4. In the H-chain, amino acid residue at position 234 is substituted by Pro. Garetosmab is produced in Chinese hamster ovary cells. Garetosmab is a glycoprotein (molecular weight: ca. 148,000) composed of 2 H-chains (γ4-chains) consisting of 453 amino acid residues each and 2 L-chains (κ-chains) consisting of 215 amino acid residues each.

登録番号 302-4-B9

JAN（日本名）：ボソリチド（遺伝子組換え）

JAN（英 名）：Vosoritide (Genetical Recombination)

アミノ酸配列及びジスルフィド結合

PGQEHPNARK YKGANKKGLS KGCFGLKLDR IGSMGSLGC

$C_{176}H_{290}N_{56}O_{51}S_3$

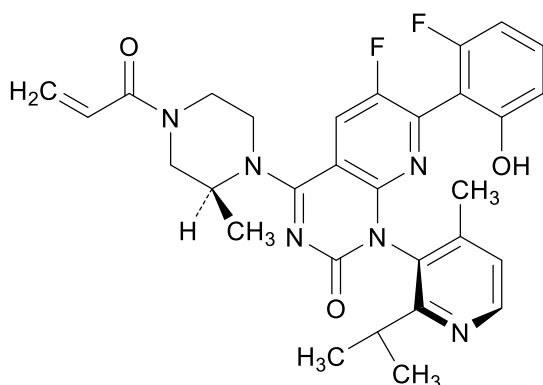
ボソリチドは、遺伝子組換えヒトC型ナトリウム利尿ペプチド（CNP）類縁体であり、ヒトCNP-53の17～53番目のアミノ酸残基に相当し、N末端に2個のアミノ酸残基（Pro-Gly）が付加している。ボソリチドは39個のアミノ酸残基からなるペプチドである。

Vosoritide is a recombinant human C-type natriuretic peptide (CNP) analog, which corresponds to amino acid residues at positions 17–53 of human CNP-53, and two amino acid residues (Pro-Gly) are attached to the N-terminus. Vosoritide is a peptide consisting of 39 amino acid residues.

登録番号 302-6-B1

JAN（日本名）：ソトラシブ

JAN（英名）：Sotorasib



C₃₀H₃₀F₂N₆O₃

(1*M*)-6-フルオロ-7-(2-フルオロ-6-ヒドロキシフェニル)-1-[4-メチル-2-(プロパン-2-イル)ピリジン-3-イル]-4-[(2*S*)-2-メチル-4-(プロパ-2-エノイル)ピペラジン-1-イル]ピリド[2,3-*d*]ピリミジン-2(1*H*)-オン

(1*M*)-6-Fluoro-7-(2-fluoro-6-hydroxyphenyl)-1-[4-methyl-2-(propan-2-yl)pyridin-3-yl]-4-[(2*S*)-2-methyl-4-(prop-2-en-1-yl)piperazin-1-yl]pyrido[2,3-*d*]pyrimidin-2(1*H*)-one

※ JAN 以外の情報は、参考として掲載しました。

International Nonproprietary Names for Pharmaceutical Substances (INN)

RECOMMENDED International Nonproprietary Names: List 85

Notice is hereby given that, in accordance with paragraph 7 of the Procedure for the Selection of Recommended International Nonproprietary Names for Pharmaceutical Substances [*Off. Rec. Wld Health Org.*, 1955, **60**, 3 (Resolution EB15.R7); 1969, **173**, 10 (Resolution EB43.R9); Resolution EB115.R4 (EB115/2005/REC/1)], the following names are selected as Recommended International Nonproprietary Names. The inclusion of a name in the lists of Recommended International Nonproprietary Names does not imply any recommendation of the use of the substance in medicine or pharmacy.

Lists of Proposed (1–117) and Recommended (1–78) International Nonproprietary Names can be found in *Cumulative List No. 17, 2017* (available in CD-ROM only).

Dénominations communes internationales des Substances pharmaceutiques (DCI)

Dénominations communes internationales RECOMMANDÉES: Liste 85

Il est notifié que, conformément aux dispositions du paragraphe 7 de la Procédure à suivre en vue du choix de Dénominations communes internationales recommandées pour les Substances pharmaceutiques [*Actes off. Org. mond. Santé*, 1955, **60**, 3 (résolution EB15.R7); 1969, **173**, 10 (résolution EB43.R9); résolution EB115.R4 (EB115/2005/REC/1)] les dénominations ci-dessous sont choisies par l'Organisation mondiale de la Santé en tant que dénominations communes internationales recommandées. L'inclusion d'une dénomination dans les listes de DCI recommandées n'implique aucune recommandation en vue de l'utilisation de la substance correspondante en médecine ou en pharmacie.

On trouvera d'autres listes de Dénominations communes internationales proposées (1–117) et recommandées (1–78) dans la *Liste récapitulative No. 17, 2017* (disponible sur CD-ROM seulement).

Denominaciones Comunes Internacionales para las Sustancias Farmacéuticas (DCI)

Denominaciones Comunes Internacionales RECOMENDADAS: Lista 85

De conformidad con lo que dispone el párrafo 7 del Procedimiento de Selección de Denominaciones Comunes Internacionales Recomendadas para las Sustancias Farmacéuticas [*Act. Of. Mund. Salud*, 1955, **60**, 3 (Resolución EB15.R7); 1969, **173**, 10 (Resolución EB43.R9); Resolución EB115.R4 (EB115/2005/REC/1) EB115.R4 (EB115/2005/REC/1)], se comunica por el presente anuncio que las denominaciones que a continuación se expresan han sido seleccionadas como Denominaciones Comunes Internacionales Recomendadas. La inclusión de una denominación en las listas de las Denominaciones Comunes Recomendadas no supone recomendación alguna en favor del empleo de la sustancia respectiva en medicina o en farmacia.

Las listas de Denominaciones Comunes Internacionales Propuestas (1–117) y Recomendadas (1–78) se encuentran reunidas en *Cumulative List No. 17, 2017* (disponible sólo en CD-ROM).

Latin , English, French, Spanish:	<i>Chemical name or description; Molecular formula;</i>
<i>Recommended INN</i>	<i>Graphic formula</i>
<i>DCI Recommandée</i>	<i>Nom chimique ou description; Formule brute;</i>
	<i>Formule développée</i>
<i>DCI Recomendada</i>	<i>Nombre químico o descripción; Fórmula molecular;</i>
	<i>Fórmula desarrollada</i>

Please find the names from proposed INN List 124 – COVID-19 (special edition) that have reached the status of recommended INN in annex.
 Veuillez trouver les noms de la Liste 124 des DCI proposées – COVID-19 (édition spéciale) ayant atteint le statut de DCI recommandées en annexe.
 Por favor, abajo se encuentran los nombres de la Lista 124 de DCI propuestas – COVID-19 (edición especial) que han alcanzado el estatus de DCI recomendadas.

acapatamabum

acapatamab

immunoglobulin scFv-scFv-scFc, anti-[*Homo sapiens* FOLH1 (folate hydrolase, prostate specific membrane antigen, PSMA)] and anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], monoclonal antibody single chain (scFv)2-scFc, bispecific; IG single chain scFv-scFv-scFc, anti FOLH1 and anti-CD3E (1-986) [scFv-VH-V-kappa anti-FOLH1 (1-243) [VH (*Homo sapiens* IGHV3-11*01 G49>C (44) (84.7%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -15-mer-tris(tetraglycyl-seryl) linker (122-136)-V-KAPPA (*Homo sapiens* IGKV1-16*01 (79.3%) -IGKJ2*01 Q120>C (236) (90.9%)) CDR-IMGT [6.3.9] (163-168.186-188.225-233) (137-243)] -6-mer seryl-tetraglycyl-seryl linker (244-249) -scFv-VH-V-lambda anti-CD3E (250-498) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87.0%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (275-282.300-309.348-363) (250-374) -15-mer-tris(tetraglycyl-seryl) linker (375-389) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (415-423.441-443.480-488) (390-498)] -4-mer- tetraglycyl linker (499-502) -scFc (h-CH2-CH3)-(h-CH2-CH3) (503-986) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (hinge 6-15 (503-512), CH2 R83>C (574), N84.4>G (579), V85>C (584) (513-622), CH3 E12 (638), M14 (640) (623-727), CHS (728-729)) (503-729) -30-mer hexakis(tetraglycyl-seryl) linker (730-759) -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (hinge 6-15 (760-769), CH2 R83>C (831), N84.4>G (836), V85>C (841) (770-879), CH3 E12 (895), M14 (897) (880-984), CHS (985-986)) (760-986)]], non-glycosylated, produced in Chinese hamster ovary (CHO) cells

acapatamab

immunoglobuline scFv-scFv-scFc, anti-[*Homo sapiens* FOLH1 (folate hydrolase, antigène membranaire spécifique de la prostate, PSMA)], et anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], anticorps monoclonal à chaîne unique (scFv)2-scFc, bispécifique;

IG à chaîne unique scFv-scFv-scFc, anti FOLH1 et anti-CD3E (1-986) [scFv-VH-V-kappa anti-FOLH1 (1-243) [VH (*Homo sapiens*IGHV3-11*01 G49>C (84.7%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -15-mer-tris(tétraglycyl-séryl) linker (122-136)-V-KAPPA (*Homo sapiens* IGKV1-16*01 (79.3%) -IGKJ2*01 Q120>C (236) (90.9%)) CDR-IMGT [6.3.9] (163-168.186-188.225-233) (137-243)] -6-mer-séryl-tétraglycyl-séryl linker (244-249) -scFv-VH-V-lambda anti-CD3E (250-498) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87.0%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (275-282.300-309.348-363) (250-374) -15-mer-tris(tétraglycyl-séryl) linker (375-389) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (415-423.441-443.480-488) (390-498)] -4-mer-tétraglycyl linker (499-502) -scFc (h-CH2-CH3)-(h-CH2-CH3) (503-986) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (charnière 6-15 (503-512), CH2 R83>C (574), N84.4>G (579), V85>C (584) (513-622), CH3 E12 (638), M14 (640) (623-727), CHS (728-729)) (503-729) -30-mer-hexakis(tétraglycyl-séryl) linker (730-759) -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (charnière 6-15 (760-769), CH2 R83>C (831), N84.4>G (836), V85>C (841) (770-879), CH3 E12 (895), M14 (897) (880-984), CHS (985-986)) (760-986)]], non-glycosylé, produit dans des cellules ovariennes de hamster chinois (CHO)

acapatamab

inmunoglobulina scFv-scFv-scFc, anti-[*Homo sapiens* FOLH1 (folato hidrolasa, antígeno membranario específico de la prostata, PSMA)], y anti-[*Homo sapiens* CD3E (CD3 épsilon, Leu-4)], anticuerpo monoclonal con cadena única (scFv)2-scFc, biespecífica;
IG con cadena única scFv-scFv-scFc, anti FOLH1 y anti-CD3E (1-986) [scFv-VH-V-kappa anti-FOLH1 (1-243) [VH (*Homo sapiens* IGHV3-11*01 G49>C (84.7%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -15-mer-tris(tetraglicil-seril) linker (122-136)-V-KAPPA (*Homo sapiens* IGKV1-16*01 (79.3%) -IGKJ2*01 Q120>C (236) (90.9%)) CDR-IMGT [6.3.9] (163-168.186-188.225-233) (137-243)] -6-mer-seril-tetraglicil-seril linker (244-249) -scFv-VH-V-lambda anti-CD3E (250-498) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87.0%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (275-282.300-309.348-363) (250-374) -15-mer-tris(tetraglicil-seril) linker (375-389) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (415-423.441-443.480-488) (390-498)] -4-mer-tetraglicil linker (499-502) -scFc (h-CH2-CH3)-(h-CH2-CH3) (503-986) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (bisagra 6-15 (503-512), CH2 R83>C (574), N84.4>G (579), V85>C (584) (513-622), CH3 E12 (638), M14 (640) (623-727), CHS (728-729)) (503-729) -30-mer-hexakis(tetraglicil-seril) linker (730-759) -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (bisagra 6-15 (760-769), CH2 R83>C (831), N84.4>G (836), V85>C (841) (770-879), CH3 E12 (895), M14 (897) (880-984), CHS (985-986)) (760-986)]], no glicosilado, producido en las células ováricas de hámster chino (CHO)

Heavy chain / Chaîne lourde / Cadena pesada			
QVQLVESGGG	LVPKPGESLR	SCAASGFTFS	DYYMYWVRQA
ISDGGYYTY	SDIIGKRFIT	SRDNAKNSLY	LQMNSLKAED
PLLRHGAMY	WGQGLVTVS	SGGGSGGGG	SGGGGSDIQM
VGDRVITICK	ASQNVDTNVA	WYQKPGQAP	KSLIYSASYV
SASGTDFTLT	ISSVQSEDF	TYQCQYDQ	LITFGCGTKL
VQLVESGGGL	VQPGGSLKLS	CAASGFTFNK	YAMNWRQAP
RSKYNNYATY	YADSVKDRFT	ISRDDSKNTA	YLQMNMLKTE
GNFGNSYISY	WAYWQGGTLV	TVSSGGGGSG	GGSGGGGGSQ
VSPGGTVTLT	CGSSTGAVTS	GNYPNWWQK	PGQAPRLIG
ARFSGSLGG	KAALTLGSGV	PEDEAEYYCV	LWYSNRWVFG
GGDKTHTCP	CPAPELLGGP	SVFLFPPKPK	DTLMISRTPE
EDPEVKFMWY	VDGVEVHNK	TKPCEEQYGS	TYRCVSLTV
YKCKVSNKAL	PAPIEKTISK	AKGQPREPQV	YTLPPSREEM
VKGFPYSDIA	VEVESNGQPE	NNYKTTTPVL	DSGDSFFLYS
QGNVFSQSV	HEALHNHYTQ	KSLSLSPGKG	GGSGGGGGSG
GGSGGGGGSD	KTHTCPPCPA	PELLGGPSVF	LFPFKPKDTL
VVVDVSHEDP	EVKFNWYVDG	VEVHNKATKP	CEEQYGSTYR
DWLNKGEYK	KVSNKALPAP	IEKTISKAKG	QPREPQVYTL
QVSLTCLVKG	FYPSDIAVEW	ESNGQPENNY	KTTTPVLDSD
VDKSRWQQGN	VFSCSVMEHA	LHNHYTQKSL	SLSPGK

Post-translational modifications			
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro			
Intra-V (C23-C104)	22-96	159-224	271-347
Intra-C (C23-C104)	543-603	649-707	800-860
Intra-CH2 (C83-C85)	574-584	831-841	
Inter-VH-VL (C49-C120)	44-236		
Inter-h11 (h 11 - h 11)	508-765		
Inter-h14 (h 14 - h 14)	511-768		

No N-glycosylation sites / pas de sites de N-glycosylation / ningún posición de N-glicosilación
CH2 N84.4-G:
579, 836
Aglycosylated

C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal
CHS K2:
986

adrenomedullinum pegolum #
adrenomedullin pegol

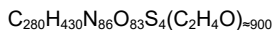
human adrenomedullin, C-terminal L-tyrosil amide, pegylated at O⁴ of Y¹ with [(3S)-3-amino-4-[(2R)-1-amino-3-[(3RS)-2,5-dioxo-1-{3-oxo-3-[α-methyl(polyoxyethylene)-ω-amino]propyl]pyrrolidin-3-yl)sulfanyl]-1-oxopropan-2-yl]amino]-4-oxobutyl]carbamoyl; O^{4.1}-[(3S)-3-amino-4-[(2R)-1-amino-3-[(3RS)-1-{3-[α-methylpoly(oxyethylene)-ω-amino]-3-oxopropyl]-2,5-dioxopyrrolidin-3-yl)sulfanyl]-1-oxopropan-2-yl]amino]-4-oxobutyl]carbamoyl]adrenomedullin (human)

adrénomedulline pégol

adrénomédulline humaine, L-tyrosil amide C-terminal, pégylée en O⁴ de Y¹ avec [(3S)-3-amino-4-[(2R)-1-amino-3-[(3RS)-2,5-dioxo-1-{3-oxo-3-[α-méthyl(polyoxyéthylène)-ω-amino]propyl]pyrrolidin-3-yl)sulfanyl]-1-oxopropan-2-yl]amino]-4-oxobutyl]carbamoyl; O^{4.1}-[(3S)-3-amino-4-[(2R)-1-amino-3-[(3RS)-1-{3-[α-méthylpoly(oxyéthylène)-ω-amino]-3-oxopropyl]-2,5-dioxopyrrolidin-3-yl)sulfanyl]-1-oxopropan-2-yl]-4-oxobutyl]carbamoyl]adrénomédulline (humaine)

adrenomedulina pegol

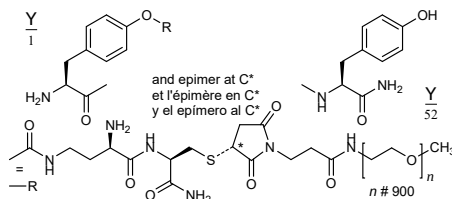
adrenomedulina humana, C-terminal L-tirosil amida, pegilada en O⁴ de Y¹ con [(3S)-3-amino-4-[(2R)-1-amino-3-[(3RS)-2,5-dioxo-1-{3-oxo-3-[α-metil(polioxi-etileno)-ω-amino]propil]pirrolidin-3-il)sulfanil]-1-oxopropan-2-il]amino]-4-oxobutil]carbamoilo; O^{4.1}-[(3S)-3-amino-4-[(2R)-1-amino-3-[(3RS)-1-{3-[α-metilpoli(oxi-etileno)-ω-amino]-3-oxopropil]-2,5-dioxopirrolidin-3-il)sulfanil]-1-oxopropan-2-il]-4-oxobutil]carbamoil]adrenomedulina (humana)



Sequence / Séquence / Secuencia

YRQSMNFGG LRSFGCRFGT CTVQKLAHQI YQFTDKDKDN VAPRSKISPQ 50
GY 52Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
36-86 43-89

Modified residues / Résidus modifiés / Restos modificados

**aldumastatum**

aldumastat

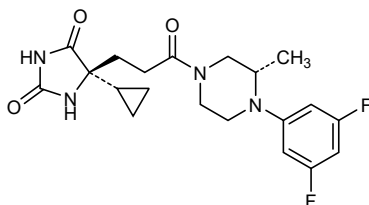
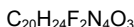
(5S)-5-cyclopropyl-5-{3-[(3S)-4-(3,5-difluorophenyl)-3-methylpiperazin-1-yl]-3-oxopropyl}imidazolidine-2,4-dione

aldumastat

(5S)-5-cyclopropyl-5-{3-[(3S)-4-(3,5-difluorophényl)-3-méthylpipérazin-1-yl]-3-oxopropyl}imidazolidine-2,4-dione

aldumastat

(5S)-5-ciclopropil-5-{3-[(3S)-4-(3,5-difluorofenil)-3-metilpiperazin-1-il]-3-oxopropil}imidazolidina-2,4-diona

**alnuctamabum #**

alnuctamab

immunoglobulin G1-kappa/lambda with domain crossover, anti-[*Homo sapiens* TNFRSF17 (TNF receptor superfamily member 17, tumor necrosis factor receptor superfamily, member 17, B cell maturation antigen, BCMA, BCM, TNFRSF13A, CD269)] and anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], monoclonal antibody, bispecific, trivalent; gamma1-lambda heavy chain anti-TNFRSF17 and anti-CD3E (VH-CH1-V-lambda-CH1-h-CH2-CH3) (1-671) [VH anti-TNFRSF17 (*Homo sapiens* IGHV3-23*01 (94.9%) -(IGHD) -IGHJ5*01 (93.3%)) CDR-IMGT [8.8.9] (26-33.51-58.97-105) (1-116) -*Homo sapiens* IGHG1*01, G1m17 (CH1 K26>E (146), K119>E (212), K120 (213) (117-214), hinge 1-6 (215-220)) (117-220) -10-mer bis(tetraglycyl-seryl) linker (221-230) -V-LAMBDA anti-CD3E (*Mus musculus* IGLV1*01 (81.2%) -IGLJ1*01 (100%)]/*Homo sapiens* IGLV7-46*01 (80.0%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (256-264.282-284.321-329) (231-339)) -2-mer diseryl linker (340-341) -*Homo sapiens* IGHG1*01 G1m17,1, G1v14 CH2 A1.3, A1.2, G1v32 CH3 W22 (CH1 K120 (438) (342-439), hinge 1-15 (440-454), CH2 L1.3>A (458), L1.2>A (459), P114>G (553) (455-564), CH3 S10>C (578), D12 (580), L14 (582), T22>W (590) (565-669), CHS (670-671)) (342-671)],

(219-216')-disulfide with kappa light chain anti-TNFRSF17 (1'-216') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (90.6%) -IGKJ1*01 (91.7%)) CDR-IMGT [7.3.10] (27-33.51-53.90-99) (1'-109') -*Homo sapiens* IGKC*01 (98.1%) E12>R (125), Q13>K (126), Km3 A45.1 (155), V101 (193) (110'-216')]; (444-232'')-disulfide with VH-C-kappa light chain anti-CD3E, (1'''-232''') [VH (*Homo sapiens* IGHV3-23*03 (87.0%) - (IGHD) -IGHJ6*01 (90.9%)) CDR-IMGT [8.10.16] (26-33.51-60.99-114) (1'''-125''') -*Homo sapiens* IGKC*01 (99.1%) T1.3>S (127), Km3 A45.1 (171), V101 (209) (126'''-232''')]; gamma1 heavy chain anti-TNFRSF17 (1''-446'') [VH (*Homo sapiens* IGHV3-23*01 (94.9%) - (IGHD) -IGHJ5*01 (93.3%)) CDR-IMGT [8.8.9] (26-33.51-58.97-105) (1''-116'') -*Homo sapiens* IGHG1*01 G1m17,1, G1v14 CH2 A1.3, A1.2, G1v33 CH3 S22, A24, V86 (CH1 K26>E (146), K119>E (212), K120 (213) (117''-214''), hinge 1-15 (215''-229''), CH2 L1.3>A (233), L1.2>A (234), P114>G (328) (230''-339''), CH3 Y5>C (348), D12 (355), L14 (357), T22>S (365), L24>A (367), Y86>V (406) (340''-444''), CHS (445''-446'') (117''-446'')]; (219''-216''')-disulfide with kappa light chain anti-TNFRSF17 (1''''-216''') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (90.6%) -IGKJ1*01 (91.7%)) CDR-IMGT [7.3.10] (27-33.51-53.90-99) (1''''-109''') -*Homo sapiens* IGKC*01 (98.1%) E12>R (125), Q13>K (126), Km3 A45.1 (155), V101 (193) (110''''-216''')]; dimer (450-225'':453-228':578-348'')-trisdysulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

alnuctamab

immunoglobuline G1-kappa/lambda avec domaines échangés, anti-[*Homo sapiens* TNFRSF17 (membre 17 de la superfamille des récepteurs du TNF, membre 17 de la superfamille des récepteurs du facteur de nécrose tumorale, antigène de maturation de cellule B, BCMA, BCM, TNFRSF13A, CD269)] et anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], anticorps monoclonal, bispécifique, trivalent; chaîne lourde gamma1 anti-TNFRSF17 et anti-CD3E (VH-CH1-V-lambda-CH1-h-CH2-CH3) (1-671) [VH anti-TNFRSF17 (*Homo sapiens* IGHV3-23*01 (94.9%) - (IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.9] (26-33.51-58.97-105) (1-116) -*Homo sapiens* IGHG1*01 G1m17 (CH1 K26>E (146), K119>E (212), K120 (213) (117-214), charnière 1-6 (215-220)) (117-220) -10-mer bis(tétraglycyl-séryl) linker (221-230) -V-LAMBDA anti-CD3E (*Mus musculus* IGLV1*01 (81.2%) -IGLJ1*01 (100%)/*Homo sapiens* (IGLV7-46*01 (80.0%) -IGLJ3*02 (100%) CDR-IMGT [9.3.9] (256-264.282-284.321-329) (231-339)) -2-merdiséryl linker (340-341) -*Homo sapiens* IGHG1*01 G1m17,1,G1v14 CH2 A1.3, A1.2, G1v32 CH3 W22 (CH1 K120 (438) (342-439), charnière 1-15 (440-454), CH2 L1.3>A (458), L1.2>A (459), P114>G (553) (455-564), CH3 S10>C (578), D12 (580), L14 (582), T22>W (590) (565-669), CHS (670-671)) (342-671)], (219-216')-disulfure avec la chaîne légère kappa anti-TNFRSF17 (1'-216') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (90.6%) -IGKJ1*01 (91.7%)) CDR-IMGT [7.3.10] (27-33.51-53.90-99) (1'-109') -*Homo sapiens* IGKC*01 (98.1%) E12>R (125), Q13>K (126), Km3 A45.1 (155), V101 (193) (110'-216')]; (444-232'')-disulfure avec la chaîne légère VH-C-kappa anti-CD3E (1'''-232''') [VH (*Homo sapiens* IGHV3-23*03 (87.0%) -IGHJ6*01 (90.9%)) CDR-IMGT [8.10.16] (26-33.51-60.99-114) (1'''-125''') -*Homo sapiens* IGKC*01 (99.1%) T1.3>S (127), Km3 A45.1 (171), V101 (209) (126'''-232''')];

alnuctamab

chaîne lourde gamma1 anti-TNFRSF17 (1"-446") [VH (*Homo sapiens* IGHV3-23*01 (94.9%) -(IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.9] (26-33.51-58.97-105) (1"-116") -*Homo sapiens* IGHG1*01 G1m17,1, G1v14 CH2 A1.3, A1.2, G1v33 CH3 S22, A24, V86 (CH1 K26>E (146), K119>E (212), K120 (213) (117"-214"), charnière 1-15 (215"-229"), CH2 L1.3>A (233), L1.2>A (234), P114>G (328) (230"-339"), CH3 Y5>C (348), D12 (355), L14 (357), T22>S (365), L24>A (367), Y86>V (406) (340"-444"), CHS (445"-446")) (117"-446")], (219"-216"")-disulfure avec la chaîne légère kappa anti-TNFRSF17 (1""-216"")) [V-KAPPA (*Homo sapiens* IGKV3-20*01 (90.6%) -IGKJ1*01 (91.7%)) CDR-IMGT [7.3.10] (27-33.51-53.90-99) (1""-109"")-*Homo sapiens* IGKC*01 (98.1%) E12>R (125), Q13>K (126), Km3 A45.1 (155), V101 (193) (110""-216"")); dimère (450-225":453-228":578-348")-trisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

immunoglobulina G1-kappa/lambda con dominios cruzados, anti-[*Homo sapiens* TNFRSF17 (miembro 17 de la superfamilia de los receptores del TNF, miembro 17 de la superfamilia de los receptores del factor de necrosis tumoral, antígeno de maduración de célula B, BCMA, BCM, TNFRSF13A, CD269)] y anti-[*Homo sapiens* CD3E (CD3 épsilon, Leu-4)], anticuerpo monoclonal, biespecífico, trivalente; cadena pesada gamma1 anti-TNFRSF17 y anti-CD3E (VH-CH1-V-lambda-CH1-h-CH2-CH3) (1-671) [VH anti-TNFRSF17 (*Homo sapiens* IGHV3-23*01 (94.9%) -(IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.9] (26-33.51-58.97-105) (1-116) -*Homo sapiens* IGHG1*01 G1m17 (CH1 K26>E (146), K119>E (212), K120 (213) (117-214), bisagra 1-6 (215-220)) (117-220) -10-mer bis(tetraglicil-seril) linker (221-230) -V-LAMBDA anti-CD3E (*Mus musculus* IGLV1*01 (81.2%) -IGLJ1*01 (100%))/*Homo sapiens* (IGLV7-46*01 (80.0%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (256-264.282-284.321-329) (231-339)) -2-merdiseril linker (340-341) -*Homo sapiens* IGHG1*01 G1m17,1,G1v14 CH2 A1.3, A1.2, G1v32 CH3 W22 (CH1 K120 (438) (342-439), bisagra 1-15 (440-454), CH2 L1.3>A (458), L1.2>A (459), P114>G (553) (455-564), CH3 S10>C (578), D12 (580), L14 (582), T22>W (590) (565-669), CHS (670-671)) (342-671)], (219-216')-disulfuro con la cadena ligera kappa anti-TNFRSF17 (1'-216') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (90.6%) -IGKJ1*01 (91.7%)) CDR-IMGT [7.3.10] (27-33.51-53.90-99) (1'-109') -*Homo sapiens* IGKC*01 (98.1%) E12>R (125), Q13>K (126), Km3 A45.1 (155), V101 (193) (110'-216')]; (444-232')-disulfuro con la cadena ligera VH-C-kappa anti-CD3E (1""-232"")) [VH (*Homo sapiens* IGHV3-23*03 (87.0%) -IGHJ6*01 (90.9%)) CDR-IMGT [8.10.16] (26-33.51-60.99-114) (1""-125"")-*Homo sapiens* IGKC*01 (99.1%) T1.3>S (127), Km3 A45.1 (171), V101 (209) (126""-232"")); cadena pesada gamma1 anti-TNFRSF17 (1"-446") [VH (*Homo sapiens* IGHV3-23*01 (94.9%) -(IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.9] (26-33.51-58.97-105) (1"-116") -*Homo sapiens* IGHG1*01 G1m17,1, G1v14 CH2 A1.3, A1.2, G1v33 CH3 S22, A24, V86 (CH1 K26>E (146), K119>E (212), K120 (213) (117"-214"), bisagra 1-15 (215"-229"), CH2 L1.3>A (233), L1.2>A (234), P114>G (328) (230"-339"), CH3 Y5>C (348), D12 (355), L14 (357), T22>S (365), L24>A (367), Y86>V (406) (340"-444"), CHS (445"-446")) (117"-446")], (219"-216"")-disulfuro con la cadena ligera kappa anti-TNFRSF17 (1""-216"")) [V-KAPPA (*Homo sapiens* IGKV3-20*01 (90.6%) -IGKJ1*01 (91.7%)) CDR-IMGT [7.3.10] (27-33.51-53.90-99) (1""-109"")-*Homo sapiens* IGKC*01 (98.1%) E12>R (125), Q13>K (126), Km3 A45.1 (155), V101 (193) (110""-216"")); dímero (450-225":453-228":578-348")-trisdisulfuro, producido en las células ováricas de hamster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada (1-671 anti-TNFRSF17 and anti-CD3)		
EVQLLESGGG	LVQPGGSLRL	SCAASGFTFS DNAMGWVRQA PGKGLEWVSA 50
ISGPGSSTYY	ADSVKGRFTI	SRDNSKNTLY LQMNSLRAED TAVYYCAKVL 100
GWFDYWGQGT	LVTVSSASTK	GPSVFPLAPS SKSTSGGTAA LGCLVEDYFP 150
EPVTVSWNSG	ALTSVGHFTF	AVLQSSGLYS LSSVTVTPSS SLGTQTYICN 200
VNHKPSNTKV	DEKVEPKSCD	GGGSGGGGGS QAVVTQEPSL TVSPGGTVTL 250
TCGSSTGAVT	TSNYANWVQE	KPQQAFLRLI GGTNKRAPGT PARFSGSLLG 300
GKAALTLSGA	QPEDEAEYYC	ALWYSNLWVF GGGTKLTVLS SASTKGPSVF 350
PLAPSSKSTS	GGTAALGCLV	KDYFPEPTVW SNWSGALTSV VHTFPAVLQS 400
SGLYSLSSVV	TVPSSSLGTQ	TYICNVNHKP SNTKVDKQVE PKSCDKTHTC 450
PPCPAPEAAG	GPSVFLFPPK	PKDTLMISRT PEVTCVVVDV SHEDPEVKFN 500
WYVDGVVEHN	AKTKPREEQY	NSTYRVVSVL TVLHQDWLNG KEYCKVSNK 550
ALGAPIEKTI	SKAKGQPREP	QVYTLPPCRD ELTKNQVSLW CLVKGFYPSD 600
IAVEWESNGQ	PENNYKTTTP	VLDSDGSFFL YSKLTVDKSR WQQGNVFCSS 650
VMHEALHNHY	TQKLSLSLSPG	K 671

Heavy chain / Chaîne lourde / Cadena pesada (1*-446* anti-TNFRSF17)		
EVQLLESGGG	LVQPGGSLRL	SCAASGFTFS DNAMGWVRQA PGKGLEWVSA 50
ISGPGSSTYY	ADSVKGRFTI	SRDNSKNTLY LQMNSLRAED TAVYYCAKVL 100
GWFDYWGQGT	LVTVSSASTK	GPSVFPLAPS SKSTSGGTAA LGCLVEDYFP 150
EPVTVSWNSG	ALTSVGHFTF	AVLQSSGLYS LSSVTVTPSS SLGTQTYICN 200
VNHKPSNTKV	DEKVEPKSCD	KHTTCCPPCA PEAGGSPSVF LFPPKPKDTL 250
MISRTPEVTC	VVVDVSHEDP	EVKFNWYVDG VEVHNAKTKP REEQYNSTYR 300
VVSVTLVLHQ	DWLNGKEYCK	KVSNKALGAP IEKTIISKAKG QPREPQVCTL 350
PPSRDELTKN	QVSLSCAVKG	FYPSDIAVEW ESNQGPENNY KTTPPVLDSD 400
GSFFLVSKLT	VDKSRWQQGN	VFSCSVMHEA LHNHYTQKSL SLSPGK 446

Light chain / Chaîne légère / Cadena ligera (1*-216* anti-TNFRSF17)		
EIVLTQSPGT	LSLSPGERAT	LSCRASQSVS DEYLSHWYQQ PGQAPRLLIH 50
SASTRATGIP	DRFSGSGSGT	DFTLAISRLE PEDFAVYYCQ QYGYPPDFTF 100
GQGTAKVEIKR	TVAAPSVFIF	PPSDRKLKSG TASVCLLNH FYPREKQVQM 150
KVDNALQSGN	SQESVTEQDS	KDSTYLSLST LTLKADYEK HKVYACEVTH 200
QGLSSPVTKS	FNRGEC	216

Light chain / Chaîne légère / Cadena ligera (1*-232* anti-CD3E)		
EVQLLESGGG	LVQPGGSLRL	SCAASGFTFS TYAMNWVRQA PGKGLEWVSR 50
IRSKYNNYAT	YYADSVKGRF	TIISRDTSKNT LYLQMNSLRA EDTAVYYCVR 100
HGNFGNSYVS	WFAYWGQGT	VTVSSASVAA PSVFIFPPSD EQLKSGTASV 150
VCLLNNFYPR	EAKVQWQVDN	ALQSGNSQES VTEQDSKSDT YLSSTLTLS 200
KADYEKHKVY	ACEVTHQGLS	SPYTKSFNRG EC 232

Post-translational modifications		
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro		
Intra-H (C23-C104)	22-96 143-199 252-320 368-424 485-545 591-649	
	22"-96" 143"-199" 260"-320" 366"-424"	
Intra-L (C23-C104)	23"-89" 136"-196"	
	22"-98" 152"-212"	
	23"-89" 136"-196"	
Inter-H-L (h 5-CL 126)	219-216* 444-232"	219"-216"
Inter-H-H (h 11, h 14)	450-225" 453-228"	
Inter-H-H engineered	578-348"	

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación		
H CH2 N84.4:		
521, 296"		
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.		
C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal		
H CHS K2:		
671, 446"		

ancremonamum
ancremonam

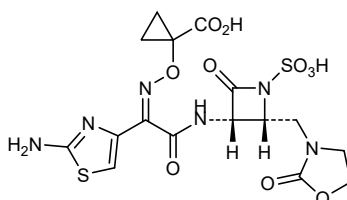
1-(((Z)-[1-(2-amino-1,3-thiazol-4-yl)-2-oxo-2-(((3S,4R)-2-oxo-4-[(2-oxo-1,3-oxazolidin-3-yl)methyl]-1-sulfoazetidid-3-yl)amino)ethylidene]amino)oxy)cyclopropane-1-carboxylic acid

ancrémonam

acide1-(((Z)-[1-(2-amino-1,3-thiazol-4-yl)-2-oxo-2-(((3S,4R)-2-oxo-4-[(2-oxo-1,3-oxazolidin-3-yl)méthyl]-1-sulfoazétidid-3-yl)amino)éthylidène]amino)oxy)cyclopropane-1-carboxylique

ancremonam

ácido1-(((Z)-[1-(2-amino-1,3-tiazol-4-il)-2-oxo-2-(((3S,4R)-2-oxo-4-[(2-oxo-1,3-oxazolidin-3-il)metil]-1-sulfoazetidid-3-il)amino)etilideno]amino)oxi)ciclopropano-1-carboxílico



apitegromabum #
apitegromab

immunoglobulin G4-lambda, anti-[*Homo sapiens* pro-MSTN (pro-growth differentiation factor 8, pro-GDF8, pro-myostatin, pro-GDF-8)], *Homo sapiens* monoclonal antibody;

gamma4 heavy chain *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV3-30*03 (100%) -(IGHD) - IGHJ6*01 (94.4%)) CDR-IMGT [8.8.19] (26-33.51-58.97-115) (1-126)-*Homo sapiens* IGHG4*01, G4v5 h P10 (CH1 (127-224), hinge 1-12 S10>P (234) (225-236), CH2 (237-346), CH3 (347-451), CHS K>del (452)) (127-452)], (140-214')-disulfide with lambda light chain *Homo sapiens* (1'-215') [V-LAMBDA (*Homo sapiens* IGLV1-44*01 (98%) -IGLJ2*01 (100%)) CDR-IMGT [8.3.10] (26-33.51-53.90-99) (1'-109') -*Homo sapiens* IGLC2*01 (100%) (110'-215')]; dimer (232-232":235-235")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

apitégromab

immunoglobuline G4-lambda, anti-[*Homo sapiens* pro-MSTN (pro-facteur de croissance et de différenciation 8, pro-GDF8, pro-myostatine, pro-GDF-8)], anticorps monoclonal *Homo sapiens*;

chaîne lourde gamma4 *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV3-30*03 (100%) -(IGHD) - IGHJ6*01 (94.4%)) CDR-IMGT [8.8.19] (26-33.51-58.97-115) (1-126)-*Homo sapiens* IGHG4*01, G4v5 h P10 (CH1 (127-224), charnière 1-12 S10>P (234) (225-236), CH2 (237-346), CH3 (347-451), CHS K>del (452)) (127-452)], (140-214')-disulfure avec la chaîne légère lambda *Homo sapiens* (1'-215') [V-LAMBDA (*Homo sapiens* IGLV1-44*01 (98%) -IGLJ2*01 (100%)) CDR-IMGT [8.3.10] (26-33.51-53.90-99) (1'-109') -*Homo sapiens* IGLC2*01 (100%) (110'-215')]; dimère (232-232":235-235")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

apitegromab

immunoglobulina G4-lambda, anti-[*Homo sapiens* pro-MSTN (pro-factor de crecimiento y de diferenciación 8, pro-GDF8, pro-miostatina, pro-GDF-8)], anticuerpo monoclonal *Homo sapiens*;

cadena pesada gamma4 *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV3-30*03 (100%) -(IGHD) - IGHJ6*01 (94.4%)) CDR-IMGT [8.8.19] (26-33.51-58.97-115) (1-126)-*Homo sapiens* IGHG4*01, G4v5 h P10 (CH1 (127-224), bisagra 1-12 S10>P (234) (225-236), CH2 (237-346), CH3 (347-451), CHS K>del (452)) (127-452)], (140-214')-disulfuro con la cadena ligera lambda *Homo sapiens* (1'-215') [V-LAMBDA (*Homo sapiens* IGLV1-44*01 (98%) -IGLJ2*01 (100%)) CDR-IMGT [8.3.10] (26-33.51-53.90-99) (1'-109') - *Homo sapiens* IGLC2*01 (100%) (110'-215')]; dímero (232-232":235-235")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
QVQLVESGGG VVQPGKSLRL SCAASGFTFS SYGMHWVRQA PGKGLEWVAV 50
TSYDGSNKYY ADSVKGRFTI SRDNSKNTLY LQMNSLRAED TAVYYCARDL 100
LVRFLEWSHY YGMDVWGQGT TTVTVSSASTK GPSVFLPAP SRSTSESTAA 150
LGCLVKDYFP EPVTVSWNSG ALTSGVHTFP AVLQSSGLYS LSSVTVTPSS 200
SLGKTITYCN VDHKPSNTKY DKRVESKYGP PCPPCAPEF LGGPSVFLFP 250
PKPKDTLMIS RTPEVTCVVV DVSQEDPEVQ FNWYVDGVEV HNAKTKPRE 300
QFNSTYRVVS VLTVLHQDWL NGKEYKCKVS NKGLPSSIEK TISKAKGQPR 350
EPQVYTLPPS QEEMTKNQVS LTCLVKGFYP SDIAVESNES GQPENNYKTT 400
PPVLDSGGSF FLYSRLTVDK SRWQEGNVFS CSVMEALHN HYTKQSLSL 450
LG 452

Light chain / Chaîne légère / Cadena ligera
QSVLTQPPSA SGTTPGQRTVI SCSGSSSNIG SNTVHWYQQL PGTAPKLLIY 50
SDNQRPSSGVP DRFSGSKSGT SASLAISGLQ SEDEADYYCA AWDLSLNGVF 100
GGGTKLTVLG QPKAAPSVTL FPPSSSEELQA NKATLVCLIS DFYPGAVTVA 150
WKADSSPVKA GVETITTPSKQ SNMKYAASSY LSLTPEQWKS HRSYSCQVTH 200
EGSTVEKTV PTECS 215

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 153-209 267-327 373-431
22"-96" 153"-209" 267"-327" 373"-431"
Intra-L (C23-C104) 22"-89" 137"-196"
22"-89" 137"-196"
Inter-H-L (h 10-CL 126) 140-214' 140"-214"
Inter-H-H (h 8, h 11) 232-232" 235-235"

N-terminal glutaminyl cyclization to pyroglutaminyl (pE, 5-oxoprolyl)
H VH Q1:
1, 1"
L VL Q1:
1', 1'"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
303, 303"
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarijos complejos fucosilados.

asundexianum
asundexian

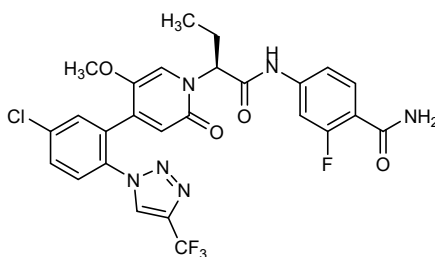
(4S)-2⁴-chloro-4-ethyl-7³-fluoro-3⁵-methoxy-3²,5-dioxo-1⁴-(trifluoromethyl)-3²H-6-aza-3(4,1)-pyridina-1(1)-[1,2,3]triazola-2(1,2),7(1)-dibenzenaheptaphane-7⁴-carboxamide

asundexian

(4S)-2⁴-chloro-4-éthyl-7³-fluoro-3⁵-méthoxy-3²,5-dioxo-1⁴-(trifluorométhyl)-3²H-6-aza-3(4,1)-pyridina-1(1)-[1,2,3]triazola-2(1,2),7(1)-dibenzénaheptaphane-7⁴-carboxamide

asundexián

(4S)-2⁴-cloro-4-etil-7³-fluoro-3⁵-metoxi-3²,5-dioxo-1⁴-(trifluorometil)-3²H-6-aza-3(4,1)-piridina-1(1)-[1,2,3]triazola-2(1,2),7(1)-dibencenaheptafano-7⁴-carboxamida



atidarsagenum autotemcelum #
atidarsagene autotemcel

Autologous CD34+ hematopoietic stem and progenitor cells transduced *ex vivo* with a replication-incompetent lentiviral vector encoding human arylsulfatase A (ARSA). Autologous CD34+ hematopoietic stem and progenitor cells (HSPC), obtained from bone marrow (BM) or mobilised peripheral blood (mPB), transduced *ex vivo* with a replication-incompetent pseudotyped HIV-1 based lentiviral vector encoding the human arylsulfatase A (ARSA) transgene under the control of a human phosphoglycerate kinase (PGK) promoter. The vector is pseudotyped with the vesicular stomatitis virus (VSV) envelope glycoprotein. The vector genome also contains a mutated Woodchuck hepatitis virus post-transcriptional regulatory element (WPRE), a fragment of the HIV-1 *gag*, the HIV-1 Rev response element (RRE), the HIV-1 central polypurine tract (cPPT) and a fragment of *nef*. The CD34+ cell population corresponds to an heterogeneous population of stem and progenitor cells, some expressing the CD90 and/or CD133 markers.

atidarsagène autotemcel

Des cellules souches et progénitrices hématopoïétiques CD34+ autologues ont été transduites *ex vivo* avec un vecteur lentiviral incompetent à la réplication codant pour l'arylsulfatase A humaine (ARSA). Des cellules souches et progénitrices hématopoïétiques CD34+ autologues (HSPC), obtenues à partir de moelle osseuse (BM) ou de sang périphérique mobilisé (mPB), ont été transduites *ex vivo* avec un vecteur lentiviral, à base de VIH-1 pseudotypé et incompetent à la réplication, codant pour le transgène de l'arylsulfatase A humaine (ARSA), sous le contrôle d'un promoteur de la phosphoglycérate kinase (PGK) humaine. Le vecteur est pseudotypé avec la glycoprotéine d'enveloppe du virus de la stomatite vésiculeuse (VSV). Le génome du vecteur contient également l'élément régulateur post-transcriptionnel Woodchuck (WPRE) du virus de l'hépatite, un fragment du *gag* du VIH-1, l'élément de réponse post-transcriptionnel (RRE) du VIH-1 et le tractus central de polypurine du VIH-1 (cPPT), ainsi qu'un fragment de *nef*. La population de cellules CD34+ correspond à une population hétérogène de cellules souches et progénitrices, certaines exprimant les marqueurs CD90 et/ou CD133.

atidarsagén autotemcel

Células troncales y progenitoras hematopoyéticas CD34+ autólogas transducidas *ex vivo* con un vector lentiviral incompetente en replicación que codifica para la arilsulfatasa A humana (ARSA). Las células troncales y progenitoras hematopoyéticas CD34+ autólogas, obtenidas de médula ósea o movilizadas de sangre periférica, se transducen *ex vivo* con un vector lentiviral basado en VIH-1 seudotipado y incompetente en replicación, que codifica el transgen de la arilsulfatasa A humana (ARSA) bajo el control de un promotor de la fosfoglicerato quinasa humana (PGK). El vector está seudotipado con la glicoproteína de la envuelta del virus de la estomatitis vesicular (VSV). El genoma del vector también contiene el elemento regulador post-transcripcional del virus de la hepatitis de la marmota (WPRE), un fragmento de *gag* del VIH-1, un elemento de respuesta post-transcripcional (RRE) del VIH-1 y un tramo central de poli-purinas (cPPT) de VIH-1, así como un fragmento de *nef*. La población de células CD34+ corresponde en una población de células troncales y progenitoras, algunas expressando los marcadores CD90 y/o CD133.

avipendekinum pegolum #
avipendekin pegol

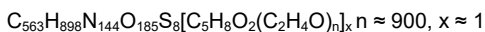
human L-methionyl-interleukin 15, produced in *Escherichia coli*, pegylated at N² of Met¹ and N⁶ of Lys residues with 4-[α-methylpoly(oxyethylene)-ω-oxy]butanoyl groups;
L-methionyl-interleukin 15 (human, non-glycosylated), produced in *Escherichia coli*, substituted on an average of one site among N² of Met¹ and/or N⁶ of lysyl residues with 4-[α-methylpoly(oxyethylene)-ω-oxy]butanoyl groups (~40 kDa each)

avipendékine pégol

L-méthionyl-interleukine 15 humaine, produite par *Escherichia coli*, pégylée en N² de Met¹ et N⁶ des résidus Lys avec des groupes 4-[α-méthylpoly(oxyéthylène)-ω-oxy]butanoyle;
L-méthionyl-interleukine 15 (humaine, non-glycosylée), produite dans *Escherichia coli*, substituée sur une moyenne d'un site parmi le N² du Met¹ et/ou N⁶ des résidus lysyl par des groupes 4-[α-méthylpoly(oxyéthylène)-ω-oxy]butanoyle (~40 kDa chacun)

avipendekina pegol

L-metionil-interleukina 15 humana, producida en *Escherichia coli*, pegilada en N² de Met¹ y N⁶ de residuos Lis con grupos 4-[α-metilpoli(oxietileno)-ω-oxi]butanoilo;
L-metionil-interleukina 15 (humana, no glicosilada), producida en *Escherichia coli*, sustituida pro término medio de un sitio entre el N² del Met¹ y/o el N⁶ de los residuos lisil por grupos 4-[α-metilpoli(oxietileno)-ω-oxi]butanoilo (~40 kDa cada uno)



Sequence / Séquence / Secuencia

MNWVNVISDL **K**KIEDLIQSM HIDATLYTES DVHPSC**K**VTA **M**KCFLLLELQV 50
 ISLESGDASI HDTVENLIIL ANNSLSSNGN VTESG**K**KECE ELEE**K**NIKEF 100
 LQSFVHIVQM FINTS 115

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
 36-86, 43-89

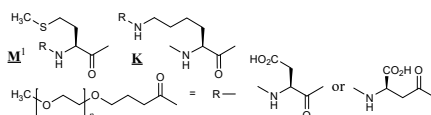
Partial deamidation / Désamidation partielle / Desamidación parcial

N78 (Asn>Asp, isoaspartyl)

Modified residues / Résidus modifiés / Restos modificados

avipendekin : pegol ~ 1:1

M¹ (Met 1: ~32-40 %), **K** (Lys 11, 37, 42, 87: ~33-42 %; Lys 12, 95, 98: ~18-35 %)

**avotaciclimum**

avotaciclib

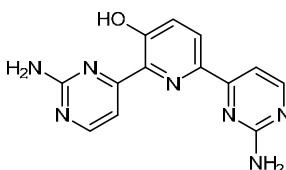
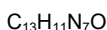
avotaciclib

avotaciclib

2,6-bis(2-aminopyrimidin-4-yl)pyridin-3-ol

2,6-bis(2-aminopyrimidin-4-yl)pyridin-3-ol

2,6-bis(2-aminopirimidin-4-il)piridin-3-ol

**azercabtagenium zapreleucelum #**

azercabtagene zapreleucel

Adult allogeneic CD4⁺ and CD8⁺ enriched T cells transduced with an anti-CD19 chimeric antigen receptor (CAR) that is site-specific integrated into the cellular genome.

Adult allogeneic CD4⁺ and CD8⁺ enriched T cells, gene edited through the electroporation of homing endonuclease mRNA, which is translated into endonuclease protein that introduces a double-stranded DNA break in exon 1 of the endogenous T cell receptor alpha constant (TRAC) locus. Cells are subsequently transduced with a recombinant non-replicating adeno-associated virus type 6 (rAAV6) vector, encoding an anti-CD19 CAR transgene consisting of a murine CD19-specific scFv (clone FMC63), a CD8 hinge/transmembrane domain, a novel 6 (N6) co-stimulatory domain and CD3 zeta signaling domain, under the control of the synthetic promoter JeT and the SV40 poly(A) signal. CAR transgene integration is achieved via homology directed recombination at the endonuclease target site (exon 1 of TRAC). Depletion of CD3⁺ cells is performed on the

	transduced cells to remove cells with functional endogenous T cell receptor (TCR). The substance is a mixture of non-effector (naive, stem cell memory, central memory) (CCR7+) and effector memory and effector (CCR7-) CD4+ and CD8+ T cells with generally $\geq 99\%$ T cell receptor knockout, and $\geq 40\%$ anti-CD19 CAR knock-in. Activation of the cells is accompanied <i>in vitro</i> by release of IFNγ, IL-2, IL-6, and TNFα in an antigen-dependent manner.
azercabtagène zapréleucel	Lymphocytes T adultes allogéniques de type CD4+ et CD8+ transduits avec un récepteur d'antigène chimérique (CAR) anti-CD19 qui est intégré de manière spécifique au site. Lymphocytes T allogéniques enrichies en cellules de type CD4+ et CD8+, dont les gènes ont été édités par l'électroporation du mRNA de l'endonucléase de ciblage, qui est traduite en une protéine endonucléase qui introduit une rupture d'ADN double brin dans l'exon 1 du locus du récepteur alpha endogène aux cellules T (TRAC). Les cellules sont transduites ensuite avec un vecteur du virus recombinant adéno-associé, non réplicatif de type 6 (rAAV6), qui code pour un transgène anti-CD19 CAR, qui consiste en un scFv spécifique à CD19 (clone FMC63), un domaine CD8 transmembranaire/charnière, en un nouveau domaine de co-stimulation 6 (N6) et en un domaine de signalisation CD3 zêta, sous le contrôle du promoteur synthétique JeT et d'un signal poly(A) SV40. L'intégration du transgène CAR est obtenue via la recombinaison homologue dirigée au site de ciblage de l'endonucléase (l'exon 1 du TRAC). La déplétion des cellules CD3+ est effectuée sur les cellules transduites pour éliminer les cellules avec des récepteurs T (TCR) endogènes. La substance est un mélange de lymphocytes T non-effecteurs (naïfs, cellules souches mémoire, mémoire centrale) (CCR7+) et lymphocytes T mémoire effecteurs et effecteurs (CCR7-) CD4+ et CD8+ avec en général $\geq 99\%$ de récepteurs des lymphocytes T knock-out et $\geq 40\%$ de CAR anti-CD19 knock-in. L'activation des cellules est accompagnée <i>in vitro</i> par le relâchement d'IFNγ, IL-2, IL-6, et TNFα d'une manière antigène-dépendante.
azercabtagén zapreleucel	Linfocitos T adultos, alogénicos, enriquecidos en CD4+ y CD8+, transducidos con un receptor de antígenos quimérico (CAR) anti-CD19 que se integra en el genoma celular de forma específica de sitio. Linfocitos T adultos, alogénicos, enriquecidos en CD4+ y CD8+, con genes editados mediante la electroporación del ARNm de una endonucleasa de homing, que se traduce a una proteína endonucleasa que introduce una rotura en el ADN de doble cadena en el exón 1 del locus de la región constante del receptor alfa de células T endógenas (TRAC). Las células son posteriormente transducidas con un vector recombinante, no replicativo, de virus adeno-asociado tipo 6 (rAAV6) que codifica para un transgén de CAR anti-CD19 que consiste en un scFv murino anti-CD19 (clon FMC63), un dominio bisagra/transmembrana

CD8, un nuevo dominio de co-estimulación 6 (N6) y el dominio de señalización CD3 zeta, bajo el control del promotor sintético JeT y la señal poly(A) del SV40. La integración del transgén CAR se consigue mediante recombinación homóloga dirigida en el sitio diana de la endonucleasa (exón 1 de TRAC). Se realiza la depleción de células CD3+ en las células transducidas para eliminar las células con un receptor de células T (TCR) endógeno funcional. La sustancia es una mezcla de linfocitos T no efectores (ingenuos, células troncales memoria, memoria central) (CCR7+) y linfocitos T memoria efectores y efectores (CCR7-) CD4+ y CD8+ con en general ≥99% de receptores de linfocitos T (TCR) knock-out y ≥40% de CAR anti-CD19 knock-in. La activación de las células está acompañada *in vitro* por la liberación de IFN γ , IL-2, IL-6, y TNF α de una manera antígeno-dependiente.

bapotulimabum #
bapotulimab

immunoglobulin G2-kappa, anti-[*Homo sapiens* ILDR2 (immunoglobulin like domain containing receptor 2, C1orf32)], *Homo sapiens* monoclonal antibody; gamma2 heavy chain *Homo sapiens* (1-445) [VH (*Homo sapiens* IGHV1-69*10 (100%) -(IGHD) - IGHJ5*01 (81.2%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG2*01, G2m.. CH2 V45.1 (CH1 (121-218), hinge 1-12 (219-230), CH2 V45.1 (281) (231-339), CH3 (340-444), CHS K>del (445)) (121-445)], (134-219')-disulfide with kappa light chain *Homo sapiens* (1'-219') [V-KAPPA (*Homo sapiens* IGKV2-28*01 (99%) -IGKJ4*01 (83.3%)) CDR-IMGT [11.3.9] (27-37.55-57.94-102) (1'-112') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (158), V101 (196) (113'-219')]; dimer (222-222":223-223":226-226":229-229")-tetrakisdisulfide, produced in a Chinese hamster ovary (CHO)-derived cell line, glycoform alfa

bapotulimab

immunoglobuline G2-kappa, anti-[*Homo sapiens* ILDR2 (récepteur 2 contenant un domaine immunoglobuline-like, C1orf32)], anticorps monoclonal *Homo sapiens*; chaîne lourde gamma2 *Homo sapiens* (1-445) [VH (*Homo sapiens* IGHV1-69*10 (100%) -(IGHD) - IGHJ5*01 (81.2%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG2*01, G2m.. CH2 V45.1 (CH1 (121-218), charnière 1-12 (219-230), CH2 V45.1 (281) (231-339), CH3 (340-444), CHS K>del (445)) (121-445)], (134-219')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-219') [V-KAPPA (*Homo sapiens* IGKV2-28*01 (99%) -IGKJ4*01 (83.3%)) CDR-IMGT [11.3.9] (27-37.55-57.94-102) (1'-112') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (158), V101 (196) (113'-219')]; dimère (222-222":223-223":226-226":229-229")-tétrakisdisulfure, produite dans une lignée cellulaire dérivée des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

bapotulimab

inmunoglobulina G2-kappa, anti-[*Homo sapiens* ILDR2 (receptor 2 que contiene un dominio inmunoglobulina-like, C1orf32)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma2 *Homo sapiens* (1-445) [VH (*Homo sapiens* IGHV1-69*10 (100%) -(IGHD) -IGHJ5*01 (81.2%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG2*01, G2m.. CH2 V45.1 (CH1 (121-218), bisagra 1-12 (219-230), CH2 V45.1 (281) (231-339), CH3 (340-444), CHS K>del (445)) (121-445)], (134-219')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-219') [V-KAPPA (*Homo sapiens* IGKV2-28*01 (99%) -IGKJ4*01 (83.3%)) CDR-IMGT [11.3.9] (27-37.55-57.94-102) (1'-112') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (158), V101 (196) (113'-219')]; dímero (222-222":223-223":226-226":229-229")-tetrakisdisulfuro, producido en una línea celular derivada de las células ováricas de hamster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada	
QVQLVQSGAE VKKPGSSVKV SCKASGGTFS SYAISWVRQA PGQGLEWMGG	50
IIPILGIANY AQKFQGRVTI TADKSTSTAY MELSSLRSED TAVYYCARGR	100
LPYGFDFWSW GQGTLLTVVSS ASTKGPSVFP LAPCSRSTSE STAALGCLVK	150
DYFPEPVTVS WNSGALTSGV HTFPAVLQSS GLYSLSSVVT VPSSNFGTQT	200
YTCNVDHKPS NTKVDKTVR KCCVECPPCP APPVAGPSVF LFPPKPKDTL	250
MISRTPEVTC VVVDVSHEDP EVQFNWYVDG VEVHNAKTPR REEQNFSTFR	300
VVSULTVVHQ DWLNGKEYKC KVSNGKLPAP IEKTIKTKG QPREPQVYTL	350
PPSREEMTKN QVSLTCLVKG FYPSDIAVEW ESNQGPENNY KTTTPMLDSD	400
GSFFLYSKLT VDKSRWQQGN VFSCSVMHFA LHNHYTQKSL SLSPG	445
Light chain / Chaîne légère / Cadena ligera	
DIVMTQSPLS LPVTPGEPAS ISCRSSQSL L YSNGNYLWD YLQKPGQSPQ	50
LLIYLGSNRA SGVPRFSGS GSGTDFTLKI SRVEAEDVG VYCMQALQTP	100
LTFGGGKLE IRRTVAAPSV FIFPPSDEQL KSGTASVVL LNNFYPREAK	150
VQWQVDNALQ SGNSQESVTE QDSKSTYSL SSTLTLSKAD YEKHKVYACE	200
VTHQGLSSPV TKSFNRGEC	219
Post-translational modifications	
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro	
Intra-H (C23-C104)	22°-96° 147°-203° 260°-320° 366°-424°
	22°-96° 147°-203° 260°-320° 366°-424°
Intra-L (C23-C104)	23°-93° 139°-199°
	23°-93° 139°-199°
Inter-H-L (CH1 10-CL 126)*	134°-219° 134°-219°
Inter-H-H (h 4, h 5, h 8, h 11)*	222°-222° 223°-223° 226°-226° 229°-229°
*A-isoform, predominant	
N-terminal glutaminyl cyclization to pyroglutaminyl (pE, 5-oxoprolyl)	
H VH Q1:	
1, 1°	
N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación	
H CH2 N84.4:	
296, 296°	
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantennarios complejos fucosilados.	

barecetamabum #
barecetamab

immunoglobulin G1-lambda, anti-[*Homo sapiens* ERBB3 (receptor tyrosine-protein kinase erbB-3, HER3)], *Homo sapiens* monoclonal antibody; gamma1 heavy chain *Homo sapiens* (1-451) [VH (*Homo sapiens* IGHV3-23*01 (90.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -*Homo sapiens* IGHG1*01, nG1m1, G1m17 (CH1 K120 (218) (122-219), hinge 1-15 (220-234), CH2 (235-344), CH3 E12 (360), M14 (362) (345-449), CHS (450-451)) (122-451)], (224-215')-disulfide with lambda light chain *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-47*02 (92.9%) -IGLJ2*01 (100%)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC3*03 (100%) (111'-216')]; dimer (230-230":233-233")-bisdisulfide, produced in Chinese hamster ovary (CHO)-K1 cell line, glycoform alfa

barécétamab

immunoglobuline G1-lambda, anti-[*Homo sapiens* ERBB3 (récepteur à activité tyrosine kinase erbB-3, HER3)], *Homo sapiens* anticorps monoclonal;

chaîne lourde gamma1 *Homo sapiens* (1-451) [VH (*Homo sapiens* IGHV3-23*01 (90.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -*Homo sapiens* IGHG1*01, nG1m1, G1m17 (CH1 K120 (218) (122-219), charnière 1-15 (220-234), CH2 (235-344), CH3 E12 (360), M14 (362) (345-449), CHS (450-451)) (122-451)], (225-215')-disulfure avec la chaîne légère lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-47*02 (92.9%) -IGLJ2*01 (100%)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC3*03 (100%) (111'-216')]; dimère (230-230":233-233")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-K1, glycoforme alfa

barecetamab

inmunoglobulina G1-lambda, anti-[*Homo sapiens* ERBB3 (receptor de la actividad tirosina kinasa erbB-3, HER3)], *Homo sapiens* anticuerpo monoclonal;

cadena pesada gamma1 *Homo sapiens* (1-451) [VH (*Homo sapiens* IGHV3-23*01 (90.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -*Homo sapiens* IGHG1*01, nG1m1, G1m17 (CH1 K120 (218) (122-219), bisagra 1-15 (220-234), CH2 (235-344), CH3 E12 (360), M14 (362) (345-449), CHS (450-451)) (122-451)], (225-215')-disulfuro con la cadena ligera lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-47*02 (92.9%) -IGLJ2*01 (100%)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC3*03 (100%) (111'-216')]; dímero (230-230":233-233")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-K1, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
 EVQLLESGGG LVQPGGSLRL SCAASGFTFS DYDMSWVRQA PGKGLEWVST 50
 IDLDSGSIYY ADSVQGRFTI SRDNSKNTLY LQMSLRRAED TAVYYCAKDL 100
 HMGPEGPFDY WQGQGLTVTS SASTKGPSVF PLAPSKSTS GGTAALGCLV 150
 KDYFPEPVTV SWNSGALTSV VHTFPAVLQS SGLYSLSVV TVPSSSLGTQ 200
 TYICNVNHKK SNTKVDKVE PKSCDKHTC PPCPAPELLG GPSVFLFPK 250
 PKDTLMISRT PEVTCVVVDV SHEDPEVKFN WYDVGVEHN AKTKPREEQY 300
 NSTYRVVSVL TVLHQDWLNG KEYKCKVSNK ALPAPIEKT SKAKGQPREP 350
 QYVTLPPSRE EMTKNQVSLT CLVKGFYPSP IAVVESNGQ PENNYKTPFP 400
 VLDSGGSFFL YSKLTVDKSR WQQGNVFSQS VMHEALHNHY TQKSLSLSPG 450
 K 451

Light chain / Chaîne légère / Cadena ligera
 QSVLTQPPSA SGTPGQRVTI SCSSGSSNIG SNSVSWYQQL PGTAPKLLIY 50
 SDNHRPSGVP DRFSGSKSGT SASLAISGLR SEDEADYYCQ GMDTSLSGHV 100
 FGGGTKLTVL GQPKAAPSVT LFPPSSEELQ ANKATLVCLI SDFYPGAVTV 150
 AWKADSSPVK AGVETTTPSK QSNKNYAASS YLSLTPEQWK SHKSYSCQVT 200
 HEGSTVEKTV APTECS 216

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 148-204 265-325 371-429
 22"-96" 148"-204" 265"-325" 371"-429"

Intra-L (C23-C104) 22"-89" 138"-197"
 22"-89" 138"-197"

Inter-H-L (h 5-CL 126) 224-215' 224"-215"
 Inter-H-H (h 11, h 14) 230-230" 233-233"

N-terminal glutaminy cyclization to pyroglutamyl (pE, 5-oxoprolyl)

L VL Q1:

I', I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

301, 301"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

451, 451"

batiraxceptum #
batiraxcept

human tyrosine-protein kinase receptor UFO (AXL oncogene) fragment (8-202, 1-195 in the current sequence), variant (G¹⁴>S⁷, A⁵⁴>V⁴⁷, D⁶⁹>G⁶², V⁷⁴>A⁶⁷, G¹⁰⁹>R¹⁰²), fused via peptidyl linker (¹⁹⁶GGGGS²⁰⁰) to a human immunoglobulin G1 Fc fragment (201-426, C-terminal Lys residue deleted), dimer, glycosylated, produced in Chinese hamster ovary (CHO) cells; [G¹⁴>S(7), A⁵⁴>V(47), D⁶⁹>G(62), V⁷⁴>A(67), G¹⁰⁹>R(102)] human AXL receptor tyrosine kinase (AXL oncogene, tyrosine-protein kinase receptor UFO) (8-202)-peptide (1-195), fused via a G₄S linker (196-200) to a human immunoglobulin G1 C-terminal K>del Fc fragment (201-426), dimer (206-206':209-209')-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

batiraxcept

récepteur de la tyrosine kinase-protéine UFO (oncogène AXL), fragment (8-202, 1-195 dans la séquence présente), variant (G¹⁴>S⁷, A⁵⁴>V⁴⁷, D⁶⁹>G⁶², V⁷⁴>A⁶⁷, G¹⁰⁹>R¹⁰²), fusionné via un peptide liant (¹⁹⁶GGGGS²⁰⁰) à un fragment Fc de l'immunoglobuline humaine G1 (201-426, résidu Lys C-terminal supprimé), dimère, glycosylé, produit par des cellules ovariennes de hamsters chinois (CHO); [G¹⁴>S(7), A⁵⁴>V(47), D⁶⁹>G(62), V⁷⁴>A(67), G¹⁰⁹>R(102)] tyrosine kinase du récepteur AXL humaine (oncogène AXL, récepteur de la tyrosine kinase-protéine UFO) (8-202)-peptide (1-195), fusionné via un peptide liant G₄S (196-200) au fragment Fc C-terminal K>del de l'immunoglobuline G1 humaine (201-426), dimère (206-206':209-209')-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

batiraxcept

receptor de la proteína tirosina kinasa humana UFO (oncogén AXL) fragmento (8-202, 1-195 en la secuencia actual), variante (G¹⁴>S⁷, A⁵⁴>V⁴⁷, D⁶⁹>G⁶², V⁷⁴>A⁶⁷, G¹⁰⁹>R¹⁰²), fusionado a través de un conector peptidil (¹⁹⁶GGGGS²⁰⁰) a una inmunoglobulina humana G1 Fc fragmento (201-426, residuo Lis C-terminal eliminado), dímero, glicosilado, producido en células ováricas de hámster Chino (CHO); [G¹⁴>S(7), A⁵⁴>V(47), D⁶⁹>G(62), V⁷⁴>A(67), G¹⁰⁹>R(102)] AXL receptor tirosina kinasa humana (oncogén AXL, receptor de la proteína tirosina-kinasa UFO) (8-202)-péptido (1-195), fusionado a través de un péptido G₄S (196-200) con un fragmento Fc C-terminal K>del de la inmunoglobulina G1 humana (201-426), dímero (206-206':209-209')-bisdisulfuro, producido en células ováricas de hámster Chino (CHO), glicosilada alfa

Sequence / Séquence / Secuencia
EESPEVSNFG NITGARGLTG TLRCQLQVQG EPPEVHWLRD GQILELVDS 50
QTQVFLGEDE QGDWIVASQL RITSLLQSDT GQYQCLVFLG HOTFVSQPGY 100
VPLEGLPYFL EEPEDRTVA NTFNLSCQA QGPEPEVDLL WLQDAVPLAT 150
APGHGPORSL HVPGLNKTSS FSCEAHNAKG VTTSRTAITI VLPQCGGGGS 200
DKTHTCFPCF APELLGGPSV FLFPKPKCT LMSRTPEVT CVVVDVSHED 250
PEVKENWYVD GVEVHNAKTK PREEQYNSTY RVVSVLTVLH QDWLNGKEYK 300
KRVSNKALPA PIEKTISKAK GQPREPQVYT LPFSREEMTK NQVSLTCLVK 350
GFYPSDIAVE WESNGQPENN YKTTFPVLD DGSFFLYSKL TVDKSRWQQG 400
NVFESCVWHE ALHNHYTQKS LSLSPG 426

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
intra-AXL 24-85 128-173 intra-Fc 241-301 347-405
24'-85' 128'-173' 241'-301' 347'-405'
inter-Fc 206-206' 209-209'

Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación
AXL: N11, N125, N166, N11', N125', N166';
Fc: N277, N277'

befotertinibum

befotertinib

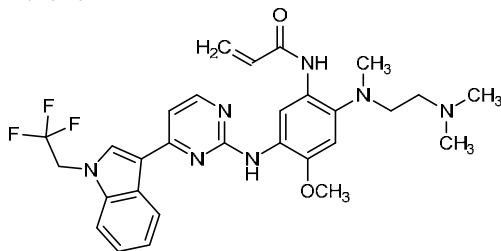
N-[2-[[2-(dimethylamino)ethyl](methyl)amino]-4-methoxy-5-({4-[1-(2,2,2-trifluoroethyl)-1*H*-indol-3-yl]pyrimidin-2-yl}amino)phenyl]prop-2-enamide

béfotertinib

N-[2-[[2-(diméthylamino)éthyl](méthyl)amino]-4-méthoxy-5-({4-[1-(2,2,2-trifluoroéthyl)-1*H*-indol-3-yl]pyrimidin-2-yl}amino)phényl]prop-2-énamide

befotertinib

N-[2-[[2-(dimetilamino)etil](metil)amino]-4-metoksi-5-({4-[1-(2,2,2-trifluoroetil)-1*H*-indol-3-il]pirimidin-2-il}amino)fenil]prop-2-enamida

C₂₉H₃₂F₃N₇O₂**belumosudilum**

belumosudil

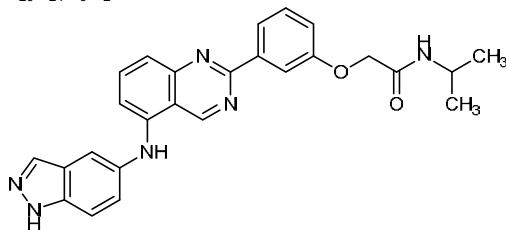
2-(3-{4-[(1*H*-indazol-5-yl)amino]quinazolin-2-yl}phenoxy)-*N*-(propan-2-yl)acetamide

bélumosudil

2-(3-{4-[(1*H*-indazol-5-yl)amino]quinazolin-2-yl}phénoxy)-*N*-(propan-2-yl)acetamide

belumosudil

2-(3-{4-[(1*H*-indazol-5-il)amino]quinazolin-2-il}fenoksi)-*N*-(propan-2-il)acetamida

C₂₆H₂₄N₆O₂**bentracimabum #**

bentracimab

immunoglobulin Fab G1-lambda, anti-[*ticagrelor* and its active metabolite AR-C124910XX], *Homo sapiens* monoclonal antibody; VH-(CH1-hinge) gamma1 heavy chain *Homo sapiens* (1-234) [VH (*Homo sapiens* IGHV1-69*06 (85.7%) -(IGHD) -IGHJ1*01 (100%)) CDR-IMGT [8.7.14] (26-33.51-58.97-118) (1-129) -*Homo sapiens* IGHG1*03 (100%) G1m3 (CH1 R120 (226) (130-227), hinge 1-7 (228-234)) (130-234)], (232-215')-disulfide with lambda light chain *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-51*01 (94.6%) -IGLJ2*01 (81.8%)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC2*01(100%) (111'-216')], non-glycosylated, produced in the bacteria *Escherichia coli* (*E. coli*)

bentracimab

immunoglobuline Fab G1-lambda, anti-[*ticagrelor* et son métabolite actif AR-C124910XX], anticorps monoclonal *Homo sapiens*;
VH-(CH1-charnière) chaîne lourde gamma1 *Homo sapiens* (1-234) [VH (*Homo sapiens* (1-234) [VH (*Homo sapiens* IGHV1-69*06 (85.7%) -(IGHD) -IGHJ1*01 (100%)) CDR-IMGT [8.7.14] (26-33.51-58.97-118) (1-129) -*Homo sapiens* IGHG1*03 (100%) G1m3 (CH1 R120 (226) (130-227), charnière 1-7 (228-234)) (130-234)], (232-215')-disulfure avec la chaîne légère lambda *Homo sapiens* (1'-216')] [V-LAMBDA (*Homo sapiens* IGLV1-51*01 (94.6%) -IGLJ2*01 (81.8%)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC2*01(100%) (111'-216')], non-glycosylé, produit dans la bactérie *Escherichia coli* (*E. coli*)

bentracimab

inmunoglobulina Fab G1-lambda, anti-[*ticagrelor* y su metabolito activo AR-C124910XX], anticuerpo monoclonal *Homo sapiens*;
VH-(CH1-bisagra) cadena pesada gamma1 *Homo sapiens* (1-234) [VH (*Homo sapiens* (1-234) [VH (*Homo sapiens* IGHV1-69*06 (85.7%) -(IGHD) -IGHJ1*01 (100%)) CDR-IMGT [8.7.14] (26-33.51-58.97-118) (1-129) -*Homo sapiens* IGHG1*03 (100%) G1m3 (CH1 R120 (226) (130-227), bisagra 1-7 (228-234)) (130-234)], (232-215')-disulfuro con la cadena ligera lambda *Homo sapiens* (1'-216')] [V-LAMBDA (*Homo sapiens* IGLV1-51*01 (94.6%) -IGLJ2*01 (81.8%)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC2*01(100%) (111'-216')], no glicosilado, producido en la bacteria *Escherichia coli* (*E. coli*)

Heavy chain / Chaîne lourde / Cadena pesada
QVQLQESGAE VVKPGSSVRV SCKASGCTFD SYSIHWVRQA PCQGLEMMGG 5C
ILPAFTLSS AQDFQAEVTI SADKSTSTAY MELSGLRSED TAVYYCARGS 100
FDYFWSASH PPNDAIAIW QGTLVTVSSA STQKSPVFPL APSSKSTSEG 150
TAALGCLVKD YFPEPTVSW NSGALTSGVH TFPAYLQSSG LYSLSSTVTV 200
FSSSLGTQTY ICNVNHRKPSN TKVDRKVERK SCDK 234

Light chain / Chaîne légère / Cadena ligera
QSVVTQFPVS SAAPGQKVTI SCSGNSDIG NNYVSMYQQL PCTAPKLLIY 5C
DNNKRPSPGIP DRESGSKST SATLAITGLQ AGDEADYVCG TWLYDRAVGL 100
FGGGRKVTYL GQPKAAFSVT LFPPSSBELQ ANKATLVCLI SDFYFGAVTV 150
AWKADSSPVK AGVETTTPSK QSNNKYAASS YLSLTPEQWK SHRSYSQVTV 200
HEGSTVERTV APTECS 216

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 156-212
Intra-L (C23-C104) 22'-89' 138'-197'
Inter-H-L (h 5-CL 126) 232-215'

N-terminal glutaminylation / cyclization to pyroglutamy (pE, 5-oxoprolyl)
L VH Q1:
I
L VL Q1:
I'

No N-glycosylation site / pas de sites de N-glycosylation / ningun posición de N-glicosilación
Agycosylated

bepirovirsenum
bepirovirsen

all-P-ambo-2'-O-(2-methoxyethyl)-P-thioguanlyl-(3'→5')-2'-O-(2-methoxyethyl)-5-methyl-P-thiocytidylyl-(3'→5')-2'-O-(2-methoxyethyl)-P-thioadenylyl-(3'→5')-2'-O-(2-methoxyethyl)-P-thioguanlyl-(3'→5')-2'-O-(2-methoxyethyl)-P-thioadenylyl-(3'→5')-2'-deoxy-P-thioguanlyl-(3'→5')-2'-deoxy-P-thioguanlyl-(3'→5')-P-

thiothymidyl-(3'→5')-2'-deoxy-*P*-thioguanyl-(3'→5')-2'-deoxy-*P*-thioadenyl-(3'→5')-2'-deoxy-*P*-thioadenyl-(3'→5')-2'-deoxy-*P*-thioguanyl-(3'→5')-2'-deoxy-5-methyl-*P*-thiocytidyl-(3'→5')-2'-deoxy-*P*-thioguanyl-(3'→5')-2'-deoxy-*P*-thioadenyl-(3'→5')-2'-*O*-(2-methoxyethyl)-*P*-thioadenyl-(3'→5')-2'-*O*-(2-methoxyethyl)-*P*-thioguanyl-(3'→5')-2'-*O*-(2-methoxyethyl)-5-methyl-*P*-thiouridyl-(3'→5')-2'-*O*-(2-methoxyethyl)-*P*-thioguanyl-(3'→5')-2'-*O*-(2-methoxyethyl)-5-methylcytidine

bépirovirsen

tout-P-ambo-2'-O-(2-méthoxyéthyl)-P-thioguanlyl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthyl-P-thiocytidyl-(3'→5')-2'-O-(2-méthoxyéthyl)-P-thioadényl-(3'→5')-2'-O-(2-méthoxyéthyl)-P-thioguanlyl-(3'→5')-2'-O-(2-méthoxyéthyl)-P-thioadényl-(3'→5')-2'-désoxy-P-thioguanlyl-(3'→5')-2'-désoxy-P-thioguanlyl-(3'→5')-P-thiothymidyl-(3'→5')-2'-désoxy-P-thioguanlyl-(3'→5')-2'-désoxy-P-thioadényl-(3'→5')-2'-désoxy-P-thioadényl-(3'→5')-2'-désoxy-P-thioguanlyl-(3'→5')-2'-désoxy-5-méthyl-P-thiocytidyl-(3'→5')-2'-désoxy-P-thioguanlyl-(3'→5')-2'-désoxy-P-thioadényl-(3'→5')-2'-O-(2-méthoxyéthyl)-P-thioadényl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthyl-P-thiouridyl-(3'→5')-2'-O-(2-méthoxyéthyl)-P-thioguanlyl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthylcytidine

bepirovirsén

todo-P-ambo-2'-O-(2-metoxietil)-P-tioguanilil-(3'→5')-5-metil-2'-O-(2-metoxietil)-P-tiocitidilil-(3'→5')-2'-O-(2-metoxietil)-P-tioadenilil-(3'→5')-2'-O-(2-metoxietil)-P-tioguanilil-(3'→5')-2'-O-(2-metoxietil)-P-tioadenilil-(3'→5')-2'-desoxi-P-tioguanilil-(3'→5')-2'-desoxi-P-tioguanilil-(3'→5')-P-tiotimidilil-(3'→5')-2'-desoxi-P-tioguanilil-(3'→5')-2'-desoxi-P-tioadenilil-(3'→5')-2'-desoxi-P-tioadenilil-(3'→5')-2'-desoxi-P-tioguanilil-(3'→5')-2'-desoxi-5-metil-P-tiocitidilil-(3'→5')-2'-desoxi-P-tioguanilil-(3'→5')-2'-desoxi-P-tioadenilil-(3'→5')-2'-O-(2-metoxietil)-P-tioadenilil-(3'→5')-2'-O-(2-metoxietil)-P-tioguanilil-(3'→5')-5-metil-2'-O-(2-metoxietil)-P-tiouridilil-(3'→5')-2'-O-(2-metoxietil)-P-tioguanilil-(3'→5')-2'-O-(2-metoxietil)-5-metilcitidina

$$\text{C}_{230}\text{H}_{309}\text{N}_{88}\text{O}_{115}\text{P}_{19}\text{S}_{19}$$

(3'-5') G=C=A=G=A=d[G=G=T=G=A=A=G=C=G=A=]A=G=U=G=C

Legend d : 2'-deoxynucleotides

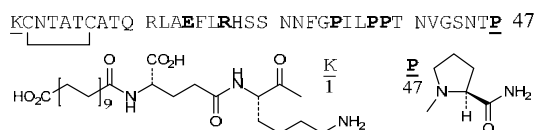
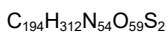
X: 2'-*O*-(2-methoxyethyl)nucleotide

C, C, U : 5-methylnucleotide

beremagenum geperpavecum #
beremagene geperpavec

A replication-defective herpes simplex virus encoding human collagen type VII alpha 1 chain (COL7A1).

	A recombinant replication-defective, non-integrating herpes simplex virus-1 (HSV-1) (KOS strain) encoding two copies of human collagen type VII alpha 1 chain (COL7A1), inserted into the immediate early (IE) infected cell protein 4 (ICP4) gene regions after deletion of both copies of ICP4 genes, under the control of the human cytomegalovirus (hCMV) immediate early promoter and bovine growth hormone polyadenylation signal (bGHpA). The infected cell protein 22 (ICP22) gene is also deleted.
béremagène géperpavec	Un virus herpes simplex à réplication défectueuse codant pour la chaîne du collagène humain de type VII alpha 1 (COL7A1). Un virus recombinant de l'herpès simplex-1 (HSV-1) (souche KOS), non intégrateur et à défaut de réplication, codant pour deux copies de la chaîne du collagène humain de type VII alpha 1 (COL7A1), insérées dans les régions du gène de la protéine d'infection cellulaire précoce 4 (ICP4) après délétion des deux copies des gènes ICP4, sous le contrôle du promoteur précoce immédiat du cytomégalovirus humain (hCMV) et du signal de polyadénylation de l'hormone de croissance bovine (bGHpA). Le gène de la protéine d'infection cellulaire précoce 22 (ICP22) est également supprimé.
beremagén geperpavec	Un virus herpes simplex deficiente para replicación que codifica para la cadena de colágeno humano de tipo VII alfa 1 (COL7A1). Un virus herpes simplex -1 (HSV-1) (cepa KOS) recombinante, no integrativo y deficiente en replicación, que codifica dos copias de la cadena de colágeno humano de tipo VII alfa 1 (COL7A1), insertadas en las regiones del gen de la proteína de infección celular inmediatamente temprana 4 (ICP4) tras la delección de las dos copias de genes ICP4, bajo el control del promotor inmediatamente temprano del citomegalovirus humano (hCMV) y la señal de poliadenilación de la hormona de crecimiento bovina (bGHpA). El gen de la proteína de infección celular 22 (ICP22) está también delecionado.
cagrilintidum cagrilintide	$N^{2,1}$-[N-(19-carboxynonadecanoyl)-L-γ-glutamyl]-[N¹⁴>E, V¹⁷>R, A²⁵>P, S²⁸>P, S²⁹>P, Y³⁷>P]-human amylin (islet amyloid polypeptide, IAPP, diabetes-associated peptide, DAP, insulinoma amyloid peptide, <i>INN</i>: <i>amlintide</i>), (35-40)-disulfide
cagrilintide	$N^{2,1}$-[N-(19-carboxynonadécanoyl)-L-γ-glutamyl]-[N¹⁴>E, V¹⁷>R, A²⁵>P, S²⁸>P, S²⁹>P, Y³⁷>P]-amyline humaine (polypeptide amyloïde des îlots, IAPP, peptide associé au diabète, DAP, polypeptide amyloïde de l'insulinome, <i>DCI</i>: <i>amlintide</i>), (35-40)-disulfure
cagrilintida	$N^{2,1}$-[N-(19-carboxinonadecanoil)-L-γ-glutamyl]-[N¹⁴>E, V¹⁷>R, A²⁵>P, S²⁸>P, S²⁹>P, Y³⁷>P]-amilina humana (polipéptido amiloide de los islotes, IAPP, péptido asociado a la diabetes, DAP, polipéptido amiloide insulinoma, <i>DCI</i>: <i>amlintida</i>), (35-40)-disulfuro

**cantharidinum**

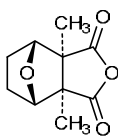
cantharidin

(3a*R*,4*S*,7*R*,7a*S*)-3a,7a-dimethylhexahydro-4,7-epoxy-2-benzofuran-1,3-dione

cantharidine

(3a*R*,4*S*,7*R*,7a*S*)-3a,7a-diméthylhexahydro-4,7-époxy-2-benzofurane-1,3-dione

cantaridina

(3a*R*,4*S*,7*R*,7a*S*)-3a,7a-dimetilhexahidro-4,7-epoxi-2-benzofurano-1,3-diona**carfloglitazarum**

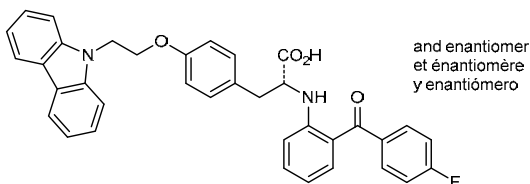
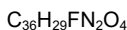
carfloglitazar

O-[2-(9*H*-carbazol-9-yl)ethyl]-*N*-[2-(4-fluorobenzoyl)phenyl]-DL-tyrosine

carfloglitazar

O-[2-(9*H*-carbazol-9-yl)éthyl]-*N*-[2-(4-fluorobenzoyl)phényl]-DL-tyrosine

carfloglitazar

O-[2-(9*H*-carbazol-9-il)etil]-*N*-[2-(4-fluorobenzoi)fenil]-DL-tirosina**cedirogantum**

cedirogant

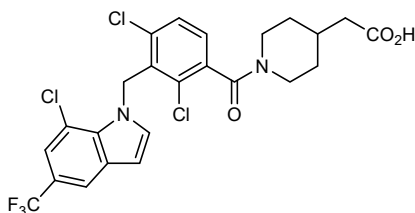
[1-(2,4-dichloro-3-[[7-chloro-5-(trifluoromethyl)-1*H*-indol-1-yl]methyl]benzoyl)piperidin-4-yl]acetic acid

cédirogant

acide [1-(2,4-dichloro-3-[[7-chloro-5-(trifluorométhyl)-1*H*-indol-1-yl]méthyl]benzoyl)pipéridin-4-yl]acétique

cedirogant

ácido [1-(2,4-dicloro-3-[[7-cloro-5-(trifluorometil)-1*H*-indol-1-il]metil]benzoi)piperidin-4-il]acético



cevaretigenum ritoparvovecum #
cevaretigene ritoparvovec

A non-replicating adeno-associated viral vector serotype 5 vector encoding human retinoid isomerohydrolase (RPE65).

A recombinant non-replicating adeno-associated viral vector serotype 5 (rAAV5), encoding codon-optimised human retinoid isomerohydrolase (retinal pigment epithelium-specific 65 kDa protein [RPE65]), under the control of the human RPE65 promoter and an SV40 poly(A) signal, flanked by AAV2 inverted terminal repeats (ITR).

cévarétigène ritoparvovec

Un vecteur adénoviral à réplication défectueuse de sérotype 5 et codant pour l'isomérohydrolase rétinienne humaine (RPE65).

Un vecteur adénoviral recombinant à réplication défectueuse de sérotype 5 (rAAV5), codant pour une isomérohydrolase rétinienne humaine, avec des codons optimisés (protéine de 65 kDa spécifique de l'épithélium pigmentaire rétinien [RPE65]), sous le contrôle du promoteur RPE65 humain et d'un signal SV40 poly(A), flanqué de répétitions terminales inversées (ITR) AAV2.

cevaretigén ritoparvovec

Un vector de virus adeno-asociado serotipo 5, no replicativo, que codifica para la isomerohidrolasa retinoide humana (RPE65).

Un vector de virus adeno-asociado recombinante serotipo 5, (rAAV5) no replicativo, que codifica para la isomerohidrolasa retinoide humana (proteína de 65 kDa específica del epitelio pigmentario retiniano [RPE65]) con los codones optimizados, bajo el control del promotor de RPE65 humano y una señal poly(A) de SV40, flanqueado por las repeticiones terminales invertidas (ITR) del AAV2.

cipaglucosidasum alfa #
cipaglucosidase alfa

human lysosomal α -glucosidase (acid alpha-glucosidase, acid maltase) fragment (57-952, 57-69 from the propeptide sequence), (1-896, 1-13 in the current sequence), produced in Chinese hamster ovary (CHO) cells, glycosylated;
human lysosomal α -glucosidase (acid alpha-glucosidase, GAA, acid maltase; EC 3.2.1.20) pre-protein (57-952)-peptide (with 13 residues (57-69) of the propeptide (28-69)), produced in Chinese hamster ovary (CHO) cells, glycoform alfa

cipaglucosidase alfa	α -glucosidase lysosomale humaine (alpha-glucosidase acide, maltase acide) fragment (57-952, 57-69 de la séquence du propeptide), (1-896, 1-13 dans la séquence actuelle), produit dans des cellules ovariennes de hamster chinois (CHO), glycosylée; α -glucosidase lysosomale humaine (alpha-glucosidase acide, GAA, maltase acide; EC 3.2.1.20) pré-pro-protéine (57-952)-peptide (avec 13 résidus (57-69) du propeptide (28-69)), produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa
cipaglucosidasa alfa	α -glucosidasa humana lisosomal (ácido alfa-glucosidasa, ácido maltasa) fragmento (57-952, 57-69 de la secuencia propérido), (1-896, 1-13 en la secuencia actual), producido en células ováricas de hámster Chino (CHO), glisosiladas; α -glucosidasa lisosomal humana (alfa glucosidasa ácida, GAA, maltasa ácida; EC 3.2.1.20) pre-pro-proteína (57-952)-péptido, (con 13 residuos (57-69) del propéptido (28-69)), producido en las células ováricas de hámster Chino (CHO), glicosilada alfa

Sequence / Séquence / Secuencia		
QGGASRPGPR	DAQAHPGRPR	AVPTQCDVPP
NSRFDCA	PDK	AITQEQCEAR
50		
GCCYIPAKQG	LQGAQMGQPW	CFPPSPYPSY
KLENLSSEM	GYTATLTRTT	100
PTFFPKDILT	LRLDVMETE	NRLHFTIKDP
ANRRYEVPLE	TPHVSRAPS	150
PLYSVEFSEE	PFGVIVRRQL	DGRVLLNTTV
APLFFADQFL	QLSTSLPSQY	200
ITGLAEHLSP	LMLSTSWTRI	TLWNRDLAPT
PGANLYGSH	FYLALEDGGS	250
AHGVFLNSN	AMDVVLQPS	ALSWRSTGGI
LDVYIFLGPE	PKSVVQQYLD	300
VVGYPFMPY	WGLGFHLRW	GYSSTAITRQ
VVENMTRAHF	PLDVQWDL	350
YMDSRDFTF	NKDGFRDFA	MVQELHQGGR
RYMMIVDPAI	SSSGPAGSYR	400
PYDEGLRRGV	FITNETGQPL	IGKVWPGSTA
FPDFTNP	TAL	AWEDMVAEF
450		
HDQVPFDGMW	IDMNEPSNFI	RGSEDCPN
ELENPPYVPG	VVGGTLQAAT	500
ICASSHQFLS	THYNLHNL	YG
LTEAIA	SHRA	LVKARGTRPF
VISRSTFAGH	550	
GRYAGHWTGD	VWSSWEQLAS	SVPEILQFNL
LGVPLVGADV	CGFLGNTSEE	600
LCYRWTLGA	FYPFMRNHNS	LLSLPQEPYS
FSEPAQQAMR	KALTTRYALL	650
PHLYTLFHQA	HVAGETVARP	LFEFFKDDSS
TWTVDHQLLW	GEALLITPVL	700
QAGKAEVTGY	FPLGTWYDLQ	TVPVEALGSL
PPPPAAPREP	AIHSEGVWVT	750
LPAPLDITNV	HLRAGYIIP	L
QPGPLTTTES	RQOPMALAVA	LTGKGARGE
800		
LFWDDGESLE	VLERGAYTQV	IFLARNNTIV
NELVRVTSEG	AGLQLQKVTV	850
LGVATAPQQV	LSNGVPVSNF	TYSPDTKVLD
ICVSLLMGEQ	FLVSWC	896
Post-translational modifications		
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro		
26-53, 36-52, 47-71, 477-502, 591-602		
Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación		
N84, N177, N334, N414, N596, N826, N869		

cirtuvivintum
cirtuvivint

2-(4-methylpiperazin-1-yl)-N-[6-(1-methyl-1H-pyrazol-4-yl)isoquinolin-3-yl]pyridine-4-carboxamide

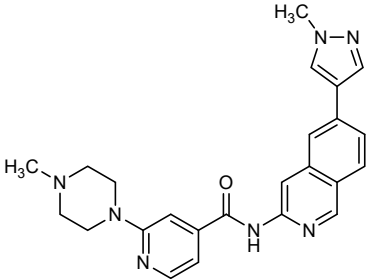
cirtuvivint

2-(4-méthylpipérazin-1-yl)-N-[6-(1-méthyl-1H-pyrazol-4-yl)isoquinoléin-3-yl]pyridine-4-carboxamide

cirtuvivint

2-(4-metilpiperazin-1-il)-N-[6-(1-metil-1H-pirazol-4-il)isoquinolein-3-il]piridina-4-carboxamida

C₂₄H₂₅N₇O



coprelotamabum

coprelotamab

immunoglobulin G1-kappa, anti-[*Homo sapiens* ERBB2 (epidermal growth factor receptor 2, receptor tyrosine protein kinase erbB-2, EGFR2, HER2, HER-2, p185cerbB2, NEU, CD340)], humanized monoclonal antibody; gamma1 heavy chain humanized (1-450) [VH humanized (*Homo sapiens* IGHV3-66*01 (81.6%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) - *Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), hinge 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-214')-disulfide with kappa light chain humanized (1'-214') [V-KAPPA humanized (*Homo sapiens* IGKV1-39*01 (86.3%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (229-229":232-232")-bisdisulfide, produced in Chinese hamster ovary (CHO)-DG44 cell line, glycoform alfa

coprélotamab

immunoglobuline G1-kappa, anti-[*Homo sapiens* ERBB2 (récepteur 2 du facteur de croissance épidermique, récepteur tyrosine-protéine kinase erbB-2, EGFR2, HER2, HER-2, p185c-erbB2, NEU, CD340)], anticorps monoclonal humanisé; chaîne lourde gamma1 humanisée (1-450) [VH humanisé (*Homo sapiens* IGHV3-66*01 (81.6%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) - *Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), charnière 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-214')-disulfure avec la chaîne légère kappa humanisée (1'-214') [V-KAPPA humanisé (*Homo sapiens* IGKV1-39*01 (86.3%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (229-229":232-232")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-DG44, glycoforme alfa

coprelotamab

inmunoglobulina G1-kappa, anti-[*Homo sapiens* ERBB2 (receptor 2 del factor de crecimiento epidérmico, receptor tirosina-proteína kinasa erbB-2, EGFR2, HER2, HER-2, p185c-erbB2, NEU, CD340)], anticuerpo monoclonal humanizado; cadena pesada gamma1 humanizada (1-450) [VH humanizado (*Homo sapiens* IGHV3-66*01 (81.6%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), bisagra 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-214')-disulfuro con la cadena ligera kappa humanizada (1'-214') [V-KAPPA humanizado (*Homo sapiens* IGKV1-39*01 (86.3%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (229-229":232-232")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-DG44, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
 EVQLVESGGG LVQPGGSLRL SCAASGFNIK DTYIHWVRQA PGKGLEWVAR 50
 IYPTNGYTRY ADSVKGRTTI SADTSKNITAY LQMNSLRAED TAVVYCSRWG 100
 GDGFYAMDYW GQGTLLTVSS ASTKGPSVFP LAPSSKSTSG GTAALGCLVK 150
 DYFPEPTVS WNSGALTSGV HTFPAVLQSS GLYSLSSVVT VPSSSLGTQT 200
 YICNVNHKPS NTKVDKKVEP KSCDKHTCP PCPAPELLGG PSVFLFPPKP 250
 KDTLMISRTPEVTCVVVDVSHEDPEVKFNW YVDGVEVHNA KTKPREEQYN 300
 STYRVVSVLT VLNQDNLNGK EYKCKVSNKA LPATIEKTIS KAKGQPREPQ 350
 VYTLPPSRDE LTKNQVSLTCLVKGFYPSDI AVEWESNGQP ENNYKTTTPV 400
 LSSDGSFFLY SKLTVDKSRW QQGNVFCSCV MHEALHNHYT QKSLSLSPGK 450

Light chain / Chaîne légère / Cadena ligera
 DIQMTQSPSS LSASVGRVIT ITCRASQDVN TAVAWYQQKPK GKAPKLLIYS 50
 ASFLYSGVPS RFSGSRSGTD FTLTISSLQPEDFATYYCQ HYTTPPTFGQ 100
 GTKVEIKRTV AAPSVFIFPP SDEQLKSGTA SVVCLLNIFY PREAKVQWVK 150
 DNALQSGNSQ ESVTEQDSKDTSTYLSSTLT LSKADYEKHK VYACEVTHQG 200
 LSSPVTKSFN RGECC 214

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 147-203 264-324 370-428
 22"-96" 147"-203" 264"-324" 370"-428"

Intra-L (C23-C104) 23"-88" 134"-194"
 23"-88" 134"-194"

Inter-H-L (h 5-CL 126) 223-214' 223"-214"

Inter-H-H (h 11, h 14) 229-229" 232-232"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

300, 300"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

450, 450"

cotoretigenum toliparvovecum

cotoretigene toliparvovec

A replication-defective adeno-associated viral vector encoding codon-optimised human retinitis pigmentosa GTPase regulator (RPGR).

A recombinant replication-defective adeno-associated viral vector serotype 8 (rAAV8) encoding codon-optimised human retinitis pigmentosa GTPase regulator (RPGR), under the control of human rhodopsin kinase (RK) promoter and bovine growth hormone poly(A) sequence, flanked by AAV2 inverted terminal repeats (ITR).

cotoretigène toliparvovec

Un vecteur adénoviral défectueux en terme de réplication et codant pour le régulateur de la GTPase de la rétinite pigmentaire humaine avec des codons optimisés (RPGR). Un vecteur adénoviral recombinant de sérotype 8 (rAAV8) à réplication défectueuse codant pour un régulateur de la GTPase de la rétinite pigmentaire humaine (RPGR) avec des codons optimisés, sous contrôle du promoteur de la kinase de la rhodopsine humaine et de la séquence poly(A) de l'hormone de croissance bovine, flanqué de répétitions terminales inversées (ITR) de l'AAV2.

cotoretigén toliparvovec

Un vector de virus adeno-asociado, deficiente de replicación, que codifica para el regulador de la GTPasa implicado en la retinosis pigmentaria humana (RPGR) con los codones optimizados.

Un vector de virus adeno-asociado recombinante de serotipo 8 (rAAV8), deficiente de replicación, que codifica para el regulador de la GTPasa implicado en la retinosis pigmentaria humana (RPGR) con los codones optimizados, bajo el control del promotor de la rodopsin quinasa (RK) humana y la secuencia poly(A) de la hormona de crecimiento bovina, flanqueado por las repeticiones terminales invertidas (ITR) del AAV2.

Recommended INN: List 85

WHO Drug Information, Vol. 35, No. 1, 2021

cotosudilum

cotosudil

6-[(2R)-2-methyl-1,4-diazocane-1-sulfonyl]isoquinoline

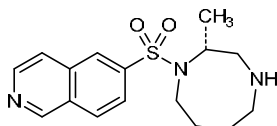
cotosudil

6-[(2R)-2-méthyl-1,4-diazocane-1-sulfonyl]isoquinoléine

cotosudil

6-[(2R)-2-metil-1,4-diazocano-1-sulfonil]isoquinoleina

C₁₆H₂₁N₃O₂S



dafsolimabum

dafsolimab

immunoglobulin G2B-kappa, anti-[CD3E (CD3 epsilon, Leu-4)], *Mus musculus* monoclonal antibody;

gamma2b heavy chain *Mus musculus* (1-456) [VH (*Mus musculus* IGHV1-4*01 (95.9%) -(IGHD) -IGHJ4*01 (94.1%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Mus musculus* IGHG2B*02 (100%) (CH1 (121-217), hinge 1-22 (218-239), CH2 (240-349), CH3 (350-454), CHS (455-456)) (121-456)], (135-213')-disulfide with kappa light chain *Mus musculus* (1'-213') [V-KAPPA (*Mus musculus* IGKV4-59*01 (100%) -IGKJ5*01 (100%)) CDR-IMGT [5.3.9] (27-31.49-51.88-96) (1'-106') -*Mus musculus* IGKC1*01 (100%) (107'-213')];

dimer (229-229'':232-232'':235-235'':238-238'')-tetrakisdisulfide, produced in SP2/0-derived mouse myeloma cells, glycoform alfa

dafsolimab

immunoglobuline G2B-kappa, anti-[CD3E (CD3 epsilon, Leu-4)], anticorps monoclonal *Mus musculus*;

chaîne lourde gamma2b *Mus musculus* ((1-456) [VH (*Mus musculus* IGHV1-4*01 (95.9%) -(IGHD) -IGHJ4*01 (94.1%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Mus musculus* IGHG2B*02 (100%) (CH1 (121-217), charnière 1-22 (218-239), CH2 (240-349), CH3 (350-454), CHS (455-456)) (121-456)], (135-213')-disulfure avec la chaîne légère kappa *Mus musculus* (1'-213') [V-KAPPA (*Mus musculus* IGKV4-59*01 (100%) -IGKJ5*01 (100%)) CDR-IMGT [5.3.9] (27-31.49-51.88-96) (1'-106') -*Mus musculus* IGKC1*01 (100%) (107'-213')];

dimère (229-229'':232-232'':235-235'':238-238'')-tétrakisdisulfure, produit par une ligne cellulaire dérivée de myélome murin SP2/0, glycoforme alfa

dafsolimab

inmunoglobulina G2B-kappa, anti-[CD3E (CD3 épsilon, Leu-4)], anticuerpo monoclonal *Mus musculus*;

cadena pesada gamma2b *Mus musculus* ((1-456) [VH (*Mus musculus* IGHV1-4*01 (95.9%) -(IGHD) -IGHJ4*01 (94.1%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Mus musculus* IGHG2B*02 (100%) (CH1 (121-217), bisagra 1-22 (218-239), CH2 (240-349), CH3 (350-454), CHS (455-456)) (121-456)], (135-213')-disulfuro con la cadena ligera kappa *Mus musculus* (1'-213') [V-KAPPA (*Mus musculus* IGKV4-59*01 (100%) -IGKJ5*01 (100%)) CDR-IMGT [5.3.9] (27-31.49-51.88-96) (1'-106') -*Mus musculus* IGKC1*01 (100%) (107'-213')];

dímero (229-229'':232-232'':235-235'':238-238'')-tetrakisdisulfuro, producido en una línea celular de mieloma murino SP2/0, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
 QVQLQSGAG LARFGASVKM SCARSTYFT SYTHRRVYKQ FGQGLEWIGY 50
 INFSSGYTNV IQREKDKATL TADKSSSTAY MQVSSLTSED SAVVYCARGS 10C
 RYDYYGHGYR GQGTSTVTVSS AKTTFPSVYP LAFGGGDTTG SSVTLGCLVK 15C
 GYFPESVTVT WNSGSLSSSV HTFFALLQSG LYTMSSSVTV PSSTWFSQTV 200
 TCSVAHFSS TTVDKKLEPS GFISTINPCP FCKECHKCPA FNLEGGPSVF 250
 IFFPNKIDVL MISLTKVTC VVVDVSEDDP DVQISWVFN VEVHTAQTTQ 30C
 HREDYNSTIR VVSTLPIQHQ DMMSGKEFKC KVNKKDLPSF IERTISKING 35C
 LVRAEQVYIL FPPAEQLRSK DVSLTCLVVG FNPGDISEV TSNHTEENY 40C
 FDTAPVLQSD GSYTIYSKLN MKTSKWKETO SFSCHVRHGS LKNYLYKKTI 45C
 SRSRGK 456

Light chain / Chaîne légère / Cadena ligera
 QIVLTQSPAI MSASPGKEVT MTCSASSSVS YHHWYQQKSG TSPKRWIYDT 50
 SKLASGVPAR FSGSGSTYS SLTISSEAE DAATYYCQW SSNPLTFGAG 10C
 TKLELRADA APTVSIFPPS SEQLTSGGAS VVCFANRYP KDINVKWKID 15C
 GSERQNGVLN SWTDQSKDS TYSNSSLTLL TKDEYERHNS YTCEATHRTS 200
 TSFIVKSFNR NEC 213

Post-translational modifications
 Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
 Intra-H (C23-C104) 22-96 147-202 270-330 376-434
 22'-96" 147'-202" 270'-330" 376'-434"
 Intra-L (C23-C104) 23'-87" 133'-193"
 23"-87" 133"-193"
 Inter-H-L (C1H 11-C1 126) 135'-213' 135"-213"
 Inter-H-L (h 12, h 15, h 18, h 21) 229'-229" 232'-232" 235'-235" 238'-238"

N-terminal glutamyl-lysine cleavage to pyroglutanyl (pL; 5-oxoprol)
 HIV Q1:
 1, 1"
 L VI Q1:
 1', 1"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
 H C102 N84,4:
 306,306"

Fucosylated complex bi-antennary SP2/0-type glycans / glycanes de type SP2/0 bi-antennaires
 complejos fucosilados / glicanos de tipo SP2/0 bi-antennarios complejos fucosilados
 C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal
 H C105 K2:
 456,456"

dafsolimabum setaritoxum #

dafsolimab setaritox

immunoglobulin G2B-kappa, anti-[CD3E (CD3 epsilon, Leu-4)], *Mus musculus* monoclonal antibody conjugated to aglycosylated ricin toxin A (RTA);

gamma2b heavy chain *Mus musculus* (1-456) [VH (*Mus musculus* IGHV1-4*01 (95.9%) - (IGHD) -IGHJ4*01 (94.1%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Mus musculus* IGHG2B*02 (100%) (CH1 (121-217), hinge 1-22 (218-239), CH2 (240-349), CH3 (350-454), CHS (455-456)) (121-456)], (135-213')-disulfide with kappa light chain *Mus musculus* (1'-213') [V-KAPPA (*Mus musculus* IGKV4-59*01 (100%) - IGKJ5*01 (100%)) CDR-IMGT [5.3.9] (27-31.49-51.88-96) (1'-106') -*Mus musculus* IGKC1*01 (100%) (107'-213')]; dimer (229-229":232-232":235-235":238-238")-tetrakisdisulfide, produced in SP2/0-derived mouse myeloma cells, glycoform alfa, substituted at N⁶ of an average of 1.6 lysyl residues with 4-[(1RS)-1-[[L-methionyl-ricin toxin A-chain (Met-RTA, non-glycosylated, produced in *Escherichia coli*)-S²⁶⁰-yl]sulfanyl]ethyl]benzoyl groups

dafsolimab sétaritox

immunoglobuline G2B-kappa, anti-[CD3E (CD3 epsilon, Leu-4)], anticorps monoclonal *Mus musculus* conjugué à la toxine A de la ricine aglycosylée (RTA); chaîne lourde gamma2b *Mus musculus* -(1-456) [VH (*Mus musculus* IGHV1-4*01 (95.9%) - (IGHD) -IGHJ4*01 (94.1%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Mus musculus* IGHG2B*02 (100%) (CH1 (121-217), charnière 1-22 (218-239), CH2 (240-349), CH3 (350-454), CHS (455-456)) (121-456)], (135-213')-disulfure avec la chaîne légère kappa *Mus musculus* (1'-213') [V-KAPPA (*Mus musculus* IGKV4-59*01 (100%) - IGKJ5*01 (100%)) CDR-IMGT [5.3.9] (27-31.49-51.88-96) (1'-106') -*Mus musculus* IGKC1*01 (100%) (107'-213')]; dimère (229-229":232-232":235-235":238-238")-tétrakisdisulfure, produit par une lignée cellulaire dérivée de myélome murin SP2/0, glycoforme alfa, substitué en N⁶ sur un moyenne de 1.6 résidus lysyl par des groupes 4-[(1RS)-1-[[L-méthionyl-chaîne A de la toxine de ricine (Met-RTA, non-glycosylée, produite par *Escherichia coli*)-S²⁶⁰-yl]sulfanyl]éthyl]benzoyl

dafsolimab setaritox

inmunoglobulina G2B-kappa, anti-[CD3E (CD3 épsilon, Leu-4)], anticuerpo monoclonal *Mus musculus* conjugado con la toxina A de la ricina aglicosilada (RTA);
cadena pesada gamma2b *Mus musculus* -(1-456) [VH (*Mus musculus* IGHV1-4*01 (95.9%) -(IGHD) -IGHJ4*01 (94.1%) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Mus musculus* IGHG2B*02 (100%) (CH1 (121-217), bisagra 1-22 (218-239), CH2 (240-349), CH3 (350-454), CHS (455-456)) (121-456)], (135-213')-disulfuro con la cadena ligera kappa *Mus musculus* (1'-213') [V-KAPPA (*Mus musculus* IGKV4-59*01 (100%) -IGKJ5*01 (100%)) CDR-IMGT [5.3.9] (27-31.49-51.88-96) (1'-106') -*Mus musculus* IGKC1*01 (100%) (107'-213')];
dímero (229-229'':232-232'':235-235'':238-238'')-tetrakisdisulfuro, producido en una línea celular de mieloma murino SP2/0, forma glicosilada alfa, sustituida en N⁶ de un promedio de 1,6 residuos de lisilo con grupos de 4-[(1RS)-1-[[L-metionil-cadena A de toxina de ricina (Met-RTA, no glicosilada, producida por *Escherichia coli*)-S²⁶⁰-il]sulfanil]etil]benzoilo

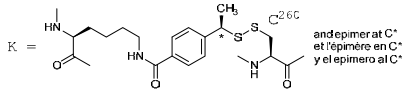
Heavy chain/Chaîne lourde/ Cadenapesada
QVQLQSGGAE LARPGASVKM SCKASGYTFT SYTMHWVKDR PGQGLEWIGY 5C
INPSSGYTNY IQRFKDKATL TADKSSSTAY MOVSSLTSED SAVYICARGS 100
RYDYIGMDYV GQGTSTVVS AKTTPPSVYP LAPGCGDTTG SSVTLGCLVK 150
GYFPESTVTV WNSGSLSSV HTFPALLQSG LYTNSSSVTV PSTWPSQTV 20C
TCSVAFPASS TTVOCKLEPS GPISINPCP ECKECHKCPA PNLEGGPSVE 25C
IYPENIKUVL MISLTPKVTG VVVDSVEDDP DVQISWFEVN VEVHTAGTQT 30C
HREDYNTIR VVSTLPIQHG DWSGKEFKC KVNKKDLPS IERTISKING 35C
LVRAPOVYLL PPPAEQLSRK DVSITCLVVG FNPGLISVEW TSNHTEENY 40C
KDTAPQVDSG GSYFIYSKLR MNTSKWEKTD SFSCNVRHEG LKNYYLKKTI 45C
SRSPGK 456

Light chain/ Chaîne légère/ Cadena ligera
QIVLTQSPAL MSASPGEKVT MTCASSSVS YMHVYQQKSG TSPKRWIYDC 5C
SKLASGVAPR FSGSGSTYS SLTISSEAE DAATYYCQQW SSNPLTFGAG 100
TKLLKPAD APTVSIFPPS SEQLTSGGAS VVCFLNFPY KDINVKWKID 150
GSEKQVGLN SWTDQSKDS TYSMSSTLT TLKDEYERHNS YTCEATHKTS 20C
TSPIVKSFNR NEC 213

Met-Ricin A chain/ Met-chaîne A de la Ricine/ Met-cadena A de la Ricina
MIFPKQYPII NPTTAGATVG SYTNFIRAVR GLTTGADVE HUIPVLNPRV 5C
GLPINQRFIL VELSNHAELS VTLLADVTNA YVVGVRAGNS AYFFEDNQOE 100
DAEAIHTLFI DVQNYTFAP GGNVDRLEQL AGNLDRENIEL GNGPLEEAI 150
ALYYSTGGT QLPTLARSFI ICIQMISEAA RFQYIEGEMR TRIRYMRSA 20C
PDESVITLEN SWERLSTAIQ ESNQGAFASP IQLQRNRGSK FSVYDVSLIL 25C
PIIALMVYMC APPPSQF 268

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure/ Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 147-202 270-330 376-434
22"-96" 147"-202" 270"-330" 376"-434"
Intra-L (C23-C104) 23-87 133-193
23"-87" 133"-193"
Inter-H-L (CH1 H-CL 126) 135-213' 135"-213"
Inter-H-H (h 12, h 15, h 18, h 21) 229-229' 232-232" 235-235" 238-238"
Met-RTA: C172 = Cys-SH,
C260 = Cys-S-S-CH(CH3)-p-C6H4-CO-[N6-Lysyl-H,L]]

Conjugation sites/ Sites de conjugaison/ Posiciones de conjugación
major: K74 K141' K206"
K74" K141" K206"
minor: K326 K329 K349 K442 K447 K18' K52' K168'
K326" K329" K349" K442" K447" K18" K52" K168"



N-terminal glutaminyI cyclization to pyrrolutamyI (pE, 5-oxoprolyl)
H V H (Q1):
I, I"
L V L (Q1):
I', I'''

N-glycosylation sites/ Sites de N-glycosylation/ Posiciones de N-glicosilación
H CH2 N84.4:
306, 306"
Fucosylated complex bi-antennary SP2/0-type glycans / glycanes de type SP2/0 bi-antennaires complexes fucosylés / glicanos de tipo SP2/0 biantenarijos complejos fucosilados.

C-terminal lysine clipping/ Coupure de la lysine C-terminale/ Recorte de lisina C-terminal
H CHS K2:
456, 456"

daleutonum

daleuton

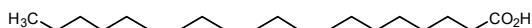
(8Z,11Z,14Z)-icosa-8,11,14-trienoic acid

daleuton

acide (8Z,11Z,14Z)-icosa-8,11,14-triénoïque

daleutón

ácido (8Z,11Z,14Z)-icosa-8,11,14-trienoico

**dalosirvatum**

dalosirvat

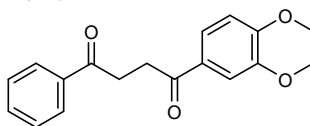
1-(2,3-dihydro-1,4-benzodioxin-6-yl)-4-phenylbutane-1,4-dione

dalosirvat

1-(2,3-dihydro-1,4-benzodioxin-6-yl)-4-phénylbutane-1,4-dione

dalosirvat

1-(2,3-dihidro-1,4-benzodioxin-6-il)-4-fenilbutano-1,4-diona

**dalpiciclibum**

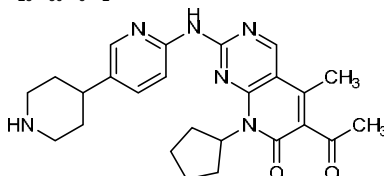
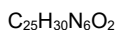
dalpiciclib

6-acetyl-8-cyclopentyl-5-methyl-2-[[5-(piperidin-4-yl)pyridin-2-yl]amino]pyrido[2,3-*d*]pyrimidin-7(8*H*)-one

dalpiciclib

6-acétyl-8-cyclopentyl-5-méthyl-2-[[5-(pipéridin-4-yl)pyridin-2-yl]amino]pyrido[2,3-*d*]pyrimidin-7(8*H*)-one

dalpiciclib

6-acetil-8-ciclopentil-5-metil-2-[[5-(piperidin-4-il)piridin-2-il]amino]pirido[2,3-*d*]pirimidin-7(8*H*)-ona**danavorextonum**

danavorexton

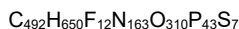
methyl (2*R*,3*S*)-3-(methanesulfonamido)-2-[[*cis*-4-phenylcyclohexyl]oxy]methyl}piperidine-1-carboxylate

danavorexton

(2*R*,3*S*)-3-(méthanesulfonamido)-2-[[*cis*-4-phénylcyclohexyl]oxy]méthyl}pipéridine-1-carboxylate de méthyle

danavorexton

(2*R*,3*S*)-2-[[*cis*-4-fenilciclohexil]oxi]metil}-3-(metanosulfonamido)piperidina-1-carboxilato de metilo



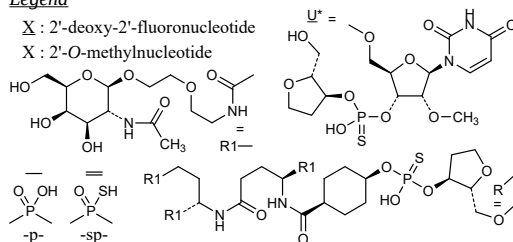
(3'-5') R=G-U-G-G-A-C-U-U-C-U-C-A-A-U-U-U-C-U*

(5'-3') C=A-C-C-U-G-A-A-G-A-G-U-U-A-A-A=A=G=A

Legend

X : 2'-deoxy-2'-fluoronucleotide

X : 2'-O-methylnucleotide



darigabatum

darigabat

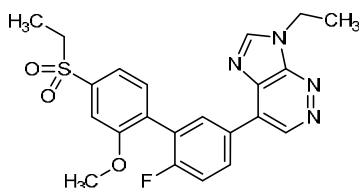
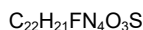
4-[4'-(ethanesulfonyl)-6-fluoro-2'-methoxy[1,1'-biphenyl]-3-yl]-7-ethyl-7H-imidazo[4,5-c]pyridazine

darigabat

4-[4'-(éthanesulfonyl)-6-fluoro-2'-méthoxy[1,1'-biphényl]-3-yl]-7-éthyl-7H-imidazo[4,5-c]pyridazine

darigabat

4-[4'-(etanosulfonyl)-6-fluoro-2'-metoxi[1,1'-bifenil]-3-il]-7-etil-7H-imidazo[4,5-c]piridazina



darovasertibum

darovasertib

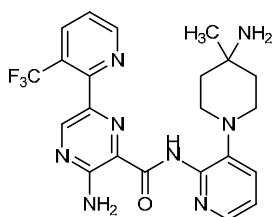
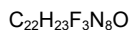
3-amino-N-[3-(4-amino-4-methylpiperidin-1-yl)pyridin-2-yl]-6-[3-(trifluoromethyl)pyridin-2-yl]pyrazine-2-carboxamide

darovasertib

3-amino-N-[3-(4-amino-4-méthylpipéridin-1-yl)pyridin-2-yl]-6-[3-(trifluorométhyl)pyridin-2-yl]pyrazine-2-carboxamide

darovasertib

3-amino-N-[3-(4-amino-4-metilpiperidin-1-il)piridin-2-il]-6-[3-(trifluorometil)piridin-2-il]pirazina-2-carboxamida



daxdilimabum #

daxdilimab

immunoglobulin G1-lambda, anti-[*Homo sapiens* LILRA4 (leukocyte immunoglobulin like receptor A4, ILT7, CD85g)], *Homo sapiens* monoclonal antibody; gamma1 heavy chain *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV1-18*01 (98.0%) -(IGHD) - IGHJ2*01 (92.9%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -*Homo sapiens* IGHG1*03 (100%) G1m3, nG1m1 (CH1 R120 (219) (123-220), hinge 1-15 (221-235), CH2 (236-345), CH3 E12 (361), M14 (363) (346-450), CHS (451-452)) (123-452)], (225-215')-disulfide with lambda light chain *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV2-14*01 (98.0%) -IGLJ2*01 (90.9%)) CDR-IMGT [9.3.10] (26-34.52-54.91-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dimer (231-231":234-234")-bisdisulfide, produced in a Chinese hamster ovary (CHO)-derived cell line, glycoform alfa

daxdilimab

immunoglobuline G1-lambda, anti-[*Homo sapiens* LILRA4 (récepteur A4 immunoglobuline-like des leucocytes, ILT7, CD85g)], anticorps monoclonal *Homo sapiens*; chaîne lourde gamma1 *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV1-18*01 (98.0%) -(IGHD) - IGHJ2*01 (92.9%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -*Homo sapiens* IGHG1*03 (100%) G1m3, nG1m1 (CH1 R120 (219) (123-220), charnière 1-15 (221-235), CH2 (236-345), CH3 E12 (361), M14 (363) (346-450), CHS (451-452)) (123-452)], (225-215')-disulfure avec la chaîne légère lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV2-14*01 (98.0%) -IGLJ2*01 (90.9%)) CDR-IMGT [9.3.10] (26-34.52-54.91-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dimère (231-231":234-234")-bisdisulfure, produite dans une lignée cellulaire dérivée des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

daxdilimab

inmunoglobulina G1-lambda, anti-[*Homo sapiens* LILRA4 (receptor A4 inmunoglobulina-like de los leucocitos, ILT7, CD85g)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma1 *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV1-18*01 (98.0%) -(IGHD) - IGHJ2*01 (92.9%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -*Homo sapiens* IGHG1*03 (100%) G1m3, nG1m1 (CH1 R120 (219) (123-220), bisagra 1-15 (221-235), CH2 (236-345), CH3 E12 (361), M14 (363) (346-450), CHS (451-452)) (123-452)], (225-215')-disulfuro con la cadena ligera lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV2-14*01 (98.0%) -IGLJ2*01 (90.9%)) CDR-IMGT [9.3.10] (26-34.52-54.91-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dímero (231-231":234-234")-bisdisulfuro, producido en una línea celular derivada de las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadenapesada
QVQLQSGAE VKKFGASVKV SCKASGYTFT SYGISWVRQA PGQGLEWMGW 50
ISAYNGNTNY AQLQGRVTM TIDTSTSTAY MELRSLRSDO TAVYYCARNG 100
LWGWDSDAFD IWGRGTLVTY SSASTKGPSV FFLAPSSKST SGGTAALGCL 150
VKDYFPEPVT VSWNSGALTIS GVHTFPAVLQ SSGLYSLSSV VTPSSSLGT 20C
QTYICNVRRHK PSNTKVDKRV EPKSCDKTHT CPFCPAPELL GGPVFLFPF 25C
KPKDTLMISR TPEVTCVVVD VSHEDPEVKF NWYVDGVEVH NAKTKPREEQ 30C
YNSTYRVVSV LTVLHQDWLN GKEYCKVSN KALPAPIEKT ISKAKGPPE 35C
PQVYTLPPSR EEMTKNQYSL TCLVKGFYPS DIAVEWESNG QPENNYKTF 400
PVLDSGSEFF LYSKLTVDKS RWQQGNVFSC SVMHEALHNN YTQKSLSLSP 450
GK 452

Light chain / Chaîne légère / Cadena ligera
QSALTQPASV SGSPGQSITI SCTGTSSDVG GYNYVSWYQQ HPGKAPKLNI 50
YDVSNRPSGV SNRPSGSKSG NTASLTISGL QAEDADYIC SSYTSSSTV 100
FGGGTKVTVL GQPKAAPSVT LFPSSSEELQ ANKATLVCLI SDFYPGAVT 150
AWKADSSPVK AGVETTTFSK QSNNKYAASS YLSLTPEQWK SHRSYSCQVT 20C
HEGSTVERVT APTTECS 216

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 149-205 266-326 372-430
22"-96" 149"-205" 266"-326" 372"-430"
Intra-L (C23-C104) 22"-90" 138"-197"
22"-90" 138"-197"
Inter-H-L (h 5-CL 126) 225-215" 225"-215"
Inter-H-H (h 11, h 14) 231-231" 234-234"

N-terminal glutaminyl cyclization to pyroglutaminyl (pE, 5-oxoprolinyl)
H VH Q1:
I, I"
L VL Q1:
I', I'''

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
302, 302"
Afucoylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires
complexes afucosylés / glicanos de tipo CHO biantennarios complejos afucosilados

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal
H CHS K2:
452, 452"

dazodalibepum #
dazodalibep

engineered binding protein (1-85) anti-(human CD40 ligand) derived from the human tenascin third fibronectin type III domain (785-869), fused via the peptidyl linker ⁸⁶GGGGGGGGGGGGGGGG¹⁰⁰ to engineered binding protein (101-190) anti-(human CD40 ligand) derived from the human tenascin third fibronectin type III domain (780-869), fused via the peptidyl linker ¹⁹¹GGGGGGGGGGGG²⁰⁰ to human albumin (201-785), variant (C³⁴>S²³⁴), produced in *Escherichia coli*; [tenascin (785-869)-peptide (1-85) (containing a third fibronectin type III domain), engineered for binding to the CD40 ligand (CD40L)]-[G₁₅ linker (86-100)]-[tenascin (780-869)-peptide (101-190) (containing a third fibronectin type III domain), engineered for binding to the CD40 ligand (CD40L)]-[G₁₀ linker (91-100)]-[(C³⁴>S)-human serum albumin (HSA)] fusion protein, produced in *Escherichia coli*

dazodalibep

protéine de liaison mise au point (1-85) anti-(ligand de CD40 humain) dérivée du troisième domaine de la fibronectine de type III de la ténascine humaine, fusionnée via un peptide liant ⁸⁶GGGGGGGGGGGGGGGG¹⁰⁰ à la protéine de liaison mise au point (101-190) anti-(ligand de CD40 humain) dérivée du troisième domaine de la fibronectine de type III de la ténascine humaine, fusionnée via un peptide liant ¹⁹¹GGGGGGGGGGGG²⁰⁰ à l'albumine humaine (201-785), variant (C³⁴>S²³⁴), produite par *Escherichia coli*; protéine de fusion de [ténascine (785-869)-peptide (1-85) (contenant un troisième domaine de fibronectine de type III), conçu pour se lier au ligand de CD40 (CD40L)]-[peptide G₁₅ de liaison (86-100)]-[ténascine (780-869)-peptide (101-190) (contenant un troisième domaine de fibronectine de type III), conçu pour se lier au ligand de CD40 (CD40L)]-[peptide G₁₀ de liaison (91-100)]-[(C³⁴>S)-albumine sérique humaine (ASH)], produite par *Escherichia coli*

dazodalibep

proteína de unión diseñada (1-85) anti-(ligando CD40 humano) derivada de un tercer dominio de fibronectina tipo III de tenascina humana (785-869), fusionada a través de un conector ⁸⁶GGGGGGGGGGGGGG¹⁰⁰ a la proteína de unión diseñada (101-190) anti-(ligando CD40 humano) derivado de un tercer dominio de fibronectina tipo III de tenascina humana (780-869), fusionado a través de un conector peptidil ¹⁹¹GGGGGGGGGG²⁰⁰ a la albúmina humana (201-785), variante (C³⁴>S²³⁴), producida en *Escherichia coli*;
proteína de fusión de [tenascina (785-869)-péptido (1-85) (que contiene un tercer dominio de fibronectina tipo III), diseñado para unirse al ligando de CD40 (CD40L)]-[péptido conector G₁₅ (86-100)]-[tenascina (780-869)-péptido (101-190) (que contiene un tercer dominio de fibronectina tipo III), diseñado para unirse al ligando de CD40 (CD40L)]-[péptido conector G₁₀ (91-100)]-[C³⁴>S)-albúmina sérica humana (ASH)], producida por *Escherichia coli*

Sequence / Séquence / Secuencia		
SQIEVKDVTDT	TTALITWSDD	FGEYVWCELT YGIKDVPGDR TTIDLWYHHA 50
HYSIGNLKPD	TEYEVSLICR	SGDMSSNPAK ETFTTGGGGG GGGGGGGGGG 100
RLDAPSQIEV	KDVTDTTALI	TWSDDFGEYV WCELTYGIKD VPGDRTTIDL 150
WYHHAHYSIG	NLKPDTEYEV	SLICRSGDMS SNPAKETFTT GGGGGGGGGG 200
DAHKSEVAHR	FKDLGEENFK	ALVLI AFAQY LQQSPFEDHV KLVNEVTEFA 250
KTCVADESAR	NCDKSLHTLF	GDKLCTVATL RETYGMADC CAKQEPERNE 300
CFLQHKDDNP	NLRLVRPEV	DVMCTAFHDN EETFLKKYLY EIARRHPYFY 350
APELLFFAKR	YKAATFECQ	AADKAACLLP KLDLRLDEGK ASSAKQRLKC 400
ASLQKFGERA	FKAWAVARLS	QRFPKAEFAE VSKLVDTLTK VHTECCGDL 450
LECADDRADL	AKYICENQDS	ISSKLKECC E KPLLEKSHCI AEVENDEMPA 500
DLPSLAADFV	ESKDVCKNYA	EAKDVF LGMF LYEYARRHPD YSVVLLRLA 550
KTYETTLKLC	CAAADPHECY	AKVDFEFKPL VEEPQNLIKQ NCELFEQLGE 600
YKFQNALVR	YTKKVPQVST	PTLVEVSRNL GKVGSCKCKH PEAKRMPCAE 650
DYLSVVLNQL	CVLHEKTPVS	DRVTKCCTES LVNRRPCFSA LEVDETYVPK 700
EFNAETFTFH	ADICTLSEKE	RQIKKQ TALV ELVKHKPKAT KEQLKAVMDD 750
FAAFVEKCKC	ADDKETCFAE	EGKKLVAASQ AALGL 785
Mutations / Mutations / Mutaciones		
F ^{18,123>S} , K ^{19,124>D} , P ^{20,125>D} , L ^{21,126>F} , A ^{22,127>G} , I ^{24,129>Y} , D ^{25,130>V} , G ^{26,131>W} , I ^{27,132>C} , T ^{46,151>W} , E ^{47,152>Y} , D ^{48,153>H} , E ^{49,154>H} , N ^{50,155>A} , Q ^{51,156>H} , S ^{69,174>C} , R ^{71,176>S} , C ^{234>S}		
Post-translational modifications		
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro		
27-69, 132-174, 253-262, 275-290, 291-301, 324-368, 369-377, 400-445, 446-453, 465-478, 479-489, 516-560, 561-569, 592-637, 638-648, 661-676, 677-687, 714-758, 759-767		
Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación		
none / aucune / ninguna		

depemokimabum #
depemokimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* IL5 (interleukin 5, IL-5)], humanized monoclonal antibody;
gamma1 heavy chain humanized (1-449) [VH humanized (*Homo sapiens* IGHV2-70*19 (75.8%) -(IGHD) -IGHJ4*01 (85.7%)) CDR-IMGT [8.7.13] (26-33.51-57.96-108) (1-119) -*Homo sapiens* IGHG1*03 G1m3, nG1m1, G1v21 CH2 Y15.1, T16, E18 (CH1 R120 (216) (120-217), hinge 1-15 (218-232), CH2 M15.1>Y (254), S16>T (256), T18>E (258) (233-342), CH3 E12 (358), M14 (360) (343-447), CHS (448-449)) (120-449)], (222-220')-disulfide with kappa light chain humanized (1'-220') [V-KAPPA (*Homo sapiens* IGKV4-1*01 (91.1%) -IGKJ2*02 (90.9%)) CDR-IMGT [12.3.9] (27-38.56-58.95-103) (1'-113') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (159), V101 (197) (114'-220')]; dimer (228-228":231-231")-bisdisulfide, produced in Chinese hamster ovary (CHO)-K1 cell line lacking the glutamine synthetase (GS) gene, glycoform alfa

dépémokimab

immunoglobuline G1-kappa, anti-[*Homo sapiens* IL5 (interleukine 5, IL-5)], anticorps monoclonal humanisé;
chaîne lourde gamma1 humanisée (1-449) [VH humanisé (*Homo sapiens* IGHV2-70*19 (75.8%) -(IGHD) -IGHJ4*01 (85.7%)) CDR-IMGT [8.7.13] (26-33.51-57.96-108) (1-119) -*Homo sapiens* IGHG1*03 G1m3, nG1m1, G1v21 CH2 Y15.1, T16, E18 (CH1 R120 (216) (120-217), charnière 1-15 (218-232), CH2 M15.1>Y (254), S16>T (256), T18>E (258) (233-342), CH3 E12 (358), M14 (360) (343-447), CHS (448-449)) (120-449)], (222-220')-disulfure avec la chaîne légère kappa humanisée (1'-220') [V-KAPPA humanisé (*Homo sapiens* IGKV4-1*01 (91.1%) -IGKJ2*02 (90.9%)) CDR-IMGT [12.3.9] (27-38.56-58.95-103) (1'-113') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (159), V101 (197) (114'-220')];
dimère (228-228":231-231")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-K1 ne présentant pas le gène de la glutamine synthétase, glycoforme alfa

depemokimab

inmunoglobulina G1-kappa, anti-[*Homo sapiens* IL5 (interleukina 5, IL-5)], anticuerpo monoclonal humanizado;
cadena pesada gamma1 humanizada (1-449) [VH humanizado (*Homo sapiens* IGHV2-70*19 (75.8%) -(IGHD) -IGHJ4*01 (85.7%)) CDR-IMGT [8.7.13] (26-33.51-57.96-108) (1-119) -*Homo sapiens* IGHG1*03 G1m3, nG1m1, G1v21 CH2 Y15.1, T16, E18 (CH1 R120 (216) (120-217), bisagra 1-15 (218-232), CH2 M15.1>Y (254), S16>T (256), T18>E (258) (233-342), CH3 E12 (358), M14 (360) (343-447), CHS (448-449)) (120-449)], (222-220')-disulfuro con la cadena ligera kappa humanizada (1'-220') [V-KAPPA humanizado (*Homo sapiens* IGKV4-1*01 (91.1%) -IGKJ2*02 (90.9%)) CDR-IMGT [12.3.9] (27-38.56-58.95-103) (1'-113') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (159), V101 (197) (114'-220')];
dímero (228-228":231-231")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-K1 en ausencia del gen glutamina sintetasa (GS), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada

QVTLRESGPA	LVKPTQTLLT	TCTVSGFSLT	GSSVHWVRQP	PGKGLEWLG	50
IWASGGTDYN	SALMSRLSIS	KDTSRNQVVL	TMTNMDPVD	ATYYCARDPP	100
SGLLRLDYG	RGLTVTSSA	STKGPSVFPL	APSSKSTSGG	TAALGCLVKD	150
YFPEPVTVSW	NSGALTSVGH	TFFPAVLQSSG	LYSLSSVTV	PSSSLGTQTY	200
ICNVNHHKPSN	TKVDRKVEPK	SCDKTHTCCP	CPAPELLGGP	SVFLFPKPK	250
DTLYITREPE	VTCVVVDVSH	EDPEVKFNWY	VDGVEVHNAK	TKPREEQYNS	300
TYRVSVSLTV	LHQDWLNGKE	YKCKVSNKAL	PAPIEKTISK	AKGQPREPQV	350
YTPLPSREEM	TKNQVSLTCL	VKGFPYSDIA	VEWESNGQPE	NNYKTTTPVL	400
DSGGSFFLYS	KLTVDKSRWQ	QGNVFSCSVW	HEALHNHYTQ	KSLSLSPGK	449

Light chain / Chaîne légère / Cadena ligera

DIVMTQSPDS	LAVSLGERAT	INCKSSQSL	NSGNQKNYLA	WYQQKPGQPP	50
KLLIYGASTR	ESGVPRDFSG	SGSGTDFTLT	ISSLQAEDVA	VYYQCNVHSF	100
PFTFGGGTKL	EIKRTVAAPS	VFIFPPSDEQ	LKSGTASVVC	LLNNFYPREA	150
KVQWKVDNAL	QSGNSQESVT	EQDSKSTYS	LSSTLTLSKA	DYEEKHKVYAC	200
EVTHQGLSSP	VTKSFNRGEC				220

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104)	22"-95	146"-202	263"-323	369"-427
	22"-95"	146"-202"	263"-323"	369"-427"
Intra-L (C23-C104)	23'-94'	140'-200'		
	23"-94"	140"-200"		
Inter-H-L (h 5-CL 126)	222-220'	222"-220"		
Inter-H-H (h 11, h 14)	228-228"	231-231"		

N-terminal glutaminyl cyclization to pyroglutamyl (pE, 5-oxoprolyl)
VH Q1:
I, I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
299, 299"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal
H CHS K2:
449, 449"

deucravacitinibum

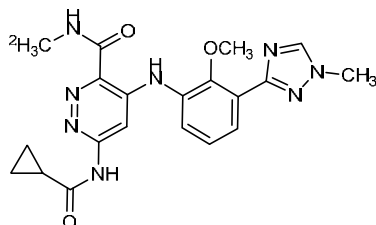
deucravacitinib

6-(cyclopropanecarboxamido)-4-[2-methoxy-3-(1-methyl-1*H*-1,2,4-triazol-3-yl)anilino]-*N*-(²H₃)methylpyridazine-3-carboxamide

deucravacitinib

6-(cyclopropanecarboxamido)-4-[2-méthoxy-3-(1-méthyl-1*H*-1,2,4-triazol-3-yl)anilino]-*N*-(²H₃)méthylpyridazine-3-carboxamide

deucravacitinib

6-(ciclopropanocarboxamido)-*N*-(²H₃)metil-4-[3-(1-metil-1*H*-1,2,4-triazol-3-il)-2-metoxianilino]piridazina-3-carboxamidaC₂₀H₁₉²H₃N₈O₃**dirocaftor**

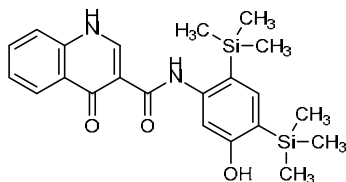
dirocaftor

N-[5-hydroxy-2,4-bis(trimethylsilyl)phenyl]-4-oxo-1,4-dihydroquinoline-3-carboxamide

dirocaftor

N-[5-hydroxy-2,4-bis(triméthylsilyl)phényl]-4-oxo-1,4-dihydroquinoléine-3-carboxamide

dirocaftor

N-[5-hidroxi-2,4-bis(trimetilsilil)fenil]-4-oxo-1,4-dihidroquinoleína-3-carboxamidaC₂₂H₂₈N₂O₃Si₂**divozilimabum #**

divozilimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* MS4A1 (membrane-spanning 4-domains subfamily A member 1, CD20)], monoclonal antibody;

gamma1 heavy chain (1-451) [VH (*Homo sapiens*IGHV1-46*01 (86.7%) - (IGHD) -IGHJ1*01 (86.7%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -*Homo sapiens*IGHG1*03 (100%) G1m3, nG1m1 (CH1 R120 (218) (122-219), hinge 1-15 (220-234), CH2 (235-344), CH3 E12 (360), M14 (362) (345-449), CHS (450-451)) (122-451)], (224-213')-disulfide with kappa light chain (1'-213') [V-KAPPA (*Mus musculus*IGKV4-72*01 (85.3%) -IGKJ1*01 (90.9%)/*Homo sapiens*IGKV3D-11*02 (73.3%) -IGKJ4*01 (100%)) CDR-IMGT [5.3.9] (27-31.49-51.88-96) (1'-106') -*Homo sapiens*IGKC*01 (100%), Km3 A45.1 (152), V101 (190) (107'-213')]; dimer (230-230''-233-233'')-bisdisulfide, produced in a Chinese hamster ovary (CHO)-S cell line, glycoform alfa

divozilimab	immunoglobuline G1-kappa, anti-[<i>Homo sapiens</i> MS4A1 (membre 1 de la sous-famille A à 4 domaines transmembranaires, CD20)], anticorps monoclonal; chaîne lourde gamma1 (1-451) [VH (<i>Homo sapiens</i> IGHV1-46*01 (86.7%) - (IGHD) -IGHJ1*01 (86.7%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) - <i>Homo sapiens</i> IGHG1*03 (100%) G1m3, nG1m1 (CH1 R120 (218) (122-219), charnière 1-15 (220-234), CH2 (235-344), CH3 E12 (360), M14 (362) (345-449), CHS (450-451)) (122-451)], (224-213')-disulfure avec la chaîne légère kappa (1'-213') [V-KAPPA (<i>Mus musculus</i> IGKV4-72*01 (85.3%) -IGKJ1*01 (90.9%)/ <i>Homo sapiens</i> IGKV3D-11*02 (73.3%) - IGKJ4*01 (100%)) CDR-IMGT [5.3.9] (27-31.49-51.88-96) (1'-106') - <i>Homo sapiens</i> IGKC*01 (100%), Km3 A45.1 (152), V101 (190) (107'-213')]; dimère (230-230":233-233")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-S, glycoforme alfa
divozilimab	immunoglobulina G1-kappa, anti-[<i>Homo sapiens</i> MS4A1 (miembro 1 de la subfamilia A con 4 dominios transmembranarios, CD20)], anticuerpo monoclonal; cadena pesada gamma1 (1-451) [VH (<i>Homo sapiens</i> IGHV1-46*01 (86.7%) - (IGHD) -IGHJ1*01 (86.7%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) - <i>Homo sapiens</i> IGHG1*03 (100%) G1m3, nG1m1 (CH1 R120 (218) (122-219), bisagra 1-15 (220-234), CH2 (235-344), CH3 E12 (360), M14 (362) (345-449), CHS (450-451)) (122-451)], (224-213')-disulfuro con la cadena ligera kappa (1'-213') [V-KAPPA (<i>Mus musculus</i> IGKV4-72*01 (85.3%) -IGKJ1*01 (90.9%)/ <i>Homo sapiens</i> IGKV3D-11*02 (73.3%) - IGKJ4*01 (100%)) CDR-IMGT [5.3.9] (27-31.49-51.88-96) (1'-106') - <i>Homo sapiens</i> IGKC*01 (100%), Km3 A45.1 (152), V101 (190) (107'-213')]; dímero (230-230":233-233")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-S, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
EVQLVQPGAE VVKPGASVKV SCKASGYTFT SYNMHWVRQA PGRGLEWMGA 50
IYPGNGDTSY NQKFGRVTM TRDKSTSTVY MELSSLRSED TAVYYCARST 100
YYGDDWYFNV WQGLTLVTVS SASTKGPSVF PLAPSSKSTS GGTAALGCLV 150
KDYFPEPVTV SWNSGALTSG VHTFPAVLQS SGLYSLSSV TTPSSSLGTQ 200
TYICNVNHKP SNTKVDKRVK PKSCDKHTHC PPCPAPELLG GPSVFLFPPK 250
PKDITLMSRT PEVTCVVVDV SHEDPEVKFN WYVDGVEVHN AKTKPREEQY 300
NSTYRVVSVL TVLHQDWLNG KEYKCKVSNK ALPAPIEKT I SKAKGQPREP 350
QVYTLPPSRE EMTKNQVSLT CLVKGFPYPSD IAVEWESNGQ PENNYKTPPP 400
VLDSGGSFFL YSKLTVDKSR WQQGNVFSCS VMHEALHNHY TQKSLSLSPG 450
K 451

Light chain / Chaîne légère / Cadena ligera
QIVLSQSPAI LSASPGERVLT LTRASSSVS YIHWFQQKPG KAPKPLIYAT 50
SNLASGVPSR FSGSGSGTDF SLTISRVEPE DFAVYYCQQW TSNPPTFGGG 100
TKVEIKRTVA APSVFIFFPS DEQLKSGTAS VVCLLNNFYF REAKVQWKVD 150
NALQSGNSQE SVTEQDSKDS TYLSSTLT TL SKADYEKKV YACEVTHQGL 200
SSPVTKSFNR GEC 213

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 148-204 265-325 371-429
22"-96" 148"-204" 265"-325" 371"-429"
Intra-L (C23-C104) 23'-87' 133'-193'
23"-87" 133"-193"
Inter-H-L (h 5-CL 126) 224-213' 224"-213"
Inter-H-H (h 11, h 14) 230-230" 233-233"

N-terminal glutaminyl cyclization to pyroglutamyl (pE, 5-oxoprolyl)
L VL Q1:
I', I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
301, 301"
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal
H CHS K2:
451, 451"

ebronucimabum #

ebronucimab

immunoglobulin G1-lambda2, anti-[*Homo sapiens* PCSK9 (proprotein convertase subtilisin/kexin type 9, neural apoptosis-regulated convertase 1, NARC1, NARC-1, proprotein convertase 9, PC9)], *Homo sapiens* monoclonal antibody; gamma1 heavy chain *Homo sapiens* (1-453) [VH (*Homo sapiens* IGHV3-21*01 (91.8%) -(IGHD) - IGHJ3*01 (100%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123)-*Homo sapiens* IGHG1*01 (100%), G1m17,1 (CH1 K120 (220) (124-221), hinge 1-15 (222-236), CH2 (237-346), CH3 D12 (362), L14 (364) (347-451), CHS (452-453)) (124-453)], (226-216')-disulfide with lambda2 light chain *Homo sapiens* (1'-217') [V-LAMBDA (*Homo sapiens* IGLV2-11*01 (81.6%) -IGLJ1*01 (90.9%)) CDR-IMGT [9.3.11] (26-34.52-54.91-101) (1'-111') -*Homo sapiens* IGLC2*01 (100%) (112'-217')]; dimer (232-232":235-235")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

ébronucimab

immunoglobuline G1-lambda2, anti-[*Homo sapiens* PCSK9 (proprotéine convertase subtilisine/kexine type 9, convertase 1 régulée par l'apoptose neuronale, NARC1, NARC-1, proprotéine convertase 9, PC9)], anticorps monoclonal *Homo sapiens*; chaîne lourde gamma1 *Homo sapiens* (1-453) [VH (*Homo sapiens* IGHV3-21*01 (91.8%) -(IGHD) - IGHJ3*01 (100%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123)-*Homo sapiens* IGHG1*01 (100%), G1m17,1 (CH1 K120 (220) (124-221), charnière 1-15 (222-236), CH2 (237-346), CH3 D12 (362), L14 (364) (347-451), CHS (452-453)) (124-453)], (226-216')-disulfure avec la chaîne légère lambda2 *Homo sapiens* (1'-217') [V-LAMBDA (*Homo sapiens* IGLV2-11*01 (81.6%) -IGLJ1*01 (90.9%)) CDR-IMGT [9.3.11] (26-34.52-54.91-101) (1'-111') -*Homo sapiens* IGLC2*01 (100%) (112'-217')]; dimère (232-232":235-235")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

ebronucimab

inmunoglobulina G1-lambda2, anti-[*Homo sapiens* PCSK9 (proproteína convertasa subtilisina/kexina tipo 9, convertasa 1 regulada por la apoptosis neuronal, NARC1, NARC-1, proproteína convertasa 9, PC9)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma1 *Homo sapiens* (1-453) [VH (*Homo sapiens* IGHV3-21*01 (91.8%) -(IGHD) - IGHJ3*01 (100%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123)-*Homo sapiens* IGHG1*01 (100%), G1m17,1 (CH1 K120 (220) (124-221), bisagra 1-15 (222-236), CH2 (237-346), CH3 D12 (362), L14 (364) (347-451), CHS (452-453)) (124-453)], (226-216')-disulfuro con la cadena ligera lambda2 *Homo sapiens* (1'-217') [V-LAMBDA (*Homo sapiens* IGLV2-11*01 (81.6%) -IGLJ1*01 (90.9%)) CDR-IMGT [9.3.11] (26-34.52-54.91-101) (1'-111') -*Homo sapiens* IGLC2*01 (100%) (112'-217')]; dímero (232-232":235-235")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
EVQLVESGGG LVQPGRSLRL SCAASGFTFS SYSMNWVROA PGKLEWVSG 5C
ISSSSSYISY ADSVQGRFTI SPDNGKNSLY LQMNSLRAED TALYFCAREY 100
DFWSAYYDAF DVWGQGTMTV VSSASTKGPS VFELAPSSKS TSGGTAALGC 150
LVKDYFPEPV TVSWNSGALT SGVHTFPAYL QSSGLYSLSS VVTVPSSSLG 200
TQTYICNVNH KPSNTKVDK VEPKSCDKTH TCPPCPAPEL LGGPSTFLEP 250
FKPKDTLMIS RTPEVTCVVV DVSHEDEPEK ENWYVDGVEV HNAKTKPREE 300
QYNSTYRVVS VLTVLHQDWL NGKEYCKVKS NKALFAPIEK TISKAKGQPR 350
EPQVYTLFPS RDELTKNQVS LTCVLKGTFP SDIAVENESN GQPENNYKTT 400
PFVLDSGDSF FLYSKLTVDK SRWQQGVFES CSVMHEALHN HYTKKSLSL 450
PGK 453

Light chain / Chaîne légère / Cadena ligera
QSELTQPRSV SGSPGQSVTI SCTGTSRNIG GGNVDVHWYQQ HPGKAPKLLI 5C
SGVIERSSGV PDRFSGSKSG NTASLTISGL QAEDADYYC QSFQDGLSGS 100
VEGTGTDVTV LGQPKAAPSV TLFPPSSEEL QANKATLVCL ISDFYPGAVT 150
VAWKAADSSPV KAGVETTPPS KQSNKKTAAS SYLSLTPEQW KSHRSYSCQV 200
THEGSTVEKT VAPTECS 217

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22°-96' 150°-206' 267°-327' 373°-431"
22°-96' 150°-206' 267°-327' 373°-431"
Intra-L (C23-C104) 22°-90' 139°-198'
22°-90' 139°-198'
Inter-H-L (h5-CL126) 226°-216' 226°-216"
Inter-H-H (h11, h14) 232°-232' 235°-235"

N-terminal glutaminyl cyclization to pyroglutamyI (pE, 5-oxopropyl)
L VL Q1:
I', I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2N84,4:
303, 303"
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires
complexes fucosylés / glicanos de tipo CHO biantennarios complejos fucosilados.

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal
H CHSK2:
453, 453"

edralbrutinibum
edralbrutinib

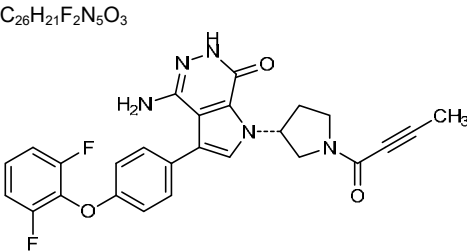
4-amino-1-[(3*R*)-1-(but-2-ynoyl)pyrrolidin-3-yl]-3-[4-(2,6-difluorophenoxy)phenyl]-1,6-dihydro-7*H*-pyrrolo[2,3-*d*]pyridazin-7-one

édralbrutinib

4-amino-1-[(3*R*)-1-(but-2-ynoyl)pyrrolidin-3-yl]-3-[4-(2,6-difluorophénoxy)phényl]-1,6-dihydro-7*H*-pyrrolo[2,3-*d*]pyridazin-7-one

edralbrutinib

4-amino-1-[(3*R*)-1-(but-2-inoil)pirrolidin-3-il]-3-[4-(2,6-difluorofenoxi)fenil]-1,6-dihidro-7*H*-pirrolo[2,3-*d*]piridazin-7-ona



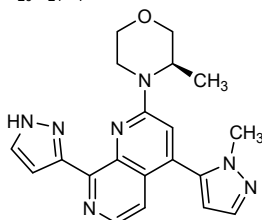
elimusertibum
elimusertib

2-[(3*R*)-3-methylmorpholin-4-yl]-4-(1-methyl-1*H*-pyrazol-5-yl)-8-(1*H*-pyrazol-3-yl)-1,7-naphthyridine

élimusertib

2-[(3*R*)-3-méthylmorpholin-4-yl]-4-(1-méthyl-1*H*-pyrazol-5-yl)-8-(1*H*-pyrazol-3-yl)-1,7-naphthyridine

elimusertib

2-[(3*R*)-3-metilmorfolin-4-il]-4-(1-metil-1*H*-pirazol-5-il)-8-(1*H*-pirazol-3-il)-1,7-naftiridina $C_{20}H_{21}N_7O$ 

eluvixtamabum #

eluvixtamab

immunoglobulin scFv-scFv, anti-[*Homo sapiens* CD33 (sialic acid binding Ig-like lectin 3, SIGLEC3, SIGLEC-3, gp67, p67)] and anti-[*Homo sapiens* CD3E (CD3 epsilon)], monoclonal antibody single chain, bispecific; IG scFv-scFv single chain, anti-CD33 and anti-CD3E (1-511) [scFv-VH-V-kappa anti-CD33 (1-250) [VH (*Homo sapiens* IGHV1-18*01 (82.7%) - (IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122)-15-mer tris(tetraglycyl-seryl) linker (123-137) -V-KAPPA (*Homo sapiens* IGKV4-1*01 (83.2%) -IGKJ5*01 (100%)) CDR-IMGT [12.3.9] (164-175.193-195.232-240) (138-250)] -6-mer seryl-tetraglycyl-seryl linker (251-256) -scFv-VH-V-lambda anti-CD3E (257-505) [VH (*Mus musculus* IGHV10-1*02 (91.9%) - (IGHD) -IGHJ3*01 (86.7%))/*Homo sapiens* IGHV3-73*01 (87%) - (IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (282-289.307-316.355-370) (257-381) -15-mer tris(tetraglycyl-seryl) linker (382-396) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (422-430.448-450.487-495) (397-505)] - hexahistidine (506-511)], non-glycosylated, produced in Chinese hamster ovary (CHO) cells

éluvixtamab

immunoglobuline scFv-scFv, anti-[*Homo sapiens* CD33 (lectine 3 de type Ig-like liant l'acide sialique, SIGLEC3, SIGLEC-3, gp67, p67)] et anti-[*Homo sapiens* CD3E (CD3 epsilon)], anticorps monoclonal à chaîne unique, bispécifique; IG scFv-scFv à chaîne unique, anti-CD33 et anti-CD3E (1-511) [scFv-VH-V-kappa anti-CD33 (1-250) [VH (*Homo sapiens* IGHV1-18*01 (82.7%) - (IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122)-15-mer tris(tétraglycyl-séryl) linker (123-137) -V-KAPPA (*Homo sapiens* IGKV4-1*01 (83.2%) -IGKJ5*01 (100%)) CDR-IMGT [12.3.9] (164-175.193-195.232-240) (138-250)] -6-mer séryl-tétraglycyl-séryl linker (251-256) -scFv-VH-V-lambda anti-CD3E (257-505) [VH (*Mus musculus* IGHV10-1*02 (91.9%) - (IGHD) -IGHJ3*01 (86.7%))/*Homo sapiens* IGHV3-73*01 (87%) - (IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (282-289.307-316.355-370) (257-381) -15-mer tris(tétraglycyl-séryl) linker (382-396) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (422-430.448-450.487-495) (397-505)] - hexahistidine (506-511)], non-glycosylé, produit dans des cellules ovariennes de hamster chinois (CHO)

eluvixtamab

immunoglobulina scFv-scFv, anti-[*Homo sapiens* CD33 (lectina 3 de tipo Ig-like aglutinando el ácido siálico, SIGLEC3, SIGLEC-3, gp67, p67)] y anti-[*Homo sapiens* CD3E (CD3 épsilon)], anticuerpo monoclonal con cadena única, biespecífico;

IG scFv-scFv con cadena única, anti-CD33 y anti-CD3E (1-511) [scFv-VH-V-kappa anti-CD33 (1-250) [VH (*Homo sapiens* IGHV1-18*01 (82.7%) -(IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122)-15-mer tris(tetraglicil-seril) linker (123-137) -V-KAPPA (*Homo sapiens* IGKV4-1*01 (83.2%) -IGKJ5*01 (100%)) CDR-IMGT [12.3.9] (164-175.193-195.232-240) (138-250)] -6-mer seril-tetraglicil-seril linker (251-256) -scFv-VH-V-lambda anti-CD3E (257-505) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (282-289.307-316.355-370) (257-381) -15-mer tris(tetraglicil-seril) linker (382-396) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%) CDR-IMGT [9.3.9] (422-430.448-450.487-495) (397-505)] -hexahistidina (506-511)], no glicosilado, producido en las células ováricas de hamster chino (CHO)

Heavy chain / Chaîne lourde / Cadena pesada
QVQLVQSGAE VKKPGESVKV SCKASGYTFT NYGMNWKQA PGQGLEWKGW 50
INTYTGEPY ADKFQGRVTM TTDSTSTAY MEIRNLGGDD TAVYYCARWS 10C
NSDGYVYVFD YAGQQTSTVT SSGGGGSGGG GSGGGGSD-V MTQSPDSLTV 15C
SLGERTTINC KSSQSVLQSS TNKNSLAWYQ QKPGQPPKLL LSWASTRESG 200
IPDRFSGSGS GTDFTLTIDS PQPDSATYY CQQAHPFPIIT FGQGTRELIK 250
SGGGGSEVQL VESGGGLVQF GGSLLKSCAA SGFTFNKYAM NNVROAPKRG 30C
LEWVARIRSK YNNYATYYAL SVKDRFTISR DDSKNTAYLQ MNNLKIEDTA 35C
VYYCVRHGNF GNSYISYWAY WQQTTLTVS SGGGGSGGGG SGGGGSQTVV 40C
TQEPSTLYSP GGTVTLTGGS STGAVTSGNY PNWVQKPGQ APRGLIGGK 45C
FLAPGTPARF SGLLGKAA LTLGVPQPED EAEYYCYLNY SNRWVFGGT 50C
KLTVLHRRHH H 511

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-V (C23C104) 22-96 160-231 278-354 418-486

N-terminal glutaminyl cyclization to pyroglutamyI (pE, 5-oxoprolyl)
VH Q1: I

No N-glycosylation sites / pas de sites de N-glycosylation / ningún posición de N-glicosilación
Aglycosylated

emerfetamabum #
emerfetamab

immunoglobulin scFv-scFv-scFc, anti-[*Homo sapiens* CD33 (sialic acid binding Ig-like lectin 3, SIGLEC3, SIGLEC-3, gp67, p67)] and anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], monoclonal antibody single chain (scFv)2-scFc, bispecific;
IG scFv-scFv-scFc single chain, anti-CD33 and anti-CD3E (1-993) [scFv-VH-V-kappa anti-CD33 (1-250) [VH (*Homo sapiens* IGHV1-18*01 G49>C (44) (81.6%) -(IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -15-mer tris(tetraglycyl-seryl) linker (123-137) -V-KAPPA (*Homo sapiens* IGKV4-1*01 (83.2%) -IGKJ5*01 Q120>C (243) (91.7%)) CDR-IMGT [12.3.9] (164-175.193-195.232-240) (138-250)] -6-mer seryl-tetraglycyl-seryl linker (251-256) -scFv-VH-V-lambda anti-CD3E (257-505) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (282-289.307-316.355-370) (257-381) -15-mer tris(tetraglycyl-seryl) linker (382-396) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (422-430.448-450.487-495) (397-505)] -4-mer tetraglycyl linker (506-509) -scFc (h-CH2-CH3)-(h-CH2-CH3) (510-993) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (hinge 6-15 (510-519), CH2 R83>C (581), N84.4>G (586), V85>C (591) (520-629), CH3 E12 (645), M14 (647) (630-734), CHS (735-736)) (510-736) -30-mer hexakis(tetraglycyl-seryl) linker (737-766) -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (hinge 6-15 (767-776), CH2 R83>C (838), N84.4>G (843), V85>C (848) (777-886), CH3 E12 (902), M14 (904) (887-991), CHS (992-993)) (767-993)], non-glycosylated, produced in Chinese hamster ovary (CHO) cells

émérfétamab

immunoglobuline scFv-scFv-scFc, anti-[*Homo sapiens* CD33 (lectine 3 de type Ig-like liant l'acide sialique, SIGLEC3, SIGLEC-3, gp67, p67)], et anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], anticorps monoclonal à chaîne unique (scFv)₂-scFc, bispécifique;

IG scFv-scFv-scFc chaîne unique, anti-CD33 et anti-CD3E (1-993) [scFv-VH-V-kappa anti-CD33 (1-250) [VH (*Homo sapiens*IGHV1-18*01 G49>C (44) (81.6%) -(IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -15-mer-tris(tétraglycyl-séryl) linker (123-137) -V-KAPPA (*Homo sapiens* IGKV4-1*01 (83.2%) -IGKJ5*01 Q120>C (243) (91.7%)) CDR-IMGT [12.3.9] (164-175.193-195.232-240) (138-250)] -6-mer séryl-tétraglycyl-séryl linker (251-256) -scFv-VH-V-lambda anti-CD3E (257-505) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (282-289.307-316.355-370) (257-381) -15-mer tris(tétraglycyl-séryl) linker (382-396) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (422-430.448-450.487-495) (397-505)] -4-mer-tétraglycyl linker (506-509) -scFc (h-CH2-CH3)-(h-CH2-CH3) (510-993) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (charnière 6-15 (510-519), CH2 R83>C (581), N84.4>G (586), V85>C (591) (520-629), CH3 E12 (645), M14 (647) (630-734), CHS (735-736)) (510-736) -30-mer hexakis(tétraglycyl-séryl) linker (737-766) -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (charnière 6-15 (767-776), CH2 R83>C (838), N84.4>G (843), V85>C (848) (777-886), CH3 E12 (902), M14 (904) (887-991), CHS (992-993)) (767-993)]], non-glycosylé, produit dans des cellules ovariennes de hamster chinois (CHO)

emerfetamab

inmunoglobulina scFv-scFv-scFc, anti-[*Homo sapiens* CD33 (lectina 3 de tipo Ig-like aglutinando el ácido siálico, SIGLEC3, SIGLEC-3, gp67, p67)], y anti-[*Homo sapiens* CD3E (CD3 épsilon, Leu-4)], anticuerpo monoclonal con cadena única (scFv)₂-scFc, biespecífico;

IG scFv-scFv-scFc cadena única, anti-CD33 y anti-CD3E (1-993) [scFv-VH-V-kappa anti-CD33 (1-250) [VH (*Homo sapiens* IGHV1-18*01 G49>C (44) (81.6%) -(IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -15-mer-tris(tetraglicil-seril) linker (123-137) -V-KAPPA (*Homo sapiens* IGKV4-1*01 (83.2%) -IGKJ5*01 Q120>C (243) (91.7%)) CDR-IMGT [12.3.9] (164-175.193-195.232-240) (138-250)] -6-mer seril-tetraglicil-seril linker (251-256) -scFv-VH-V-lambda anti-CD3E (257-505) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (282-289.307-316.355-370) (257-381) -15-mer tris(tetraglicil-seril) linker (382-396) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (422-430.448-450.487-495) (397-505)] -4-mer-tetraglicil linker (506-509) -scFc (h-CH2-CH3)-(h-CH2-CH3) (510-993) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (bisagra 6-15 (767-776), CH2 R83>C (838), N84.4>G (843), V85>C (848) (777-886), CH3 E12 (902), M14 (904) (887-991), CHS (992-993)) (767-993)]], no glicosilado, producido en las células ováricas de hamster chino (CHO)

Heavy chain / Chaîne lourde / Cadena pesada
QVQLVQSGAE VKKFGESYEV SKKASGITFT NYGMNWKQA PGQCLEWNGW 50
INTITGEPFY ADKFQGRVTM TTDSTSTAY MEIRNLGGDD TAVYICARMS 100
NSDGYVYIFD YWQGGTSYTV SSGGGGSGGG GSGGGGSDIV MTQSPDSLTV 150
SLGERITING KSSQSVLDS TNRNSLAWIQ QKFGQFPKLL LSWASTRESG 200
IDRFSGSGG CDPFLTIDS PQEDSATYY CQQAHPFIT FCGTRLEIK 250
SGGGGSEYQL VESGGGLVQF GSKLGLSCA SGTENKYRM NWVRQAFKGG 300
LEWVARIRSK YNNYATYYAD SVKDRFTISR DSKNTAYLQ MNHLKIEVTA 350
VYVCVRHGNF GNSYISYWAY MGQCTLVTVS SGGGGSGGGG SGGGSGTVV 400
TQEPSLTYSF GGTVTLTGGG STGAVTSGNY FNNWQKFGQ APFGLIGCTK 450
FLAPGTPARF SGLLGGKAA LTLGGVQPED EAEYCVLWY SNRWVFGGT 500
KLTVLGGGGC KTHTCPFPCA PELLGGPSVF LKPKPKDIL MISRTPEVTC 550
VVDVDSHEDP EVKFNWYVDG VETHNAKTKP CEEQYGSTYR CVSVLTVCAC 600
DWLNGREYKC KVSNKALPAP IEKTISKARG QPREPOVITL PPSREMTKN 650
QVSLTCLYKG FYPSDIAVEW ESNQGPENNY KTTTPVLDSD GSFFLYSKIT 700
VDKSRWQGN VFSQSVWHEA LHNHYTKSL SLSPGKGGGG SGGGGSGGG 750
SGGGSGGGG SGGGGGSKTH TCPFCAPEL LGGPSVFLP PKPKDTLMIS 800
RTPEVTCVTV DYSHEDPEVK FNNWYVDGEV HNAKTKPCRE QYGYTRCVS 850
VLTVLHQWL NGKEYCKVS NKALPAPIEK TISKAKGQPR EPOVYTL?PS 900
REEMTKNQVS LTCVLKGFYP SDIAVEWESN GQPENNYKTT PPVLDSDGSGF 950
FLYSKLTVDK SRWQGNVFS CSMVHEALHN HYTKSLSLS PGK 993

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-V (C23-C104) 22-96 160-231 278-354 418-486
Intra-C (C23-C104) 550-610 656-714 807-867 913-971
Intra-CH2 (C83-C85) 581-591 838-848
Inter-VH-VL (C49-C120) 44-243
Inter-h11 (h 11 - h 11) 515-772
Inter-h14 (h 14 - h 14) 518-775

N-terminal glutaminylation cyclization to pyroglutamylation (pE, 5-oxoproline)
VH Q1:
I

No N-glycosylation sites / pas de sites de N-glycosylation / ningún posición de N-glicosilación
CH2 N84.4>G:
586, 843
Aglycosylated

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal
CHS K2:
993

enibarcimabum #
enibarcimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* ADM (adrenomedullin)], monoclonal antibody;
gamma1 heavy chain (1-448) [VH (*Homo sapiens* IGHV1-69*01 (86.5%) - (IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens* IGHG1*01 (100%) G1m17.1 (CH1 K120 (215) (119-216), hinge 1-15 (217-231), CH2 (232-341), CH3 D12 (357), L14 (359) (342-446), CHS (447-448)) (119-448)], (221-219')-disulfide with kappa light chain (1'-219') [V-KAPPA (*Mus musculus* IGKV1-117*01 (88.0%) -IGKJ2*01 (100%)/*Homo sapiens* IGKV2-30*01 (88.0%) -IGKJ2*01 (91.7%)) CDR-IMGT [11.3.9] (27-37.55-57.94-102) (1'-112') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (158), V101 (196) (113'-219')];
dimer (227-227'':230-230'')-bisdisulfide, produced in Chinese hamster ovary (CHO)-DG44 cell line, glycoform alfa

énibarcimab

immunoglobuline G1-kappa, anti-[*Homo sapiens* ADM (adrénomédulline)], anticorps monoclonal;
chaîne lourde gamma1 (1-448) [VH (*Homo sapiens* IGHV1-69*01 (86.5%) - (IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens* IGHG1*01 (100%) G1m17.1 (CH1 K120 (215) (119-216), charnière 1-15 (217-231), CH2 (232-341), CH3 D12 (357), L14 (359) (342-446), CHS (447-448)) (119-448)], (221-219')-disulfure avec la chaîne légère kappa (1'-219') [V-KAPPA (*Mus musculus* IGKV1-117*01 (88.0%) -IGKJ2*01 (100%)/*Homo sapiens* IGKV2-30*01 (88.0%) -IGKJ2*01 (91.7%)) CDR-IMGT [11.3.9] (27-37.55-57.94-102) (1'-112') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (158), V101 (196) (113'-219')];
dimère (227-227'':230-230'')-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-DG44, glycoforme alfa

enibarcimab

inmunoglobulina G1-kappa, anti-[*Homo sapiens* ADM (adrenomedulina)], anticuerpo monoclonal; cadena pesada gamma1 (1-448) [VH (*Homo sapiens* IGHV1-69*01 (86.5%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (215) (119-216), bisagra 1-15 (217-231), CH2 (232-341), CH3 D12 (357), L14 (359) (342-446), CHS (447-448)) (119-448)], (221-219')-disulfuro con la cadena ligera kappa (1'-219') [V-KAPPA (*Mus musculus* IGKV1-117*01 (88.0%) -IGKJ2*01 (100%)/*Homo sapiens* IGKV2-30*01 (88.0%) -IGKJ2*01 (91.7%)) CDR-IMGT [11.3.9] (27-37.55-57.94-102) (1'-112') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (158), V101 (196) (113'-219')]; dímero (227-227"-230-230")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-DG44, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
 QVQLVQSGAE VKKPGSSVKV SCKASGYTFS RYIEWVRQA PGQGLEWIGE 50
 ILPGSGSTNY NQKFGGRVTI TADTSTSTAY MELSSLRSED TAVYYCTEGY 100
 EYDGFYWGQ GTTVTVSSAS TKGPSVFPLA PSSKSTSGGT AALGCLVKDY 150
 FPEPVTYSWN SGALTSGVHT FPAVLQSSGL YSLSSVTVTP SSSLGTQTYI 200
 CNVNHKPSNT KVDKKVEPKS CDKTHTCPPC PAPELLGGPS VFLFPPKPKD 250
 TLMISRTPEV TCVVVDVSHE DPEVKFNWYV DGVEVHNAKT KPREEQYNST 300
 YRVVSVLTVL HQDWLNGKEY KCKVSNKALP APIEKTISKA KGQPREPQVY 350
 TLPPSRDELT KNQVSLTCLV KGFYPSDIAV EWESNGQPEN NYKTTPPVLD 400
 SDGSFFFLYSK LTVDKSRWQQ GNVFSCSVMH EALHNHYTQK SLSLSPGK 448

Light chain / Chaîne légère / Cadena ligera
 DVVLQTQSPLS LPVTILGPAS ISCRSSQSIV YSNGNTYLEW YLQRPQGSPR 50
 LLITYRVSNRF SGVPDRFSGS GSGTDFTLKI SRVEAEDVG Y YCFQGSHP 100
 YTFGGGKTLK IKRTVAAPSV FIFPPSDEQL KSGTASVCL LNNFYPREAK 150
 VQWVKVDNALQ SGNSQESVTE QDSKDSYSL SSTLTLSKAD YEKHKVYACE 200
 VTHQGLSSPV TKSFNREGC 219

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 145-201 262-322 368-426
 22'-96" 145"-201" 262"-322" 368"-426"

Intra-L (C23-C104) 23'-93' 139'-199"
 23"-93" 139"-199"

Inter-H-L (h 5-CL 126) 221-219' 221"-219"

Inter-H-H (h 11, h 14) 227-227" 230-230"

N-terminal glutaminyl cyclization to pyroglutamyl (pE, 5-oxoprolyl)

H VH Q1:

1, 1"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

298, 298"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantennarios complejos fucosilados.

enomimeranum

enomimeran

Messenger RNA encoding melanoma-associated antigen 3 (MAGE-A3).

Messenger RNA (mRNA), 5'-capped, encoding codon-optimised melanoma-associated antigen 3 (MAGE-A3, cancer/testis antigen 1.3) expressed as a fusion protein comprising a secretory signal peptide, MAGE-A3, P2P16 tetanus toxoid-derived helper epitopes and a major histocompatibility complex (MHC) class I transmembrane and cytoplasmic domain (MITD), connected by three glycine/serine-rich (GS) linker peptides, flanked by 5' and 3' untranslated regions and a 3' poly(A) tail.

énomiméran

ARN messenger codant pour l'antigène 3 associé au mélanome (MAGE-A3).

	Un ARN messager (ARNm), protégé en 5', codant pour l'antigène 3 associé au mélanome (MAGE-A3, antigène 1.3 du cancer testiculaire) , avec des codons optimisés, exprimé sous la forme d'une protéine de fusion comprenant un peptide signal de sécrétion, MAGE-A3, les épitopes auxiliaires P2P16 dérivés de l'anatoxine tétanique et un domaine transmembranaire et cytoplasmique (MITD) du complexe majeur d'histocompatibilité (CMH) de classe I, reliés par trois peptides de liaison riches en glycine/sérine (GS), flanqués de régions non traduites en 5' et 3' et d'une queue poly(A) en 3'.
enomimerán	ARN mensajero que codifica para el antígeno asociado a melanoma 3 (MAGE-A3). ARN mensajero (ARNm), protegido en 5', que codifica para el antígeno asociado a melanoma 3 (MAGE-A3, antígeno de cáncer/testículo 1.3), con codones optimizados, expresado como una proteína de fusión que consta de un péptido señal de secreción, MAGE-A3, los epítomos auxiliares P2P16 derivados del toxoide tetánico y un dominio citoplásmico y transmembrana del complejo principal de histocompatibilidad (MHC) de clase I, conectados mediante tres péptidos de enlace ricos en glicina/serina (GS), flanqueado por las regiones 5' y 3' no traducidas y una cola poly(A) en 3'.
entacingenum turiparvovecum #	
entacingene turiparvovec	A replication-defective adeno-associated viral vector encoding cone photoreceptor cyclic nucleotide-gated channel $\beta 3$ (CNGB3). A recombinant non-replicating adeno-associated viral vector serotype 8 (rAAV8), encoding codon-optimised human cone photoreceptor cyclic nucleotide gated channel $\beta 3$ (CNGB3), under the control of the human cone arrestin (hCAR) promoter and an SV40 poly(A) sequence, flanked by AAV2 inverted terminal repeats (ITR).
entacingène turiparvovec	Un vecteur adénoviral à réplication défectueuse codant pour le canal du photorécepteur conique $\beta 3$ (CNGB3), dépendant des nucléotides cycliques. Un vecteur adénoviral recombinant non réplcatif de sérotype 8 (rAAV8), codant pour le canal du photorécepteur du cône humain $\beta 3$ (CNGB3), dépendant des nucléotides cycliques, avec des codons optimisés, sous le contrôle du promoteur hCAR (arrestine du cône humain) et d'une séquence poly(A) SV40, flanquée de répétitions terminales inversées (ITR) AAV2.
entacingén turiparvovec	Un vector de virus adeno-asociado, deficiente de replicación, que codifica para el canal $\beta 3$ regulado por nucleótidos cíclicos (CNGB3) del fotorreceptor de los conos. Un vector de virus adeno-asociado recombinante serotipo 8 (rAAV8), deficiente de replicación, que codifica para el canal $\beta 3$ regulado por nucleótidos cíclicos (CNGB3) del fotorreceptor de los conos, con codones optimizados, bajo el control del promotor de la arrestina de cono humana (hCAR) y una secuencia poly(A) de SV40, flanqueado por las repeticiones invertidas terminales (ITR) del AAV2.

eplontersenum

eplontersen

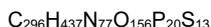
all-P-ambo-5'-O-(28-[(2-acetamido-2-deoxy-β-D-galactopyranosyl)oxy]-16,16-bis[[3-({6-[(2-acetamido-2-deoxy-β-D-galactopyranosyl)oxy]hexyl)amino}-3-oxopropoxy]methyl]-1-hydroxy-1,10,14,21-tetraoxo-2,18-dioxo-9,15,22-triaza-1λ⁶-phosphaoctacosan-1-yl)-2'-O-(2-methoxyethyl)-5-methyl-P-thiouridylyl-(3'→5')-2'-O-(2-methoxyethyl)-5-methylcytidylyl-(3'→5')-2'-O-(2-methoxyethyl)-5-methyluridylyl-(3'→5')-2'-O-(2-methoxyethyl)-5-methyluridylyl-(3'→5')-2'-O-(2-methoxyethyl)guanylyl-(3'→5')-2'-deoxy-P-thioguanilyl-(3'→5')-P-thiothymidylyl-(3'→5')-P-thiothymidylyl-(3'→5')-2'-deoxy-P-thioadenilyl-(3'→5')-2'-deoxy-5-methyl-P-thiocytidylyl-(3'→5')-2'-deoxy-P-thioadenilyl-(3'→5')-P-thiothymidylyl-(3'→5')-2'-deoxy-P-thioguanilyl-(3'→5')-2'-deoxy-P-thioadenilyl-(3'→5')-2'-deoxy-P-thioadenilyl-(3'→5')-2'-O-(2-methoxyethyl)adenilyl-(3'→5')-2'-O-(2-methoxyethyl)-5-methyluridylyl-(3'→5')-2'-O-(2-methoxyethyl)-5-methyl-P-thiocytidylyl-(3'→5')-2'-O-(2-methoxyethyl)-5-methyl-P-thiocytidylyl-(3'→5')-2'-O-(2-methoxyethyl)-5-methylcytidine

éplontersen

tout-P-ambo-5'-O-(28-[(2-acétamido-2-désoxy-β-D-galactopyranosyl)oxy]-16,16-bis[[3-({6-[(2-acétamido-2-désoxy-β-D-galactopyranosyl)oxy]hexyl)-amino}-3-oxopropoxy]méthyl]-1-hydroxy-1,10,14,21-tétraoxo-2,18-dioxo-9,15,22-triaza-1λ⁶-phosphaoctacosan-1-yl)-2'-O-(2-méthoxyéthyl)-5-méthyl-P-thiouridylyl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthylcytidylyl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthyluridylyl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthyluridylyl-(3'→5')-2'-O-(2-méthoxyéthyl)guanylyl-(3'→5')-2'-désoxy-P-thioguanilyl-(3'→5')-P-thiothymidylyl-(3'→5')-P-thiothymidylyl-(3'→5')-2'-désoxy-P-thioadénylyl-(3'→5')-2'-désoxy-5-méthyl-P-thiocytidylyl-(3'→5')-2'-désoxy-P-thioadénylyl-(3'→5')-P-thiothymidylyl-(3'→5')-2'-désoxy-P-thioguanilyl-(3'→5')-2'-désoxy-P-thioadénylyl-(3'→5')-2'-désoxy-P-thioadénylyl-(3'→5')-2'-O-(2-méthoxyéthyl)adénylyl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthyluridylyl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthyl-P-thiocytidylyl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthyl-P-thiocytidylyl-(3'→5')-2'-O-(2-méthoxyéthyl)-5-méthylcytidine

eplontersén

todo-P-ambo-5'-O-(28-[(2-acetamido-2-desoxi-β-D-galactopiranosil)oxi]-16,16-bis[[3-({6-[(2-acetamido-2-desoxi-β-D-galactopiranosil)oxi]hexil)amino}-3-oxopropoxi]metil]-1-hidroxi-1,10,14,21-tetraoxo-2,18-dioxo-9,15,22-triaza-1λ⁶-fosfaoctacosan-1-il)-2'-O-(2-metoxietil)-5-metil-P-tiuridilil-(3'→5')-2'-O-(2-metoxietil)-5-metiluridilil-(3'→5')-2'-O-(2-metoxietil)-5-metiluridilil-(3'→5')-2'-O-(2-metoxietil)guanilil-(3'→5')-2'-desoxi-P-tioguanilil-(3'→5')-P-tiotimidilil-(3'→5')-P-tiotimidilil-(3'→5')-2'-desoxi-P-tioadenilil-(3'→5')-2'-desoxi-5-metil-P-tiocitidilil-(3'→5')-2'-desoxi-P-tioadenilil-(3'→5')-P-tiotimidilil-(3'→5')-2'-desoxi-P-tioguanilil-(3'→5')-2'-desoxi-P-tioadenilil-(3'→5')-2'-desoxi-P-tioadenilil-(3'→5')-2'-O-(2-metoxietil)adenilil-(3'→5')-2'-O-(2-metoxietil)-5-metiluridilil-(3'→5')-2'-O-(2-metoxietil)-5-metil-P-tiocitidilil-(3'→5')-2'-O-(2-metoxietil)-5-metil-P-tiocitidilil-(3'→5')-2'-O-(2-metoxietil)-5-metilcytidina



(3'-5') R1 -U=C-U-U-G-d(G=T=A=C=A=T=G=A=A=)A-U-C=C=C

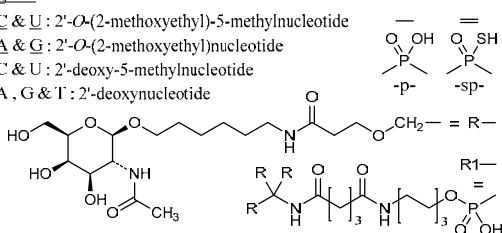
Legend:

C & U : 2'-O-(2-methoxyethyl)-5-methylnucleotide

A & G : 2'-O-(2-methoxyethyl)nucleotide

C & U : 2'-deoxy-5-methylnucleotide

A, G & T : 2'-deoxynucleotide



eprenetapoptum

eprenetapopt

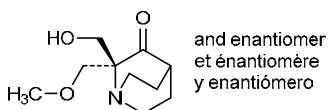
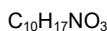
rac-(2*R*)-2-(hydroxymethyl)-2-(methoxymethyl)-1-azabicyclo[2.2.2]octan-3-one

éprénétapopt

rac-(2*R*)-2-(hydroxyméthyl)-2-(méthoxyméthyl)-1-azabicyclo[2.2.2]octan-3-one

eprenetapopt

rac-(2*R*)-2-(hidroximetil)-2-(metoximetil)-1-azabicyclo[2.2.2]octan-3-ona



etevritamabum #

etevritamab

immunoglobulin scFv-scFv, anti-[*Homo sapiens* EGFR (epidermal growth factor receptor, receptor tyrosine-protein kinase erbB-1, ERBB1, HER1, HER-1, ERBB) variant III (EGFRvIII)] and anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], monoclonal antibody single chain, bispecific;

IG scFv-scFv single chain (1-512) [scFv-VH-V-kappa anti-EGFRvIII(1-251)] [VH (*Homo sapiens* IGHV3-33*01 G49>C (44) (93.9%) -IGHD) -IGHJ4*01 (100%)] CDR-IMGT [8.8.17] (26-33.51-58.97-113) (1-124) -15-mer tris(tetraglycyl-seryl) linker (125-139) -V-KAPPA (*Homo sapiens* IGKV2-24*01 (92.0%) -IGKJ1*01 Q120>C (244) (90.9%)] CDR-IMGT [11.3.9] (166-176.194-196.233-241) (140-251)] -6-mer seryl-tetraglycyl-seryl linker (252-257) -scFv-VH-V-lambda anti-CD3E (258-506) [VH (*Mus musculus* IGHV10-1*02 (91.9%)-(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87.0%)-(IGHD) -IGHJ5*01 (100%)] [8.10.16] (258-382) -15-mer tris(tetraglycyl-seryl) linker (383-397) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)] CDR-IMGT [9.3.9] (423-431.449-451.488-496) (398-506)] -hexahistidine (507-512)], non-glycosylated, produced in Chinese hamster ovary (CHO) cells

étévritamab

immunoglobuline scFv-scFv, anti-[*Homo sapiens* EGFR (récepteur du facteur de croissance épidermique, récepteur tyrosine-protéine kinase erb-1, ERBB1, HER1, HER-1, ERBB) variant III (EGFRvIII)] et anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], anticorps monoclonal à chaîne unique, bispécifique;
IG scFv-scFv chaîne unique (1-512) [scFv-VH-V-kappa anti-EGFRvIII(1-251) [VH (*Homo sapiens* IGHV3-33*01 G49>C (44) (93.9%) -IGHD) -IGHJ4*01 (100%)] CDR-IMGT [8.8.17] (26-33.51-58.97-113) (1-124) -15-mer tris(tétraglycyl-séryl) linker (125-139) – V-KAPPA (*Homo sapiens* IGKV2-24*01 (92.0%) -IGKJ1*01 Q120>C (244) (90.9%)] CDR-IMGT [11.3.9] (166-176.194-196.233-241) (140-251)] -6-mer séryl-tétraglycyl-séryl linker (252-257) -scFv-VH-V-lambda anti-CD3E (258-506) [VH (*Mus musculus* IGHV10-1*02 (91.9%)-(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87.0%)-(IGHD) -IGHJ5*01 (100%)] [8.10.16] (258-382) -15-mer tris(tétraglycyl-séryl) linker (383-397) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%) CDR-IMGT [9.3.9] (423-431.449-451.488-496) (398-506)] -hexahistidine (507-512)], non-glycosylé, produit dans des cellules ovariennes de hamsters chinois (CHO)

etevritamab

inmunoglobulina scFv-scFv, anti-[*Homo sapiens* EGFR (receptor del factor de crecimiento epidérmico, receptor tirosina-proteína kinasa erb-1, ERBB1, HER1, HER-1, ERBB) variante III (EGFRvIII)] y anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], anticuerpo monoclonal de cadena única, biespecífico;
IG scFv-scFv cadena única (1-512) [scFv-VH-V-kappa anti-EGFRvIII(1-251) [VH (*Homo sapiens* IGHV3-33*01 G49>C (44) (93.9%) -IGHD) -IGHJ4*01 (100%)] CDR-IMGT [8.8.17] (26-33.51-58.97-113) (1-124) -15-mer tris(tetraglicil-seril) linker (125-139) – V-KAPPA (*Homo sapiens* IGKV2-24*01 (92.0%) -IGKJ1*01 Q120>C (244) (90.9%)] CDR-IMGT [11.3.9] (166-176.194-196.233-241) (140-251)] -6-mer seril-tetraglicil-seril linker (252-257) -scFv-VH-V-lambda anti-CD3E (258-506) [VH (*Mus musculus* IGHV10-1*02 (91.9%)-(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87.0%)-(IGHD) -IGHJ5*01 (100%)] [8.10.16] (258-382) -15-mer tris(tetraglicil-seril) linker (383-397) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%) CDR-IMGT [9.3.9] (423-431.449-451.488-496) (398-506)] -hexahistidina (507-512)], no glicosilado, producido en las células ováricas de hámster chino (CHO)

Sequence / Séquence / Secuencia

```

QVQLVESGGG VVQSGRSLRL SCAASGFTFR NYGMHWVRQA PGKCLEWVAV 50
IWYDGSCKYY ADSVRGRFTI SRDNKNTLY LQMNSLRAED TAVYYCARDG 100
YDILTGNPRD FDYWGQGLV TVSSGGGSG 66GSGGGSGD TVMTQTPLSS 150
HVTLGQPAIS SCRSSQSLVH SDGNTLYSWL QRRGQPPRL LIYRISRRFS 200
GVPDRFSGSG AGDTFTLEIS RVEAEDVGVY YCMQSTHVPY TFGCGTKAEI 250
KSGGGGSEVQ LVESGGGLVQ PGSLKLSKA ASGFTFNKYA MNWVRQAPEK 300
GLEWVARIRS KYNWYATYYA DSVKDRFTIS RDSKNTAYL QMNNLKTEDT 350
AVYYCVRRHG FGNYSIYYA YWGQGLTVTV SSGGGGSGGG 6SGGGGSGQTV 400
VTQEPSLTVS PGSTVTLTCG SSTGAVTSGN YPNWVQKPKG QAPRGLIGGT 450
KFLAPGTAPR FSGSLGGKA ALTLGSGVQPE DEAEYYCVLW YSNRWVFGGG 500
TKLTVLHHHH HH 512

```

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-chain (C23 C104):V 22-96 162-232 279-355 419-487

(C49-C120) V 44-244

N-terminal glutaminylation cyclization to pyroglutamylation (pE, 5-oxoprolyl)

VH Q1:

1

No N-glycosylation sites / pas de sites de N-glycosylation / ningún posición de N-glicosilación

Aglycosylated

fanotaprimum

fanotaprim

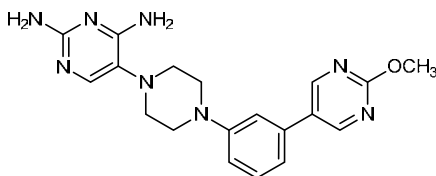
5-{4-[3-(2-methoxypyrimidin-5-yl)phenyl]piperazin-1-yl}pyrimidine-2,4-diamine

fanotaprim

5-{4-[3-(2-méthoxypyrimidin-5-yl)phényl]pipérazin-1-yl}pyrimidine-2,4-diamine

fanotaprim

5-{4-[3-(2-metoxipirimidin-5-il)fenil]piperazin-1-il}pirimidina-2,4-diamina

 $C_{19}H_{22}N_8O$ **favezelimabum #**

favezelimab

immunoglobulin G4-kappa, anti-[*Homo sapiens* LAG3 (lymphocyte activating 3, lymphocyte-activation 3, CD223)], humanized monoclonal antibody; gamma4 heavy chain humanized (1-446) [VH (*Homo sapiens* IGHV1-58*01 (85.6%) -(IGHD) -IGHJ1*01 (91.7%)) [8.8.12] (1-119) -*Homo sapiens* IGHG4*01 (CH1 (120-217), hinge 1-12 (218-229), CH2 (230-339), CH3 (340-444), CHS (445-446)) (120-446)], (133-218')-disulfide with kappa light chain humanized (1'-218') [V-KAPPA (*Homo sapiens* IGKV2D-29*01 (82.0%) -IGKJ4*01 (100%)) [10.3.9] (1'-111') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (157), V101 (195) (112'-218')]; dimer (225-225":228-228")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

favézélimab

immunoglobuline G4-kappa, anti-[*Homo sapiens* LAG3 (activateur 3 des lymphocytes, lymphocyte-activation 3, CD223)], anticorps monoclonal humanisé; chaîne lourde gamma4 humanisée (1-446) [VH (*Homo sapiens* IGHV1-58*01 (85.6%) -(IGHD) -IGHJ1*01 (91.7%)) [8.8.12] (1-119) -*Homo sapiens* IGHG4*01 (CH1 (120-217), charnière 1-12 (218-229), CH2 (230-339), CH3 (340-444), CHS (445-446)) (120-446)], (133-218')-disulfure avec la chaîne légère kappa humanisée (1'-218') [V-KAPPA (*Homo sapiens* IGKV2D-29*01 (82.0%) -IGKJ4*01 (100%)) [10.3.9] (1'-111') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (157), V101 (195) (112'-218')]; dimère (225-225":228-228")-bisdisulfure, produit dans des cellules ovariennes de hamsters chinois (CHO), glycoforme alfa

favezelimab

inmunoglobulina G4-kappa, anti-[*Homo sapiens* LAG3 (activador 3 de los linfocitos, linfocito-activación 3, CD223)], anticuerpo monoclonal humanizado; cadena pesada gamma4 humanizada (1-446) [VH (*Homo sapiens* IGHV1-58*01 (85.6%) -(IGHD) -IGHJ1*01 (91.7%)) [8.8.12] (1-119) -*Homo sapiens* IGHG4*01 (CH1 (120-217), bisagra 1-12 (218-229), CH2 (230-339), CH3 (340-444), CHS (445-446)) (120-446)], (133-218')-disulfuro con la cadena ligera kappa humanizada (1'-218') [V-KAPPA (*Homo sapiens* IGKV2D-29*01 (82.0%) -IGKJ4*01 (100%)) [10.3.9] (1'-111') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (157), V101 (195) (112'-218')]; dímero (225-225":228-228")-bisdisulfuro, producido en las células ováricas de hamsters chinos (CHO), glicofoma alfa

Heavy chain / Chaîne lourde / Cadena pesada
QMQLVQSGPE VKKPGTTSVKV SCKASGYTFT DYNVDWVRQA RGQRLEWIGD 50
INPNDGGTIY AQKFQERVITI TVDKSTSTAY MELSSLRSED TAVYYCARNY 100
RWFGAMDHWG QGTTVTVSSA STKGPSVFPL APCSRSTSES TAALGCLVKD 150
YFPEPVTVSW NSGALTSGVH TFPVQLQSSG LYSLSVVTV PSSSLGKTTY 200
TCNVHDKPSN TKVDKRVESK YGPPCPPCPA PEFLLGGPSVF LFPPKPKDTL 250
MISRTPVETC VVVDVSQEDP EVQFNWYVDG VEVHNAKTKP REEQFNSTYR 300
VVSVLTVLHQ DWLNGKEYKC KVSNGKLPS SIEKTIISKAKG QPREPVYTL 350
PPSQEEMTKN QVSLTCLVKG FYPSTIAVEW ESNQGPENNY KTTTPVLDSD 400
GSFFLYSRLT VDKSRWQEGN VFSCSVMEHA LHNHYTQKSL SLSLGG 446

Light chain / Chaîne légère / Cadena ligera
DIVMTQTPLS LSVTPGQPAS ISCKASQSLD YEGSDSMNWY LQKPGQPPQL 50
LIYGASNLES GVPDRFSGSG SGTDFTLKIS RVEAEDGVY YCQQSTEDPR 100
TFGGGTKEVI KRTVAAPSVF IFPPSDEQLK SGTASVCLL NNFYPREAKV 150
QWKVDNALQS GNSQESVTEQ DSKDSTYSLS STLTLKADY EKHKVYACEV 200
THQGLSSPVT KSFNRGEC 218

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfu
Intra-H (C23-C104) 22"-96" 146"-202" 260"-320' 366"-424"
22"-96" 146"-202" 260"-320' 366"-424"
Intra-L (C23-C104) 23"-92" 138"-198"
23"-92" 138"-198"
Inter-H-L (CH1 10-CL 126) 133-218' 133"-218"
Inter-H-H (h 11, h 14) 225-225" 228-228"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
296, 296"
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennai
complexes fucosylés / glicanos de tipo CHO biantenarijos complejos fucosilados.
C-terminal lysine clipping:
H CHS K2:
446, 446"

firazorextonum
firazorexton

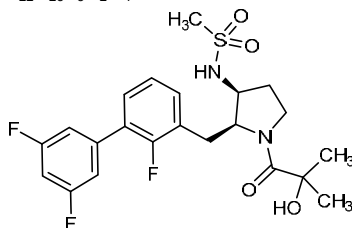
N-((2S,3S)-1-(2-hydroxy-2-methylpropanoyl)-2-
[(2,3',5'-trifluoro[1,1'-biphenyl]-3-yl)methyl]pyrrolidin-3-
yl)methanesulfonamide

firazorexton

N-((2S,3S)-1-(2-hydroxy-2-méthylpropanoyl)-2-
[(2,3',5'-trifluoro[1,1'-biphényl]-3-yl)méthyl]pyrrolidin-3-
yl)méthanesulfonamide

firazorextón

N-((2S,3S)-1-(2-hidroxi-2-metilpropanoil)-2-[(2,3',5'-
trifluoro[1,1'-bifenil]-3-il)metil]pirrolidin-3-
il)metanosulfonamida

C₂₂H₂₅F₃N₂O₄S**flutufolastatium (¹⁸F)**flutufolastat (¹⁸F)

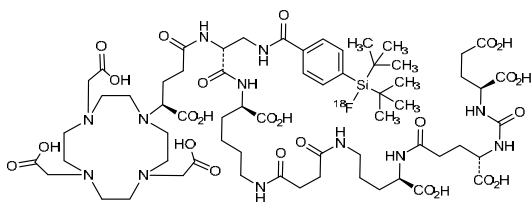
N²-(N-((4S)-4-carboxy-4-[4,7,10-tris(carboxymethyl)-1,4,7,10-tetraaza-cyclododecan-1-yl]butanoyl)-3-[4-(di-*tert*-butyl(¹⁸F)fluorosilyl]benzamido)-D-alanyl)-N⁶-[4-(N²-{N-[(L-glutamic acid-N-yl)carbonyl]-L-γ-glutamyl}-D-ornithin-N⁵-yl)-4-oxobutanoyl]-D-lysine

flutufolastat (¹⁸F)

N²-(N-((4S)-4-carboxy-4-[4,7,10-tris(carboxyméthyl)-1,4,7,10-tétraaza-cyclododécan-1-yl]butanoyl)-3-[4-(di-*tert*-butyl(¹⁸F)fluorosilil]benzamido)-D-alanyl)-N⁶-[4-(N²-{N-[(acide L-glutamique-N-yl)carbonyl]-L-γ-glutamyl}-D-ornithin-N⁵-yl)-4-oxobutanoyl]-D-lysine

flutufolastat (¹⁸F)

N²-(N-((4S)-4-carboxi-4-[4,7,10-tris(carboximetil)-1,4,7,10-tetraazaciclo-dodecan-1-il]butanoil)-3-[4-(di-*tert*-butil(¹⁸F)fluorosilil]benzamido)-D-alanyl)-N⁶-[4-(N²-{N-[(ácido L-glutámico-N-il)carbonil]-L-γ-glutamil}-D-ornitin-N⁵-il)-4-oxobutanoil]-D-lisina

C₆₃H₉₉¹⁸FN₁₂O₂₅Si**flubentylosinum**

flubentylosin

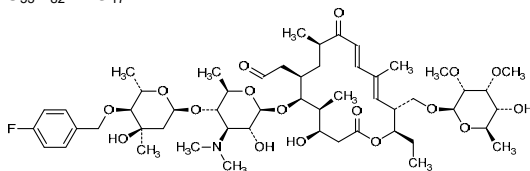
[(4*R*,5*S*,6*S*,7*R*,9*R*,11*E*,13*E*,15*R*,16*R*)-15-[[[(6-deoxy-2,3-di-*O*-methyl-β-D-allopyranosyl)oxy]methyl]-6-[[[3,6-dideoxy-4-*O*-{2,6-dideoxy-4-*O*-[(4-fluorophenyl)methyl]-3-*C*-methyl-α-L-*ribo*-hexopyranosyl]-3-(dimethylamino)-β-D-glucopyranosyl]oxy]-16-ethyl-4-hydroxy-5,9,13-trimethyl-2,10-dioxo-1-oxacyclohexadeca-11,13-dien-7-yl]acetaldehyde; 4^B-*O*-(4-fluorobenzyl)tylosin

flubentylosine

[(4*R*,5*S*,6*S*,7*R*,9*R*,11*E*,13*E*,15*R*,16*R*)-15-[[[(6-désoxy-2,3-di-*O*-méthyl-β-D-allopyranosyl)oxy]methyl]-6-[[[3,6-didésoxy-4-*O*-{2,6-didésoxy-4-*O*-[(4-fluorophényl)méthyl]-3-*C*-méthyl-α-L-*ribo*-hexopyranosyl]-3-(diméthylamino)-β-D-glucopyranosyl]oxy]-16-éthyl-4-hydroxy-5,9,13-triméthyl-2,10-dioxo-1-oxacyclohexadéca-11,13-dié-7-yl]acétaldéhyde; 4^B-*O*-(4-fluorobenzyl)tylosine

flubentilosina

[(4*R*,5*S*,6*S*,7*R*,9*R*,11*E*,13*E*,15*R*,16*R*)-15-[[[(6-desoxi-2,3-di-*O*-metil-β-D-alopiranosil)oxi]metil]-6-[[[3,6-didesoxi-4-*O*-{2,6-didesoxi-4-*O*-[(4-fluorofenil)metil]-3-*C*-metil-α-*L*-ribohexopiranosil]-3-(dimetilamino)-β-D-glucopiranosil]oxi]-16-etil-4-hidroxi-5,9,13-trimetil-2,10-dioxo-1-oxaciclohexadeca-11,13-dien-7-il]acetaldehído; 4^B-*O*-(4-fluorobencil)tilosina

C₅₃H₈₂FNO₁₇**fordadistrogenum movaparvovecum #**

fordadistrogene movaparvovec

A replication-defective adeno-associated viral vector encoding a codon-optimised mini-dystrophin (DMD). A recombinant replication-defective adeno-associated viral vector serotype 9 (rAAV9), encoding a codon-optimised mini-dystrophin (DMD) gene comprising the N-terminus, hinge regions H1, H3 and H4, spectrin-like repeats R1, R2 and R22-24, and the C-terminal cysteine-rich region, under the control of a human muscle-specific synthetic enhancer/promoter and a synthetic poly(A) sequence, flanked by AAV2 inverted terminal repeats (ITR).

fordadistrogène movaparvovec

Un vecteur adénoviral à réplication défectueuse et codant pour une mini-dystrophine (DMD) avec des codons optimisés.

Un vecteur adénoviral recombinant de sérotype 9 (rAAV9) à réplication défectueuse, codant pour un gène de mini-dystrophine (DMD) avec des codons optimisés, comprenant l'extrémité N-terminale, les régions charnières H1, H3 et H4, les répétitions de type spectrine R1, R2 et R22-24, et la région C-terminale riche en cystéines, sous le contrôle d'un promoteur/amplificateur synthétique spécifique du muscle humain et d'une séquence synthétique poly(A), flanquée de répétitions terminales inversées (ITR) de l'AAV2.

fordadistrogén movaparvovec

Un vector de virus adeno-asociado, deficiente de replicación, que codifica para una mini-distrofina (DMD) con los codones optimizados.

Un vector de virus adeno-asociado recombinante de serotipo 9 (rAAV9), deficiente de replicación, que codifica para un gen de mini-distrofina (DMD) con los codones optimizados que contiene el extremo N-terminal, las regiones bisagra H1, H3 y H4, las repeticiones de tipo espectrina R1, R2 y R22-24, y la región C-terminal rica en cisteínas, bajo el control de un promotor/amplificador (enhancer) sintético específico de músculo humano y una secuencia poly(A) sintética, flanqueado por las repeticiones terminales invertidas (ITR) del AAV2.

gallium (⁶⁸Ga) gozetotidumgallium (⁶⁸Ga) gozetotide

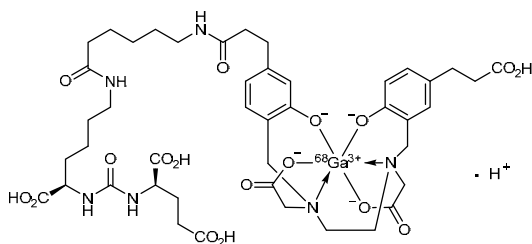
{N-[(N⁶-{6-[3-(3-[[{2-[[{5-(2-carboxyethyl)-2-hydroxy-κO-phenyl]methyl}(carboxy-κO-methyl)amino-κN]ethyl}(carboxy-κO-methyl)amino-κN]methyl)-4-hydroxy-κO-phenyl]propanamido]hexanoyl]-L-lysine-N²-yl)carbonyl]-L-glutamato(3-)}(⁶⁸Ga)gallium

gallium (⁶⁸Ga) gozétotide

{N-[(N⁶-{6-[3-(3-[[{2-[[{5-(2-carboxyéthyl)-2-hydroxy-κO-phényl]méthyl}(carboxy-κO-méthyl)amino-κN]éthyl}(carboxy-κO-méthyl)amino-κN]méthyl)-4-hydroxy-κO-phényl]propanamido]hexanoyl]-L-lysine-N²-yl)carbonyl]-L-glutamato(3-)}(⁶⁸Ga)gallium

galio (⁶⁸Ga) gozetótida

{N-[(N⁶-{6-[3-(3-[[{2-[[{5-(2-carboxietil)-2-hidroxi-κO-fenil]metil}(carboxi-κO-metil)amino-κN]etil}(carboxi-κO-metil)amino-κN]metil)-4-hidroxi-κO-fenil]propanamido]hexanoil]-L-lisine-N²-il)carbonil]-L-glutamato(3-)}(⁶⁸Ga)galio

**garivulimabum #**

garivulimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* CD274 (programmed death ligand 1, PDL1, PD-L1, B7 homolog 1, B7H1)], humanized monoclonal antibody; gamma1 heavy chain humanized (1-446) [VH humanized (*Homo sapiens* IGHV3-NL1*01 (84.7%) - (IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.7.11] (26-33.51-57.96-106) (1-117) -*Homo sapiens* IGHG1*01 G1m17,1, G1v4 CH2 A114 (CH1 K120 (214) (118-215), hinge 1-15 (216-230) -CH2 V1.2>A (234), P114>A (329) (231-339), CH3 D12 (355), L14 (357) (340-444), CHS (445-446)) (118-446)], (220-215')-disulfide with kappa light chain humanized (1'-215') [V-KAPPA humanized (*Homo sapiens* IGKV1-5*03 (84.0%) -IGKJ2*01 (91.7%)) CDR-IMGT [6.3.10] (27-32.50-52.89-98) (1'-108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dimer (226-226":229-229")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alpha

garivulimab

immunoglobuline G1-kappa, anti-[*Homo sapiens* CD274 (ligand 1 de mort programmée, PDL1, PD-L1, homologue 1 de B7, B7H1)], anticorps monoclonal humanisé;

chaîne lourde gamma1 humanisée (1-446) [VH humanisé (*Homo sapiens* IGHV3-NL1*01 (84.7%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.7.11] (26-33.51-57.96-106) (1-117) - *Homo sapiens* IGHG1*01 G1m17,1, G1v4 CH2 A114 (CH1 K120 (214) (118-215), charnière 1-15 (216-230) -CH2 V1.2>A (234), P114>A (329) (231-339), CH3 D12 (355), L14 (357) (340-444), CHS (445-446)) (118-446)], (220-215')-disulfure avec la chaîne légère kappa humanisée (1'-215') [V-KAPPA humanisé (*Homo sapiens* IGKV1-5*03 (84.0%) -IGKJ2*01 (91.7%)) CDR-IMGT [6.3.10] (27-32.50-52.89-98) (1'-108') - *Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dimère (226-226":229-229")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

garivulimab

inmunoglobulina G1-kappa, anti-[*Homo sapiens* CD274 (ligando 1 de muerte programada, PDL1, PD-L1, homólogo 1 de B7, B7H1)], anticuerpo monoclonal humanizado; cadena pesada gamma1 humanizada (1-446) [VH humanizado (*Homo sapiens* IGHV3-NL1*01 (84.7%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.7.11] (26-33.51-57.96-106) (1-117) -*Homo sapiens* IGHG1*01 G1m17,1, G1v4 CH2 A114 (CH1 K120 (214) (118-215), bisagra 1-15 (216-230) -CH2 V1.2>A (234), P114>A (329) (231-339), CH3 D12 (355), L14 (357) (340-444), CHS (445-446)) (118-446)], (220-215')-disulfuro con la cadena ligera kappa humanizada (1'-215') [V-KAPPA humanizado (*Homo sapiens* IGKV1-5*03 (84.0%) -IGKJ2*01 (91.7%)) CDR-IMGT [6.3.10] (27-32.50-52.89-98) (1'-108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dímero (226-226":229-229")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada

EVQLVESGGG	LVQPGGSLRL	SCAVSGFSLT	SYGVHWVRQA	PGKLEWVAV	50
IWAGGSNTYA	DSVKGRTFTIS	KDTSKNTVYL	QMNSLRAEDT	AVYYCAKPYG	100
TSAMDYWGQG	TLTVTSAST	KGPSVFPLAP	SSKSTSGGTA	ALGCLVKDYF	150
PEPTVTSWNS	GALTSGVHTF	PAVLQSSGLY	SLSSVTVTPS	SSLGTQTYIC	200
NVNHKPSNTK	VDDKVEPKSC	DKTHTCPPCP	APPAAGPSVF	LFPPKPKDTL	250
MISRTPTEVC	VVDVSHEDP	EVKFNWYVDG	VEVHNAKTKP	REEQYNSTYR	300
VVSVLTVLHQ	DWLNGKEYKC	KVSNKALAAP	IEKTSKAKG	QPREPVYTL	350
PPSRDELTKN	QVSLTCLVKG	FYPSDIAVEW	ESNGQPENNY	KTTPPVLDSD	400
GSFFLYSKLT	VDKSRWQGN	VFSCSVMEHA	LHNHYTQKSL	SLSPGK	446

Light chain / Chaîne légère / Cadena ligera

DIQMTQSPSS	LSASVGDRVT	ITCKASQDVG	IVVAWYQQK	GKAPKLLIYW	50
ASIRHTGVP	RFSGSGSGTE	FTLTISSLQP	DDFATYYCQ	YSNYPLYTFG	100
QGTKEVIKRT	VAAPSVFIFP	PSDEQLKSGT	ASVCLNNF	YPREAKVQWK	150
VDNALQSGNS	QESVTEQDSK	DSTYSLSTL	TLISKADYEK	KVYACEVTHQ	200
GLSSPVTKSF	NRGEC				215

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104)	22-95	144-200	260-320	366-424
	22"-95"	144"-200"	260"-320"	366"-424"

Intra-L (C23-C104)	23'-88"	135'-195"
	23"-88"	135"-195"

Inter-H-L (h 5-CL 126)	220-215'	220"-215"
Inter-H-H (h 11, h 14)	226-226"	229-229"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

296, 296"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenararios complejos fucosilados.

C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

446, 446"

garveleucelum
garveleucel

Allogeneic T cell-enriched leukocyte preparation, devoid of alloreactive T cells.
The cells are derived from peripheral blood mononuclear cells (PBMCs) collected by apheresis from a haploidentical stem cell donor.
The cells are lymphocyte enriched, and during processing patient and donor cells are co-cultured *ex vivo* in an MLR (mixed lymphocyte reaction) to stimulate activation of host alloreactive T-cells and subsequent depletion of these cells using a proprietary rhodamine-based photodynamic treatment. In this process, monocytes are largely eliminated from the cell mixture.
The cells are primarily T cells (average ~90%) consisting of both CD4+ T-helper cells and CD8+ T-cytotoxic cells (mean CD4:CD8 ratio of 2.1), and other leucocytes (average ~10%, with B-cells (~ 6-7%) and NK cells (~ 2-3%).

garvéleucel

Préparation de leucocytes enrichis en cellules T allogéniques, dépourvus de cellules T alloréactives.
Les cellules sont dérivées de cellules mononucléaires du sang périphérique (PBM) prélevées par aphérèse chez un donneur de cellules souches partiellement compatible pour les antigènes HLA.
Les cellules sont enrichies en lymphocytes et, pendant le traitement, les cellules du patient et du donneur sont co-cultivées *ex vivo* selon une RML (réaction lymphocytaire mixte) pour stimuler l'activation des cellules T allo-réactives de l'hôte et l'épuisement ultérieur de ces cellules grâce à un traitement photodynamique exclusif à base de rhodamine. Dans ce processus, les monocytes sont largement éliminés du mélange de cellules.
Les cellules sont principalement des cellules T (en moyenne ~90%) composées à la fois de cellules T auxiliaires CD4+ et de cellules T cytotoxiques CD8+ (rapport moyen CD4:CD8 de 2.1), et d'autres leucocytes (en moyenne ~10%, avec des cellules B (~6-7%) et des cellules NK (~ 2-3%).

garveleucel

Preparación de leucocitos enriquecidos en células T alogénicos, desprovistos de células T aloreactivas.
Las células se obtienen a partir de células mononucleares de sangre periférica (PBMCs) recogidas por aféresis de un donante de células madre haploidentico.
Las células se enriquecen en linfocitos, y durante el procesamiento, las células del paciente y del donante se co-cultivan *ex vivo* en una MLR (reacción mixta de linfocitos) para estimular la activación de células T aloreactivas frente al huésped y la subsiguiente depleción de estas células usando un tratamiento fotodinámico basado en la rodamina. En este proceso, los monocitos en su mayoría se eliminan de la mezcla celular.
Las células son principalmente células T (~90% como media) y consisten en células T colaboradoras CD4+ y células T citotóxicas CD8+ (ratio medio CD4:CD8 de 2.1), y otros leucocitos (media ~10%, con células B (~6-7%) y células NK (~2-3%).

gavocabtagenum autoleucelum #

gavocabtagene autoleucel

Autologous T cells transduced *ex vivo* with a self-inactivating lentiviral vector encoding an anti-mesothelin (MSLN) chimeric antigen receptor

Autologous T cells transduced *ex vivo* with a non-replicating, self-inactivating (SIN) lentiviral vector, encoding an anti-mesothelin (MSLN) single-domain antibody (sdAb) based chimeric antigen receptor with a leader sequence derived from granulocyte-macrophage colony-stimulating factor receptor alpha (GMCSFR-alpha), fused to the CD3ε subunit of the T-cell receptor (TCR) via an A3(G4S)3LE flexible linker, which is incorporated into the endogenous TCR complex upon expression. The transgene is under control of elongation factor-1 alpha (EF-1α) promoter and woodchuck hepatitis virus post-transcriptional regulatory element. The vector genome also contains a packaging signal, a partial *gag* sequence, a Rev Response Element (RRE), a central polypurine tract (cPPT) and a 3' polypurine tract (3'PPT).

gavocabtagène autoleucel

Cellules T autologues transduites *ex vivo* avec un vecteur lentiviral non répliquatif codant pour un récepteur chimérique anti-mésothéline (MSLN).

Cellules T autologues transduites *ex vivo* avec un vecteur lentiviral non répliquatif et auto-inactivable (SIN), codant pour un récepteur antigénique chimérique basé sur les anticorps anti-mésothéline (MSLN) à domaine unique (sdAb). Il comprend une séquence leader dérivée du récepteur alpha du facteur de stimulation des colonies de granulocytes-macrophages (GMCSFR-alpha), fusionné à la sous-unité CD3ε du récepteur des cellules T (TCR) via un polypeptide lier flexible nommé A3(G4S)3LE, qui est incorporé dans le complexe TCR endogène lors de l'expression. Le transgène est sous le contrôle du promoteur du facteur d'élongation 1 alpha (EF-1α) ainsi que de l'élément régulateur post-transcriptionnel du virus de l'hépatite de la marmotte. Le génome du vecteur contient aussi un signal de conditionnement des gènes, une séquence *gag* partielle, un élément de réponse Rev (RRE), une région centrale avec tractus polypurine (cPPT) et un tractus polypurine 3' (3'PPT).

gavocabtagén autoleucel

Células T autólogas transducidas *ex vivo* con un vector lentiviral auto-inactivante que codifica para un receptor de antígenos quimérico anti-mesotelina (MSLN).

Células T autólogas transducidas *ex vivo* con vector lentiviral auto-inactivante, no replicativo, que codifica para un receptor de antígenos quimérico basado en un anticuerpo de dominio único (sdAb) anti-mesotelina (MSLN) con una secuencia líder derivada del receptor alfa del factor estimulador de colonias de granulocitos-macrófagos (GMCSFR-alfa), fusionado a la subunidad CD3ε del receptor de la célula T (TCR) mediante un conector (linker) flexible A3(G4S)3LE, que se incorpora en el complejo TCR endógeno al expresarse. El transgen está bajo el control del promotor del factor de elongación 1 alfa (EF-1α) y elemento regulador post-transcripcional del virus de la hepatitis de la marmota. El genoma del vector también contiene una señal empaquetadora, un secuencia *gag* parcial, un elemento de respuesta Rev (RRE), un segmento central de polipurinas (cPPT) y un segmento 3' de polipurinas (3'PPT).

geptanolimabum

geptanolimab

immunoglobulin G4-kappa, anti-[*Homo sapiens* PDCD1 (programmed cell death 1, PD-1, PD1, CD279)], monoclonal antibody;
gamma4 heavy chain (1-440) [VH (*Homo sapiens*IGHV7-4-1*02 (85.7%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.7] (26-33.51-58.97-103) (1-114)-*Homo sapiens* IGHG4*01, G4v5 h P10 (CH1 (115-212), hinge 1-12 S10>P (222) (213-224), CH2 (225-334), CH3 (335-439), CHS K>del (440) (115-440)], (28-217')-disulfide with kappa light chain (1'-217') [V-KAPPA (*Mus musculus* IGKV3-5*01 (89.9%) -IGKJ1*01 (90.9%)/*Homo sapiens* IGKV3D-11*02 (64.9%) -IGKJ2*01 (100%)) CDR-IMGT [10.3.8] (27-36.54-56.93-100) (1'-110') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (156), V101 (194) (111'-217')]; dimer (220-220":223-223")-bisdisulfide, produced in Chinese hamster ovary (CHO)-S cell line, glycoform alfa

geptanolimab

immunoglobuline G4-kappa, anti-[*Homo sapiens* PDCD1 (protéine 1 de mort cellulaire programmée, PD-1, PD1, CD279)], anticorps monoclonal;
chaîne lourde gamma4 (1-440) [VH (*Homo sapiens* IGHV7-4-1*02 (85.7%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.7] (26-33.51-58.97-103) (1-114) -*Homo sapiens* IGHG4*01, G4v5 h P10 (CH1 (115-212), charnière 1-12 S10>P (222) (213-224), CH2 (225-334), CH3 (335-439), CHS K>del (440) (115-440)], (128-217')-disulfure avec la chaîne légère kappa (1'-217') [V-KAPPA (*Mus musculus* IGKV3-5*01 (89.9%) -IGKJ1*01 (90.9%)/*Homo sapiens* IGKV3D-11*02 (64.9%) -IGKJ2*01 (100%)) CDR-IMGT [10.3.8] (27-36.54-56.93-100) (1'-110') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (156), V101 (194) (111'-217')]; dimère (220-220":223-223")-bisdisulfure, produit dans des cellules ovariennes de hamsters chinois (CHO) lignée cellulaire CHO-S, glycoforme alfa

geptanolimab

inmunoglobulina G4-kappa, anti-[*Homo sapiens* PDCD1 (proteína 1 de muerte celular programada, PD-1, PD1, CD279)], anticuerpo monoclonal;
cadena pesada gamma4 (1-440) [VH (*Homo sapiens* IGHV7-4-1*02 (85.7%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.7] (26-33.51-58.97-103) (1-114) -*Homo sapiens* IGHG4*01, G4v5 h P10 (CH1 (115-212), bisagra 1-12 S10>P (222) (213-224), CH2 (225-334), CH3 (335-439), CHS K>del (440) (115-440)], (128-217')-disulfuro con la cadena ligera kappa (1'-217') [V-KAPPA (*Mus musculus* IGKV3-5*01 (89.9%) -IGKJ1*01 (90.9%)/*Homo sapiens* IGKV3D-11*02 (64.9%) -IGKJ2*01 (100%)) CDR-IMGT [10.3.8] (27-36.54-56.93-100) (1'-110') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (156), V101 (194) (111'-217')]; dímero (220-220":223-223")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-S, forma glicosilada alfa

gindameranum #
gindameran

gindaméran

Heavy chain / Chaîne lourde / Cadena pesada
QIQLVQSGSE LKKFGASVKV SCKASGYTFT NFGMNWVRQA PQGGLKWMGW 50
ISGYTREPTY AADFKGRFVL SLDTSVSTAY LQISSLKAEQ TAVYYCARDV 100
FDYWGQCTLV TVSSASTKGP SVFPLAPCSR STSESTAALG CLVKDYFPEP 150
VTVSWSNGAL TSGVHTFFAV LQSSGLYSLG SVTVTFSSSL GTKTYTCNDV 200
HKPSNTKVDK RVESKYGPPC PFCPAPEFLG GPSVLEFPK PKDTLMISRT 250
FEVTCVVVDV SQEDPEVQFN WYVDGVEVHN ARTKPREEQF NSTYRVVSVL 300
TVLHQDWLNG KEYKCKVSNK GLPSSIEKTI SKAKGQPREP QVYTLPSQE 350
EMTKNQVSLT CLVKGFYPSD LAVERESNGQ PENNYKTTPE VLDSGDSFEEL 400
YSRLTVDSKR WQEGNVFSCS VMHEALHNNY TQKLSLSLGL 440

Light chain / Chaîne légère / Cadena ligera
DIVLTQSPAS LAVSPGGRAT ITCRASEVD NYGYSFMNWF QQKPGQFFKL 50
LIYRASNLSE GVPARFSGSG SRTDFTLTIN FVEADDTANY YQQQSNADPT 100
FGQGTLEIK RTVAAPSVFL FPFSDQLKS GTASVVCCLN NRYPREARVQ 150
WKVDNALQSG NSQESVTEQD SKDSTYLSLS TLTLTKADYE KHKVYACEVT 200
HQGLSSPVTK SFNRGEC 217

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22°-96° 141°-197° 255°-315° 361°-419°
22°-96° 141°-197° 255°-315° 361°-419°
Intra-L (C23-C104) 23°-92° 137°-197°
23°-92° 137°-197°
Inter-H-L (h10-CL126) 128°-217° 128°-217°
Inter-H-H (h8.h11) 220°-220° 223°-223°

N-terminal glutaminyl cyclization to pyroglutaminyl (pE, 5-oxoprolinyl)
H V H Q I:
I, I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
HCH2N84.4:
291, 291"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenaricos complejos fucosilados.

Messenger RNA encoding cancer/testis antigen 1 (New York oesophageal squamous cell carcinoma antigen-1).
Messenger RNA (mRNA), 5'-capped, encoding codon-optimised New York esophageal squamous cell carcinoma antigen-1 (NY-ESO-1; cancer/testis antigen 1, CTAG1A) expressed as a fusion protein comprising a secretory signal peptide, NY-ESO-1, P2P16 tetanus toxoid-derived helper epitopes and a major histocompatibility complex (MHC) class I transmembrane and cytoplasmic domain (MITD), connected by three glycine/serine-rich (GS) linker peptides, flanked by 5' and 3' untranslated regions and a 3' poly(A) tail.

ARN messenger codant pour l'antigène 1 du cancer testiculaire (antigène 1 du carcinome squameux de l'œsophage, dit de New York).
ARN messenger (ARNm), protégé en 5', codant pour l'antigène 1 du carcinome squameux de l'œsophage, dit de New York (NY-ESO-1; antigène 1 du cancer testiculaire, CTAG1A), avec des codons optimisés, exprimé sous la forme d'une protéine de fusion comprenant un peptide signal de sécrétion, NY-ESO-1, les épitopes auxiliaires P2P16 dérivés de l'anatoxine tétanique et un domaine transmembranaire et cytoplasmique (MITD) du complexe majeur d'histocompatibilité (CMH) de classe I, reliés par trois peptides de liaison riches en glycine/sérine (GS), flanqués de régions non traduites en 5' et 3' et d'une queue poly(A) en 3'.

Recommended INN: List 85

gindamerán

WHO Drug Information, Vol. 35, No. 1, 2021

ARN mensajero que codifica para el antígeno 1 de cáncer/testículo (antígeno 1 de carcinoma escamoso del esófago Nueva York).

ARN mensajero (ARNm), protegido en 5', que codifica para el antígeno 1 de carcinoma escamoso del esófago Nueva York (NY-ESO-1; antígeno cáncer/testículo 1, CTAG1A) con codones optimizados, que se expresa como una proteína de fusión que consta de un péptido señal de secreción, NY-ESO-1, los epítomos auxiliares P2P16 derivados del toxoide tetánico y un dominio citoplásmico y transmembrana del complejo principal de histocompatibilidad (MHC) de clase I (MITD), conectados mediante tres péptidos de enlace ricos en glicina/serina (GS), flanqueado por las regiones 5' y 3' no traducidas y una cola poly(A) en 3'.

grisnilimabum #

grisnilimab

immunoglobulin G2A-lambda, anti-[CD7 (CD7 antigen (p41), GP40, LEU-9, TP41, Tp40)], *Mus musculus* monoclonal antibody;
gamma2a heavy chain *Mus musculus* (1-453) [VH (*Mus musculus* IGHV9-3-1*01 (98%) -(IGHD) - IGHJ2*01 (93.3%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Mus musculus* IGHG2A*01 (100%) (CH1 (124-220), hinge 1-16 (221-236), CH2 (237-346), CH3 (347-451), CHS (452-453)) (124-453)], (138-214')-disulfide with lambda light chain *Mus musculus* (1'-215') [V-LAMBDA (*Mus musculus* IGLV1*01 (98%) -IGLJ1*01 (100%)) CDR-IMGT [9.3.9] (26-34.52-54.91-99) (1'-109') -*Mus musculus* IGLC1*01 (100%) (110'-215')];
dimer (230-230":233-233":235-235")-trisulfide, produced in SP2/0-derived mouse myeloma cells, glycoform alfa

grisnilimab

immunoglobuline G2A-lambda, anti-[CD7 (CD7 antigène (p41), GP40, LEU-9, TP41, Tp40)], anticorps monoclonal *Mus musculus*;
chaîne lourde gamma2a *Mus musculus* (1-453) [VH (*Mus musculus* IGHV9-3-1*01 (98%) -(IGHD) - IGHJ2*01 (93.3%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Mus musculus* IGHG2A*01 (100%) (CH1 (124-220), charnière 1-16 (221-236), CH2 (237-346), CH3 (347-451), CHS (452-453)) (124-453)], (138-214')-disulfure avec la chaîne légère lambda *Mus musculus* (1'-215') [V-LAMBDA (*Mus musculus* IGLV1*01 (98%) -IGLJ1*01 (100%)) CDR-IMGT [9.3.9] (26-34.52-54.91-99) (1'-109') -*Mus musculus* IGLC1*01 (100%) (110'-215')];
dimère (230-230":233-233":235-235")-trisulfure, produit par une ligne cellulaire dérivée de myélome murin SP2/0, glycoforme alfa

grisnilimab

inmunoglobulina G2A-lambda, anti-[CD7 (CD7 antígeno (p41), GP40, LEU-9, TP41, Tp40)], anticuerpo monoclonal *Mus musculus*;

cadena pesada gamma2a *Mus musculus* (1-453) [VH (*Mus musculus*IGHV9-3-1*01 (98%) -(IGHD) -IGHJ2*01 (93.3%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Mus musculus*IGHG2A*01 (100%) (CH1 (124-220), bisagra 1-16 (221-236), CH2 (237-346), CH3 (347-451), CHS (452-453)) (124-453)], (138-214')-disulfuro con la cadena ligera lambda *Mus musculus* (1'-215') [V-LAMBDA (*Mus musculus*IGLV1*01 (98%) -IGLJ1*01 (100%)) CDR-IMGT [9.3.9] (26-34.52-54.91-99) (1'-109') -*Mus musculus*IGLC1*01 (100%) (110'-215')];
dímero (230-230":233-233":235-235")-trisulfuro, producido en una línea celular de mieloma murino SP2/0, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
 QIQLVQSGPE LKPGGETVKI SCKASGYTFT NYGMNWKQA PGKGLMWLGW 50
 INTYTGETPY ADDFKGRFAF SLETSASTAY LQINNKNED TATYFCARWA 100
 YFYGSSPYFF DYWGQGTLLT VSSAKTTAPS VYPLAPVCGD TTGSSVTLGC 150
 LVKGYFPEPV TLTWNSGSL SSGVHTFPAVL QSDLYTLSSS VTVTSSTWPS 200
 QSITCNVAHP ASSTKVDKKI EPRGPTIKPC PPCKCPAPNL LGGSPVFIFP 250
 PKIKDLVMIS LSPITVTCVV DVSEDDPDVQ ISWVFNNEV HTAQQTTHRE 300
 DYNSTLRVVS ALPIQHQQWM SGKEFKCKVN NKDLPAPIER TISKPKGSVR 350
 APQVYVLPFP EEEMTKQVLT LTCMTDFMP EDIYVEWTNN GKTELNYKNT 400
 EPVLDSGDSY FMYSKLRVEK KMWVERNSYS CSVVHEGLHN HHTTKSFSRT 450
 PGK 453

Light chain / Chaîne légère / Cadena ligera
 QAVVTQESAL TTSPGETVTL TCRSGTAVT TSNYANWQEQ KPDHLFTGLI 50
 GGTNNRAGPV PARFSGSLIG DKAALITIGA QTEDEAIYFC ALWCSNHLVF 100
 GGGTKLTVLG QPKSSPSVTL FPPSSELET NKATLVCTIT DFYPGVVTVTD 150
 WKVDGTPVTQ GMETTQPSKQ SNKYMMASSY LTLTARAWER HSSYSCQVTH 200
 EGHTEKSL S RADCS 215

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 150-205 267-327 373-431
 22"-96" 150"-205" 267"-327" 373"-431"

Intra-L (C23-C104) 22'-90' 137'-196'
 22'"-90'" 137'"-196'"

Inter-H-L (CH1 11-CL 126) 138-214' 138"-214"

Inter-H-H (h 12, h 15, h 18) 230-230" 233-233" 235-235"

N-terminal glutaminyl cyclization to pyroglutamy1 (pE, 5-oxoprolyl)

H VH Q1:

1, 1"

L VL Q1:

1', 1'"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

303, 303"

Fucosylated complex bi-antennary SP2/0-type glycans / glycanes de type SP2/0 bi-antennaires complexes fucosylés / glicanos de tipo SP2/0 biantenarijos complejos fucosilados

C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

453, 453"

grisnilimabum setaritoxum #
 grisnilimab setaritox

immunoglobulin G2A-lambda, anti-[CD7 (CD7 antigen (p41), GP40, LEU-9, TP41, Tp40)], *Mus musculus* monoclonal antibody conjugated to ricin toxin A (RTA); gamma2a heavy chain *Mus musculus* (1-453) [VH (*Mus musculus*IGHV9-3-1*01 (98%) -(IGHD) -IGHJ2*01 (93.3%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Mus musculus*IGHG2A*01 (100%) (CH1 (124-220), hinge 1-16 (221-236), CH2 (237-346), CH3 (347-451), CHS (452-453)) (124-453)], (138-214')-disulfide

	with lambda light chain <i>Mus musculus</i> (1'-215') [V-LAMBDA (<i>Mus musculus</i> IGLV1*01 (98%) - IGLJ1*01 (100%)) CDR-IMGT [9.3.9] (26-34.52-54.91-99) (1'-109') -<i>Mus musculus</i> IGLC1*01 (100%) (110'-215')]; dimer (230-230":233-233":235-235")-trisulfide, produced in SP2/0-derived mouse myeloma cells, glycoform alfa, substituted at N⁶ of an average of 1.6 lysyl residues with 4-[(1<i>RS</i>)-1-[[L-methionyl-ricin toxin A-chain (Met-RTA, non-glycosylated, produced in <i>Escherichia coli</i>)-S²⁶⁰-yl]sulfanyl]ethyl]benzoyl groups
grisnilimab sétaritox	immunoglobuline G2A-lambda, anti-[CD7 (CD7 antigène (p41), GP40, LEU-9, TP41, Tp40)], anticorps monoclonal <i>Mus musculus</i> conjugué à la toxine A de la ricine (RTA); chaîne lourde gamma2a <i>Mus musculus</i> (1-453) [VH (<i>Mus musculus</i> IGHV9-3-1*01 (98%) - (IGHD) - IGHJ2*01 (93.3%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -<i>Mus musculus</i> IGHG2A*01 (100%) (CH1 (124-220), charnière 1-16 (221-236), CH2 (237-346), CH3 (347-451), CHS (452-453)) (124-453)], (138-214')-disulfure avec la chaîne légère lambda <i>Mus musculus</i> (1'-215') [V-LAMBDA (<i>Mus musculus</i> IGLV1*01 (98%) - IGLJ1*01 (100%)) CDR-IMGT [9.3.9] (26-34.52-54.91-99) (1'-109') -<i>Mus musculus</i> IGLC1*01 (100%) (110'-215')]; dimère (230-230":233-233":235-235")-trisulfure, produit par une lignée cellulaire dérivée de myélome murin SP2/0, glycoforme alfa, substitué en N⁶ sur un moyenne de 1.6 résidus lysyl par des groupes 4-[(1<i>RS</i>)-1-[[L-méthionyl-chaîne A de la toxine de ricine (Met-RTA, non-glycosylée, produite par <i>Escherichia coli</i>)-S²⁶⁰-yl]sulfanyl]éthyl]benzoyle
grisnilimab setaritox	inmunoglobulina G2A-lambda, anti-[CD7 (CD7 antígeno (p41), GP40, LEU-9, TP41, Tp40)], anticuerpo monoclonal <i>Mus musculus</i> conjugado con la toxina A de la ricina (RTA); cadena pesada gamma2a <i>Mus musculus</i> (1-453) [VH (<i>Mus musculus</i> IGHV9-3-1*01 (98%) - (IGHD) - IGHJ2*01 (93.3%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -<i>Mus musculus</i> IGHG2A*01 (100%) (CH1 (124-220), bisagra 1-16 (221-236), CH2 (237-346), CH3 (347-451), CHS (452-453)) (124-453)], (138-214')-disulfuro con la cadena ligera lambda <i>Mus musculus</i> (1'-215') [V-LAMBDA (<i>Mus musculus</i> IGLV1*01 (98%) - IGLJ1*01 (100%)) CDR-IMGT [9.3.9] (26-34.52-54.91-99) (1'-109') -<i>Mus musculus</i> IGLC1*01 (100%) (110'-215')]; dímero (230-230":233-233":235-235")-trisulfuro, producido en una línea celular de mieloma murino SP2/0, forma glicosilada alfa, sustituida en N⁶ de un promedio de 1,6 residuos de lisilo con grupos de 4-[(1<i>RS</i>)-1-[[L-metionil-cadena A de toxina de ricina (Met-RTA, no glicosilada, producida por <i>Escherichia coli</i>)-S²⁶⁰-il]sulfanil]etil]benzoilo

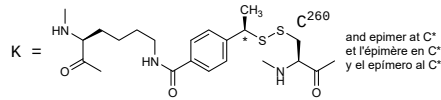
Heavy chain / Chaîne lourde / Cadena pesada
QIQLVQSGPE LKKPGETVKI SCKASGYTFT NYGMNWKQA PGKGLMWLGW 50
INTYTGEPTY ADDFKGRFAF SLETSASTAY LQINNLKNE TATYFCARWA 100
YFYGSSPYFF DYWGQGTTLT VSSAKTTAPS VYPLAPVCGD TTGSSVTLGC 150
LVKGYFPEPV TLWNVSGSL SSVHTFPAVL QSDLYTLSSS VTVTSSTWPS 200
QSITCNVAHP ASSTKVDDKI EPRGPTIKPC PPCKCPAPNL LGGPSVFIFP 250
PKIKDVLMS LSPIVTCVVV DVSEDDPDVQ ISWFVNNEV HTAQQTQTHRE 300
DYNSTLRVVS ALPIQHQQDM SGKEFKCKVN NKDLPAPIER TISKPKGSVR 350
APQVYVLPFP EEEMTKQVT LTCMVTDFMP EDIYVEWNN GKTELNYKNT 400
EPVLDSDGSY FMYSKLRVEK KMWERNYSY CSVVHEGLHN HHTTKSFSRT 450
PGK 453

Light chain / Chaîne légère / Cadena ligera
QAVVTQESAL TTSPGETVTL TCRSSTGAVT TSNYANWVQE KPDHLFTGLI 50
GGTNRAPGV PARFSGSLG DKAALTTTGA QTEDEAIYFC ALWCSNHLVF 100
GGGTLKLVLG QPKSSPSVTL FPPSSELET NKATLVCTIT DFYPGVVTVD 150
WKVDGTPVTQ GMETTQPSKQ SNNKYMSSY LTLTARAWER HSSYSCQVTH 200
EGHTVEKSLS RADCS 215

Met-Ricin A chain / Met-chaîne A de la Ricine / Met-cadena A de la Ricina
MIFPKQYPII NFFTAGATVY SYTNFIRAVR GRLTTGADV HDIVPLPNRV 50
GLPINQRFIL VELSNHAELS VTLALDVNTA YVVGYRAGNS AYFFHPDNQE 100
DAEAIHLFT DVQNYTFAP G6NYDRLEQL AGNLRENIEI GNGPLEEAIS 150
ALYYSTGGT QLPTLARSFI ICIQMISEAA RFQYIEGEMR TRIRYNRRSA 200
PDPSVITLEN SWGRLSTAIQ ESNQGAFAFP IQLQRRNGSK FSVYDVSILI 250
PIIALMVYRC APPSSQF 268

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 150-205 267-327 373-431
22"-96" 150"-205" 267"-327" 373"-431"
Intra-L (C23-C104) 22"-90" 137"-196"
22"-90" 137"-196"
Inter-H-L (CH1 11-CL 126) 138-214 138"-214"
Inter-H-H (h 12, h 15, h 18) 230-230 233-233 235-235
heterogeneous patterns involving 94" and 94" e.g.:
96-94 138-94 150-94 94-137
96"-94" 138"-94" 150"-94" 94"-137"
Met-RTA: C172 = Cys-SH, C260 = Cys-S-S-CH(CH3)-p-C6H4-CO-[N6-Lys(H,L)]

Conjugation sites / Sites de conjugaison / Posiciones de coniugación
major: K87 K228 K72 K113
K87" K228" K72" K113"
minor: K43 K215 K323 K332 K398 K421 K132
K43" K215" K323" K332" K398" K421" K132"



N-terminal glutaminy cyclization to pyroglutamyl (pE, 5-oxoprolyl)
H VH Q1:
1, 1"
L VL Q1:
1', 1"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84,4:
303, 303"
Fucosylated complex bi-antennary SP2/0-type glycans / glycanes de type SP2/0 bi-antennaires
complexes fucosylés / glicanos de tipo SP2/0 biantennarios complejos fucosilados.

C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal
H CHS K2:
453, 453"

hepcidinum
hepcidin

hepcidin (human) (hepatic bactericidal protein, hepcidin-25, liver-expressed antimicrobial peptide 1, LEAP-1, hepatic antimicrobial peptide, HAMP, ferroportin regulator protein):
S⁷,S²³:S¹⁰,S¹³:S¹¹,S¹⁹:S¹⁴,S²²-tetracyclo(L-α-aspartyl-L-threonyl-L-histidyl-L-phenylalanyl-L-prolyl-L-isoleucyl-L-cysteinyl-L-isoleucyl-L-phenylalanyl-L-cysteinyl-L-cysteinylglycyl-L-cysteinyl-L-cysteinyl-L-histidyl-L-arginyl-L-seryl-L-lysyl-L-cysteinylglycyl-L-methionyl-L-cysteinyl-L-cysteinyl-L-lysyl-L-threonine)

hepcidine

hepcidine (humaine) (protéine bactéricide hépatique, hepcidine-25, peptide antimicrobien exprimé par le foie 1, LEAP-1, peptide antimicrobien hépatique, HAMP, protéine régulatrice de la ferroportine):

Recommended INN: List 85

WHO Drug Information, Vol. 35, No. 1, 2021

$S^7, S^{23}, S^{10}, S^{13}, S^{11}, S^{19}, S^{14}, S^{22}$ -tétracyclo(L- α -aspartyl-L-thréonyl-L-histidyl-L-phénylalanil-L-prolyl-L-isoleucyl-L-cystéinyl-L-isoleucyl-L-phénylalanil-L-cystéinyl-L-cystéinylglycyl-L-cystéinyl-L-cystéinyl-L-histidyl-L-arginyl-L-séryl-L-lysyl-L-cystéinylglycyl-L-méthionyl-L-cystéinyl-L-cystéinyl-L-lysyl-L-thréonine)

hepcidina

hepcidina (humana) (proteína bactericida hepática, hepcidina-25, péptido antimicrobiano expresado en el hígado 1, LEAP-1, péptido antimicrobiano hepático, HAMP, proteína reguladora de ferroportina): $S^7, S^{23}, S^{10}, S^{13}, S^{11}, S^{19}, S^{14}, S^{22}$ -tetraciclo(L- α -aspartil-L-treonil-L-histidil-L-fenilalanil-L-prolil-L-isoleucil-L-cisteinil-L-isoleucil-L-fenilalanil-L-cisteinil-L-cisteinilglicil-L-cisteinil-L-cisteinil-L-histidil-L-arginil-L-seril-L-lisil-L-cisteinilglicil-L-metionil-L-cisteinil-L-cisteinil-L-lisil-L-treonina)

$C_{113}H_{170}N_{34}O_{31}S_9$

Sequence / Séquence / Secuencia
DTHTFICIFC CGCCHRSKCG MCKKT 25

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
7-23, 10-13, 11-19, 14-22

ibezapolstatum

ibezapolstat

2-[[[(3,4-dichlorophenyl)methyl]amino]-7-[2-(morpholin-4-yl)ethyl]-1,7-dihydro-6H-purin-6-one

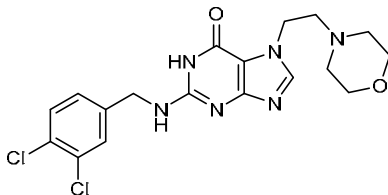
ibézapolstat

2-[[[(3,4-dichlorophényl)méthyl]amino]-7-[2-(morpholin-4-yl)éthyl]-1,7-dihydro-6H-purin-6-one

ibezapolstat

2-[[[(3,4-diclorofenil)metil]amino]-7-[2-(morfolin-4-il)etil]-1,7-dihidro-6H-purin-6-ona

$C_{18}H_{20}Cl_2N_6O_2$



icapamespibum

icapamespib

9-[2-[(2,2-dimethylpropyl)amino]ethyl]-8-[(6-iodo-2H-1,3-benzodioxol-5-yl)sulfanyl]-9H-purin-6-amine

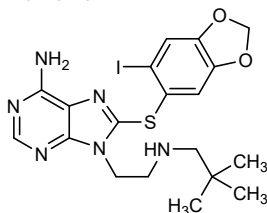
icapamespib

9-[2-[(2,2-diméthylpropyl)amino]éthyl]-8-[(6-iodo-2H-1,3-benzodioxol-5-yl)sulfanyl]-9H-purin-6-amine

icapamespib

9-[2-[(2,2-dimetilpropil)amino]etil]-8-[(6-iodo-2H-1,3-benzodioxol-5-il)sulfanil]-9H-purin-6-amina

$C_{19}H_{23}IN_6O_2S$



idactamabum #

idactamab

immunoglobulin G1-kappa, anti-[*Homo sapiens* SLC1A5 (solute carrier family 1 member 5, neutral amino acid transporter, RDRC, M7V1, AAAT, ASCT2)], *Homo sapiens* monoclonal antibody;
 gamma1 heavy chain *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV4-34*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) [8.7.15] (1-121) -*Homo sapiens* IGHG1*03 G1m3, nG1m1 (CH1 R120 (218) (122-219), hinge 1-15 (220-234), CH2 3⁴>ins^{cysteiny}l (244) (235-345), CH3 E12 (361), M14 (363) (346-450), CHS (451-452)) (122-452)], (224-214')-disulfide with kappa light chain *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-5*03 (94.7%) -IGKJ1*01 (100%)) [6.3.9] (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')];
 dimer (230-230":233-233")-bisdisulfide, produced in a Chinese hamster ovary (CHO) cell line derived from CHO-K1, glycoform alfa

idactamab

immunoglobuline G1-kappa, anti-[*Homo sapiens* SLC1A5 (famille du porteur soluté 1 membre 5, transporteur d'acide aminé neutre, RDRC, M7V1, AAAT, ASCT2)], anticorps monoclonal *Homo sapiens*;
 chaîne lourde gamma1 *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV4-34*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) [8.7.15] (1-121) -*Homo sapiens* IGHG1*03 G1m3, nG1m1 (CH1 R120 (218) (122-219), charnière 1-15 (220-234), CH2 3⁴>ins^{cysteiny}l (244) (235-345), CH3 E12 (361), M14 (363) (346-450), CHS (451-452)) (122-452)], (224-214')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-5*03 (94.7%) -IGKJ1*01 (100%)) [6.3.9] (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')];
 dimère (230-230":233-233")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) dérivant de la lignée cellulaire CHO-K1, glycoforme alfa

idactamab

inmunoglobulina G1-kappa, anti-[*Homo sapiens* SLC1A5 (familia del portador de soluto 1 miembro 5, transportador de ácido amino neutral, RDRC, M7V1, AAAT, ASCT2)], anticuerpo monoclonal *Homo sapiens*;
 cadena pesada gamma1 *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV4-34*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) [8.7.15] (1-121) -*Homo sapiens* IGHG1*03 G1m3, nG1m1 (CH1 R120 (218) (122-219), bisagra 1-15 (220-234), CH2 3⁴>ins^{cisteinil} (244) (235-345), CH3 E12 (361), M14 (363) (346-450), CHS (451-452)) (122-452)], (224-214')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-5*03 (94.7%) -IGKJ1*01 (100%)) [6.3.9] (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')];
 dímero (230-230":233-233")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular derivada de CHO-K1, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
QVQLQQWAG LLKPSETLSL TCAYVGGSEF GYYWSWIRQP PGKGLEWIGE 5C
IHHSGGANY FSLKSRVTIS VDTSKNQFSL KLSSVTAADT AVYYCARGQG 100
KNWHYDYFDY WGGGLTVTS SASTKGPSVF FLAPSSKSTG GGTAAALGLV 150
KDYFPEPVTV SWNSGALTSG VHTFPAVLQS SGLYSLSSVV TFPSSSLGTQ 200
TYICNVNHKP SNTKVDKRV EKSCKDTHC PFCPAPELLG GPSCVFLFPP 250
KPKDTLMISR TPEVTCVVVD VSHEDPEVKF NWYVDGVEVH NAKTKPREEQ 300
YNSTYRVSV LTVLHQDWLN GKELYCKVSN KALPAPIET ISKAKGQPRE 350
EQVYTLFESR EEMTKNQVSL TCLVKGFYPS DIAVEWESNG QPENNYKTP 400
PVLDSGGSEF LYSKLTVDKS RWQQGNVESC SVMHEALHNN YTQKSLSLSP 450
GK 452

Light chain / Chaîne légère / Cadena ligera
DIQMTQSPST LSASVGDRVT ITCRASQSI SWLAWYQQKPK GKAPKLLIYK 5C
ASILKIGVPS RFSGSGSGTE FTLTISSLQP DDFATYYCQQ YYYSRTFGQ 100
GTKVEIKRTV AAPSVFIFFP SDEQLKSGTA SVVCLNNFY BREAKVQWKV 150
DNALQSGNSQ ESVTEDQSKD STYLSLSLTI LSKADYEKHK VYACEVTHQG 200
LSSPVTKSEN RGEC 214

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22°-95° 148°-204° 266°-326° 372°-430°
22°-95° 148°-204° 266°-326° 372°-430°
Intra-L (C23-C104) 23°-88° 134°-194°
23°-88° 134°-194°
Inter-H-L (h 5-CL 126) 224°-214° 224°-214°
Inter-H-H (h 11, h 14) 230°-230° 233°-233°

Cysteinyln inserted for conjugation capped by a cysteinyl
HCH2C3'4):
244, 244"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
HCH2N84.4:
302, 302"
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires
complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

C-terminal lysine clipping:
HCHS K2:
452, 452"

ilacirnonum
ilacirnon

4-chloro-*N*-[5-methyl-2-(7*H*-pyrrolo[2,3-*d*]pyrimidine-4-carbonyl)pyridin-3-yl]-3-(trifluoromethyl)benzene-1-sulfonamide

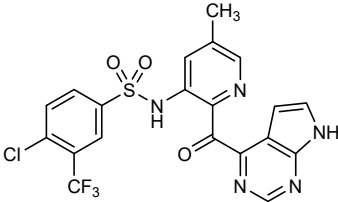
ilacirnon

4-chloro-*N*-[5-méthyl-2-(7*H*-pyrrolo[2,3-*d*]pyrimidine-4-carbonyl)pyridin-3-yl]-3-(trifluorométhyl)benzène-1-sulfonamide

ilacirnón

4-cloro-*N*-[5-metil-2-(7*H*-pirrolo[2,3-*d*]pirimidina-4-carbonil)piridin-3-il]-3-(trifluorometil)bencono-1-sulfonamida

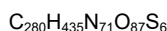
C₂₀H₁₃ClF₃N₅O₃S



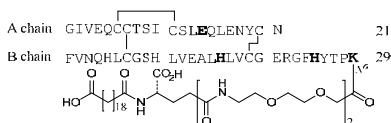
insulinum icodecum
insulin icodec

human insulin A-chain (1-21) variant (Y>E¹⁴), human insulin B-chain (1-29) variant (Y>H¹⁶, F>H²⁵) conjugated at K^{29B} at its N⁶ to (22*S*)-22,42-dicarboxy-10,19,24-trioxa-3,6,12,15-tetraoxa-9,18,23-triazadotetracontan-1-oyl; N^{6.29B}[(22*S*)-22,42-dicarboxy-10,19,24-trioxa-3,6,12,15-tetraoxa-9,18,23-triazadotetracontan-1-oyl]-[Tyr14^A>Glu, Tyr16^B>His, Phe25^B>His]-des-Thr30^B-human insulin

insuline icodéc	chaîne A de l'insuline humaine (1-21) variant (Y>E ¹⁴), chaîne B de l'insuline humaine (1-29) variant (Y>H ¹⁶ , F>H ²⁵), conjuguée en K ^{29B} par son N ⁶ à (22S)-22,42-dicarboxy-10,19,24-trioxo-3,6,12,15-tétraoxa-9,18,23-triazadotétracontan-1-oyle; N ^{6,29B} -[(22S)-22,42-dicarboxy-10,19,24-trioxo-3,6,12,15-tétraoxa-9,18,23-triazadotétracontan-1-oyl]-[Tyr14 ^A >Glu, Tyr16 ^B >His, Phé25 ^B >His]-des-Thr30 ^B -insuline humaine
insulina icodéc	insulina humana cadena-A (1-21) variante (Y>E ¹⁴), insulina humana cadena-B (1-29) variante (Y>H ¹⁶ , F>H ²⁵) conjugada con K ^{29B} con su N ⁶ al (22S)-22,42-dicarboxi-10,19,24-trioxo-3,6,12,15-tetraoxa-9,18,23-triazadotetracontan-1-oilo; N ^{6,29B} -[(22S)-22,42-dicarboxi-10,19,24-trioxo-3,6,12,15-tetraoxa-9,18,23-triazadotetracontan-1-oil]-[Tir14 ^A >Glu, Tir16 ^B >His, Fen25 ^B >His]-des-Tre30 ^B -insulina humana



Sequences / Séquences / Secuencias



Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

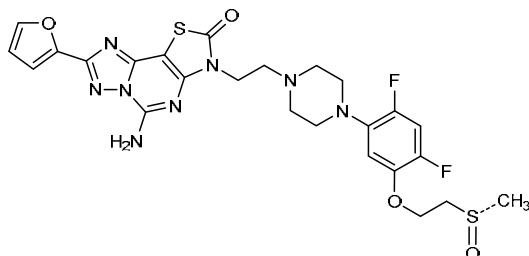
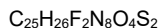
Intra-chain 6^A-11^A

Inter-chain $7^A-7^B, 20^A-19^B$

Modifications / Modifications / Modificaciones

Lys29¹³ conjugated at N⁶

inupadenantum	
inupadenant	(+)-5-amino-3-[2-[4-(2,4-difluoro-5-[2-[(S)-methanesulfinyl]ethoxy)phenyl]piperazin-1-yl]ethyl]-8-(furan-2-yl)[1,3]thiazolo[5,4-e][1,2,4]triazolo[1,5-c]pyrimidin-2(3 <i>H</i>)-one
inupadénant	(+)-5-amino-3-[2-[4-(2,4-difluoro-5-[2-[(S)-méthanesulfinyl]éthoxy]phényl]pipérazin-1-yl]éthyl]-8-(furan-2-yl)[1,3]thiazolo[5,4-e][1,2,4]triazolo[1,5-c]pyrimidin-2(3 <i>H</i>)-one
inupadenant	(+)-5-amino-3-[2-[4-(2,4-difluoro-5-[2-[(S)-metanosulfinil]etoxi]fenil]piperazin-1-il]etil]-8-(furan-2-il)[1,3]tiazolo[5,4-e][1,2,4]triazolo[1,5-c]pirimidin-2(3 <i>H</i>)-ona



invimestrocelum

invimestrocel

Allogeneic human multipotent adult progenitor cells (MAPC) derived from bone marrow and isolated by density gradient centrifugation. The cells are expanded *ex vivo* on fibronectin-coated surfaces under low-oxygen conditions and in the presence of growth media for the creation of a master cell bank (passage 4), a working cell bank (passage 6) and active substance (passage 9). The cells are adherent, secrete proangiogenic factors (including VEGF, IL-8 and CXCL5) and of mesenchymal progenitor lineage (*in vitro* differentiation to endothelial, hepatic and neuronal lineages). The cells express CD49c and CD90 ($\geq 95\%$ of the cells), and lack expression of CD45 and HLA II ($< 5\%$). Additional markers include $>90\%$ CD13, CD44, CD73, CD105 and $<5\%$ CD34. The cells have also been shown to induce T cell suppression.

invimestrocel

Cellules humaines adultes progénitrices multipotentes qui sont allogéniques (MAPC), dérivées de la moelle osseuse et isolées par centrifugation à gradient de densité. Les cellules sont amplifiées *ex vivo* sur des surfaces recouvertes de fibronectine dans des conditions de faible teneur en oxygène et en présence de milieux de croissance pour la création d'une banque de cellules primaires (passage 4) et d'une banque de cellules de travail (passage 6) et d'une substance active produit fini (passage 9). Les cellules sont adhérentes, sécrètent des facteurs pro-angiogéniques (notamment le VEGF, l'IL-8 et la CXCL5) et sont de la lignée progénitrice mésenchymateuse (différenciation *in vitro* en lignées endothéliales, hépatiques et neuronales). Les cellules expriment CD49c et CD90 ($\geq 95\%$ des cellules), et sont négatives pour CD45 et HLA II ($< 5\%$). Les marqueurs supplémentaires comprennent $> 90\%$ de CD13, CD44, CD73, CD105 et $< 5\%$ de CD34. Il a également été démontré que les cellules induisent la suppression des lymphocytes T.

invimestrocel

Células progenitoras adultas multipotentes humanas alogénicas (MAPC) derivadas de médula ósea y aisladas mediante centrifugación en gradiente de densidad. Las células se expanden *ex vivo* en superficies cubiertas de fibronectina en condiciones de oxígeno bajo y en presencia de medio de crecimiento para la creación de un banco de células maestro (pase 4) y un banco de células de trabajo (pase 6) y el principio activo (pase 9). Las células son adherentes, secretan factores proangiogénicos (incluyendo VEGF, IL-8 y CXCL5) y son de la línea progenitora mesenquimal (diferenciación *in vitro* a las líneas endotelial, hepática y neuronal). Las células expresan CD49c ($\geq 95\%$ de las células), y carecen de expresión de CD45 y HLA II ($<5\%$). Otros marcadores adicionales incluyen $>90\%$ de CD13, CD44, CD73, CD105 y $<5\%$ de CD34. Se ha demostrado también que las células inducen supresión de linfocitos T.

ipivivintum

ipivivint

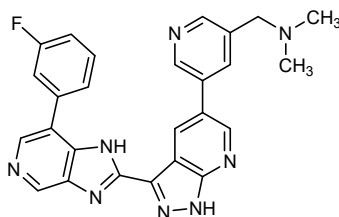
1-(5-{3-[7-(3-fluorophenyl)-1*H*-imidazo[4,5-*c*]pyridin-2-yl]-1*H*-pyrazolo[3,4-*b*]pyridin-5-yl}pyridin-3-yl)-*N,N*-dimethylmethanamine

ipivivint

1-(5-{3-[7-(3-fluorophényl)-1*H*-imidazo[4,5-*c*]pyridin-2-yl]-1*H*-pyrazolo[3,4-*b*]pyridin-5-yl}pyridin-3-yl)-*N,N*-diméthylméthanamine

ipivivint

1-(5-{3-[7-(3-fluorofenil)-1*H*-imidazo[4,5-*c*]piridin-2-il]-1*H*-pirazolo[3,4-*b*]piridin-5-il}piridin-3-il)-*N,N*-dimetilmetanamina

C₂₆H₂₁FN₈**izuralimabum #**

izuralimab

immunoglobulin half-IG G1-kappa/scFv-h-CH2-CH3, anti-[*Homo sapiens* ICOS (inductible T-cell costimulatory, activation-inducible lymphocyte immunomediatory molecule, AILIM, CD278)] and anti-[*Homo sapiens* PDCD1 (programmed cell death 1, PD-1, PD1, CD279)], monoclonal antibody, bispecific; gamma1 heavy chain anti-ICOS (1-454) [VH (*Homo sapiens* IGHV1-2*02 (99%) -(IGHD) -IGHJ3*02 (100%)) CDR-IMGT [8.8.18] (26-33.51-58.97-114) (1-125) -*Homo sapiens* IGHG1*03v, G1m3>G1m17, nG1m1 (CH1 N114>D (216) R120>K (222) (126-223), hinge 1-15 (224-238), CH2 E1.4.L1.3>P (241), L1.2>V (242), G1.1>A (243), S29>K (274), Q84.2>E (302) (239-347), CH3 E12 (363), M14 (365), L24>D (375), K26>S (377), N44>D (391), Q97>E (425), N100>D (428), M107>L (435), N114>S (441) (348-452), CHS (453-454)) (126-454)], (228-214')-disulfide with kappa light chain anti-ICOS (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-12*01 (96.8%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; IG scFv-h-CH2-CH3 single chain, anti-PDCD1 (1"-480") [scFv-V-kappa-VH anti-PDCD1 (1"-249")][V-KAPPA (*Homo sapiens* IGKV3-15*01 (80.0%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1"-107") -20-mer tetrakis(glycyl-lysyl-prolyl-glycyl-seryl) linker (108"-127") -VH (*Mus musculus* IGHV6-6*02 (87.0%) -(IGHD) -IGHJ1*03 (86.7%)/*Homo sapiens* IGHV3-15*07 (81.8%) -(IGHD) -IGHJ2*01 (93.3%)) CDR-IMGT [8.10.13] (153-160.178-187.226-238) (128"-249")] -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (250"-480") [hinge 1-15 C5>S (254) (250-264) -CH2 E1.4.L1.3>P (267), L1.2>V (268), G1.1>A (269), S29>K (300) (265-373), CH3 E12 (389), E13>Q (390), M14 (391), S20>K (397), M107>L (461), N114>S (467) (374-478), CHS (479-480)]]; dimer (234-260":237-263")-bisdisulfide, produced in Chinese hamster ovary (CHO)-S cell line, glycoform alfa

izuralimab immunoglobuline demi-IG G1-kappa/scFv-h-CH2-CH3, anti-[*Homo sapiens* ICOS (costimulateur inducible du lymphocyte T, molécule immunomédiateur lymphocytaire inducible par activation, AILIM, CD278)] et anti-[*Homo sapiens* PDCD1 (protéine 1 de mort cellulaire programmée, PD-1, PD1, CD279)], anticorps monoclonal, bispécifique; chaîne lourde gamma1 anti-ICOS (1-454) [VH (*Homo sapiens*IGHV1-2*02 (99%) -(IGHD) -IGHJ3*02 (100%)) CDR-IMGT [8.8.18] (26-33.51-58.97-114) (1-125) -*Homo sapiens*IGHG1*03v, G1m3>G1m17, nG1m1 (CH1 N114>D (216) R120>K (222) (126-223), charnière 1-15 (224-238), CH2 E1.4.L1.3>P (241), L1.2>V (242), G1.1>A (243), S29>K (274), Q84.2>E (302) (239-347), CH3 E12 (363), M14 (365), L24>D (375), K26>S (377), N44>D (391), Q97>E (425), N100>D (428), M107>L (435), N114>S (441) (348-452), CHS (453-454)) (126-454)], (228-214')-disulfure avec la chaîne légère kappa anti-ICOS (1'-214') [V-KAPPA (*Homo sapiens*IGKV1-12*01 (96.8%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens*IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; IG scFv-h-CH2-CH3 chaîne unique, anti-PDCD1 (1"-480") [scFv V-kappa-VH anti-PDCD1 (1"-249") [V-KAPPA (*Homo sapiens*IGKV3-15*01 (80.0%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1"-107") -20-mer tétrakis(glycyl-lysyl-prolyl-glycyl-séryl) linker (126"-145") -VH (*Mus musculus*IGHV6-6*02 (87.0%) -(IGHD) -IGHJ1*03 (86.7%)/*Homo sapiens*IGHV3-15*07 (81.8%) -(IGHD) -IGHJ2*01 (93.3%)) CDR-IMGT [8.10.13] (153-160.178-187.226-238) (128"-249")]] -*Homo sapiens*IGHG1*03 h-CH2-CH3, nG1m1 (250"-480") [charnière 1-15 C5>S (254) (250-264) -CH2 E1.4.L1.3>P (267), L1.2>V (268), G1.1>A (269), S29>K (300) (265-373), CH3 E12 (389), E13>Q (390), M14 (391), S20>K (397), M107>L (461), N114>S (467) (374-478), CHS (479-480)]]; dimère (234-260":237-263")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-S, glycoforme alfa

izuralimab inmunoglobulina demi-IG G1-kappa/scFv-h-CH2-CH3, anti-[*Homo sapiens* ICOS (coestimulador inducible del linfocito T, molécula inmunomediadora linfocitaria inducible por activación, AILIM, CD278)] y anti-[*Homo sapiens* PDCD1 (proteína 1 de muerte celular programada, PD-1, PD1, CD279)], anticuerpo monoclonal, biespecífico; cadena pesada gamma1 anti-ICOS (1-454) [VH (*Homo sapiens*IGHV1-2*02 (99%) -(IGHD) -IGHJ3*02 (100%)) CDR-IMGT [8.8.18] (26-33.51-58.97-114) (1-125) -*Homo sapiens*IGHG1*03v, G1m3>G1m17, nG1m1 (CH1 N114>D (216) R120>K (222) (126-223), bisagra 1-15 (224-238), CH2 E1.4.L1.3>P (241), L1.2>V (242), G1.1>A (243), S29>K (274), Q84.2>E (302) (239-347), CH3 E12 (363), M14 (365), L24>D (375), K26>S (377), N44>D (391), Q97>E (425), N100>D (428), M107>L (435), N114>S (441) (348-452), CHS (453-454)) (126-454)], (228-214')-disulfuro con la cadena ligera kappa anti-ICOS (1'-214') [V-KAPPA (*Homo sapiens*IGKV1-12*01 (96.8%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens*IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; IG scFv-h-CH2-CH3 cadena única, anti-PDCD1 (1"-480") [scFv V-kappa-VH anti-PDCD1 (1"-249") [V-KAPPA (*Homo sapiens*IGKV3-15*01 (80.0%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1"-107") -20-mer tétrakis(glicil-lisil-prolil-glicil-seril) linker (126"-145") -VH (*Mus musculus*IGHV6-6*02 (87.0%) -(IGHD) -IGHJ1*03 (86.7%)/*Homo sapiens*IGHV3-15*07 (81.8%) -(IGHD) -IGHJ2*01 (93.3%)) CDR-IMGT [8.10.13] (153-160.178-187.226-238) (128"-249")]] -*Homo sapiens*IGHG1*03 h-CH2-CH3, nG1m1 (250"-480") [bisagra 1-15 C5>S (254) (250-264) -CH2 E1.4.L1.3>P (267), L1.2>V (268), G1.1>A (269), S29>K (300) (265-373), CH3 E12 (389), E13>Q (390), M14 (391), S20>K (397), M107>L (461), N114>S (467) (374-478), CHS (479-480)]]; dímero (234-260":237-263")-bisdisulfuro, producido en las células ováricas de hamster chino (CHO) línea celular CHO-S, forma glicosilada alfa

1. Heavy chain / Chaîne lourde / Cadena pesada (anti-ICOS)	
QVQLVQSGAE VVKPGASVKV SCKASGYTFT GYMHVVRQA PGQGLEWMGW	50
INPHSGGTNY AQKFQGRVTM TRDTSISTAY MELSRLRSDD TAVYYCARTY	100
YDSSGGYYHD AFDIWQGTM VTVSSASTKG PSVFPLAPSS KSTSGGTAAL	150
GCLVKDYFPE PVTVSWNSGA LTSGVHTFPA VLQSSGLYSL SSVVTVPSSS	200
LGQTQYICNV NHHKPSDTKVD KKVPEKCDK THTCPPCAP PVAGPSVFLF	250
PPKPKDITLMI SRTPEVTCVV VDVKHEDPEV KFNWYVDGVE VHNAKTKPRE	300
EEYNSTYRVV SVLTVLHQDW LNKKEYCKV SNKALPAIE KTISKAKGQP	350
REPQVYTLPP SREEMTKNQV SLTCDVSGFY PSDIAVEWES DGQPENNYKT	400
TPPYLDSDGS FFLYSKLTVD KSRWEQGVDF SCSVLHEALH SHYTQKSLSL	450
SPGK	454
2. Light chain / Chaîne légère / Cadena ligera (anti-ICOS)	
DIQMTQSPSS VSASVGDRTV ITCRASQGIS RLLAWYQQKP GKAPKLLIYV	50
ASSLQSGVPS RFGSGSGGTD FTLTISSLQP EDFATYYCQQ ANSFPWTFGQ	100
GTKVEIKRTV AAPSVEIFPP SDEQLKSGTA SVVCLLNIFY PREAKVQWKV	150
DNALQSGNSQ ESVTEQDSKD STYLSSTLT LSKADYEKKH VYACEVTHQG	200
LSSPVTKSFN RGEK	214
3. Chain / Chaîne / Cadena IG scFv-h-CH2-CH3 (anti-PDCD1)	
EIVLTQSPAT LSASPGERVV LTCRASQSVG NDVAWYQQKP GQAPRLLINY	50
ASHRYTGVPD RFTGSGYGTE FTLTISSVQS EDFGVYYCQQ DFSSPRTFGG	100
GTKVEIKGKP GSGKPGSGKP GSGKPGSEVQ LVESGGGLVK PGGSRLRSCV	150
ASGFTFSNYW MNWVRQAPGK GLEWVAEIRL YSNNYATHYA ESVKGRFTIS	200
RDDSKSTLYL QMNNLKTEDT GVYYCTRYG NYGGYFDWVG RGLTVTVSSE	250
PKSSDKTHTC PPCAPPVAG PSVFLFPPKP KDTLMISRTV EVTCVVVDVK	300
HEDPEVKFNW YVDGVEVHNA KTKPREEQYN STYRVVSVLT VLHQDWLNGK	350
EYKCKVSNKA LPAPIEKTIS KAKGQPREPQ VYTLPPSREQ MTKNQVKLTC	400
LVKGFYPSDI AVEWESNGQP ENNYKTTTPV LDDSGSFFLY SKLTVDKSRW	450
QQGNVFCSCV LHEALSHYT QKSLSLSPGK	480
Post-translational modifications	
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro	
Intra-H (C23-C104)	22-96 152-208 268-328 374-432
Intra-L (C23-C104)	23"-88" 134"-194"
Intra-chain 3	23"-88" 149"-225" 294"-354" 400"-458"
Inter-H-L (h 5-CL 126)	228-214"
Inter-H-chain 3 (h 11, h 14)	234-260" 237-263"
N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación	
H CH2 N84.4:	
304, 330"	
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenaricos complejos fucosilados.	

laduviglusibum
laduviglusib

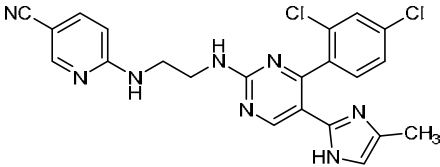
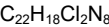
6-[(2-[[4-(2,4-dichlorophenyl)-5-(4-methyl-1H-imidazol-2-yl)pyrimidin-2-yl]amino]ethyl)amino]pyridine-3-carbonitrile

laduviglusib

6-[(2-[[4-(2,4-dichlorophényl)-5-(4-méthyl-1H-imidazol-2-yl)pyrimidin-2-yl]amino]éthyl)amino]pyridine-3-carbonitrile

laduviglusib

6-[(2-[[4-(2,4-diclorofenil)-5-(4-metil-1H-imidazol-2-il)]pirimidin-2-il]amino]etil)amino]piridina-3-carbonitrilo



lazucirnonum
lazucirnon

(4²R)-1⁴-chloro-*N,N*,1³,7⁶-tetramethyl-4⁵,5-dioxo-6-aza-7(2)-pyridina-3(1,4)-piperidina-4(1,2)-pyrrolidina-1(1)-benzenaheptaphane-7⁴-carboxamide

Recommended INN: List 85

lazucirmon

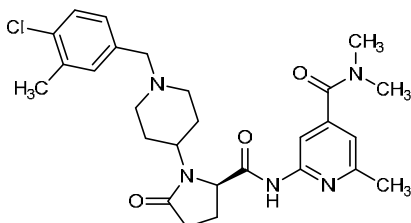
lazucirmon

WHO Drug Information, Vol. 35, No. 1, 2021

(4²R)-1⁴-chloro-*N,N*,1³,7⁶-tétraméthyl-4⁵,5-dioxo-6-aza-7(2)-pyridina-3(1,4)-pipéridina-4(1,2)-pyrrolidina-1(1)-benzénaheptaphane-7⁴-carboxamide

(4²R)-1⁴-cloro-*N,N*,1³,7⁶-tetrametil-4⁵,5-dioxo-6-aza-7(2)-piridina-3(1,4)-piperidina-4(1,2)-pirrolidina-1(1)-bencenaheptafano-7⁴-carboxamida

C₂₇H₃₄ClN₅O₃



Ierodalcibepum #

Ierodalcibep

Iérodalcibep

Ierodalcibep

engineered binding protein (1-96) anti-(human proprotein convertase subtilisin/kexin type 9), derived from human tenth fibronectin type III domain, fused via peptidyl linker (⁹⁷GSGSGS¹⁰²) to human serum albumin (103-687), variant (C³⁴>A¹³⁶), produced in Chinese hamster ovary (CHO) cells; human fibronectin tenth type III domain variant anti-[human PCSK9 (pro-protein convertase subtilisin/kexin type 9, neural apoptosis-regulated convertase 1, NARC-1, proprotein convertase 9, PC9)] (1-96) fused via a (GS)₃ peptide linker (97-102) with [Cys³⁴>Ala (136)]-human serum albumin (HSA) (103-687), produced in Chinese hamster ovary (CHO) cells, non-glycosylated

protéine de liaison mise au point (1-96) anti-(proprotéine convertase subtilisine/kexine type 9), dérivée du dixième domaine de la fibronectine humaine de type III, fusionnée via un peptide de liaison (⁹⁷GSGSGS¹⁰²) à l'albumine sérique humaine (103-687), variant (C³⁴>A¹³⁶), produit dans des cellules ovariennes de hamster chinois (CHO); peptide variant du dixième domaine de la fibronectine humaine de type III anti-[PCSK9 humain (proprotéine convertase subtilisine/kexine type 9, convertase 1 régulée par l'apoptose neuronale, NARC-1, proprotéine convertase 9, PC9)] (1-96) fusionné via un peptide (GS)₃ (97-102) à la [Cys³⁴>Ala (136)]-albumine sérique humaine (ASH) (103-687), produit dans des cellules ovariennes de hamsters chinois (CHO), non-glycosylé

proteína de unión diseñada (1-96) anti-(proproteína convertasa humana subtilisina/kexina tipo 9), derivada del décimo dominio de la fibronectina humana del tipo III, fusionada a través de un conector peptidil (⁹⁷GSGSGS¹⁰²) a la albúmina sérica humana (103-687), variante (C³⁴>A¹³⁶), producido en células ováricas de hámster Chino (CHO);

variante del décimo dominio de la fibronectina humana del tipo III anti-[PCSK9 humano (proproteína convertasa subtilisina/kexina tipo 9, convertasa 1 regulada por la apoptosa neuronal, NARC-1, proproteína convertasa 9, PC9)] fusionada mediante un péptido (GS)₃ (97-102) a la [Cis³⁴>Ala (136)]-albúmina sérica humana (ASH) (103-687), producido en células ováricas de hámster Chino (CHO), no glicosilado

Sequence / Séquence / Secuencia			
VSDVPRDLEV	VAATPTSLLI	SWDAPAEGYG	YYRITYGETG GNSPVQEFTV 50
PVSKGTATIS	GLKPGVDYTI	TVYAVEFDFF	GAGYHRPIS INYRTEGS6S 100
GSDAHKSEVA	HRFKDLGEEN	FKALVLIIFA	QYLQQA ¹³⁶ PFED HVKLVNEVTE 150
FAKTCVADES	AENCDKSLHT	LFGDKLCTVA	TLRETYGEMA DCCAKQEPER 200
NECFLQHKDD	NPNLRLVRP	EVDVMCTAFH	DNEETFLKKY LYEIARRHPY 250
FYAPELLFFA	KRYKAAFTEC	CQAADKAACL	LPKLDLDRDE GKASSAKQRL 300
KCASLQKFGE	RAFKAWAVAR	LSQRFPKAEF	AEVSKLVTDL TKVHTECCHG 350
DLLECADDRA	DLAKYICENQ	DSISSKLKEC	CEKPLLEKSH CIAEVENDEM 400
PADLPSLAAD	FVESKDVCNK	YAEAKDVLFG	MFLYEYARRH PDYSVLLLR 450
LAKTYETTLT	KCCAAADPHE	CYAKVFDEFK	PLVEEPQNLI KQNCELFEQL 500
GEYKFQNAL	VRYTKKVPQV	STPTLVEVSR	NLGKVGSKCC KHPEAKRMPC 550
AEDYLSVVLN	QLCVLHEKTP	VSDRVTKCCT	ESLVNRRPCF SALEVDETYV 600
PKEFNAETFT	FHADICTLSE	KERQIKKQTA	LVELVKHKPK ATKEQLKAVM 650
DDFAAFVEKC	CKADDKETCF	AEEGKKLVAA	SQAALGL 687
Mutation / Mutation / Mutación			
C ¹³⁶ >A			
Post-translational modifications			
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro			
155-164, 177-192, 193-203, 226-270, 271-279, 302-347, 348-355, 367-380, 381-391, 418-462, 463-471, 494-539, 540-550, 563-578, 579-589, 616-660, 661-669			
Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación			
none / aucune / ninguna			

letaplimabum #
letaplimab

immunoglobulin G4-kappa, anti-[*Homo sapiens* CD47 (integrin associated protein, IAP, MER6, OA3)], *Homo sapiens* monoclonal antibody; gamma4 heavy chain *Homo sapiens* (1-441) [VH (*Homo sapiens* IGHV4-59*01 (100%) -(IGHD) -IGHJ1*01 (100%)) CDR-IMGT [8.7.9] (26-33.51-57.96-104) (1-115)-*Homo sapiens* IGHG4*01, G4v4 CH2 A1.3, A1.2 (CH1 (116-213), hinge 1-12 S10>P (223) (214-225), CH2 F1.3>A (229), L1.2>A (230) (226-335), CH3 (336-440), CHS K>del (441) (116-441)], (129-214')-disulfide with kappa light chain *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-12*01 (96.8%) -IGKJ4*01 (91.7%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (221-221":224-224")-bisdisulfide, produced in Chinese hamster ovary (CHO)-S cell line, glycoform alfa

létaplimab

immunoglobuline G4-kappa, anti-[*Homo sapiens* CD47 (protéine associée à l'intégrine, IAP, MER6, OA3)], anticorps monoclonal *Homo sapiens*;

letaplimab

chaîne lourde gamma4 *Homo sapiens*(1-441) [VH (*Homo sapiens* IGHV4-59*01 (100%) -(IGHD) - IGHJ1*01 (100%)) CDR-IMGT [8.7.9] (26-33.51-57.96-104) (1-115) -*Homo sapiens* IGHG4*01, G4v4 CH2 A1.3, A1.2 (CH1 (116-213), charnière 1-12 S10>P (223) (214-225), CH2 F1.3>A (229), L1.2>A (230) (226-335), CH3 (336-440), CHS K>del (441) (116-441)], (129-214')-disulfure avec la chaîne légère kappa *Homo sapiens*(1'-214') [V-KAPPA (*Homo sapiens* IGKV1-12*01 (96.8%) -IGKJ4*01 (91.7%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (221-221":224-224")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-S, glycoforme alfa

immunoglobulina G4-kappa, anti-[*Homo sapiens* CD47 (proteína asociada a la integrina, IAP, MER6, OA3)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma4 *Homo sapiens*(1-441) [VH (*Homo sapiens* IGHV4-59*01 (100%) -(IGHD) - IGHJ1*01 (100%)) CDR-IMGT [8.7.9] (26-33.51-57.96-104) (1-115) -*Homo sapiens* IGHG4*01, G4v4 CH2 A1.3, A1.2 (CH1 (116-213), bisagra 1-12 S10>P (223) (214-225), CH2 F1.3>A (229), L1.2>A (230) (226-335), CH3 (336-440), CHS K>del (441) (116-441)], (129-214')-disulfuro con la cadena ligera kappa *Homo sapiens*(1'-214') [V-KAPPA (*Homo sapiens* IGKV1-12*01 (96.8%) -IGKJ4*01 (91.7%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (221-221":224-224")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-S, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada			
QVQLQESGPG	LVKPSSETLSL	TCTVSGGSGIS	SYYSWIRQP PGKGLEWIGY 50
IYYSGSTNYN	PSLKSRTVTS	VDTSKNQFSL	KLSSVTAADT AVYYCARGKT 100
GSAAWGQGTL	VTVSASTKG	PSVFPLAPCS	RSTSESTAAL GCLVKDYFPE 150
PVTVSWNSGA	LTSGVHTFPA	VLQSSGLYSL	SSVTVTPSSS LGTKTYTCNV 200
DHKPSNTKVD	KRVESKYGPP	CPPCPAPEAA	GGPSVFLFPP KPKDTLMISR 250
TPEVTCVVVD	VSQEDPEVQF	NWYVDGVEVH	NAKTKPREEQ FNSTYRVVSV 300
LTVLHQDWLN	GKEYKCKVSN	KGLPSSIEKT	ISKAKGQPRE PQVYTLPPSQ 350
EEMTKNQVSL	TCLVKGFYPS	DIAVEWESNG	QPENNYKTTT PVLDSGGSFF 400
LYSRLTVDKS	RWQEGNVFSC	SVMHEALHNN	YTKSLSLSL G 441

Light chain / Chaîne légère / Cadena ligera			
DIQMTQSPSS	VSASVGRVIT	ITCRASQGIS	RWLAWYQKPK GKAPKLLIYA 50
ASSLQSGVPS	RFGSGSGSTD	FTLTISSLQP	EDFATYYCQK TVSFPIITFGG 100
GTKVEIKRTV	AAPSVFIIPP	SDEQLKSGTA	SVVCLLNFFY PREAKVQWVK 150
DNALQSGNSQ	ESVTEQDSKD	STYLSSTLT	LSKADYEHKK VYACEVTHQG 200
LSSPVTKSFN	RGEC		214

Post-translational modifications			
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro			
Intra-H (C23-C104)	22"-95"	142"-198"	256"-316" 362"-420"
	22"-95"	142"-198"	256"-316" 362"-420"
Intra-L (C23-C104)	23'"-88'"	134'"-194'"	
	23'"-88'"	134'"-194'"	
Inter-H-L (CH1 10-CL 126)	129-214'	129"-214"	
Inter-H-H (h 8, h 11)	221-221"	224-224"	

N-terminal glutaminyl cyclization to pyroglutamyl (pE, 5-oxoprolyl)
L VH Q1:
I, I*

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
292, 292*
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenariós complejos fucosilados.

lirametostat

lirametostat

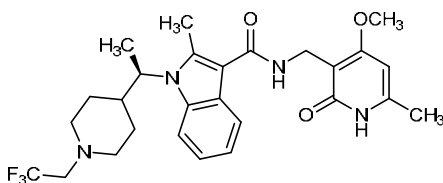
N-[(4-methoxy-6-methyl-2-oxo-1,2-dihydropyridin-3-yl)methyl]-2-methyl-1-[(1*R*)-1-[1-(2,2,2-trifluoroethyl)piperidin-4-yl]ethyl]-1*H*-indole-3-carboxamide

liramétostat

N-[(4-méthoxy-6-méthyl-2-oxo-1,2-dihydropyridin-3-yl)méthyl]-2-méthyl-1-[(1*R*)-1-[1-(2,2,2-trifluoroéthyl)pipéridin-4-yl]éthyl]-1*H*-indole-3-carboxamide

lirametostat

2-metil-*N*-[(6-metil-4-metoxi-2-oxo-1,2-dihidropiridin-3-il)metil]-1-[(1*R*)-1-[1-(2,2,2-trifluoroetil)piperidin-4-il]etil]-1*H*-indol-3-carboxamida

C₂₇H₃₃F₃N₄O₃**ludateronum**

ludaterone

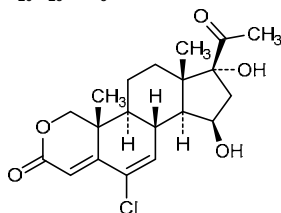
6-chloro-15 β ,17-dihydroxy-2-oxapregna-4,6-diene-3,20-dione

ludatérone

6-chloro-15 β ,17-dihydroxy-2-oxapréгна-4,6-diène-3,20-dione

ludaterona

6-cloro-15 β ,17-dihidroxi-2-oxapregna-4,6-dieno-3,20-diona

C₂₀H₂₅ClO₅**lutetium (¹⁷⁷Lu) vipivotidum tetraxetanum**lutetium (¹⁷⁷Lu) vipivotide tetraxetan

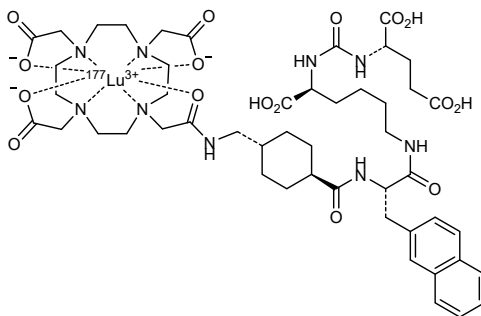
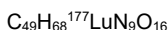
{*N*-[(*N*⁶-{3-(naphthalen-2-yl)-*N*-[*trans*-4-({2-[4,7,10-tris(carboxy- κ^3 O⁴,O⁷,O¹⁰-methyl)-1,4,7,10-tetraazacyclododecan-1-yl- κ^4 N¹,N⁴,N⁷,N¹⁰]acetamido- κ O)methyl)cyclohexane-1-carbonyl]-L-alanyl]-L-lysine-N²-yl)carbonyl]-L-glutamato(3-))](¹⁷⁷Lu)lutetium

lutécium (^{177}Lu) vipivotide tétraxetan

{N-[(N⁶-{3-(naphtalén-2-yl)-N-[trans-4-{(2-[4,7,10-tris(carboxy-κ³O⁴,O⁷,O¹⁰-méthyl)-1,4,7,10-tétrazacyclododécan-1-yl-κ⁴N¹,N⁴,N⁷,N¹⁰]acétamido-κO)méthyl)cyclohexane-1-carbonyl]-L-alanyl]-L-lysine-N²-yl)carbonyl]-L-glutamato(3-)}(^{177}Lu)lutecium

lutecio (^{177}Lu) vipivotida tetraxetan

{N-[(N⁶-{3-(naftalen-2-il)-N-[trans-4-{(2-[4,7,10-tris(carboxy-κ³O⁴,O⁷,O¹⁰-metil)-1,4,7,10-tetraazaciclododecan-1-il-κ⁴N¹,N⁴,N⁷,N¹⁰]acetamido-κO)metil)ciclohexano-1-carbonil]-L-alanil]-L-lisin-N²-il)carbonil]-L-glutamato(3-)}(^{177}Lu)lutecio

melrilimabum #
melrilimab

immunoglobulin G2-kappa, anti-[*Homo sapiens* IL1RL1 (interleukin 1 receptor like 1, DER4, FIT-1, growth stimulation expressed 2 gene, ST2, IL33R)], *Homo sapiens* monoclonal antibody;
gamma2 heavy chain *Homo sapiens* (1-449) [VH (*Homo sapiens* IGHV3-23*01 (91.8%) -(IGHD) - IGHJ4*01(92.9%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Homo sapiens* IGHG2*02v G2m23>G2m..., G2v3 CH2 A1.2, A1, S2, A30, L92, S115, S116 (CH1 T92 (195) (124-221), hinge 1-12 (222-233), CH2 V1.2>A (237), G1>A (239), P2>S (240), H30>A (270), M45.1>V (284), V92>L (311), A115>S (332), P116>S (333) (234-342), CH3 (343-447), CHS (448-449)) (124-449)], (137-214')-disulfide with kappa light chain *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV3D-11*02 (95.6%) - IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (225-225'':226-226'':229-229'':232-232'')-tetrakisdisulfide, produced in Chinese hamster ovary (CHO)-K1SV cell line lacking the Glutamine Synthetase (GS) gene, glycoform alfa

melrilimab

immunoglobuline G2-kappa, anti-[*Homo sapiens* IL1RL1 (récepteur like 1 de l'interleukine 1, DER4, FIT1, gène 2 exprimé lors de la stimulation de la croissance, ST2, IL33R)], anticorps monoclonal *Homo sapiens*;

chaîne lourde gamma2 *Homo sapiens* (1-449) [VH (*Homo sapiens* IGHV3-23*01 (91.8%) - (IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Homo sapiens* IGHG2*02v G2m23>G2m..., G2v3 CH2 A1.2, A1, S2, A30, L92, S115, S116 (CH1 T92 (195) (124-221), charnière 1-12 (222-233), CH2 V1.2>A (237), G1>A (239), P2>S (240), H30>A (270), M45.1>V (284), V92>L (311), A115>S (332), P116>S (333) (234-342), CH3 (343-447), CHS (448-449)) (124-449)], (137-214')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV3D-11*02 (95.6%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (225-225":226-226":229-229":232-232")-tétrakisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-K1SV ne présentant pas le gène de la glutamine synthétase, glycoforme alfa

melrilimab

immunoglobulina G2-kappa, anti-[*Homo sapiens* IL1RL1 (receptor like 1 de la interleukina 1, DER4, FIT1, gen 2 expresado durante la estimulación del crecimiento, ST2, IL33R)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma2 *Homo sapiens* (1-449) [VH (*Homo sapiens* IGHV3-23*01 (91.8%) - (IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Homo sapiens* IGHG2*02v G2m23>G2m..., G2v3 CH2 A1.2, A1, S2, A30, L92, S115, S116 (CH1 T92 (195) (124-221), bisagra 1-12 (222-233), CH2 V1.2>A (237), G1>A (239), P2>S (240), H30>A (270), M45.1>V (284), V92>L (311), A115>S (332), P116>S (333) (234-342), CH3 (343-447), CHS (448-449)) (124-449)], (137-214')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV3D-11*02 (95.6%) - IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (225-225":226-226":229-229":232-232")-tetraakisdisulfuro, producido en las células ováricas de hámsters chinos (CHO) línea celular CHO-K1SV en ausencia del gen Glutamina Sintetasa (GS)forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
EVQLLESQGGG LVQPGGSLRL SCAASGFTFS IYDNIWVRQA PGKGLEWSS 50
IRGEGGGTYY ADSVKGRTTI SRDNSKNTLY LQMNLSRAED TAVYYCARDP 100
WSTEGSFFVL DYWGQTLVT VSSASTKGPS VFPLAPCSRS TSESTAALGG 150
LVKDYFPEPV TVSNNSGALT SGVHTFPVAL QSSGLYSLSS VVTYTSSNFG 200
TQTYTCNVDD KPSNTKVDKT VERKCCVECP PCAPAPAAAS SVFLFPPKPK 250
DTLMSIRTPV VTCVVVDVSA EDPEVQFNWY VDGVEVHNAK TKPREQFNS 300
TFRVSVLTV LHQDMLNGKE YKCKVSNKGL PSSTIEKISK TKGQPREPQV 350
YTLPPSREEM TKNQVSLTCL VKGFYPSDIA VEWESNGQPE NNYKTTTPML 400
DSGSGFFLYS KLTVDKSRWQ QGNVYSCSVM HEALHNHYQT KSLSLSPGK 449

Light chain / Chaîne légère / Cadena ligera
EIVLTQSPAT LSLSPGERAT LSCRASQSDV DDLAWYQKP GQAPRLTIYD 50
ASNRATGIPA RFGSGSGSDT FTLTISSLEP EDFAVYYCQY YITAPLTFGQ 100
GTKVEIKRTV AAPSVFIFPP SDEQLKSGTA SVVCLNNFY PREAKVQMKV 150
DNALQSGNSQ ESVTQDSKD STYLSLSTLT LSKADYERKH VYACEVTHQG 200
LSSPVTKSFN RGE 214

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 150-206 263-323 369-427

22"-96" 150"-206" 263"-323" 369"-427"

Intra-L (C23-C104) 23"-88" 134"-194"

23"-88" 134"-194"

Inter-H-L (CH1 10-CL 126)* 137'-214' 137"-214"

Inter-H-H (h 4, h 5, h 8, h 11)* 225'-225" 226'-226" 229'-229" 232'-232"

*In addition to the isoform A, isoform A/B characterized by an inter-H-H (h 4 - CH1 10)

225'-137" and an inter-H-L (h 4 - CL 126) 225"-214", instead of the inter-H-H (h 4 - h 4)

225'-225" and of one of the two inter-H-L (CH1 10-CL 126) 137"-214";

isoform B characterized by an inter-H-H (h 5 - CH1 10) 226'-137" and an inter-H-L (h 5 - CL 126)

226"-214", instead of the inter-H-H (h 5 - h 5) 226'-226" and of the inter-H-L (CH1 10-CL 126) 137'-214";

* En plus de l'isoforme A, isoforme A/B caractérisée par un inter-H-H (h 4 - CH1 10) 225'-137"

et un inter-H-L (h 4 - CL 126) 225"-214", au lieu de l'inter-H-H (h 4 - h 4) 225'-225" et de

l'un des deux inter-H-L (CH1 10-CL 126) 137"-214";

isoforme B caractérisée par un inter-H-H (h 5 - CH1 10) 226'-137" et un inter-H-L (h 5 - CL 126)

226"-214", au lieu de l'inter-H-H (h 5 - h 5) 226'-226" et de l'inter-H-L (CH1 10-CL 126) 137'-214";

* Además de la isoforma A, isoforma A/B caracterizada por un inter-H-H (h 4 - CH1 10) 225'-137"

y un inter-H-L (h 4 - CL 126) 225"-214", en lugar del inter-H-H (h 4 - h 4) 225'-225" y uno de los

dos inter-H-L (CH1 10-CL 126) 137"-214";

isoforma B caracterizada por un inter-H-H (h 5 - CH1 10) 226'-137" y un inter-H-L (h 5 - CL 126)

226"-214", en lugar del inter-H-H (h 5 - h 5) 226'-226" y del inter-H-L (CH1 10-CL 126) 137'-214";

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

299, 299"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires

complexes fucosylés / glicanos de tipo CHO biantennarios complejos fucosilados.

C-terminal lysine clipping / Coupe de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

449, 449"

milvexianum

milvexian

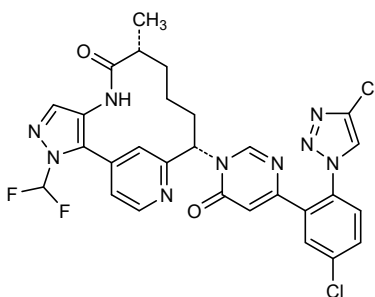
(5*R*,9*S*)-9-{4-[5-chloro-2-(4-chloro-1*H*-1,2,3-triazol-1-yl)phényl]-6-oxypyrimidin-1(6*H*)-yl}-2²-(difluorométhyl)-5-méthyl-2²*H*-3-aza-1(4,2)-pyridina-2(3,4)-pyrazolacyclononaphan-4-one

milvexian

(5*R*,9*S*)-9-{4-[5-chloro-2-(4-chloro-1*H*-1,2,3-triazol-1-yl)phényl]-6-oxypyrimidin-1(6*H*)-yl}-2²-(difluorométhyl)-5-méthyl-2²*H*-3-aza-1(4,2)-pyridina-2(3,4)-pyrazolacyclononaphan-4-one

milvexián

(5*R*,9*S*)-9-{4-[5-cloro-2-(4-cloro-1*H*-1,2,3-triazol-1-il)fenil]-6-oxopirimidin-1(6*H*)-il}-2²-(difluorometil)-5-metil-2²*H*-3-aza-1(4,2)-piridina-2(3,4)-pirazolaciclonoanfan-4-ona

C₂₈H₂₃Cl₂F₂N₉O₂**mipasetaamabum #**

mipasetaamab

immunoglobulin G1-kappa, anti-[*Homo sapiens* AXL (AXL receptor tyrosine kinase, tyrosine-protein kinase receptor UFO)], humanized monoclonal antibody; gamma1 heavy chain humanized (1-451) [VH humanized (*Homo sapiens* IGHV3-30*03 (92.9%) - (IGHD) -IGHJ4*01 (92.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -*Homo sapiens* IGHG1*03, G1m3>G1m17, nG1m1 (CH1 R>K 120 (219) (123-220), hinge 1-15 (221-235), CH2 (236-345), CH3 E12 (361), M14 (363) (346-450), CHS K>del (451)) (123-451)], (225-215')-disulfide with kappa light chain humanized (1'-215') [V-KAPPA humanized (*Homo sapiens* IGKV3-11*01 (84.4%) -IGKJ4*01 (90.9%)) CDR-IMGT [7.3.9] (27-33.51-53.90-98) (1'-108') - *Homo sapiens* IGKC*01 (100%), Km3 A45.1 (154), V101 (192) (109'-215')]; dimer (231-231":234-234")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

mipasétamab

immunoglobuline G1-kappa, anti-[*Homo sapiens* AXL (récepteur tyrosine kinase AXL, récepteur tyrosine-protéine kinase UFO)], anticorps monoclonal humanisé;

chaîne lourde gamma1 humanisée (1-451) [VH humanisé (*Homo sapiens* IGHV3-30*03 (92.9%) -(IGHD) -IGHJ4*01 (92.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) - *Homo sapiens* IGHG1*03, G1m3>G1m17, nG1m1 (CH1 R>K 120 (219) (123-220), charnière 1-15 (221-235), CH2 (236-345), CH3 E12 (361), M14 (363) (346-450), CHS K>del (451)) (123-451)], (225-215')-disulfure avec la chaîne légère kappa humanisée (1'-215') [V-KAPPA humanisé (*Homo sapiens* IGKV3-11*01 (84.4%) -IGKJ4*01 (90.9%)) CDR-IMGT [7.3.9] (27-33.51-53.90-98) (1'-108') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (154), V101 (192) (109'-215')]; dimère (231-231":234-234")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

mipasetamab

immunoglobulina G1-kappa, anti-[*Homo sapiens* AXL (receptor tirosina kinasa AXL, receptor tirosina-proteína kinasa UFO)], anticuerpo monoclonal humanizado; cadena pesada gamma1 humanizada (1-451) [VH humanizado (*Homo sapiens* IGHV3-30*03 (92.9%) -(IGHD) -IGHJ4*01 (92.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -*Homo sapiens* IGHG1*03, G1m3>G1m17, nG1m1 (CH1 R>K 120 (219) (123-220), bisagra 1-15 (221-235), CH2 (236-345), CH3 E12 (361), M14 (363) (346-450), CHS K>del (451)) (123-451)], (225-215')-disulfuro con la cadena ligera kappa humanizada (1'-215') [V-KAPPA humanizado (*Homo sapiens* IGKV3-11*01 (84.4%) -IGKJ4*01 (90.9%)) CDR-IMGT [7.3.9] (27-33.51-53.90-98) (1'-108') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (154), V101 (192) (109'-215')]; dímero (231-231":234-234")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada

QVQLVESGGG VVQPGKSLRL SCAASGFTFS SYGMSWVRQA PGKGLEWVAT 50
ISSGGSYTY Y PDSVKGRFTI SRDNSKNTLY LQMNSLRAED TAVYYCARHP 100
IYYTYDDTMD YWGQGTITVTV SSASTKGPSV FPLAPSSKST SGGTAALGCL 150
VKDYFPEPVTV VSWNSGALTS GVHTFPAVLV SSGLYSLSSV VTPVSSSLGT 200
QTYICNVNHK PSNTKVDKKV EPKSCDKTHT CPPCPAPELL GGPSVFLFPP 250
KPKDITLMISR TPEVTCVVVD VSHEDPEVKF NWYVDGVEVH NAKTKPREEQ 300
YNSTYRVVSV LTVLHQDWLNL GKEYYCKVSN KALPAPIEK ISKAKGQPRE 350
PQVYTLPPSR EEMTKNQVSL TCVKGFPYS DIAVEWESNG QPENNYKTP 400
PVLDSGGSFF LYSKLTVDKS RWQKGNVFSC SVMHEALHNN YTQKSLSLSP 450
6 451

Light chain / Chaîne légère / Cadena ligera

EIVLTQSPGT LSLSPGERAT LSCSASSSVS SGNFHWYQQK PGLAPRLLIY 50
RTSNLASGIP ARFSGSGSGT DFTLTISLLE PEDFAVYYCQ QWSGYPWTFG 100
GGTKLEIKRT VAAPSVFIFP PSDEQLKSGT ASVVCLNNF YPREAKVQWK 150
VDNALQSGNS QESVTEQDSK DSTYLSSTLT TSLKADYKEH KYACEVTHQ 200
GLSSPVTKSF NRGEC 215

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 149-205 266-326 372-430
22"-96" 149"-205" 266"-326" 372"-430"

Intra-L (C23-C104) 23"-89" 135'-195'
23'''-89''' 135'''-195'''

Inter-H-L (h 5-CL 126) 225'-215' 225"-215'"

Inter-H-H (h 11, h 14) 231-231" 234-234"

N-terminal glutaminy cyclization to pyroglutamyl (pE, 5-oxoprolyl)

H VH Q1:

I, I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4 [glycoengineered GalNAcN3 (N-azidoacetyl-D-galactosamine)]:
302, 302"-bis[N-(O-2-(acetilamino)-6-azido-2,6-dideoxy- β-D-galactopyranosyl-(1->4)-
O-[6-deoxy-α-L-galactopyranosyl-(1->6)]-2-(acetilamino)-2-deoxy- β-D-glucopyranosyl]-
L-asparagine]

mipasetamabum uzoptirinum #

mipasetamab uzoptirine

immunoglobulin G1-kappa, anti-[*Homo sapiens* AXL (AXL receptor tyrosine kinase, tyrosine-protein kinase receptor UFO)], humanized monoclonal antibody conjugated to the pyrrolobenzodiazepine (PBD) dimer, SG3199; gamma1 heavy chain humanized (1-451) [VH humanized (*Homo sapiens* IGHV3-30*03 (92.9%) -(IGHD) -IGHJ4*01 (92.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) - *Homo sapiens* IGHG1*03 nG1m1 (CH1 R120>K (219) (123-220), hinge 1-15 (221-235), CH2 (236-345), CH3 E12 (361), M14 (363) (346-450), CHS K>del (451)) (123-451)], (225-215')-disulfide with kappa light chain humanized (1'-215') [V-KAPPA (*Homo sapiens* IGKV3-11*01 (84.4%) -IGKJ4*01 (90.9%)) CDR-IMGT [7.3.9] (27-33.51-53.90-98) (1'-108') - *Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dimer (231-231":234-234")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa; conjugated on the two glycoengineered N84.4, via a spacer and a cleavable valine-alanine linker, to the cytotoxic pyrrolobenzodiazepine (PBD) dimer, SG3199

mipasétamab uzoptirine

immunoglobuline G1-kappa, anti-[*Homo sapiens* AXL (récepteur tyrosine kinase AXL, récepteur tyrosine-protéine kinase UFO)], anticorps monoclonal humanisé conjugué au dimère de pyrrolobenzodiazépine (PDB), SG3199; chaîne lourde gamma1 humanisée (1-451) [VH (*Homo sapiens* IGHV3-30*03 (92.9%) -(IGHD) -IGHJ4*01 (92.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -*Homo sapiens* IGHG1*03 nG1m1 (CH1 R120>K (219) (123-220), charnière 1-15 (221-235), CH2 (236-345), CH3 E12 (361), M14 (363) (346-450), CHS K>del (451)) (123-451)], (225-215')-disulfure avec la chaîne légère kappa humanisée (1'-215') [V-KAPPA (*Homo sapiens* IGKV3-11*01 (84.4%) -IGKJ4*01 (90.9%)) CDR-IMGT [7.3.9] (27-33.51-53.90-98) (1'-108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dimère (231-231":234-234")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa; conjugué sur les deux N84.4 glycomodifiés, via un espaceur et un linker clivable valine-alanine, au dimère pyrrolobenzodiazépine (PBD) cytotoxique, SG3199

mipasetamab uzoptirina

immunoglobulina G1-kappa, anti-[*Homo sapiens* AXL (receptor tirosina kinasa AXL, receptor tirosina-proteína kinasa UFO)], anticuerpo monoclonal humanizado conjugado con el dímero de pirrolobenzodiazepina (PDB), SG3199; cadena pesada gamma1 humanizada (1-451) [VH (*Homo sapiens* IGHV3-30*03 (92.9%) -(IGHD) -IGHJ4*01 (92.3%)) CDR-IMGT [8.8.15] (26-33.51-58.97-111) (1-122) -*Homo sapiens* IGHG1*03 nG1m1 (CH1 R120>K (219) (123-220), bisagra 1-15 (221-235), CH2 (236-345), CH3 E12 (361), M14 (363) (346-450), CHS K>del (451)) (123-451)], (225-215')-disulfuro con la cadena ligera kappa humanizada (1'-215') [V-KAPPA (*Homo sapiens* IGKV3-11*01 (84.4%) -IGKJ4*01 (90.9%)) CDR-IMGT [7.3.9] (27-33.51-53.90-98) (1'-108') - *Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')];

dímero (231-231":234-234")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa; conjugado sobre los dos N84.4 glicomodificados, mediante un espaciador y un conector escindible valina-alanina, al dímero pirrolbenzodiazepina (PBD) citotóxico, SG3199

Heavy chain / Chaîne lourde / Cadena pesada

QVQLVESGGG VVQPGKSLRL SCAASGFTFS SYGMSWVRQA PGKLEWVAT 50
 ISSGGSTYY PDSVKGRFTI SRDNTKNTLY LQMNSLRAED TAVYYCARHP 100
 IYYTYDDTMD YWGQGTITTVV SSASTKGPV FPLAPSSKST SGGTAALGCL 150
 VKDYFPEPVT VSWNSGALTS GVHTFPAVLQ SSGLYSLSSV VTFVSSSLGT 200
 QTYICNVNHR PSNTKVDKRV EPKSCDKTHT CPCCPAPELL GGPVFLFPP 250
 KPKDTLMISR TPEVTCVVVD VSHEDPEVKF NWYVDGVEVH NAKTKPREQ 300
 YNSTYRVVSV LTVLHQDWLN GKEYCKVSN KALPAPIET ISKAKGQPRE 350
 PQVYTLPPSR EEMTKNQSLS TCLVKGFPYS DIAVEWESNG QPENNYKTP 400
 PVLDSGSGFF LYSKLTVDXS RWQGNVFSC SVMHEALHHH YTKQSLSLSP 450
 G 451

Light chain / Chaîne légère / Cadena ligera

EIVLTQSPGT LSLSPGERAT LSCASSSVS SGNEHWYQK PGLAPRLLIY 50
 RTSNLASGIP ARFSGSGSGT DFTLTISSE PEDFAVYYCQ QWSGYPTWFG 100
 GGTGLEIKRT VAAPSVFIFP PSDEQLKSGT ASVCLLNLF YPREAKVQWK 150
 VDNALQSGNS QESVTECDISK DSTYLSSTL TSKADYKXK KVIACVTHQ 200
 GLSPVTKSF NRGE 215

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 149-205 266-326 372-430
 22"-96" 149"-205" 266"-326" 372"-430"
 Intra-L (C23-C104) 23"-89" 135"-195"
 23"-89" 135"-195"

Inter-H-L (h 5-CL 126) 225-215" 225"-215"

Inter-H-H (h 11, h 14) 231-231" 234-234"

N-terminal glutamyl cyclization to pyroglutamyl (pE, 5-oxopropyl)

H V H Q I:

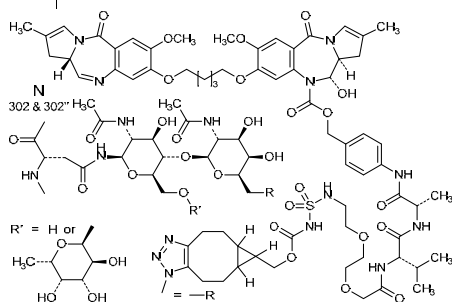
I, I"

Chemically modified N-glycosylation sites / Sites de N-glycosylation chimiquement modifiés /

Posiciones de N-glicosilación modificadas químicamente

H CH2 N84.4:

302, 302"



modoflanerum

modoflaner

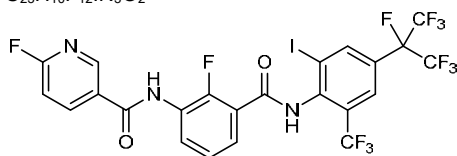
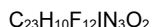
6-fluoro-N-(2-fluoro-3-[[4-(heptafluoropropan-2-yl)-2-iodo-6-(trifluoromethyl)phenyl]carbamoyl]phenyl)pyridine-3-carboxamide

modoflaner

6-fluoro-N-(2-fluoro-3-[[4-(heptafluoropropan-2-yl)-2-iodo-6-(trifluorométhyl)phényl]carbamoyl]phényl)pyridine-3-carboxamide

modoflaner

6-fluoro-N-(2-fluoro-3-[[4-(heptafluoropropan-2-il)-2-iodo-6-(trifluorometil)fenil]carbamoi]fenil)piridina-3-carboxamida

**mosliciguatum**

mosliciguat

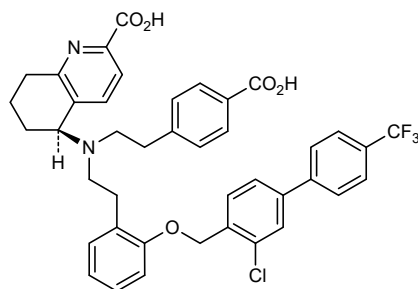
(9^S)-8-[2-(4-carboxyphenyl)ethyl]-2³-chloro-1⁴-(trifluoromethyl)-9^{5,6,7,8}-tetrahydro-4-oxa-8-aza-9(5)-quinolína-1(1),2(1,4),5(1,2)-tribenzenanonaphane-9²-carboxylic acid

mosliciguat

acide (9^S)-8-[2-(4-carboxyphényl)éthyl]-2³-chloro-1⁴-(trifluorométhyl)-9^{5,6,7,8}-tétrahydro-4-oxa-8-aza-9(5)-quinoléína-1(1),2(1,4),5(1,2)-tribenzénanonaphane-9²-carboxylique

mosliciguat

ácido (9^S)-8-[2-(4-carboxifenil)etil]-2³-cloro-1⁴-(trifluorometil)-9^{5,6,7,8}-tetrahidro-4-oxa-8-aza-9(5)-quinoleína-1(1),2(1,4),5(1,2)-tribencenanonafano-9²-carboxílico

**naporafenibum**

naporafenib

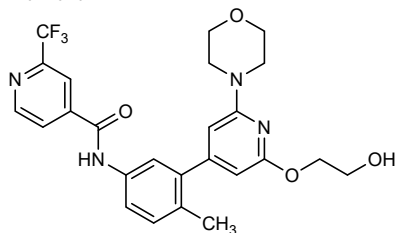
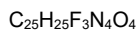
N-{3-[2-(2-hydroxyethoxy)-6-(morpholin-4-yl)pyridin-4-yl]-4-methylphenyl}-2-(trifluoromethyl)pyridine-4-carboxamide

naporafénib

N-{3-[2-(2-hydroxyéthoxy)-6-(morpholin-4-yl)pyridin-4-yl]-4-méthylphényl}-2-(trifluorométhyl)pyridine-4-carboxamide

naporafenib

N-{3-[2-(2-hidroxietoxi)-6-(morfolin-4-il)piridin-4-il]-4-metilfenil}-2-(trifluorometil)piridina-4-carboxamida



nedosiranum

nedosiran

*all-P-ambo-2'-O-methyl-P-thioadenylyl-(3'→5')-2'-O-methyluridylyl-(3'→5')-2'-deoxy-2'-fluoroguanlyl-(3'→5')-2'-O-methyluridylyl-(3'→5')-2'-deoxy-2'-fluorouridylyl-(3'→5')-2'-O-methylguanylyl-(3'→5')-2'-O-methyluridylyl-(3'→5')-2'-deoxy-2'-fluorocytidylyl-(3'→5')-2'-deoxy-2'-fluorouridylyl-(3'→5')-2'-deoxy-2'-fluorouridylyl-(3'→5')-2'-O-methyluridylyl-(3'→5')-2'-deoxy-2'-fluorouridylyl-(3'→5')-2'-O-methyluridylyl-(3'→5')-2'-deoxy-2'-fluorocytidylyl-(3'→5')-2'-O-methyluridylyl-(3'→5')-2'-O-methylguanylyl-(3'→5')-2'-O-methyladenylyl-(3'→5')-2'-O-methylguanylyl-(3'→5')-2'-O-methylcytidylyl-(3'→5')-2'-O-methyladenylyl-(3'→5')-2'-O-methylguanylyl-(3'→5')-2'-O-methylcytidylyl-(3'→5')-2'-O-[[2-(2-{5-[(2-acetamido-2-deoxy-β-D-galactopyranosyl)oxy]pentanamido}ethoxy)ethoxy]methyl]guanylyl-(3'→5')-2'-O-[[2-(2-{5-[(2-acetamido-2-deoxy-β-D-galactopyranosyl)oxy]pentanamido}ethoxy)ethoxy]methyl]adenylyl-(3'→5')-2'-O-[[2-(2-{5-[(2-acetamido-2-deoxy-β-D-galactopyranosyl)oxy]pentanamido}ethoxy)ethoxy]methyl]adenylyl-(3'→5')-2'-O-[[2-(2-{5-[(2-acetamido-2-deoxy-β-D-galactopyranosyl)oxy]pentanamido}ethoxy)ethoxy]methyl]adenylyl-(3'→5')-2'-O-methylguanylyl-(3'→5')-2'-O-methylguanylyl-(3'→5')-2'-O-methylcytidylyl-(3'→5')-2'-O-methyluridylyl-(3'→5')-2'-O-methylguanylyl-(3'→5')-2'-O-methylcytidine duplex with *all-P-ambo-2'-O-methyl-P-thioguanlyl-(5'→3')-2'-O-methyluridylyl-(5'→3')-2'-O-methyladenylyl-(5'→3')-2'-deoxy-2'-fluorocytidylyl-(5'→3')-2'-O-methyladenylyl-(5'→3')-2'-deoxy-2'-fluoroadenylyl-(5'→3')-2'-O-methylcytidylyl-(5'→3')-2'-deoxy-2'-fluoroadenylyl-(5'→3')-2'-O-methylguanylyl-(5'→3')-2'-O-methylguanylyl-(5'→3')-2'-O-methyladenylyl-(5'→3')-2'-deoxy-2'-fluoroadenylyl-(5'→3')-2'-deoxy-2'-fluoroadenylyl-(5'→3')-2'-O-methyladenylyl-(5'→3')-2'-deoxy-2'-fluoroadenylyl-(5'→3')-2'-O-methyluridylyl-(5'→3')-2'-deoxy-2'-fluoroadenylyl-(5'→3')-2'-deoxy-2'-fluoro-*P*-thioguanlyl-(5'→3')-2'-deoxy-2'-fluoro-*P*-thioadenylyl-(5'→3')-2'-deoxy-2'-fluoro-*P*-thiocytidylyl-(5'→3')-methyl hydrogen 2'-O-methyl-5'-oxa-O-5'-carba-5'-uridylate**

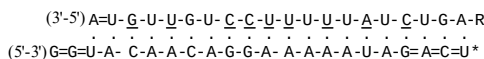
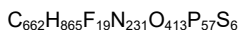
nédosiran

tout-P-ambo-2'-O-méthyl-P-thioadénylyl-(3'→5')-2'-O-méthyluridylyl-(3'→5')-2'-désoxy-2'-fluoroguanlyl-(3'→5')-2'-O-méthyluridylyl-(3'→5')-2'-désoxy-2'-fluorouridylyl-(3'→5')-2'-O-méthylguanylyl-(3'→5')-2'-O-méthyluridylyl-(3'→5')-2'-désoxy-2'-fluorocytidylyl-(3'→5')-2'-désoxy-2'-fluorouridylyl-(3'→5')-2'-désoxy-2'-fluorouridylyl-(3'→5')-2'-O-méthyluridylyl-(3'→5')-2'-désoxy-2'-fluorouridylyl-(3'→5')-2'-O-méthyluridylyl-(3'→5')-2'-désoxy-2'-fluoroadénylyl-(3'→5')-2'-O-méthyluridylyl-(3'→5')-2'-désoxy-2'-fluorocytidylyl-(3'→5')-2'-O-méthyluridylyl-(3'→5')-2'-O-méthylguanylyl-(3'→5')-2'-O-méthyladénylyl-(3'→5')-2'-O-méthylguanylyl-(3'→5')-2'-O-méthylcytidylyl-(3'→5')-2'-O-méthyladénylyl-(3'→5')-2'-O-méthylguanylyl-(3'→5')-2'-O-méthylcytidylyl-(3'→5')-2'-O-méthylcytidylyl-(3'→5')-2'-O-[[2-(2-{5-[(2-acétamido-2-désoxy-β-D-galactopyranosyl)oxy]pentanamido}éthoxy)éthoxy]méthyl]guanylyl-(3'→5')-2'-O-[[2-(2-{5-[(2-acétamido-2-désoxy-β-D-galactopyranosyl)oxy]pentanamido}éthoxy)éthoxy]méthyl]adénylyl-(3'→5')-2'-O-[[2-(2-{5-[(2-acétamido-2-désoxy-β-D-galactopyranosyl)oxy]pentanamido}éthoxy)éthoxy]méthyl]adénylyl-(3'→5')-2'-O-[[2-(2-{5-[(2-acétamido-2-désoxy-β-D-galactopyranosyl)oxy]pentanamido}éthoxy)éthoxy]méthyl]adénylyl-(3'→5')-2'-O-méthylguanylyl-(3'→5')-2'-O-méthylguanylyl-(3'→5')-2'-O-méthylcytidylyl-(3'→5')-2'-O-méthyluridylyl-(3'→5')-2'-O-méthylguanylyl-(3'→5')-2'-O-méthylcytidine

duplex avec acide *tout-P-ambo-2'-O-méthyl-P-thioguanilyl-(5'→3')-2'-O-méthyl-P-thioguanilyl-(5'→3')-2'-O-méthyluridylyl-(5'→3')-2'-O-méthyl-adénylyl-(5'→3')-2'-désoxy-2'-fluorocytidylyl-(5'→3')-2'-O-méthyladénylyl-(5'→3')-2'-désoxy-2'-fluoroadénylyl-(5'→3')-2'-O-méthylcytidylyl-(5'→3')-2'-désoxy-2'-fluoroadénylyl-(5'→3')-2'-O-méthylguanylyl-(5'→3')-2'-O-méthylguanylyl-(5'→3')-2'-O-méthyladénylyl-(5'→3')-2'-désoxy-2'-fluoroadénylyl-(5'→3')-2'-désoxy-2'-fluoroadénylyl-(5'→3')-2'-O-méthyladénylyl-(5'→3')-2'-désoxy-2'-fluoroadénylyl-(5'→3')-2'-O-méthyladénylyl-(5'→3')-2'-désoxy-2'-fluoroadénylyl-(5'→3')-2'-fluoro-*P*-thioguanilyl-(5'→3')-2'-désoxy-2'-fluoro-*P*-thioadénylyl-(5'→3')-2'-désoxy-2'-fluoro-*P*-thiocytidylyl-(5'→3')-hydrogène-2'-O-méthyl-5'-oxa-*O*-5'-carba-5'-uridylyl de méthyle*

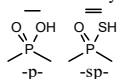
nedosirán

todo-P-ambo-2'-O-metil-P-tiadenilil-(3'→5')-2'-O-metiluridilil-(3'→5')-2'-desoxi-2'-fluoroguanilil-(3'→5')-2'-O-metiluridilil-(3'→5')-2'-desoxi-2'-fluorouridilil-(3'→5')-2'-O-metilguanilil-(3'→5')-2'-O-metiluridilil-(3'→5')-2'-desoxi-2'-fluorocitidilil-(3'→5')-2'-desoxi-2'-fluorocitidilil-(3'→5')-2'-desoxi-2'-fluorouridilil-(3'→5')-2'-O-metiluridilil-(3'→5')-2'-desoxi-2'-fluoro-citidilil-(3'→5')-2'-O-metiluridilil-(3'→5')-2'-O-metilguanilil-(3'→5')-2'-O-metiladenilil-(3'→5')-2'-O-metilguanilil-(3'→5')-2'-O-metilcitidilil-(3'→5')-2'-O-metiladenilil-(3'→5')-2'-O-metilguanilil-(3'→5')-2'-O-metilcitidilil-(3'→5')-2'-O-metilcitidilil-(3'→5')-2'-O-[[2-(2-{5-[(2-acetamido-2-desoxi-β-D-galactopiranosil)oxi]-pentanamido}etoxi)etoxi]metil]guanilil-(3'→5')-2'-O-[[2-(2-{5-[(2-acetamido-2-desoxi-β-D-galactopiranosil)oxi]pentanamido}etoxi)etoxi]metil]adenilil-(3'→5')-2'-O-[[2-(2-{5-[(2-acetamido-2-desoxi-β-D-galactopiranosil)oxi]pentanamido}etoxi)etoxi]metil]adenilil-(3'→5')-2'-O-metilguanilil-(3'→5')-2'-O-metiluridilil-(3'→5')-2'-O-metilguanilil-(3'→5')-2'-O-metilcitidina
*dúplex con todo-P-ambo-2'-O-metil-P-tioguanilil-(5'→3')-2'-O-metil-P-tioguanilil-(5'→3')-2'-O-metiluridilil-(5'→3')-2'-O-metiladenilil-(5'→3')-2'-desoxi-2'-fluorocitidilil-(5'→3')-2'-O-metiladenilil-(5'→3')-2'-desoxi-2'-fluoroadenilil-(5'→3')-2'-O-metilcitidilil-(5'→3')-2'-desoxi-2'-fluoroadenilil-(5'→3')-2'-O-metilguanilil-(5'→3')-2'-O-metilguanilil-(5'→3')-2'-O-metiladenilil-(5'→3')-2'-desoxi-2'-fluoroadenilil-(5'→3')-2'-desoxi-2'-fluoroadenilil-(5'→3')-2'-O-metiladenilil-(5'→3')-2'-desoxi-2'-fluoroadenilil-(5'→3')-2'-desoxi-2'-O-metiluridilil-(5'→3')-2'-desoxi-2'-fluoroadenilil-(5'→3')-2'-desoxi-2'-fluoro-*P*-tioguanilil-(5'→3')-2'-desoxi-2'-fluoro-*P*-tiadenilil-(5'→3')-2'-desoxi-2'-fluoro-*P*-tiocitidilil-(5'→3')-hidrógeno-2'-O-metil-5'-oxa-*O*-5'-carba-5'-uridilato de metilo*

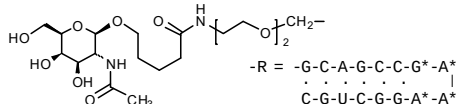
**Legend**

X: 2'-deoxy-2'-fluoronucleotide

X: 2'-O-methylnucleotide



2'-O-substituent of nucleotides A* & G*



nemvaleukinum alfa #**nemvaleukin alfa**

human interleukin 2 fragment (1-59), variant (Cys¹²⁵>Ser⁵¹), fused via peptidyl linker (⁶⁰GG⁶¹) to human interleukin 2 fragment (62-132), fused via peptidyl linker (¹³³GSGGGS¹³⁸) to human interleukin 2 receptor α-chain fragment (139-303), produced in Chinese hamster ovary (CHO) cells, glycosylated; human interleukin 2 (IL-2) (75-133)-peptide [Cys¹²⁵⁽⁵¹⁾>Ser]-mutant (1-59), fused via a G₂ peptide linker (60-61) to human interleukin 2 (IL-2) (4-74)-peptide (62-132) and via a GSG₃S peptide linker (133-138) to human interleukin 2 receptor α-chain (IL2R subunit alpha, IL2Rα, IL2RA) (1-165)-peptide (139-303), produced in Chinese hamster ovary (CHO) cells, glycoform alfa

nemvaleukine alfa

fragment de l'interleukine 2 humaine (1-59), variant (Cys¹²⁵>Ser⁵¹), fusionné via un peptide liant (⁶⁰GG⁶¹) à un fragment de l'interleukine 2 humaine (62-132), fusionné via un peptide liant (¹³³GSGGGS¹³⁸) au fragment de la chaîne α du récepteur de l'interleukine 2 (139-303), produit dans des cellules ovariennes de hamsters chinois (CHO), glycosylé; peptide 75-133 d'interleukine 2 humaine (IL-2) mutante [Cys¹²⁵⁽⁵¹⁾>Ser] (1-59), fusionnée via un peptide G₂ (60-61) au peptide 4-74 d'inter-leukine humaine 2 (IL-2) (62-132) et via un peptide GSG₃S (133-138) au peptide 1-165 de la chaîne α du récepteur de l'interleukine 2 humaine (sous-unité alpha de IL2R, IL2Rα, IL2RA) (139-303), produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

nemvaleukina alfa

fragmento de interleukina 2 humana (1-59), variante (Cis¹²⁵>Ser⁵¹), fusionado a través de un conector peptidil (⁶⁰GG⁶¹) al fragmento de interleukina 2 humana (62-132), fusionado a través de un conector peptidil (¹³³GSGGGS¹³⁸) al fragmento humano de la cadena α del receptor de interleukina 2 (139-303), producido en células ováricas de hámster Chino (CHO), glicosilado; péptido 75-133 de la interleukina 2 humana (IL-2) mutante [Cis¹²⁵⁽⁵¹⁾>Ser] (1-59), fusionado a través de un péptido G₂ (60-61) al péptido 4-74 de la interleukina 2 humana (IL-2) (62-132) y a través de un péptido GSG₃S (133-138) al péptido 1-165 de la cadena α del receptor de interleukina 2 humano (subunidad alfa de IL2R, IL2Rα, IL2RA) (139-303), producido en células ováricas de hámster Chino (CHO), glicosilada alfa

Sequence / Séquence / Secuencia

```
SKNFHLRPRD LISINIVIVL ELKGSETTFM CEYADETATI VEFLNRWITF 50
SQSIISTLTG GSSSTKKTQL QLEHLLLDLQ MILNGINNYK NPKLTRMLTF 100
KIFYMPKKATE LKHLQCLEEE LKPLEEVLNL AQSGGGGSEL CDDDPPEIPH 150
ATFKAMAYKE GTMLNCECKR GFRRIKSGSL YMLCTGNSSH SSWDNQCQCT 200
SSATRNTTKQ VTPQPEEQKE RKTTEMQSPM QPVDQASLPG HCREPPPWEN 250
EATERIYHFV VGQMVVYQCV QGYRALHRGP AESVCKMTHG KTRWTQPQLI 300
CTG 303
```

Mutation / Mutation / Mutación

C⁵¹>S

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
31-116, 141-285, 184-242, 269-301, 166-197 or 166-199, 168-199 or 168-197

Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación

N187, N206, T212

obecabtagenum autoleucelum #

obecabtagene autoleucel

Autologous CD4+ and CD8+ T cells transduced *ex vivo* with a self-inactivating lentiviral vector encoding an anti-CD19 chimeric antigen receptor. Autologous CD4+ and CD8+ T cells, enriched from peripheral blood mononuclear cells (PBMCs) obtained by apheresis transduced *ex vivo* with a non-replicating, self-inactivating (SIN) lentiviral vector, encoding a chimeric antigen receptor (CAR) consisting of an anti-CD19 specific scFv, a CD8 α stalk, transmembrane domain and anchor, a 41BB endodomain and a CD3 zeta endodomain, under the control of a human PGK1 promoter and a modified Woodchuck post transcriptional response element. The vector genome also contains a 5' terminal CMV enhancer/promoter, a Ψ packaging signal, a Rev response element (RRE) and a central polypurine tract (cPPT).

obécabtagène autoleucel

Cellules T CD4+ et CD8+ autologues transduites *ex vivo* avec un vecteur lentiviral auto-inactivable codant pour un récepteur chimérique contre l'antigène CD19. Cellules T CD4+ et CD8+ autologues, enrichies de cellules mononucléées de sang périphérique (PBMC) et obtenues par aphérèse, transduites *ex vivo* avec un vecteur lentiviral non réplcatif et auto-inactivable (SIN), codant pour un récepteur chimérique d'antigène (CAR) consistant en un scFv spécifique de CD19, un fragment extracellulaire, un domaine transmembranaire et la partie d'ancrage de CD8 α , un endo-domaine de 41BB et un endo-domaine de CD3 zêta, sous le contrôle d'un promoteur PGK1 humain et d'un élément de réponse post-transcriptionnel Woodchuck modifié. Le génome du vecteur contient également un promoteur/amplificateur CMV placé en position terminale 5', un signal d'empaquetage Ψ , un élément de réponse Rev (RRE) et un segment central de polypurine (cPPT).

obecabtagén autoleucel

Células T CD4+ y CD8+ autólogas transducidas *ex vivo* con un vector lentiviral auto-inactivante que codifica para un receptor de antígenos quimérico anti-CD19. Células T CD4+ y CD8+ autólogas, enriquecidas a partir de células mononucleares de sangre periférica (PBMCs) obtenidas por aféresis, transducidas *ex vivo* con un vector lentiviral no replicativo, auto-inactivante (SIN), que codifica para un receptor de antígenos quimérico (CAR) que consta de scFv específico anti-CD19, un tallo CD8 α , un dominio transmembranal y de anclaje, un endodominio 41BB y un endodominio CD3 zeta, bajo el control de un promotor PGK1 humano y un elemento de respuesta post transcripcional de marmota modificado. El genoma del vector también contiene un potenciador (enhancer)/promotor de CMV en 5', una señal de empaquetamiento Ψ , un elemento de respuesta Rev (RRE) y un tramo central de polipurinas (cPPT).

obrindatamabum

obrindatamab

immunoglobulin G1 scFv-h-CH2-CH3(scFv) h-CH2-CH3, anti-[*Homo sapiens* CD276 (B7H3, B7-H3, B7RP-2)] and anti-[*Homo sapiens* CD3E (CD3 epsilon)], monoclonal antibody, bispecific; IG scFv-h-CH2-CH3 single chain, anti-CD276 and anti-CD3E (1-504) [scFv-V-kappa-VH anti-CD276 and anti-CD3E (1-240) [V-KAPPA anti-CD276 (*Homo sapiens* IGKV1D-13*01 (85.1%) -IGHJ2*02 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1-107) -8-mer triglycyl-seryl-tetraglycyl linker (108-115) -VH anti-CD3E (*Mus musculus* IGHV10-1*02 (89.9%) -(IGHD) -GHJ3*01 (93.3%)/*Homo sapiens* IGHV3-72*01 (87.0%) -(IGHD) -IGHJ6*01 (90.9%)) CDR-IMGT [8.10.16] (141-148.166-175.214-229) (116-240)] -linker-E-coil (241-277) [6-mer diglycyl-cysteinyl-triglycyl linker (241-246) -28-mer tetrakis(glutamyl-valyl-dialanyl-leucyl-glutamyl-lysyl) E-coil motif (247-274) -3-mer triglycyl linker (275-277)] -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1, G1v14 CH2 A1.3, A1.2, G1v32 CH3 W22 (knob) (278-504) [hinge 6-15 (278-287), CH2 L1.3>A (291), L1.2>A (292) (288-397), CH3 E12 (413), M14 (415), T22>W (423) (398-502), CHS (503-504)]; (243-243')-disulfide with the IG scFv single chain, anti-CD3E and anti-CD276 (1'-274') [scFv V-lambda-VH anti-CD3E and anti-CD276 (1'-240') [V-LAMBDA anti-CD3 (*Mus musculus* IGLV1*01 (81.2%) -IGLJ1*01 (100%)/*Homo sapiens* IGLV7-46*01 (77.9%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (26-34.52-54.91-99) (1'-109') -9-mer tetraglycyl-seryl-tetraglycyl linker (110'-118') -VH anti-CD276 (*Homo sapiens* IGHV3-48*02 (91.8%) -(IGHD) -IGHJ4*01 (85.7%)) CDR-IMGT [8.8.15] (144-151.169-176.215-229) (119'-240')] -6-mer diglycyl-cysteinyl-triglycyl linker (241'-246') -28-mer tetrakis(lysyl-valyl-dialanyl-leucyl-lysyl-glutamyl) K-coil motif (247'-274')]; (283-6"-286-9")-bisdisulfide with the IG h-CH2-CH3 chain, IGHG1*03 nG1m1, G1v14 CH2 A1.3, A1.2, IGHG1v33 CH3 S22, A24, V86 (hole) (1"-227") [hinge 6-15 (1"-10"), CH2 L1.3>A (14), L1.2>A (15) (11"-120"), CH3 E12 (136), M14 (138), T22>S (146), L24>A (148), Y86>V (187), H115>R (215) (121"-225"), CHS (226"-227")], produced in Chinese hamster ovary (CHO) cells, glycoform alfa

obrindatamab

immunoglobuline G1 scFv-h-CH2-CH3(scFv) h-CH2-CH3, anti-[*Homo sapiens* CD276 (B7H3, B7-H3, B7RP-2)] et anti-[*Homo sapiens* CD3E (CD3 epsilon)], anticorps monoclonal, bispécifique; IG scFv-h-CH2-CH3 chaîne unique, anti-CD276 et anti-CD3E (1-504) [scFv-V-kappa-VH anti-CD276 et anti-CD3E (1-240) [V-KAPPA anti-CD276 (*Homo sapiens* IGKV1D-13*01 (85.1%) -IGHJ2*02 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1-107) -8-mer triglycyl-séryl-tétraglycyl linker (108-115) -VH anti-CD3E (*Mus musculus* IGHV10-1*02 (89.9%) -(IGHD) -GHJ3*01 (93.3%)/*Homo sapiens* IGHV3-72*01 (87.0%) -(IGHD) -IGHJ6*01 (90.9%)) CDR-IMGT [8.10.16] (141-148.166-175.214-229) (116-240)] -linker-E-coil (241-277) [-6-mer diglycyl-cystéinyl-triglycyl linker (241-246) -28-mer tétrakis(glutamyl-valyl-dialanyl-leucyl-glutamyl-lysyl) E-coil motif (247-274) -3-mer triglycyl linker (275-277)] -*Homo sapiens* IGHG1*03 h-CH2-CH3 nG1m1, G1v14 CH2 A1.3, A1.2, G1v32 CH3 W22 (knob) (278-504) [charnière 6-15 (278-287), CH2 L1.3>A (291), L1.2>A (292) (288-397), CH3 E12 (413), M14 (415), T22>W (423) (398-502), CHS (503-504)]; (243-243')-disulfure avec la chaîne unique IG scFv, anti-CD3E et

anti-CD276 (1'-274') [scFv V-lambda-VH anti-CD3E et anti-CD276 (1'-240') [V-LAMBDA anti-CD3 (*Mus musculus* IGLV1*01 (81.2%) -IGLJ1*01 (100%)/*Homo sapiens* IGLV7-46*01 (77.9%) -IGLJ3*02 (100%))] CDR-IMGT [9.3.9] (26-34.52-54.91-99) (1'-109') -9-mer tétraglycyl-séryl-tétraglycyl linker (110'-118') -VH anti-CD276 (*Homo sapiens* IGHV3-48*02 (91.8%) -(IGHD) -IGHJ4*01 (85.7%)) CDR-IMGT [8.8.15] (144-151.169-176.215-229) (119'-240') -6-mer diglycyl-cystéinyl-triglycyl linker (241'-246') -28-mer tétrakis(lysyl-valyl-dialanil-leucyl-lysyl-glutamyl) K-coil motif (247'-274')]; (283-6":286-9")-bisdisulfure avec la chaîne IG h-CH2-CH3, IGHG1*03 nG1m1, G1v14 CH2 A1.3, A1.2, IGHG1v33 CH3 S22, A24, V86 (hole) (1"-227") [charnière 6-15 (1"-10"), CH2 L1.3>A (14), L1.2>A (15) (11"-120"), CH3 E12 (136), M14 (138), T22>S (146), L24>A (148), Y86>V (187), H115>R (215) (121"-225"), CHS (226"-227")], produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

obrindatamab

inmunoglobulina G1 scFv-h-CH2-CH3(scFv)_h-CH2-CH3, anti-[*Homo sapiens* CD276 (B7H3, B7-H3, B7RP-2)] y anti-[*Homo sapiens* CD3E (CD3 épsilon)], anticuerpo monoclonal, biespecífico; IG scFv-h-CH2-CH3 cadena única, anti-CD276 y anti-CD3E (1-504) [scFv-V-kappa-VH anti-CD276 y anti-CD3E (1-240) [V-KAPPA anti-CD276 (*Homo sapiens* IGKV1D-13*01 (85.1%) -IGHJ2*02 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1-107) -8-mer triglicil-seril-tetraglicil linker (108-115) -VH anti-CD3E (*Mus musculus* IGHV10-1*02 (89.9%) -(IGHD) -GHJ3*01 (93.3%)/*Homo sapiens* IGHV3-72*01 (87.0%) -(IGHD) -IGHJ6*01 (90.9%)) CDR-IMGT [8.10.16] (141-148.166-175.214-229) (116-240)] -linker-E-coil (241-277) [-6-mer diglicil-cisteinil-triglicil linker (241-246) -28-mer tetrakis(glutamyl-valil-dialanil-leucil-glutamyl-lisil) E-coli motif (247-274) -3-mer triglicil linker (275-277)] -*Homo sapiens* IGHG1*03 h-CH2-CH3 nG1m1, G1v14 CH2 A1.3, A1.2, G1v32 CH3 W22 (knob) (278-504) [bisagra 6-15 (278-287), CH2 L1.3>A (291), L1.2>A (292) (288-397), CH3 E12 (413), M14 (415), T22>W (423) (398-502), CHS (503-504)]; (243-243')-disulfuro con la cadena única IG scFv, anti-CD3E y anti-CD276 (1'-274') [scFv V-lambda-VH anti-CD3E y anti-CD276 (1'-240') [V-LAMBDA anti-CD3 (*Mus musculus* IGLV1*01 (81.2%) -IGLJ1*01 (100%)/*Homo sapiens* IGLV7-46*01 (77.9%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (26-34.52-54.91-99) (1'-109') -9-mer tetraglicil-seril-tetraglicil linker (110'-118') -VH anti-CD276 (*Homo sapiens* IGHV3-48*02 (91.8%) -(IGHD) -IGHJ4*01 (85.7%)) CDR-IMGT [8.8.15] (144-151.169-176.215-229) (119'-240') -6-mer diglicil-cisteinil-triglicil linker (241'-246') -28-mer tetrakis(lisil-valil-dialanil-leucil-lisil-glutamyl) K-coli motif (247'-274')]; (283-6":286-9")-bisdisulfuro con la cadena IG h-CH2-CH3, IGHG1*03 nG1m1, G1v14 CH2 A1.3, A1.2, IGHG1v33 CH3 S22, A24, V86 (hole) (1"-227") [bisagra 6-15 (1"-10"), CH2 L1.3>A (14), L1.2>A (15) (11"-120"), CH3 E12 (136), M14 (138), T22>S (146), L24>A (148), Y86>V (187), H115>R (215) (121"-225"), CHS (226"-227")], producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

ociperlimabum #
ociperlimab

ociperlimab

1. Chain / Chaîne / Cadena: IG scFv h-CH2-CH3 (V-kappa anti-CD276, VH anti-CD3E)	
DIQLTQSPSF	LSASVGDRTV ITCKASQND TNVAVYQKP GKAPKALIYS 50
ASYRYSQVPS	RFSGSGSGTD FTLTISSLPQ EDFATYYCQQ YNNYPFTFGQ 100
GTKLEIKGGG	SGGGGEVQLV ESGGGLVQPG GSLRLSCAAS GFTFSTYAMN 150
WVRQAPKGKL	EWVGRIIRSKY NNYATYYADS VKDRFTISRQ DSKNSLYLQM 200
NSLKTEDTAV	YYCVRHGNFG NSYVSWFAYW GQGLTVTVSS GCGGGGEVAA 250
LEKEVAALEK	EVAALEKEVA ALEKGGGDKT HTCPPCPAPE AAGGSPVFLF 300
PPKPKDTLMI	SRTPEVTCTV VDVSHEDPEV KFNWYVDGVE VHNAKTKPRE 350
EQYNSTYRVV	SVLTVLHQDW LNKKEYKCKV SNKALPAPIE KTISKAKGQP 400
REPQVYTLPP	SREEMTKNQV SLWCLVKGYF PSDIAVEWES NGQPENNYKT 450
TPPVLDSDGS	FFLYSKLTVD KSRWQQGNVF SCSVMHEALH NHYTKSKLSL 500
SPGK	
2. Chain / Chaîne / Cadena: IG scFv (V-lambda anti-CD3E, VH anti-CD276)	
QAVVTQEPST	TVSPGGTVTL TCRSSTGAVT TSNYANWVQQ KPGQAPRGLI 50
GGTNKRAPWT	PARFSGSLLG GKAALTITGA QAEDEADYYC ALWYSNLWVF 100
GGGKTLTVLG	GGGSGGGGEV QLVESGGGLV QPGLSLRLSC AASGFTFSSF 150
GMHWVRQAPG	KGLEWVAYIS SDSSAIYYAD TVKGRFTISR DNAKNSLYLQ 200
MNSLRDIEDTA	VYYCGRGREN IYYGSRLDYW GQGTTVTVSS GCGGGGKVA 250
LKEKVAALKE	KVAALKEKVA ALKE 274
3. Chain / Chaîne / Cadena: IG h-CH2-CH3	
DKTHTCPPCP	APEAAGGPSV FLFPPKPKDT LMISRTPEVT CVWVDVSHED 50
PEVKFNWYVD	GVEVHNAKTK PREEQYNSTY RVVSVLTVLH QDWLNGKEYK 100
CKVSNKALPA	PIEKTISKAK GQPREPQVYT LPDSREEMTK NQVSLSCAVK 150
GFYPDSIAVE	WESNGQPENN YKTTTPVLDV DGSFFLVSKL TVDKSRWQQG 200
NVFSQCSVMHE	ALHNRYTQKS LSLSPGK 227
Post-translational modifications	
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro	
Intra-chain 1 (C23-C104)	23-88 137-213 318-378 424-482
Intra-chain 2 (C23-C104)	22'-90' 140'-214'
Intra-chain 3 (C23-C104)	41"-101" 147"-205"
Inter-chains 1 - 2	243-243'
Inter-chains 1 - 3 (h 11 -h 11)	283-6"
Inter-chains 1 - 3 (h 14 -h 14)	286-9"
N-terminal glutamine cyclization to pyroglutamate (pE, 5-oxoproline)	
V-lambda Q1:	
I'	
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantennarios complejos fucosilados.	
N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación	
CH2 N84.4:	
354, 77"	
C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal	
CHS K2:	
504, 227"	

immunoglobulin G1-kappa, anti-[*Homo sapiens* TIGIT (T-cell immunoreceptor with Ig domain and ITIM, V-set Ig member 9, VSIG9, V-set and transmembrane member 3, VSTM3)], humanized monoclonal antibody; gamma1 heavy chain humanized (1-449) [VH (*Homo sapiens* IGHV3-66*01 (88.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.12] (26-33.51-58.97-108) (1-119) -*Homo sapiens* IGHG1*01 nG1m1, G1m17 (CH1 K120 (216) (120-217), hinge 1-15 (218-232), CH2 (233-342), CH3 E12 (358), M14 (360) (343-447), CHS (448-449)) (120-449)], (222-214')-disulfide with kappa light chain humanized (1'-214') [V-KAPPA (*Homo sapiens* IGKV3-15*01 (87.4%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (228-228":231-231")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

immunoglobuline G1-kappa, anti-[*Homo sapiens* TIGIT (immunorécepteur des lymphocytes T avec domaine Ig et ITIM, membre 9 de l'Ig V-set, VSIG9, membre 3 de l'Ig V-set et région transmembrane, VSTM3)], anticorps monoclonal humanisé;

ociperlimab

chaîne lourde gamma1 humanisée (1-449) [VH (*Homo sapiens*IGHV3-66*01 (88.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.12] (26-33.51-58.97-108) (1-119) -*Homo sapiens*IGHG1*01 nG1m1, G1m17 (CH1 K120 (216) (120-217), charnière 1-15 (218-232), CH2 (233-342), CH3 E12 (358), M14 (360) (343-447), CHS (448-449)) (120-449)], (222-214')-disulfure avec la chaîne légère kappa humanisée (1'-214') [V-KAPPA (*Homo sapiens*IGKV3-15*01 (87.4%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens*IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (228-228":231-231")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

inmunoglobulina G1-kappa, anti-[*Homo sapiens* TIGIT (immunoreceptor de los linfocitos T con dominio Ig y ITIM, miembro 9 de la Ig V-set, VSIG9, miembro 3 de la Ig V-set y región transmembrana, VSTM3)], anticuerpo monoclonal humanizado; cadena pesada gamma1 humanizada (1-449) [VH (*Homo sapiens*IGHV3-66*01 (88.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.12] (26-33.51-58.97-108) (1-119) -*Homo sapiens*IGHG1*01 nG1m1, G1m17 (CH1 K120 (216) (120-217), bisagra 1-15 (218-232), CH2 (233-342), CH3 E12 (358), M14 (360) (343-447), CHS (448-449)) (120-449)], (222-214')-disulfuro con la cadena ligera kappa humanizada (1'-214') [V-KAPPA (*Homo sapiens*IGKV3-15*01 (87.4%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens*IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (228-228":231-231")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
EVQLVESGGG LVQPGGSLRL SCAASGFTFS DYYMYWVRQA PGKGLEWVAY 50
ITKGGGSTYY PDSVKGRFTI SPDNAKNTLY LQMNSLRAED TAVYYCARQT 100
NYDFTMDYWG OGTLVTVSSA STKGFSVFPL APSSKSTSGG TAALGCLVKD 150
YFPEFVTVSW NSGALTSGVH TFFAVLQSSG LYSLSSTVIV FSSSLGTQTY 200
ICNVNHNKFSN TKVDKKVEPK SCDKTHTCPP CPAPELLGGP SVFLFPPKPK 250
DTLMISPTPE VTCVVKVDSH EDPEVKFNHY VDGVEVHNNA TKPREEQYNS 300
TYRVVSVLTV LHQDMLNGKE YKCKVSNKAL FAPIEKTISK AKGQFREPQV 350
YTLPPFSREEM TKNQVSLTCL VKGFYPSDIA VEWESNGQPE NNYKTTPEVL 400
DSGDSFFLYS KLTVDKSRNQ QGNVFCSVM HEALHNHYTQ KSLSLSPGK 449

Light chain / Chaîne légère / Cadena ligera
EIVMTQSPAT LSVSPGERAT LSCAKASQDVG TSVAWYQQPK GPAPRLLIYW 50
ASARHTGIPA RFSGSGSGTE FTLTISLSQS EDFAVYYCQ YSSYPLTEGG 100
GTKVEIKRTV AAPSVEIFPP SDEQLKSGTA SVVCLLNNEY PREAKVQWKV 150
DNALQSGMSQ ESVTEQDSKD STISLSSTLI LSKADYERKK VYACEVTHQG 200
LSSPFTKSFN RGEK 214

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 146-202 263-323 369-427
22'-96" 146"-202" 263"-323" 369"-427"
Intra-L (C23-C104) 23'-88" 134"-194"
23"-88" 134"-194"
Inter-H-L (h 5-CL 126) 222-214' 222"-214"
Inter-H-H (h 11, h 14) 228-228" 231-231"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
299,299"
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

C-terminal lysine clipping:
H CHS K2:
449,449"

omodenbamabum

omodenbamab

immunoglobulin G3-kappa, anti-[*Staphylococcus aureus* SpA (Staphylococcal protein A)], *Homo sapiens* monoclonal antibody;
 gamma3 heavy chain *Homo sapiens* (1-502) [VH (*Homo sapiens* IGHV3-23*04 (90.8%) -(IGHD) -IGHJ5*01 (93.3%)) CDR-IMGT [8.8.18] (26-33.51-58.97-114) (1-125) -*Homo sapiens* IGHG3*01, G3m5* (G3m5,10,11,13,14,26,27) (CH1 (126-223), hinge 1-42 (224-285), CH2 (286-395), CH3 S44 (439), M84 (452), Q98 (474), I101 (477), R115 (490), F116 (491) (396-500), CHS (501-502)) (126-502)], (139-214')-disulfide with kappa light chain *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-39*01 (90.5%) -IGKJ5*01 (91.7%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')];
 dimer (236-236":239-239":245-245":251-251":254-254":260-260":266-266":269-269":275-275":281-281":284-284")-undecakisdisulfide, produced in Chinese hamster ovary (CHO)-K1SV cells, glycoform alfa

omodenbamab

immunoglobuline G3-kappa, anti-[*Staphylococcus aureus* SpA (protéine A de Staphylocoque)], anticorps monoclonal *Homo sapiens*;
 chaîne lourde gamma3 *Homo sapiens* (1-502) [VH (*Homo sapiens* IGHV3-23*04 (90.8%) -(IGHD) -IGHJ5*01 (93.3%)) CDR-IMGT [8.8.18] (26-33.51-58.97-114) (1-125) -*Homo sapiens* IGHG3*01, G3m5* (G3m5,10,11,13,14,26,27) (CH1 (126-223), charnière 1-42 (224-285), CH2 (286-395), CH3 S44 (439), M84 (452), Q98 (474), I101 (477), R115 (490), F116 (491) (396-500), CHS (501-502)) (126-502)], (139-214')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-39*01 (90.5%) -IGKJ5*01 (91.7%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')];
 dimère (236-236":239-239":245-245":251-251":254-254":260-260":266-266":269-269":275-275":281-281":284-284")-undecakisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO)-K1SV, glycoforme alfa

omodenbamab

inmunoglobulina G3-kappa, anti-[*Staphylococcus aureus* SpA (proteína A de Estafilococo)], anticuerpo monoclonal *Homo sapiens*;
 cadena pesada gamma3 *Homo sapiens* (1-502) [VH (*Homo sapiens* IGHV3-23*04 (90.8%) -(IGHD) -IGHJ5*01 (93.3%)) CDR-IMGT [8.8.18] (26-33.51-58.97-114) (1-125) -*Homo sapiens* IGHG3*01, G3m5* (G3m5,10,11,13,14,26,27) (CH1 (126-223), bisagra 1-42 (224-285), CH2 (286-395), CH3 S44 (439), M84 (452), Q98 (474), I101 (477), R115 (490), F116 (491) (396-500), CHS (501-502)) (126-502)], (139-214')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-39*01 (90.5%) -IGKJ5*01 (91.7%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')];
 dímero (236-236":239-239":245-245":251-251":254-254":260-260":266-266":269-269":275-275":281-281":284-284")-undecakisdisulfuro, producido en las células ováricas de hámster chino (CHO)-K1SV, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada	
EVQLVESGGG	LVQPGGSLRL SCAASGFTFS TYGMSWRQA PGKLEWVSS 50
ITGSGRSTFY	ADSVKGRFTI SRDNTNTLS LQMNSLRAED TAVYYCARSP 100
ADIVTGYYPW	WFDLWQGQTL VTVSSASTKG PSVFPLAPCS RSTSGGTAAL 150
GCLVKDYFPE	PVTVSWNSGA LTSVGTFFPA VLQSSGLYSL SSVVTVPSSS 200
LGTQTYTCNV	NHKPSNTKVD KRVELKTPLG DTTHTCPRCP EPKSCDTPPP 250
CPRCPPEKSC	DTPPPCPRCP EPKSCDTPPP CPRCPAPELL GGPSVFLFPP 300
KPKDITLMISR	TPEVTCVVVD VSHEDPEVQF KYYVDGVEVH NAKTKPREEQ 350
YNSTFRVSV	LTVLHQDWLN GKEYKCKVSN KALPAIEKT ISKTKGQPRE 400
PQVYTLPPSR	EEMTKNQVSL TCLVKGFYPS DIAVEWESSG QPENNYNTTP 450
PMLDSGGSFF	LYSKLTVDKS RWQQGNIFSC SVMHEALHNR FTQKSLSLSP 500
GK	502
Light chain / Chaîne légère / Cadena ligera	
DIQMOTSPSS	LSASIGDRVT ITCRASQIN TYLNWYQKQP GKAPRLLIAG 50
ASSLQSGVPS	RFGSGSGSDT FTLTISSLQP EDFATYYCQ ANSFPLTFGQ 100
GTRLEIKRTV	AAPSVFIFPP SDEQLKSGTA SVVCLLNIFY PREAKVQMKV 150
DNALQSGNSQ	ESVTEQDSKD STYLSSTLT LSKADYEKHK VYACEVTHQG 200
LSSPVTKSFN	RGEC 214
Post-translational modifications	
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro	
Intra-H (C23-C104)	22-96 152-208 316-376 422-480
	22"-96" 152"-208" 316"-376" 422"-480"
Intra-L (C23-C104)	23"-88" 134"-194"
	23'''-88''' 134'''-194'''
Inter-H-L (CH1 10-CL 126)	139-214' 139"-214"
Inter-H-H (h1 13, h1 16,	236-236" 239-239"
h2 5, h2 11, h2 14	245-245" 251-251" 254-254"
h3 5, h3 11, h3 14,	260-260" 266-266" 269-269"
h4 5, h4 11, h4 14)	275-275" 281-281" 284-284"
N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación	
H CH2 N84.4:	
352, 352"	
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.	
O-glycosylation sites / Sites de O-glycosylation / Posiciones de O-glicosilación	
h2 S7 h3 S7 h4 S7	
247, 247" 262, 262" 277, 277"	
Sialylated glycans regularly present / glycanes sialylés régulièrement présents / glicanos sialilados regularmente presentes	
C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal	
H CHS K2:	
502, 502"	

onametostatium

onametostat

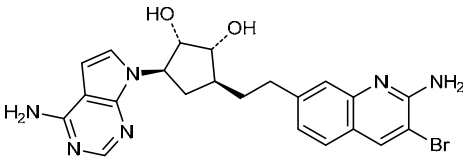
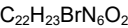
(1*S*,2*R*,3*S*,5*R*)-3-[2-(2-amino-3-bromoquinolin-7-yl)ethyl]-5-(4-amino-7*H*-pyrrolo[2,3-*d*]pyrimidin-7-yl)cyclopentane-1,2-diol

onamétostat

(1*S*,2*R*,3*S*,5*R*)-3-[2-(2-amino-3-bromoquinoléin-7-yl)éthyl]-5-(4-amino-7*H*-pyrrolo[2,3-*d*]pyrimidin-7-yl)cyclopentane-1,2-diol

onametostat

(1*S*,2*R*,3*S*,5*R*)-3-[2-(2-amino-3-bromoquinolein-7-il)etil]-5-(4-amino-7*H*-pirrolo[2,3-*d*]pirimidin-7-il)ciclopentano-1,2-diol



ontasameranum #

ontasameran

Messenger RNA encoding transmembrane phosphatase with tensin homology (TPTE).

	Messenger RNA (mRNA), 5'-capped, encoding codon-optimised transmembrane phosphatase with tensin homology (TPTE; cancer/testis antigen 44) expressed as a fusion protein comprising a secretory signal peptide, TPTE, P2P16 tetanus toxoid-derived helper epitopes and a major histocompatibility complex (MHC) class I transmembrane and cytoplasmic domain (MITD), connected by three glycine/serine-rich (GS) linker peptides, flanked by 5' and 3' untranslated regions and a 3' poly(A) tail.
ontasamérán	ARN messenger codant pour la phosphatase transmembranaire homologue à la tensine (TPTE). Un ARN messenger (ARNm), protégé en 5', codant pour la phosphatase transmembranaire homologue à la tensine (TPTE ; antigène 44 du cancer testiculaire), avec des codons optimisés, exprimé sous la forme d'une protéine de fusion comprenant un peptide signal de sécrétion, TPTE, les épitopes auxiliaires P2P16 dérivés de l'anatoxine tétanique et un domaine transmembranaire et cytoplasmique (MITD) du complexe majeur d'histocompatibilité (CMH) de classe I, reliés par trois peptides de liaison riches en glycine/sérine (GS), flanqués de régions non traduites en 5' et 3' et d'une queue poly(A) en 3'.
ontasamerán	ARN mensajero que codifica para la fosfatasa transmembranal con homología con la tensina (TPTE). ARN mensajero (ARNm), protegido en 5', que codifica para la fosfatasa transmembranal con homología con la tensina (TPTE; antígeno de cáncer/testículo 44), con codones optimizados, expresada como una proteína de fusión que consta de un péptido señal de secreción, TPTE, los epítomos auxiliares P2P16 derivados del toxoide tetánico y un dominio citoplásmico y transmembrana del complejo principal de histocompatibilidad (MHC) de clase I, conectados mediante tres péptidos de enlace ricos en glicina/serina (GS), flanqueado por las regiones 5' y 3' no traducidas y una cola poly(A) en 3'.
oplunofuspum # oplunofusp	<i>Actinomyces viscosus</i> L-methionyl sialidase (exo-α-sialidase) catalytic domain fragment (239-631, 2-394 in the current sequence), fused to human amphiregulin heparin binding domain fragment (25-45, 395-415 in the current sequence) anti-(binding to human heparin and cell surface glycosaminoglycans), produced in <i>Escherichia coli</i>; L-methionyl (1)-sialidase (exo-α-sialidase, EC:3.2.1.18) (<i>Actinomyces viscosus</i>) (239-631)-peptide (containing the sialidase domain) (2-394), fused to human amphiregulin (25-45)-peptide (binding to human heparin and glycosaminoglycans) (395-415), produced in <i>Escherichia coli</i>
oplunofusp	L-méthionyl, fragment du domaine catalytique (239-631, 2-394 dans la séquence actuelle) de la sialidase d'<i>Actinomyces viscosus</i> (exo-α-sialidase), fusionné à un fragment du domaine liant l'héparine de l'amphiréguline humaine (25-45, 395-415 dans la séquence actuelle) anti-(liant aux glycosaminoglycanes de surface cellulaire et à l'héparine, humains), produit dans <i>Escherichia coli</i>;

oplonufusp

L-méthionyl (1)-peptide 239-631 de sialidase (exo- α -sialidase, EC:3.2.1.18) (*Actinomyces viscosus*) (contenant le domaine sialidase) (2-394), fusionné au peptide 25-45 de l'amphiréguline humaine (se liant aux glycosaminoglycanes et à l'héparine) (395-415), produit dans *Escherichia coli*

L-metionil sialidasa(exo- α -sialidasa) *Actinomyces viscosus* fragmento con dominio catalítico (239-631, 2-394 en la secuencia actual), fusionado al fragmento del dominio de unión a la heparina de la anfiregulina humana (25-45, 395-415 en la secuencia actual) anti-(unión a la heparina humana y a los glicosaminoglicanos de superficie celular, humanos), producido in *Escherichia coli*;
L-metionil (1)-péptido 239-631 de sialidasa (exo- α -sialidasa, EC:3.2.1.18) (*Actinomyces viscosus*) (que contiene el dominio de sialidasa) (2-394), fusionado al péptido 25-45 de anfiregulina humana (se une a glicosaminoglicanos y heparina) (395-415), producido en *Escherichia coli*

Sequence / Séquence / Secuencia	
MDHPQATFA PAPDASTELP ASMSQAQHLA ANTATONYRI PAITTA PNGD	50
LLISYDERPK DNGNGGSDAP NPNHIVQRRS TDGKKTWSAP TYHQGTETG	100
KKVGYSDPSY VVDHQGTGTF NFHVKS YDQG WGGSRGGTDP ENRGIIQAEV	150
STSTDNGWTW THRTITADIT KDKPWTARFA ASGGGIQIQH GPAGRLVQQ	200
YTIRTAGGAV QAVSVYSDDH GKTWQAGTFI GTGMDENKVV ELSDGSMLIN	250
SRASDGS3FR KVAHSTDGGQ TWSEPVSDKN LPDSVDNAQI IRAFPNAAF	300
DPAKVLILLH HSPNFRPWSR DRGTISMSCD DGASWTTSKV FHEPFVGYTT	350
IAVQSDGSIG LLEDAHNGA DYGGIWRNF TMNWLGEQCG QKPAKKKKK	400
GRNGFNRRNR KKKNP	415

Post-translational modifications
Disulfide bridge location / Position du pont disulfure / Posición del puente disulfuro
329-389

Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación
none / aucune / ninguna

pacanalotamabum #
pacanalotamab

immunoglobulin scFv-scFv, anti-[*Homo sapiens* TNFRSF17 (TNF receptor superfamily member 17, tumor necrosis factor receptor superfamily, member 17, B cell maturation antigen, BCMA, BCM, TNFRSF13A, CD269)] and anti-[*Homo sapiens* CD3E (CD3 epsilon)], monoclonal antibody single chain, bispecific;
IG scFv-scFv single chain, anti-TNFRSF17 and anti-CD3E (1-504) [scFv-VH-V-kappa anti-TNFRSF17 (1-243) [VH (*Homo sapiens* IGHV1-46*01 (85.7%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) - 15-mer tris(tetraglycyl-seryl) linker (122-136) -V-KAPPA (*Homo sapiens* IGKV1-33*01 (91.6%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (163-168.186-188.225-233) (137-243)] - 6-mer seryl-tetraglycyl-seryl linker (244-249) -scFv-VH-V-lambda anti-CD3E (250-498) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (275-282.300-309.348-363) (250-374) - 15-mer tris(tetraglycyl-seryl) linker (375-389) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9 (415-423.441-443.480-488) (390-498)] - hexahistidine (499-504)], non-glycosylated, produced in Chinese hamster ovary (CHO) cells

pacanalotamab

immunoglobuline scFv-scFv, anti-[*Homo sapiens* TNFRSF17 (membre 17 de la superfamille des récepteurs du TNF, membre 17 de la superfamille des récepteurs du facteur de nécrose tumorale, antigène de maturation de cellule B, BCMA, BCM, TNFRSF13A, CD269)] et anti-[*Homo sapiens* CD3E (CD3 epsilon)], anticorps monoclonal à chaîne unique, bispécifique; IG scFv-scFv chaîne unique, anti-TNFRSF17 et anti-CD3E (1-504) [scFv-VH-V-kappa anti-TNFRSF17 (1-243) [VH (*Homo sapiens* IGHV1-46*01 (85.7%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121)-15-mer tris(tétraglycyl-séryl) linker (122-136) -V-KAPPA (*Homo sapiens* IGKV1-33*01 (91.6%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (163-168.186-188.225-233) (137-243)] -6-mer séryl-tétraglycyl-séryl linker (244-249) -scFv-V-lambda anti-CD3E (250-498) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%))/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (275-282.300-309.348-363) (250-374) -15-mer tris(tétraglycyl-séryl) linker (375-389) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%) CDR-IMGT [9.3.9] (415-423.441-443.480-488) (390-498)] -hexahistidine (499-504)], non-glycosylé, produit dans des cellules ovariennes de hamster chinois (CHO)

pacanalotamab

inmunoglobulina scFv-scFv, anti-[*Homo sapiens* TNFRSF17 (miembro 17 de la superfamilia de los receptores del TNF, miembro 17 de la superfamilia de los receptores del factor de necrosis tumoral, antígeno de maduración de célula B, BCMA, BCM, TNFRSF13A, CD269)] y anti-[*Homo sapiens* CD3E (CD3 épsilon)], anticuerpo monoclonal con cadena única, biespecífico; IG scFv-scFv cadena única, anti-TNFRSF17 y anti-CD3E (1-504) [scFv-VH-V-kappa anti-TNFRSF17 (1-243) [VH (*Homo sapiens* IGHV1-46*01 (85.7%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121)-15-mer tris(tetraglicil-seril) linker (122-136) -V-KAPPA (*Homo sapiens* IGKV1-33*01 (91.6%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (163-168.186-188.225-233) (137-243)] -6-mer seril-tetraglicil-seril linker (244-249) -scFv-V-lambda anti-CD3E (250-498) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%))/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (275-282.300-309.348-363) (250-374) -15-mer tris(tetraglicil-seril) linker (375-389) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%) CDR-IMGT [9.3.9] (415-423.441-443.480-488) (390-498)] -hexahistidina (499-504)], no glicosilado, producido en las células ováricas de hamster chino (CHO)

Heavy chain / Chaîne lourde / Cadena pesada

QVQLVQSGAE VKKPGASVKV SCKASGYTFT NHHIHWVRQA PGQGLEWMGY 5C
INPYPGYHAY NEKFGQRAIM TSDTSTSTVY MELSSLRSED TAVYYCARDG 100
YYRDTVDLDY WGQGTITVTS SGGGGSGGGG SGGGGSDIQM TQSPSSLAS 150
VGDRTVITCQ ASDISNYLN WYQQKPKGAF KLLIYYTSRL HTGVPSRFSG 200
SGSGTDFPT ISSLEPEDIA TYQCQQNTL PWTFGQGTKV EIKSGGGSE 250
VQLVESGGGL VPGGSLKLS CAASGFTFNK YAMNWRQAP GKGLEWVAR 300
RSKNNYATY YADSVKDRFT ISRDDSKNTA YLQMNLRKTE DTAVYYCVRH 350
GNFGNSYISY WAYWGQQLV TVSSGGGGSG GGGSGGGSGQ TVVTQEFSLT 400
VSPGGTITLT CGSSTGAVTS GNYPNWVQK PGQAPRGLIG GTKFLAPGTP 450
ARFSGSLGCG KAAITLSGVQ PEDAEYYCV LWYSNRWVFG GOTKLTPLVH 500
HHHH 504

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-V (C23 C104) 22-96 159-224 271-347 411-479

N-terminal glutamyl cyclization to pyroglutamyl (pE, 5-oxoprolyl)
VH Q1: I

No N-glycosylation sites / pas de sites de N-glycosylation / ningún posición de N-glicosilación
Aglycosylated

pariglasgenum breccaparvovecum #

pariglasgene breccaparvovec

A non-replicating adeno-associated viral vector encoding codon-optimized human glucose-6-phosphatase (G6PC). A recombinant non-replicating adeno-associated viral vector serotype 8 (rAAV8), encoding codon-optimized human glucose-6-phosphatase (G6PC) under the control of the native human G6PC promoter/enhancer and an SV40 polyadenylation sequence, containing a chimeric intron in the 5' UTR between the promoter/enhancer and the transgene, flanked by AAV2 inverted terminal repeats (ITR).

pariglasgène bréccaparvovec

Un vecteur adéno-viral non-répliatif codant pour la glucose-6-phosphatase (G6PC) humaine avec des codons optimisés. Un vecteur recombinant adéno-viral non-répliatif de sérotype 8 (rAAV8), codant pour la glucose-6-phosphatase (G6PC) humaine avec des codons optimisés, sous le contrôle du promoteur/activateur de la G6PC humaine native et d'une séquence de polyadénylation du SV40, contenant un intron chimérique dans l'UTR 5' entre le promoteur/activateur et le transgène, flanqué de répétitions terminales inversées (ITR) AAV2.

pariglasgén breccaparvovec

Un vector viral adeno-asociado, no replicativo, que codifica para la glucosa-6-fosfatasa (G6PC) humana, con codones optimizados. Un vector viral adeno-asociado recombinante de serotipo 8 (rAAV8) no replicativo, que codifica para la glucosa-6-fosfatasa (G6PC) humana, con codones optimizados, bajo el control del promotor/potenciador (enhancer) nativo de la G6PC humana y una secuencia de poliadenilación del SV40, conteniendo un intron quimérico en la región 5' UTR entre el promotor/potenciador y el transgén, flanqueado por repeticiones invertidas terminales (ITR) del AAV2.

pavurutamabum #

pavurutamab

immunoglobulin scFv-scFv-scFc, anti-[*Homo sapiens* TNFRSF17 (TNF receptor superfamily member 17, tumor necrosis factor receptor superfamily, member 17, B cell maturation antigen, BCMA, BCM, TNFRSF13A, CD269)] and anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], monoclonal antibody single chain (scFv)2-scFc, bispecific; IG single chain scFv-scFv-scFc, anti-TNFRSF17 and anti-CD3E (1-986) [scFv-VH-V-kappa anti-TNFRSF17 (1-243) [VH (*Homo sapiens* IGHV1-46*01 G49>C (44) (84.7%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -15-mer tris(tetraglycyl-seryl) linker (122-136)-V-KAPPA (*Homo sapiens* IGKV1-33*01 (91.6%) -IGKJ1*01 Q120>C (236) (91.7%)) CDR-IMGT [6.3.9] (163-168.186-188.225-233) (137-243)] -6 mer seryl-tetraglycyl-seryl linker (244-249) -scFv-VH-V-lambda anti-CD3E (250-498) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (275-282.300-309.348-363) (250-374) -15-mer tris(tetraglycyl-seryl) linker (375-389) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (415-423.441-443.480-488) (390-498)] -4-mer tetraglycyl linker

	(499-502) -scFc (h-CH2-CH3)-(h-CH2-CH3) (503-986) [<i>Homo sapiens</i> IGHG1*03 h-CH2-CH3, nG1m1 (hinge 6-15 (503-512), CH2 R83>C (574), N84.4>G (579), V85>C (584) (513-622), CH3 E12 (638), M14 (640) (623-727), CHS (728-729)) (503-729) -30-mer hexakis(tétraglycyl-séryl) linker (730-759) -<i>Homo sapiens</i> IGHG1*03 h-CH2-CH3, nG1m1 (hinge 6-15 (760-769), CH2 R83>C (831), N84.4>G (836), V85>C (841) (770-879), CH3 E12 (895), M14 (897) (880-984), CHS (985-986)) (760-986)], non-glycosylated, produced in Chinese hamster ovary (CHO) cells
pavurutamab	immunoglobuline scFv-scFv-scFc, anti-[<i>Homo sapiens</i> TNFRSF17 (membre 17 de la superfamille des récepteurs du TNF, membre 17 de la superfamille des récepteurs du facteur de nécrose tumorale, antigène de maturation de cellule B, BCMA, BCM, TNFRSF13A, CD269)] et anti-[<i>Homo sapiens</i> CD3E (CD3 epsilon, Leu-4)], anticorps monoclonal à chaîne unique (scFv)2-scFc, bispécifique; IG à chaîne unique scFv-scFv-scFc, anti-TNFRSF17 et anti-CD3E (1-986) [scFv-VH-V-kappa anti-TNFRSF17 (1-243) [VH (<i>Homo sapiens</i> IGHV1-46*01 G49>C (44) (84.7%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -15-mer tris(tétraglycyl-séryl) linker (122-136)-V-KAPPA (<i>Homo sapiens</i> IGKV1-33*01 (91.6%) -IGKJ1*01 Q120>C (236) (91.7%)) CDR-IMGT [6.3.9] (163-168.186-188.225-233) (137-243)] -6-mer séryl-tétraglycyl-séryl linker (244-249) -scFv-VH-V-lambda anti-CD3E (250-498) [VH (<i>Mus musculus</i> IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/<i>Homo sapiens</i> IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (275-282.300-309.348-363) (250-374) -15-mer tris(tétraglycyl-séryl) linker (375-389) -V-LAMBDA (<i>Homo sapiens</i> IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (415-423.441-443.480-488) (390-498)] -4-mer tétraglycyl linker (499-502) -scFc (h-CH2-CH3)-(h-CH2-CH3) (503-986) [<i>Homo sapiens</i> IGHG1*03 h-CH2-CH3, nG1m1 (503-729) (charnière 6-15 (503-512), CH2 R83>C (574), N84.4>G (579), V85>C (584) (513-622), CH3 E12 (638), M14 (640) (623-727), CHS (728-729)) -30-mer hexakis(tétraglycyl-séryl) linker (730-759) -<i>Homo sapiens</i> IGHG1*03 h-CH2-CH3, nG1m1 (charnière 6-15 (760-769), CH2 R83>C (831), N84.4>G (836), V85>C (841) (770-879), CH3 E12 (895), M14 (897) (880-984), CHS (985-986)) (760-986)], non-glycosylé, produit dans des cellules ovariennes de hamster chinois (CHO)
pavurutamab	immunoglobulina scFv-scFv-scFc, anti-[<i>Homo sapiens</i> TNFRSF17 (miembro 17 de la superfamilia de los receptores del TNF, miembro 17 de la superfamilia de los receptores del factor de necrosis tumoral, antígeno de maduración de célula B, BCMA, BCM, TNFRSF13A, CD269)] y anti-[<i>Homo sapiens</i> CD3E (CD3 épsilon, Leu-4)], anticuerpo monoclonal con cadena única (scFv)2-scFc, biespecífico;

IG con cadena única scFv-scFv-scFc, anti-TNFRSF17 y anti-CD3E (1-986) [scFv-VH-V-kappa anti-TNFRSF17 (1-243) [VH (*Homo sapiens* IGHV1-46*01 G49>C (44) (84.7%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -15-mer tris(tetraglicil-seril) linker (122-136)-V-KAPPA (*Homo sapiens* IGKV1-33*01 (91.6%) -IGKJ1*01 Q120>C (236) (91.7%)) CDR-IMGT [6.3.9] (163-168.186-188.225-233) (137-243)] -6-mer seril-tetraglicil-seril linker (244-249) -scFv-VH-V-lambda anti-CD3E (250-498) [VH (*Mus musculus* IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (275-282.300-309.348-363) (250-374) -15-mer tris(tetraglicil-seril) linker (375-389) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (415-423.441-443.480-488) (390-498)] -4-mer tetraglicil linker (499-502) -scFc (h-CH2-CH3)-(h-CH2-CH3) (503-986) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (503-729) (bisagra 6-15 (503-512), CH2 R83>C (574), N84.4>G (579), V85>C (584) (513-622), CH3 E12 (638), M14 (640) (623-727), CHS (728-729)) -30-mer hexakis(tetraglicil-seril) linker (730-759) -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (bisagra 6-15 (760-769), CH2 R83>C (831), N84.4>G (836), V85>C (841) (770-879), CH3 E12 (895), M14 (897) (880-984), CHS (985-986)) (760-986)], no glicosilado, producido en las células ováricas de hamster chino (CHO)

Heavy chain / Chaîne lourde / Cadena pesada	
QVQLVQSGAE	50
INPYPGYHAY	100
YVRDITVLDY	150
VGDRVTITCQ	200
SGSGTDFITFT	250
VQLVESGGGL	300
RSKYNNYATY	350
GNFNSYISY	400
VSPGGTVTLT	450
ARFSGSLTGG	500
GGDKTHTCPP	550
EDPEVKFNMY	600
YKCKVSNKAL	650
VKGFYPSDIA	700
QGNVFSCSVM	750
GGSGGGGGSD	800
VVVDVSHEDP	850
DWLNGKEYKC	900
QVSLTCLVKG	950
VDKSRWQQGN	986

Post-translational modifications	
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro	
Intra-V (C23-C104)	22-96 159-224 271-347 411-479
Intra-C (C23-C104)	543-603 649-707 800-860 906-964
Intra-CH2 (C83-C85)	574-584 831-841
Inter VH-VL (C49-C120)	44-236
Inter h11 (h 11-h 11)	508-765
Inter h14 (h 14-h 14)	511-768

N-terminal glutaminy cyclization to pyroglutamyl (pE, 5-oxopropyl)
VH Q1: 1

No N-glycosylation sites / pas de sites de N-glycosylation / ningún posición de N-glicosilación
H CH2 N84.4>G:
579, 836
Aglycosylated

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal
CHS K2:
986

pegtibatinasum #**pegtibatinase**

human cystathionine β -synthase fragment (2-413, M¹ cleaved) mutant (C¹⁵>S), homodimer, produced in *Escherichia coli*, pegylated at N⁶ of lysine residues with an average of five 5-[α -methylpoly(oxyethylene)- ω -oxy]-5-oxopentanoyl groups per protein monomer; des-Met¹-[Cys¹⁵>Ser] human cystathionine β -synthase (CBS, beta-thionase, serine sulfhydrase, EC:4.2.1.22) (2-413)-peptide, non-covalent dimer, produced in *Escherichia coli*, substituted with approximately five 5-[α -methylpoly(oxyethylene)- ω -oxy]-5-oxopentanoyl groups (20 kDa each) per protein monomer at N⁶ of lysine residues

pegtibatinase

fragment de la cystathionine β -synthase humaine (2-413, M¹ supprimée) mutant (C¹⁵>S), homodimère, produit dans *Escherichia coli*, pégylé en N⁶ des résidus lysine avec une moyenne de cinq groupes 5-[α -méthylpoly(oxyéthylène)- ω -oxy]-5-oxopentanoyle par monomère de protéine; peptide 2-413 de des-Mét¹-[Cys¹⁵>Sér] cystathionine β -synthase humaine (CBS, bêta-thionase, sérine sulfhydrase, EC: 4.2.1.22), dimère non covalent, produit dans *Escherichia coli*, substitué par environ cinq groupes 5-[α -méthylpoly(oxyéthylène)- ω -oxy]-5-oxopentanoyle (~20 kDa chacun) par monomère de protéine au N⁶ des résidus de lysine

pegtibatinasa

fragmento de la cistationina β -sintasa humana (2-413, M¹ escindido) mutante (C¹⁵>S), homodímero, producido en *Escherichia coli*, pegilada en N⁶ residuos de lisina con una medida de cinco grupos 5-[α -metilpoli(oxietileno)- ω -oxi]-5-oxopentanoilo por monómero de proteína; péptido 2-413 de des-Met¹-[Cis¹⁵>Ser] cistationina β -sintasa humana (CBS, beta-tionasa, serina sulfhidrasa, EC: 4.2.1.22), dímero no covalente, producido en *Escherichia coli*, sustituido con aproximadamente cinco grupos 5-[α -metilpoli(oxietileno)- ω -oxi]-5-oxopentanoilo (~20 kDa cada uno) por proteína monomérica en N⁶ de residuos de lisina

Sequence / Séquence / Secuencia

```
-PSETFQAEV GPTCSHRSG PHSAGSLEK GSPEDKEAKE PLNIRPDAPS 5C
RCTWQLGRFA SESPHRHTAP AKSPKILPDI LKKIGDTFMV RINKIGKKFG 100
LKCELLAKCE FFNAGGSVKD RISLRMIEDA ERDGLKPGD TIIPTSGNT 150
GIGLALAAAV RGYRCIIVMP EKMSSEKVDV LRALGAELVR TPTNARFOSP 20C
ESHVGVAVRL KNEIPNSHIL DQYRNASNPL AHYUTTADEI LQQCDGKLDH 25C
LVASVGTGGT ITGIARKLKE KCPGCRIGV DPEGSILAEP EELNQTQETT 30C
YEVEGIGYDF IPTVLDRITV DKWFKSNDEE APTFARMLIA QEGLLCGGSA 35C
GSTVAVAVKA AQELQEGQRC VVILPDSVRN YMTKFLSDRW MLQKGFLKEE 40C
DLTENKPPWW HLR 415
```

Mutation / Mutation / Mutación

C⁴⁵>S

Post-translational modifications

Removal of Met¹ / Suppression de Met¹ / Eliminación de Met¹

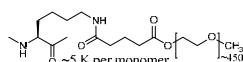
Disulfide bridges location / Position des ponts disulfure / Posición del puentes disulfuro
272-275, 272'-275' (and further variable potential bridges)

Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación
none / aucune / ninguna

Cofactor binding sites / Sites de liaison des cofacteurs / Sitios de unión de cofactores

N⁶-pyridoxylidene 5'-phosphate / 5'-phosphate de N⁶-pyridoxylidene / 5'-fosfato de N⁶-piridoxilideno
K119, K119'
heme B (protoheme IX) / hémé b (protohème IX) / hemo B (protohemo IX)
Cys52-S -Fe(II)- N¹-His65, Cys52'-S -Fe(II)- N¹-His65'

Chemical modification / Modification chimique / Modificación química



pegulicianinum

pegulicianine

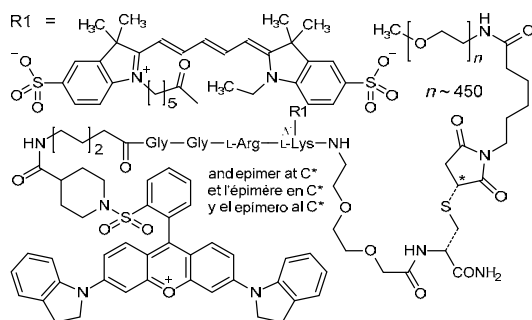
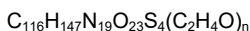
N-[6-(1-{2-[3,6-bis(2,3-dihydro-1*H*-indol-1-yl)xanthylium-9-yl]benzene-1-sulfonyl}piperidine-4-carboxamido)hexanoyl]glycylglycyl-L-arginyl-*N*⁶-(6-{2-[(1*E*,3*E*,5*Z*)-5-(1-ethyl-3,3-dimethyl-5-sulfonato-1,3-dihydro-2*H*-indol-2-ylidene)penta-1,3-dien-1-yl]-3,3-dimethyl-5-sulfonato-3*H*-indol-1-ium-1-yl}hexanoyl)-L-lysyl-[2-(2-aminoethoxy)ethoxy]acetyl-*S*-[(3*RS*)-1-{6-[α-methylpoly(oxyethylene)-ω-amino]-6-oxohexyl}-2,5-dioxopyrrolidin-3-yl]-L-cysteinamide

pégulicianine

N-[6-(1-{2-[3,6-bis(2,3-dihydro-1*H*-indol-1-yl)xanthylium-9-yl]benzène-1-sulfonyl}pipéridine-4-carboxamido)hexanoyl]glycylglycyl-L-arginyl-*N*⁶-(6-{2-[(1*E*,3*E*,5*Z*)-5-(1-éthyl-3,3-diméthyl-5-sulfonato-1,3-dihydro-2*H*-indol-2-ylidène)penta-1,3-diène-1-yl]-3,3-diméthyl-5-sulfonato-3*H*-indol-1-ium-1-yl}hexanoyl)-L-lysyl-[2-(2-aminoéthoxy)éthoxy]acétyl-*S*-[(3*RS*)-1-{6-[α-méthylpoly(oxyéthylène)-ω-amino]-6-oxohexyl}-2,5-dioxopyrrolidin-3-yl]-L-cystéinamide

pegulicianina

N-[6-(1-{2-[3,6-bis(2,3-dihydro-1*H*-indol-1-il)xantilio-9-il]benceno-1-sulfoñil}piperidina-4-carboxamido)hexanoil]glicilglicil-L-arginil-*N*⁶-(6-{2-[(1*E*,3*E*,5*Z*)-5-(1-etil-3,3-dimetil-5-sulfonato-1,3-dihidro-2*H*-indol-2-ilideno)penta-1,3-dien-1-il]-3,3-dimetil-5-sulfonato-3*H*-indol-1-io-1-il}hexanoil)-L-lisil-[2-(2-aminoetoxi)etoxi]acetil-*S*-[(3*RS*)-1-{6-[α-metilpoli(oxietileno)-ω-amino]-6-oxohexil}-2,5-dioxopirrolidin-3-il]-L-cisteinamida

**pelabresibum**

pelabresib

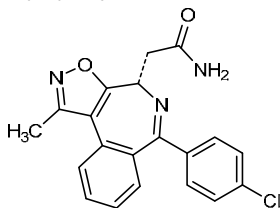
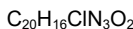
2-[(4*S*)-6-(4-chlorophenyl)-1-methyl-4*H*-[1,2]oxazolo[5,4-*d*][2]benzazepin-4-yl]acetamide

pélabrésib

2-[(4*S*)-6-(4-chlorophényl)-1-méthyl-4*H*-[1,2]oxazolo[5,4-*d*][2]benzazépin-4-yl]acetamide

pelabresib

2-[(4*S*)-6-(4-clorofenil)-1-metil-4*H*-[1,2]oxazolo[5,4-*d*][2]benzazepin-4-il]acetamida

**pemrametostat**

pemrametostat

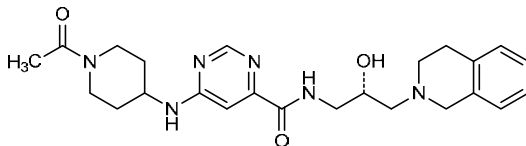
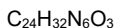
6-[(1-acetylpiperidin-4-yl)amino]-N-[(2S)-3-(3,4-dihydroisoquinolin-2(1H)-yl)-2-hydroxypropyl]pyrimidine-4-carboxamide

pemramétostat

6-[(1-acétylpipéridin-4-yl)amino]-N-[(2S)-3-(3,4-dihydroisoquinoléin-2(1H)-yl)-2-hydroxypropyl]pyrimidine-4-carboxamide

pemrametostat

6-[(1-acetilpiperidin-4-il)amino]-N-[(2S)-3-(3,4-dihidroisoquinolein-2(1H)-il)-2-hidroxiampil]pirimidina-4-carboxamida

**penpulimabum #**

penpulimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* PDCD1 (programmed cell death 1, PD-1, PD1, CD279)], monoclonal antibody; gamma1 heavy chain (1-448) [VH (*Homo sapiens* IGHV3-23*04 (88.7%) -(IGHD) -IGHJ6*01 (90.9%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) - *Homo sapiens* IGHG1*01 G1m17,1, G1v14 CH2 A1.3, A1.2 (CH1 K120 (215) (119-216), hinge 1-15 (217-231), CH2 L1.3>A (235), L1.2>A (236), G1>A (238) (232-341), CH3 D12 (357), L14 (359) (342-446), CHS (447-448)) (119-448)], (221-214')-disulfide with kappa light chain (1'-214') [V-KAPPA (*Mus musculus* IGKV14-111*01 (86.3%) -IGKJ5*01 (100%)/*Homo sapiens* IGKV1-16*01 (80%) -IGKJ2*01 (81.8%) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107) -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (227-227":230-230")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

penpulimab

immunoglobuline G1-kappa, anti-[*Homo sapiens* PDCD1 (protéine 1 de mort cellulaire programmée, PD-1, PD1, CD279)], anticorps monoclonal;

chaîne lourde gamma1 (1-448) [VH (*Homo sapiens* IGHV3-23*04 (88.7%) -(IGHD) -IGHJ6*01 (90.9%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens* IGHG1*01 G1m17,1, G1v14 CH2 A1.3, A1.2 (CH1 K120 (215) (119-216), charnière 1-15 (217-231), CH2 L1.3>A (235), L1.2>A (236), G1>A (238) (232-341), CH3 D12 (357), L14 (359) (342-446), CHS (447-448)) (119-448)], (221-214')-disulfure avec la chaîne légère kappa (1'-214') [V-KAPPA (*Mus musculus* IGKV14-111*01 (86.3%) -IGKJ5*01 (100%)/*Homo sapiens* IGKV1-16*01 (80%) -IGKJ2*01 (81.8%) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]]; dimère (227-227":230-230")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

penpulimab

inmunoglobulina G1-kappa, anti-[*Homo sapiens* PDCD1 (proteína 1 de muerte celular programada, PD-1, PD1, CD279)], anticuerpo monoclonal; cadena pesada gamma1 (1-448) [VH (*Homo sapiens* IGHV3-23*04 (88.7%) -(IGHD) -IGHJ6*01 (90.9%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens* IGHG1*01 G1m17,1, G1v14 CH2 A1.3, A1.2 (CH1 K120 (215) (119-216), bisagra 1-15 (217-231), CH2 L1.3>A (235), L1.2>A (236), G1>A (238) (232-341), CH3 D12 (357), L14 (359) (342-446), CHS (447-448)) (119-448)], (221-214')-disulfuro con la cadena ligera kappa (1'-214') [V-KAPPA (*Mus musculus* IGKV14-111*01 (86.3%) -IGKJ5*01 (100%)/*Homo sapiens* IGKV1-16*01 (80%) -IGKJ2*01 (81.8%) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]]; dímero (227-227":230-230")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada

EVQLVESGGG LVQPGGSLRL SCAASGFAFS SYDMSWVRQA PGKGLDWAT 050
ISGGGRITYY PDSVKGRFTI SRDNSKNNLY LQMSLRAED TALYYCANRY 100
GEAWFAYWQO GTLVTVSAS TKGPSVFPLA PSSKSTSGGT AALGCLVKDY 150
FPEPVTVSWN SGALTSQVHT FPAVLQSSGL YSLSSVVTVP SSSLGTQTYI 200
CNVNHKPSNT KVDKKVEPKS CDKHTCTPPC PAPEAAGAPS VFLFPPKPKD 250
TLMISRTPEV TCVVVDVSHS DPEVKFNWYV DGEVHNNAKT KPREEQYNST 300
YRVVSVLTVL HQDWLNGKEY KCKVSNKALP APIEKTISKA KGQPREPQVY 350
TLPPSRDEL TKNQVSLTCLV KGFYPSDIAV EWESNGQPEN NYKTTTPPVL 400
SDGSFFLYSK LTVDKSRWQO GNVFSCSVMH EALHNHYTQK SLSLSPGK 448

Light chain / Chaîne légère / Cadena ligera

DIQMTQSPSS MSASVGDRTV FTRASQDIN TYLSWFQKQP GKSPKTLIYR 050
ANRLVSGVPS RFSGSQSGQD YTLTISSLQP EDMATYYCLQ YDEFPLTFGA 100
GTKLELKRTV AAPSVFIFPP SDEQLKSGTA SVVCLLNINFP PREAKVQWKV 150
DNALQSGNSQ ESVTEQDSKD STYLSSTLT LSKADYEHKK VYACEVTHQG 200
LSSPVTKSFN RGEK 214

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 145-201 262-322 368-426
22"-96" 145"-201" 262"-322" 368"-426"

Intra-L (C23-C104) 23"-88" 134"-194"
23"-88" 134"-194"

Inter-H-L (h 5-CL 126) 221-214' 221"-214"

Inter-H-H (h 11, h 14) 227-227" 230-230"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

298, 298"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantennarios complejos fucosilados.

C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

448, 448"

pidnarulexum

pidnarulex

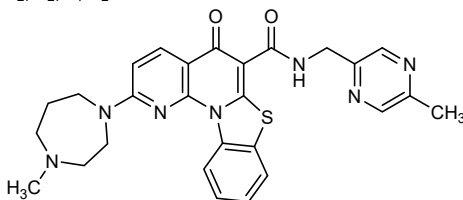
2-(4-methyl-1,4-diazepan-1-yl)-N-[(5-methylpyrazin-2-yl)methyl]-5-oxo-5H-[1,3]benzothiazolo[3,2-a][1,8]naphthyridine-6-carboxamide

pidnarulex

2-(4-méthyl-1,4-diazépan-1-yl)-N-[(5-méthylpyrazin-2-yl)méthyl]-5-oxo-5H-[1,3]benzothiazolo[3,2-a][1,8]naphtyridine-6-carboxamide

pidnarulex

2-(4-metil-1,4-diazepan-1-il)-N-[(5-metilpirazin-2-il)metil]-5-oxo-5H-[1,3]benzotiazolo[3,2-a][1,8]naftiridina-6-carboxamida

 $C_{27}H_{27}N_7O_2S$ **pimivalimabum #**

pimivalimab

immunoglobulin G4-kappa, anti-[*Homo sapiens* PDCD1 (programmed cell death 1, PD-1, PD1, CD279)], *Homo sapiens* monoclonal antibody; gamma4 heavy chain *Homo sapiens* (1-445) [VH (*Homo sapiens* IGHV1-46*01 (96.9%) -(IGHD) - IGHJ5*02 (100%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens* IGHG4*01 (CH1 (119-216), hinge 1-12 S10>P (226) (217-228), CH2 (229-338), CH3 (339-443), CHS (444-445)) (119-445)], (132-214')-disulfide with kappa light chain *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-5*01 (97.9%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (224-224":227-227")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

pimivalimab

immunoglobuline G4-kappa, anti-[*Homo sapiens* PDCD1 (protéine 1 de mort cellulaire programmée, PD-1, PD1, CD279)], anticorps monoclonal *Homo sapiens*; chaîne lourde gamma4 *Homo sapiens* (1-445) [VH (*Homo sapiens* IGHV1-46*01 (96.9%) -(IGHD) - IGHJ5*02 (100%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens* IGHG4*01 (CH1 (119-216), charnière 1-12 S10>P (226) (217-228), CH2 (229-338), CH3 (339-443), CHS (444-445)) (119-445)], (132-214')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-5*01 (97.9%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (224-224":227-227")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

pimivalimab

inmunoglobulina G4-kappa, anti-[*Homo sapiens* PDCD1 (proteína 1 de muerte celular programada, PD-1, PD1, CD279)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma4 *Homo sapiens* (1-445) [VH (*Homo sapiens* IGHV1-46*01 (96.9%) -(IGHD) -IGHJ5*02 (100%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens* IGHG4*01 (CH1 (119-216), bisagra 1-12 S10>P (226) (217-228), CH2 (229-338), CH3 (339-443), CHS (444-445)) (119-445)], (132-214')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-5*01 (97.9%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214'')]; dímero (224-224":227-227")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
QVQLVQSGAE VKKPGASVKV SCKASGYTFP SYMHMVRQA PGQGLEWMGI 50
INPEGGSTAY AQKFQGRVTM TRDTSTSTVY MELSSLRSED TAVYYCARGG 100
TYDYTYWGQ GTLVTVSSAS TKGPSVFPLA PCSRSTSEST AALGCLVKDY 150
FPEPVTVSWN SGALTSGVHT FPAVLQSSGL YSLSSVWVTP SSSLGKTITY 200
CNVDHKPSNT KVDKRVESKY GPCCPPCPAP EFLGSPSVFL FPPKPKDTLM 250
ISRTPVETCV VVDVSQEDPE VQFNWYDGV EVHNAKTKPR EEQFNSTYRV 300
VSVLTVLHQD WLNQKEYKCK VSNKGLPSSI EKTISKAKGQ PREPQVYTL 350
PSQEEMTKNQ VSLTCLVKGF YPSDIAVEWE SNGQPENNYK TTPPVLDSDG 400
SFFLYSRLTV DKSRWQEGNV FSCSVMEAL HNHYTQKSLS LSLGK 445

Light chain / Chaîne légère / Cadena ligera
DIQMTQSPST LSASVGDRVT ITCRASQSI SWLAWYQQKPK GKAPKLLIYE 50
ASSLESGVPS RFGSGSGSGTE FTLTISSLQP DDFATYYCQ YNSFPPPTFGG 100
GTKVEIKRTV AAPSVFIFPP SDEQLKSGTA SVVCLLNNFY PREAKVKQWK 150
DNALQSGNSQ ESVTEQDSKD STYLSLSTLT LSKADYEKHK VYACEVTHQG 200
LSSPVTKSFN RGEK 214

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 145-201 259-319 365-423
22"-96" 145"-201" 259"-319" 365"-423"
Intra-L (C23-C104) 23'-88' 134'-194'
23'''-88''' 134'''-194'''
Inter-H-L (h 10-CL 126) 132-214' 132"-214"
Inter-H-H (h 8, h 11) 224-224" 227-227"

N-terminal glutaminyl cyclization to pyroglutamyl (pE, 5-oxoprolyl)
H VH Q1:
I, I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
295, 295"
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenaricos complejos fucosilados.

C-terminal lysine clipping:
H CHS K2:
445, 445"

pirepematum
pirepemat

(3S)-3-(2,3-difluorophenyl)-3-methoxypyrrolidine

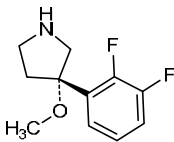
pirépémat

(3S)-3-(2,3-difluorophényl)-3-méthoxypyrrolidine

pirepemat

(3S)-3-(2,3-difluorofenil)-3-metoxipirrolidina

C₁₁H₁₃F₂NO



pomulmeranum #

pomulmeran

Messenger RNA encoding cystic fibrosis transmembrane conductance regulator (CFTR).
 Messenger RNA (mRNA), 5' capped, encoding codon-optimised human cystic fibrosis transmembrane conductance regulator (CFTR; channel conductance-controlling ATPase, cAMP-dependent chloride channel, ATP-binding cassette sub-family C member 7, ABCC7), flanked by 5' and 3' untranslated regions and a 3' poly(A) tail (200-1000 A).

pomulméran

ARN messager codant pour le régulateur de conductance transmembranaire de la mucoviscidose (CFTR).
 ARN messager (ARNm), protégé en 5', codant pour le régulateur de conductance transmembranaire de la mucoviscidose humaine avec des codons optimisés (CFTR ; canal ATPase contrôlant la conductance, canal chlorure dépendant de l'AMPc, sous-famille C membre 7, ABCC7), flanqué de régions 5' et 3' non traduites et d'une queue 3' poly(A) (200-1000 A).

pomulmerán

RNA mensajero que codifica para el regulador de la conductancia transmembrana de la fibrosis quística (CFTR).
 RNA mensajero (RNAm), protegido en 5', que codifica para el regulador de la conductancia transmembrana de la fibrosis quística (CFTR; canal ATPasa controlador de la conductancia, canal de cloro dependiente de AMPc, subfamilia C miembro 7 del casete de unión a ATP, ABCC7) con los codones optimizados, flanqueado por regiones 5' y 3' no traducidas y una cola poly(A) (200-1000 A) en 3'.

posoleucelum

posoleucel

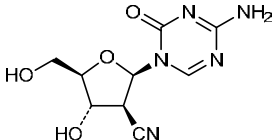
Allogeneic human leukocyte antigen (HLA)-matched virus-specific T cells reactive to cytomegalovirus (CMV), EBV, BK virus, adenovirus and human herpesvirus-6 (HHV-6). The cells are derived from peripheral blood mononuclear cells (PBMCs) from donors seropositive for all five of these viruses.

The cells are expanded and co-cultured with 15-mer peptides overlapping by 11 amino acids spanning antigenic proteins from polyomaviruses BK and JC (VP1 and large T), from adenovirus (hexon and penton), from cytomegalovirus (IE1 and pp65), from Epstein-Barr virus (LMP2, EBNA1, BZLF1), and from human herpesvirus-6 (U90, U11 and U14). The cells have specific anti-virus reactivity for each virus above a pre-specified potency threshold, and predominantly express CD3 (generally >95%) with a mixture of CD4+ (average 60%) and CD8+ (average 34%) subsets, including both central (CD45RA-/62L+/CCR7+) and effector (CD45RA-/62L-/CCR7-) memory markers.

posoleucel

Lymphocytes T allogéniques, compatibles pour les antigènes leucocytaires humains (HLA) et spécifiques pour le cytomégalovirus (CMV), l'EBV, le virus BK, l'adénovirus et le virus herpès humain 6 (HHV-6). Les cellules sont dérivées de cellules mononucléaires du sang périphérique (PBMC) provenant de donneurs séropositifs pour ces cinq virus.

Les cellules sont amplifiées et cultivées avec des peptides de 15 résidus se chevauchant par 11 acides aminés couvrant les protéines antigéniques du polyomavirus BK et JC (VP1 et grand T), de l'adénovirus (hexon et penton), du cytomégalovirus (IE1 et pp65), du virus d'Epstein-Barr (LMP2, EBNA1, BZLF1), et du virus herpès humain 6 (U90, U11 et U14).

	Les cellules ont une réactivité antivirus spécifique pour chacun des virus au-delà d'un seuil prédéfini, et expriment principalement CD3 (généralement >95%) avec un mélange de sous-populations de CD4+ (60% en moyenne) et de CD8+ (34% en moyenne), comprenant des marqueurs des mémoires centrale (CD45RA-/62L+/CCR7+) et effectrice (CD45RA-/62L-/CCR7-).
posoleucel	Células T alogénicas específicas de virus con compatibilidad para el antígeno leucocitario humano (HLA) reactivas frente a citomegalovirus (CMV), EBV, virus BK, adenovirus y herpesvirus humano 6 (HHV-6). Las células se obtienen a partir de células mononucleares de sangre periférica (PBMCs) de donantes seropositivos para estos cinco virus. Las células tienen una reactividad antivirus específica para cada de estos virus por encima de un umbral de potencia preestablecido, y se expanden y co-cultivan con péptidos de 15 residuos que solapan en 11 amino ácidos cubriendo proteínas antigénicas de poliomavirus BK y JC (VP1 y T grande), de adenovirus (hexón y pentón), de citomegalovirus (IE1 y pp65), de virus de Epstein-Barr (LMP2, EBNA1, BZLF1), y de herpesvirus humano 6 (U90, U11 and U14). Las células expresan predominantemente CD3 (generalmente >95%) con una mezcla de subtipos CD4+ (60% como media) y CD8+ (34% como media), incluyendo marcadores de memoria central (CD45RA-/62L+/CCR7+) y efectora (CD45RA-/62L-/CCR7-).
radgocitabinum radgocitabine	4-amino-1-(2-cyano-2-deoxy-β-D-arabinofuranosyl)pyrimidin-2(1<i>H</i>)-one; 2'-cyano-2'-deoxy-D-<i>arabino</i>-cytidine
radgocitabine	4-amino-1-(2-cyano-2-désoxy-β-D-arabinofuranosyl)pyrimidin-2(1<i>H</i>)-one; 2'-cyano-2'-désoxy-D-<i>arabino</i>-cytidine
radgocitabina	4-amino-1-(2-ciano-2-desoxi-β-D-arabinofuranosil)pirimidin-2(1<i>H</i>)-ona; 2'-ciano-2'-desoxi-D-<i>arabino</i>-citidina
	$C_{10}H_{12}N_4O_4$ 
rebonuputemcelum rebonuputemcel	Allogenic human adult nucleus pulposus cells. The cells are derived from the central region of intervertebral discs and isolated from the tissue using mechanical and enzymatic (collagenase) procedures. Isolated cells are expanded in culture involving

	adherent and non- adherent conditions until clusters of cells, 100-200 µm in diameter, are generated. Cells comprising the cell substance are dissociated out of these clusters and stored frozen. Cells (on average >95%) express the surface markers CD73, CD90, CD44 and HLA-ABC, and release proteoglycan, collagen and anti-inflammatory cytokines. Cells (on average <3%) lack expression of CD24 (classically associated with native nucleus pulposus cells), CD34, CD45, CD271, Stro-1 or HLA-DR/DP/DQ.
rébonuputemcel	Cellules allogènes du noyau pulpeux humain adulte. Les cellules sont dérivées de la région centrale des disques intervertébraux et isolées du tissu à l'aide de procédures mécaniques et enzymatiques (collagénase). Les cellules isolées sont multipliées en culture dans des conditions adhérentes et non adhérentes jusqu'à ce que des groupes de cellules, de 100 à 200 µm de diamètre, soient générés. Les cellules composant la substance cellulaire sont dissociées de ces amas et conservées congelées. Les cellules (en moyenne > 95%) expriment les marqueurs de surface CD73, CD90, CD44 et HLA-ABC, et libèrent du protéoglycane, du collagène et des cytokines anti-inflammatoires. Les cellules (en moyenne <3%) manquent d'expression de CD24 (classiquement associé aux cellules pulpeuses du noyau natif), CD34, CD45, CD271, Stro-1 ou HLA-DR/DP/DQ.
rebonuputemcel	Células alogénicas del núcleo pulposo adulto humano. Las células son derivadas de la región central de los discos intervertebrales y se aíslan del tejido usando procedimientos mecánicos y enzimáticos (colagenasa). Las células aisladas se expanden en cultivo en condiciones adherentes y no adherentes hasta que se generan grumos de células de 100-200 mm de diámetro. Las células que componen el principio activo se disocian de estos grumos y se almacenan congeladas. >95% de las células expresan los marcadores de superficie CD73, CD90, CD44 y HLA-ABC, y liberan proteoglicano, colágeno y citoquinas anti-inflamatorias. <3% de las células expresan CD24 (asociado clásicamente con células del núcleo pulposo nativo), CD34, CD45, CD271, Stro-1 o HLA-DR/DP/DQ.
recaticimabum # recaticimab	immunoglobulin G1-kappa, anti-[<i>Homo sapiens</i> PCSK9 (proprotein convertase subtilisin/kexin type 9, neural apoptosis-regulated convertase 1, NARC1, NARC-1, proprotein convertase 9, PC9)], monoclonal antibody;

	gamma1 heavy chain (1-451) [VH (<i>Homo sapiens</i> IGHV1-2*02 (89.8%) -(IGHD) -IGHJ6*01 (100%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -<i>Homo sapiens</i> IGHG1*01, G1m17,1, G1v21 CH2 Y15.1, T16, E18 (CH1 K120 (218) (122-219), hinge 1-15 (220-234), CH2 M15.1>Y (256), S16>T (258), T18>E (260) (235-344), CH3 D12 (360), L14 (362) (345-449), CHS (450-451)) (122-451)], (224-219')-disulfide with kappa light chain (1'-219') [V-KAPPA (<i>Mus musculus</i> IGKV8-21*01 (85%) -IGKJ4*01 (91.7%)/<i>Homo sapiens</i> IGKV4-1*01 (76.8%) -IGKJ2*02 (100%)) CDR-IMGT [12.3.8] (27-38.56-58.95-102) (1'-112') -<i>Homo sapiens</i> IGKC*01 (100%) Km3 A45.1 (158), V101 (196) (113-219')]; dimer (230-230":233-233")-bisdisulfide, produced in a Chinese hamster ovary (CHO) cell line (CHOK1SV GS-KO), glycoform alfa
récatcimab	immunoglobuline G1-kappa, anti-[<i>Homo sapiens</i> PCSK9 (proprotéine convertase subtilisine/kexine type 9, convertase 1 régulée par l'apoptose neuronale, NARC1, NARC-1, proprotéine convertase 9, PC9)], anticorps monoclonal; chaîne lourde gamma1 (1-451) [VH (<i>Homo sapiens</i> IGHV1-2*02 (89.8%) -(IGHD) -IGHJ6*01 (100%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -<i>Homo sapiens</i> IGHG1*01, G1m17,1, G1v21 CH2 Y15.1, T16, E18 (CH1 K120 (218) (122-219), charnière 1-15 (220-234), CH2 M15.1>Y (256), S16>T (258), T18>E (260) (235-344), CH3 D12 (360), L14 (362) (345-449), CHS (450-451)) (122-451)], (224-219')-disulfure avec la chaîne légère kappa (1'-219') [V-KAPPA (<i>Mus musculus</i> IGKV8-21*01 (85%) -IGKJ4*01 (91.7%)/<i>Homo sapiens</i> IGKV4-1*01 (76.8%) -IGKJ2*02 (100%)) CDR-IMGT [12.3.8] (27-38.56-58.95-102) (1'-112') -<i>Homo sapiens</i> IGKC*01 (100%) Km3 A45.1 (158), V101 (196) (113-219')]; dimère (230-230":233-233")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHOK1SV GS-KO, glycoforme alfa
recaticimab	inmunoglobulina G1-kappa, anti-[<i>Homo sapiens</i> PCSK9 (proproteína convertasa subtilisina/kexina tipo 9, convertasa 1 regulada por la apoptosis neuronal, NARC1, NARC-1, proproteína convertasa 9, PC9)], anticuerpo monoclonal; cadena pesada gamma1 (1-451) [VH (<i>Homo sapiens</i> IGHV1-2*02 (89.8%) -(IGHD) -IGHJ6*01 (100%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -<i>Homo sapiens</i> IGHG1*01, G1m17,1, G1v21 CH2 Y15.1, T16, E18 (CH1 K120 (218) (122-219), bisagra 1-15 (220-234), CH2 M15.1>Y (256), S16>T (258), T18>E (260) (235-344), CH3 D12 (360), L14 (362) (345-449), CHS (450-451)) (122-451)], (224-219')-disulfuro con la cadena ligera kappa (1'-219') [V-KAPPA (<i>Mus musculus</i> IGKV8-21*01 (85%) -IGKJ4*01 (91.7%)/<i>Homo sapiens</i> IGKV4-1*01 (76.8%) -IGKJ2*02 (100%)) CDR-IMGT [12.3.8] (27-38.56-58.95-102) (1'-112') -<i>Homo sapiens</i> IGKC*01 (100%) Km3 A45.1 (158), V101 (196) (113-219')]; dímero (230-230":233-233")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHOK1SV GS-KO, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada	
QVQLVQSGAE VVKPGASVKV SCKASGYTFT DYWMHWVRQA PGQGLEWMGY	50
INPSSSGFTKY HQNFKDRVTM TRDTSISTAY MELSRLRSDD TAVYYCARQY	100
DYDEDMYFDV WGQGTITVTVS SASTKGPSVF PLAPSSKSTS GGTAALGLV	150
KDYFPEPVTV SWNSGALTSG VHTFPAVLQS SGLYSLSSVY TVPSSSLGTQ	200
TYICNVNHKP SNTKVDKKVE PKSCDKTHTC PPCPAPELLG GPSVFLFPKP	250
PKDTLYITRE PEVTCVVVDV SHEDPEVKFN WYVDGVEVHN AKTKPREEQY	300
NSTYRVSVL TVLHQDWLNG KEYCKVSNK ALPAPIEIKI SKAKGQPREP	350
QVYTLPPSRD ELTKNQVSLT CLVKGFYPSD IAEWESNGQ PENNYKTTTP	400
VLDSDGSEFFL YSKLTVDKSR WQGNVVFSCS VMHEALHNHY TQKSLSLSPG	450
K	451
Light chain / Chaîne légère / Cadena ligera	
DIVMSQSPSS LSASVGDRVT ITCKSSQSLN NSRTRKNFLA WYQKPGKSP	50
KLLIYWASTR ESGVPDRFSG SGSGDTFTLT ISSLPQEDFA TYYCKQSFNL	100
FTFGQGTKLE IKRTVAAPSV FIFPPSDEQL KSGTASVCL LNNFYPREAK	150
VQWVKVDNALQ SGNSQESVTE QDSKDSYSL SSTLTLSKAD YEKHKVYACE	200
VTHQGLSSPV TKSFNRGEC	219
Post-translational modifications	
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro	
Intra-H (C23-C104)	22-96 148-204 265-325 371-429
	22"-96" 148"-204" 265"-325" 371"-429"
Intra-L (C23-C104)	23"-94" 139"-199"
	23"-94" 139"-199"
Inter-H-L (h 5-CL 126)	224-219" 224"-219"
Inter-H-H (h 11, h 14)	230-230" 233-233"
N-terminal glutaminyl cyclization to pyroglutamyl (pE, 5-oxoprolyl)	
H VH Q1:	
I, I"	
N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación	
H CH2 N84.4:	
301, 301"	
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.	
C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal	
H CHS K2:	
451, 451"	

relmacabtagenum autoleu celum #
relmacabtagene autoleu cel

Autologous CD4+ and CD8+ enriched T cells transduced ex vivo with a self-inactivating lentivirus vector encoding an anti-CD19 chimeric antigen receptor.

Autologous CD4+ and CD8+ T cells, enriched from peripheral blood mononuclear cells (PBMCs) transduced with a non-replicative self-inactivating (SIN) lentiviral vector encoding a chimeric antigen receptor (CAR) consisting of the human CD19-specific scFv, an IgG4 hinge region, CD28 transmembrane domain, CD137 (4-1BB) co-stimulatory domain and CD3 zeta signalling domain, under the control of a hybrid Elongation Factor 1 alpha (EF1α) and Human T cell Leukemia Virus type 1 (HTLV-1) regulatory element promoter and woodchuck hepatitis virus post-transcriptional regulatory element (WPRE). The vector genome also encodes a truncated human epidermal growth factor receptor (EGFRt) that is co-expressed with the CAR and cleaved from it by the T2A self-cleaving peptide. The vector genome also contains a splice donor, a fragment of gag, a Rev response element (RRE), a mutated central polypurine tract (cPPT), and a splice acceptor.

relmacabtagène autoleu cel

Cellules T autologues enrichies en CD4+ et CD8+ transduites ex vivo avec un vecteur lentiviral auto-inactifant codant pour un récepteur chimérique dirigé contre l'antigène CD19.

Cellules T CD4+ et CD8+ autologues, enrichies de cellules mononucléaires du sang périphérique (PBMC), transduites avec un vecteur lentiviral auto-inactivable et non répliquatif (SIN) codant pour un récepteur chimérique (CAR) constitué du scFv spécifique de l'antigène CD19 humain, d'une région charnière IgG4, d'un domaine transmembranaire CD28, du domaine co-stimulateur de CD137 (4-1BB) et du domaine de signalisation CD3 zeta, sous le contrôle d'un hybride du facteur d'allongement 1 alpha (EF1α) et du promoteur de l'élément régulateur du virus de la leucémie humaine à cellules T de type 1 (HTLV-1), ainsi que de l'élément régulateur post-transcriptionnel du virus de l'hépatite de la marmotte (WPRE). Le génome du vecteur code également pour un récepteur tronqué du facteur de croissance épidermique humain (EGFRt) qui est co-exprimé avec le CAR et en est relâché par le peptide T2A auto-coupant. Le génome du vecteur contient également un donneur d'épissage, un fragment de gag, un élément de réponse Rev (RRE), un segment central polypurine muté (cPPT) et un accepteur d'épissage.

relmacabtagén autoleucel

Células T autólogas enriquecidas en CD4+ y CD8+ transducidas ex vivo con un lentivirus auto-inactivante que codifica un receptor de antígenos quiméricos anti-CD19.

Las células T autólogas CD4+ y CD8+, enriquecidas a partir de células mononucleares de sangre periférica (PBMCs), transducidas con un vector lentiviral no replicativo y auto-inactivante (SIN) que codifica para un receptor de antígenos quimérico (CAR) consistente en un scFv específico de CD19 humano, una región bisagra IgG4, un dominio transmembrana CD28, un dominio co-estimulador CD137 (4-1BB) y un domino de señalización CD3 zeta, bajo el control de un promotor híbrido del factor de elongación 1 alfa (EF1α) y el elemento regulador del virus de la leucemia de células T de tipo 1 humano (HTLV-1), y elemento regulador post-transcripcional del virus de la hepatitis de la marmota (WPRE). El genoma del vector también codifica para un receptor del factor de crecimiento epidérmico humano truncado (EGFRt) que se coexpresa con el CAR y luego se escinde de él por el péptido de auto escisión T2A. El genoma del vector también contiene un sitio donante de procesamiento (splicing), un fragmento de gag, un elemento de respuesta Rev (RRE), un segmento central de polipurinas (cPPT) y un sitio aceptor de procesamiento (splicing).

resiniferatoxinum
resiniferatoxin

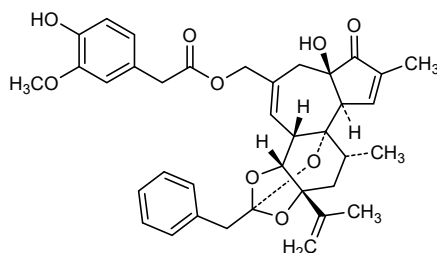
[(2S,3aR,3bS,6aR,9aR,9bR,10R,11aR)-2-benzyl-6a-hydroxy-8,10-dimethyl-7-oxo-11a-(prop-1-en-2-yl)-3a,3b,6,6a,9a,10,11,11a-octahydro-2H,7H-2,9b-epoxyazuleno[5,4-e][1,3]benzodioxol-5-yl)methyl (4-hydroxy-3-methoxyphenyl)acetate

résinifératoxine

(4-hydroxy-3-méthoxyphényl)acétate de
[(2*S*,3*aR*,3*bS*,6*aR*,9*aR*,9*bR*,10*R*,11*aR*)-2-benzyl-6*a*-hydroxy-8,10-
diméthyl-7-oxo-11*a*-(prop-1-én-2-yl)-3*a*,3*b*,6,6*a*,9*a*,10,11,11*a*-
octahydro-2*H*,7*H*-2,9*b*-époxyazuléno[5,4-*e*][1,3]benzodioxol-5-
yl]méthyle

resiniferatoxina

(4-hidroxi-3-metoxifenil)acetato de
[(2*S*,3*aR*,3*bS*,6*aR*,9*aR*,9*bR*,10*R*,11*aR*)-2-bencil-6*a*-hidroxi-8,10-
dimetil-7-oxo-11*a*-(prop-1-en-2-il)-3*a*,3*b*,6,6*a*,9*a*,10,11,11*a*-
octahidro-2*H*,7*H*-2,9*b*-epoxiazuleno[5,4-*e*][1,3]benzodioxol-5-il]metilo

C₃₇H₄₀O₉**revakinagenum tarorectelum #**

revakinagene tarorectel

Allogeneic human adult retinal pigment epithelial (RPE) cell line stably transfected with a plasmid encoding human ciliary neurotrophic factor (CNTF).

An allogeneic human adult retinal pigment epithelial (RPE) cell line stably transfected with a plasmid encoding human ciliary neurotrophic factor (CNTF) fused in frame to the murine immunoglobulin mu-factor VDJ4C1 signal peptide, under the control of a mouse metallothionein promoter and a human growth hormone polyadenylation signal. The plasmid also contains three eukaryotic expression cassettes for the selectable markers neomycin resistance, mouse DHFR (dihydrofolate reductase) and HSV-TK (thymidine kinase), plus the bacterial components of the pUC18 backbone, including the ampicillin resistance gene (AmpR), origin of replication (ORI) and LacZ gene (alpha fragment of β -galactosidase).

révakinagène tarorectel

Lignée cellulaire épithéliale allogénique du pigment rétinien humain adulte (EPR) transfectée de façon stable avec un plasmide codant pour le facteur neurotrophique ciliaire humain (CNTF).

Une lignée cellulaire épithéliale allogénique du pigment rétinien humain adulte (RPE) transfectée de manière stable avec un plasmide codant pour le facteur neurotrophique ciliaire humain (CNTF) fusionné dans le cadre de lecture du peptide signal VDJ4C1 du facteur mu d'immunoglobuline murine, sous le contrôle d'un promoteur de la métallothionéine de souris et d'un signal de polyadénylation de l'hormone de croissance humaine. Le plasmide contient également des cassettes d'expression eucaryotes pour trois marqueurs de sélection, à savoir la résistance à la néomycine, la DHFR (réductase du dihydrofolate) et la HSV-TK (kinase de la thymidine) de souris, ainsi que les composants bactériens du plasmide pUC18, dont le gène de résistance à l'ampicilline (AmpR), l'origine de réplication (ORI) et le gène du LacZ (fragment alpha de la β -galactosidase).

revakinagén taroretcel

Línea celular epitelial alogénica de pigmento retiniano adulto (RPE) humano transfectada de forma estable con un plásmido que codifica para el factor neurotrófico ciliar (CNTF) humano.

Una línea celular epitelial alogénica de pigmento retiniano adulto (RPE) humano transfectada de forma estable con un plásmido que codifica para el factor neurotrófico ciliar (CNTF) humano fusionado en marco de lectura al péptido señal VDJ4AC1 del factor mu de la inmunoglobulina murina, bajo el control de un promotor de la metalotionina de ratón y una señal de poliadenilación de la hormona de crecimiento humana. El plásmido contiene también tres casetes de expresión eucariotas para los marcadores de selección de resistencia a neomicina, DHFR (dihidrofolato reductasa) de ratón y HSV-TK (timidín quinasa), más los componentes bacterianos del esqueleto de pUC18, incluyendo el gen de resistencia a ampicilina (AmpR), el origen de replicación (ORI) y el gen LacZ (fragmento alfa de la β -galactosidasa).

rezvilutamidum

rezvilutamide

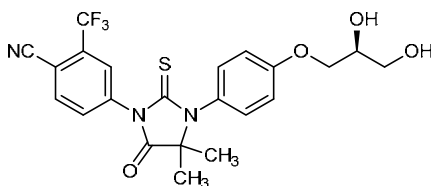
4-(3-{4-[(2S)-2,3-dihydroxypropoxy]phenyl}-4,4-dimethyl-5-oxo-2-sulfanylideneimidazolidin-1-yl)-2-(trifluoromethyl)benzonitrile

rezvilutamide

4-(3-{4-[(2S)-2,3-dihydroxypropoxy]phényl}-4,4-diméthyl-5-oxo-2-sulfanylidèneimidazolidin-1-yl)-2-(trifluorométhyl)benzonitrile

rezvilutamida

4-(3-{4-[(2S)-2,3-dihidroxiopropoxi]fenil}-4,4-dimetil-5-oxo-2-sulfanilidenoimidazolidin-1-il)-2-(trifluorometil)benzonitrilo

$$\text{C}_{22}\text{H}_{20}\text{F}_3\text{N}_3\text{O}_4\text{S}$$


rineterkibum

rineterkib

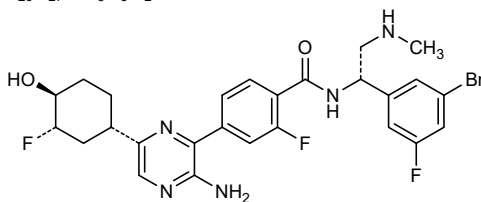
4-{3-amino-6-[(1S,3S,4S)-3-fluoro-4-hydroxycyclohexyl]pyrazin-2-yl}-N-[(1S)-1-(3-bromo-5-fluorophenyl)-2-(methylamino)ethyl]-2-fluorobenzamide

rinéterkib

4-{3-amino-6-[(1S,3S,4S)-3-fluoro-4-hydroxycyclohexyl]pirazin-2-yl}-N-[(1S)-1-(3-bromo-5-fluorophényl)-2-(méthylamino)éthyl]-2-fluorobenzamide

rineterkib

4-{3-amino-6-[(1S,3S,4S)-3-fluoro-4-hidroxiciclohexil]pirazin-2-il}-N-[(1S)-1-(3-bromo-5-fluorofenil)-2-(metilamino)etil]-2-fluorobenzamida

$C_{26}H_{27}BrF_3N_5O_2$ **rintodestrantum**

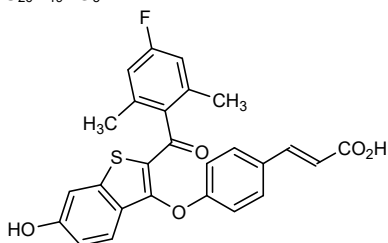
rintodestrant

(2*E*)-3-(4-([2-(4-fluoro-2,6-diméthylbenzoyl)-6-hydroxy-1-benzothiophen-3-yl]oxy)phényl)prop-2-énoïque

rintodestrant

acide (2*E*)-3-(4-([2-(4-fluoro-2,6-diméthylbenzoyl)-6-hydroxy-1-benzothiophén-3-yl]oxy)phényl)prop-2-énoïque

rintodestrant

ácido (2*E*)-3-(4-([2-(4-fluoro-2,6-dimetilbenzoi)-6-hidroxí-1-benzotiofen-3-il]oxi)fenil)prop-2-énoico $C_{26}H_{19}FO_5S$ **roducitabinum**

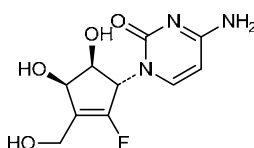
roducitabine

4-amino-1-[(1*S*,4*R*,5*S*)-2-fluoro-4,5-dihydroxy-3-(hydroxyméthyl)cyclopent-2-en-1-yl]pyrimidin-2(1*H*)-one;
4'-a-fluoro-4',4'-a-didehydro-4'-O-carbacytidine

roducitabine

4-amino-1-[(1*S*,4*R*,5*S*)-2-fluoro-4,5-dihydroxy-3-(hydroxyméthyl)cyclopent-2-én-1-yl]pyrimidin-2(1*H*)-one;
4'-a-fluoro-4',4'-a-didéhydro-4'-O-carbacytidine

roducitabina

4-amino-1-[(1*S*,4*R*,5*S*)-2-fluoro-4,5-dihidroxi-3-(hidroximetil)ciclopent-2-en-1-il]pirimidin-2(1*H*)-ona;
4'-a-fluoro-4',4'-a-didehido-4'-O-carbacitidina $C_{10}H_{12}FN_3O_4$ 

senaparibum

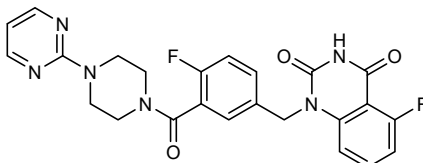
senaparib

5-fluoro-1-({4-fluoro-3-[4-(pyrimidin-2-yl)piperazine-1-carbonyl]phenyl}methyl)quinazoline-2,4(1*H*,3*H*)-dione

sénaparib

5-fluoro-1-({4-fluoro-3-[4-(pyrimidin-2-yl)pipérazine-1-carbonyl]phényl}méthyl)quinazoline-2,4(1*H*,3*H*)-dione

senaparib

5-fluoro-1-({4-fluoro-3-[4-(pirimidin-2-il)piperazina-1-carbonil]fenil}metil)quinazolina-2,4(1*H*,3*H*)-diona $C_{24}H_{20}F_2N_6O_3$ **sivopixantum**

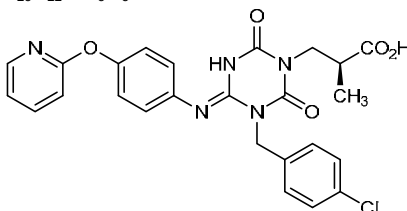
sivopixant

(2*S*)-3-[(4*E*)-3-[(4-chlorophenyl)methyl]-2,6-dioxo-4-({4-[(pyridin-2-yl)oxy]phenyl}imino)-1,3,5-triazinan-1-yl]-2-methylpropanoic acid

sivopixant

acide (2*S*)-3-[(4*E*)-3-[(4-chlorophényl)méthyl]-2,6-dioxo-4-({4-[(pyridin-2-yl)oxy]phényl}imino)-1,3,5-triazinan-1-yl]-2-méthylpropanoïque

sivopixant

ácido (2*S*)-3-[(4*E*)-3-[(4-clorofenil)metil]-2,6-dioxo-4-({4-[(piridin-2-il)oxil]fenil}imino)-1,3,5-triazinan-1-il]-2-metilpropanoico $C_{25}H_{22}ClN_5O_5$ **sizavaleucelum**

sizavaleucl

Allogeneic, HLA-matched CD3+ T-cells, derived from G-CSF mobilized peripheral blood stem cells (PBSC) collected by apheresis. Cells are transplant organ donor-derived, non-cultured and cryopreserved with CD34+ hematopoietic stem cells (HSC) (*firzotemcel* (121)(83)) obtained from the same HLA-matched donor. Intended for delivery to a single HLA-matched recipient.

sizavaleucel

Des cellules T CD3+ allogéniques, exprimant des antigènes HLA compatibles, dérivées de cellules souches du sang périphérique (PBSC) mobilisées par le G-CSF et collectées par aphérèse. Les cellules sont dérivées de donneurs d'organes de transplantation, ne sont pas cultivées et sont cryoconservées avec des cellules souches hématopoïétiques (HSC) CD34+ (*firzotemcel* (121)(83)) qui proviennent du même donneur compatible pour les antigènes HLA. Destinées à être données à un seul receveur compatible pour les antigènes HLA.

sizavaleucel

Células T CD3+ alogénicas, con HLA compatible, derivadas de células madre de sangre periférica (PBSC) movilizadas con G-CSF y recogidas por aféresis. Las células proceden de donantes de órganos para trasplante, no se cultivan y se criopreservan con células madre hematopoyéticas (HSC) CD34+ (*firzotemcel* (121)(83)) obtenidas del mismo donante HLA compatible. Destinadas para administrarse a un sólo receptor HLA compatible.

sotigalimabum #

sotigalimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* CD40 (tumor necrosis factor receptor superfamily member 5, TNFRSF5)], humanized monoclonal antibody; gamma1 heavy chain humanized (1-450) [VH (*Homo sapiens* IGHV3-30*01 (79.8%) -(IGHD) -IGHJ1*01 (90.9%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 G1m17,1 (CH1 K120 (217) (121-218), hinge 1-15 (219-233), CH2 S29>E (270) (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-215')-disulfide with kappa light chain humanized (1'-215') [V-KAPPA (*Homo sapiens* IGKV1-27*01 (91%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.13] (27-32.50-52.89-101) (1'-108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dimer (229-229":232-232")-bisdisulfide, produced in Chinese hamster ovary (CHO)-derived cell line, glycoform alfa

sotigalimab

immunoglobuline G1-kappa, anti-[*Homo sapiens* CD40 (membre 5 de la superfamille des récepteurs du TNF, TNFRSF5)], anticorps monoclonal humanisé; chaîne lourde gamma1 humanisée (1-450) [VH (*Homo sapiens* IGHV3-30*01 (79.8%) -(IGHD) -IGHJ1*01 (90.9%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 G1m17,1 (CH1 K120 (217) (121-218), charnière 1-15 (219-233), CH2 S29>E (270) (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-215')-disulfure avec la chaîne légère kappa humanisée (1'-215') [V-KAPPA (*Homo sapiens* IGKV1-27*01 (91%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.13] (27-32.50-52.89-101) (1'-108') (1'-108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dimère (229-229":232-232")-bisdisulfure, produite dans une lignée cellulaire dérivée des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

Recommended INN: List 85
sotigalimab

WHO Drug Information, Vol. 35, No. 1, 2021

inmunoglobulina G1-kappa, anti-[*Homo sapiens* CD40 (miembro 5 de la superfamilia de los receptores del TNF, TNFRSF5)], anticuerpo monoclonal humanizado; cadena pesada gamma1 humanizada (1-450) [VH (*Homo sapiens* IGHV3-30*01 (79.8%) -(IGHD) - IGHJ1*01 (90.9%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 G1m17,1 (CH1 K120 (217) (121-218), bisagra 1-15 (219-233), CH2 S29>E (270) (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-215')-disulfuro con la cadena ligera kappa humanizada (1'-215') [V-KAPPA (*Homo sapiens* IGKV1-27*01 (91%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.13] (27-32.50-52.89-101) (1'-108') (1'-108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dímero (229-229":232-232")-bisdisulfuro, producido en una línea celular derivada de las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada					
QVQLVESGGG	VVQPGRSRLR	SCAASGFSFS	STYVCWVRQA	PGKGLEWIAIC	50
IYTGdGTNYS	ASWAKGRFTI	SKDSSKNVTY	LQMNSLRAED	TAVYFCARPD	100
ITYGFaINFW	PGPTLVTVSS	ASTKGPSVF	LAPSSKSTSG	GTAALGCLVK	150
DYFPEPVTVS	WNSGALTSGV	HTFPAVLQSS	GLYSLSSVVT	VPSSSLGTQT	200
YICNVNHKPS	NTKVDKKVEP	KSCDKTHTCP	PCPAPELLGG	PSVFLFPPKP	250
KDTLMISRTP	EVTCTVVVDVE	HEDPEVKFNW	YVDGVEVHNA	KTKPREEQYN	300
STYRVVSVLT	VLHQDWLNGK	EYKCKVSNKA	LPAPIEKTIIS	KAKGQPREPQ	350
VYTLPPSRDE	LTKNQVSLTC	LVKGFYPSDI	AVEWESNGQP	ENNYKTTTPPV	400
LDSDGSFFLY	SKLTVDKSRW	QQGNVFSCSV	MHEALHNHYT	QKSLSLSPGK	450
Light chain / Chaîne légère / Cadena ligera					
DIQMTQSPSS	LSASVGDRVT	IKCQASQSI	SRLAWYQQKP	GKPPKLLIYR	50
ASTLASGVPS	RFGSGSGSDT	FTLTISSLQ	EDVATYYCQC	TGYGISWPIG	100
GGTKVEIKRT	VAAPSFIIFP	PSDEQLKSGT	ASVVCLLNNF	YPREAKVQWK	150
VDNALQSGNS	QESVTEQDSK	DSTYLSLSTL	TLSKADYEKH	KVYACEVTHQ	200
GLSSPVTKSF	NRGEC				215
Post-translational modifications					
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro					
Intra-H (C23-C104)	22-96	35-50	147-203	264-324	370-428
	22"-96"	35"-50"	147"-203"	264"-324"	370"-428"
Intra-L (C23-C104)	23'-88'	135'-195'			
	23"-88"	135"-195"			
Inter-H-L (h 5-CL 126)	223'-215'	223"-215"			
Inter-H-H (h 11, h 14)	229-229"	232-232"			
N-terminal glutaminyl cyclization to pyroglutamyl (pE, 5-oxoprolyl)					
H VH Q1:					
1, 1"					
N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación					
H VH N65:					
58, 58"					
H CH2 N84.4:					
300, 300"					
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.					
C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal					
H CHS K2:					
450, 450"					

sotorasibum
sotorasib

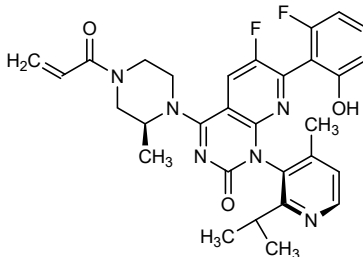
(1*M*)-6-fluoro-7-(2-fluoro-6-hydroxyphenyl)-1-[4-methyl-2-(propan-2-yl)pyridin-3-yl]-4-[(2*S*)-2-methyl-4-(prop-2-enoyl)piperazin-1-yl]pyrido[2,3-*d*]pyrimidin-2(1*H*)-one

sotorasib

(1*M*)-6-fluoro-7-(2-fluoro-6-hydroxyphényl)-1-[4-méthyl-2-(propan-2-yl)pyridin-3-yl]-4-[(2*S*)-2-méthyl-4-(prop-2-énoyl)pipérazin-1-yl]pyrido[2,3-*d*]pyrimidin-2(1*H*)-one

sotorasib

(1*M*)-6-fluoro-7-(2-fluoro-6-hydroxifenil)-1-[4-metil-2-(propan-2-il)piridin-3-il]-4-[(2*S*)-2-metil-4-(prop-2-enoil)piperazin-1-il]pirido[2,3-*d*]pirimidin-2(1*H*)-ona

C₃₀H₃₀F₂N₆O₃

sulanemadlinum

sulanemadlin

C^{2.11}, C^{2.4}-[(4*E*)-undec-4-ene-1,11-diyl](*N*-acetyl-L-leucyl-L-threonyl-L-phenylalanyl-L-alanyl-L-α-glutamyl-L-tyrosyl-L-tryptophyl-L-alanyl-L-glutamyl-L-leucyl-D-alanyl-L-alanyl-L-alanyl-L-alanyl-L-alanyl-L-alanyl-D-alaninamide)

sulanémadline

C^{2.11}, C^{2.4}-[(4*E*)-undéc-4-ène-1,11-diyl](*N*-acétyl-L-leucyl-L-thréonyl-L-phénylalanyl-L-alanyl-L-α-glutamyl-L-tyrosyl-L-tryptophyl-L-alanyl-L-glutamyl-L-leucyl-D-alanyl-L-alanyl-L-alanyl-L-alanyl-L-alanyl-L-alanyl-D-alaninamide)

sulanemadlina

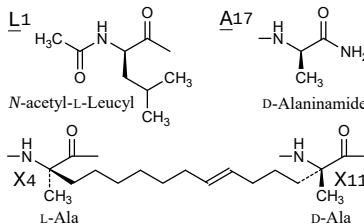
C^{2.11}, C^{2.4}-[(4*E*)-undec-4-eno-1,11-diil](*N*-acetyl-L-leucyl-L-threonyl-L-phenylalanyl-L-alanyl-L-α-glutamyl-L-tyrosyl-L-tryptophyl-L-alanyl-L-glutamyl-L-leucyl-D-alanyl-L-alanyl-L-alanyl-L-alanyl-L-alanyl-L-alanyl-D-alaninamide)

C₉₅H₁₄₀N₂₀O₂₃

Sequence / Séquence / Secuencia

LTFXEYWAQL XAAAAAA

Modified residues / Résidus modifiés / Restos modificados



suntinorextonum

suntinorexton

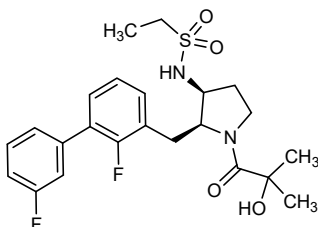
N-[(2*S*,3*S*)-2-[(2,3'-difluoro[1,1'-biphenyl]-3-yl)methyl]-1-(2-hydroxy-2-methylpropanoyl)pyrrolidin-3-yl]ethanesulfonamide

suntinorexton

N-[(2*S*,3*S*)-2-[(2,3'-difluoro[1,1'-biphényl]-3-yl)méthyl]-1-(2-hydroxy-2-méthylpropanoyl)pyrrolidin-3-yl]éthanesulfonamide

suntinorextón

N-[[(2*S*,3*S*)-2-[(2,3'-difluoro[1,1'-bifenil]-3-il)metil]-1-(2-hidroxi-2-metilpropanoil)pirrolidin-3-il]etanosulfonamida

$$\text{C}_{23}\text{H}_{28}\text{F}_2\text{N}_2\text{O}_4\text{S}$$


suratadenoturevum #
suratadenoturev

A replication-competent adenovirus type 5 in which the E1A and E1B genes are under the control of a human telomerase reverse transcriptase (hTERT) and IRES, respectively.

A recombinant replication-competent adenovirus type 5 (Ad-5) in which the E1 gene, including its normal transcriptional regulatory element, has been deleted and replaced by an expression cassette containing an hTERT (human telomerase reverse transcriptase) promoter, the Ad E1A gene, an IRES (internal ribosome entry site) and the Ad E1B gene. E1A expression is controlled by the hTERT promoter; E1B translation is effected by the IRES.

suratadénoturev

Un adénovirus de type 5 réplcatif dans lequel les gènes E1A et E1B sont, respectivement, sous le contrôle de la transcriptase inverse de la télomérase humaine (hTERT) et d'IRES.

Un adénovirus recombinant réplcatif de type 5 (Ad-5) dans lequel le gène E1, y compris son élément normal de régulation de la transcription, a été supprimé et remplacé par une cassette d'expression contenant un promoteur hTERT (transcriptase inverse de la télomérase humaine), le gène Ad E1A, un IRES (site d'entrée interne du ribosome) et le gène Ad E1B. L'expression de E1A est contrôlée par le promoteur hTERT; la traduction de E1B est effectuée par l'IRES.

suratadenoturev

Un adenovirus tipo 5 competente de replicación en el que los genes E1A y E1B están bajo el control de la telomerasa transcriptasa inversa humana (hTERT) e IRES, respectivamente.

Un adenovirus tipo 5 (Ad-5) recombinante competente de replicación en el que el gen E1, incluyendo su elemento de regulación transcripcional normal, ha sido delecionado y reemplazado por un casete de expresión que contiene un promotor de hTERT (telomerasa transcriptasa inversa humana), el gen Ad E1A, un IRES (sitio interno de entrada al ribosoma) y el gen Ad E1B. La expresión de E1A esta controlada por el promotor hTERT; La traducción de E1B se efectúa por el IRES.

taletrectinibum

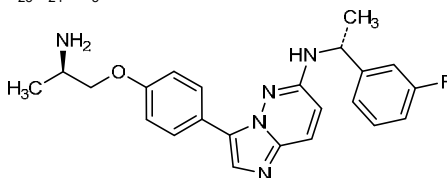
taletrectinib

3-{4-[(2*R*)-2-aminopropoxy]phenyl}-*N*-[(1*R*)-1-(3-fluorophenyl)ethyl]imidazo[1,2-*b*]pyridazin-6-amine

talétrectinib

3-{4-[(2*R*)-2-aminopropoxy]phényl}-*N*-[(1*R*)-1-(3-fluorophényl)éthyl]imidazo[1,2-*b*]pyridazin-6-amine

taletrectinib

3-{4-[(2*R*)-2-aminopropoxi]fenil}-*N*-[(1*R*)-1-(3-fluorofenil)etil]imidazo[1,2-*b*]piridazin-6-amina $C_{23}H_{24}FN_5O$ **taniraleucelum**

taniraleucel

Allogeneic natural killer (NK) cells derived from CD34+ umbilical cord blood (UCB) hematopoietic stem/progenitor cells.

The cells are expanded and differentiated in the presence of cytokines including thrombopoietin, SCF, Flt3 ligand, IL-7, IL-15 and IL-2 (average population doublings 15) to generate the NK cell population. Cells (on average >95% of the cells) are CD56+/CD3- and have detectable levels of CD94, NKG2D and CD16. The cells secrete IFN- γ , perforin and granzyme B, and have cytolytic activity against human tumour cell lines.

taniraleucel

Cellules tueuses naturelles (NK) allogéniques dérivées de cellules souches/progénitrices hématopoïétiques CD34+ du sang de cordon ombilical (UCB). Les cellules sont amplifiées et différenciées en présence de cytokines, notamment la thrombopoïétine, le SCF, le ligand de Flt3, l'IL-7, l'IL-15 et l'IL-2 (la population double en moyenne 15 fois) pour générer la population de cellules NK. Les cellules (en moyenne >95% des cellules) sont CD56+/CD3- et ont des niveaux détectables de CD94, NKG2D et CD16. Les cellules sécrètent l'IFN- γ , la perforine et la granzyme B, et ont une activité cytolytique contre les lignées de cellules tumorales humaines.

taniraleucel

Células natural killer (NK) alogénicas derivadas de células madre/progenitoras hematopoyéticas de sangre de cordón umbilical (UCB). Las células se expanden y diferencian en presencia de citoquinas incluyendo trombopoyetina, SCF, ligando de Flt3, IL-7, IL-15 e IL-2 (la media de duplicaciones de la población es 15 veces) hasta generar la población de células NK. Las células (como media >95% de las células) son CD56+/CD3- y tienen niveles detectables de CD94, NKG2D y CD16. Las células secretan IFN- γ , perforina y granzima B, y tienen actividad citolítica frente a líneas celulares tumorales humanas.

tarlatamabum

tarlatamab

immunoglobulin scFv-scFv-scFc, anti-[*Homo sapiens* DLL3 (delta-like ligand 3)] and anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], monoclonal antibody single chain (scFv)2-scFc, bispecific; IG single chain scFv-scFv-scFc, anti-DLL3 and anti-CD3E (1-982) [scFv-VH-V-kappa anti-DLL3 (1-241) [VH (*Homo sapiens*IGHV4-59*01 G49>C (44) (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.7.12] (26-33.51-57.96-107) (1-118) -15-mer-tris(tetraglycyl-seryl) linker (119-133)-V-KAPPA (*Homo sapiens* IGKV3-20*01 (91.7%) -IGKJ2*01 Q120>C (234) (90.9%)) CDR-IMGT [7.3.9] (160-166.184-186.223-231) (134-241)] -6-mer-seryl-tetraglycyl-seryl linker (242-247) -scFv-VH-V-lambda anti-CD3E (248-496) [VH (*Mus musculus*IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87.0%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (273-280.298-307.346-361) (248-372) -15-mer-tris(tetraglycyl-seryl) linker (373-387) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (413-421.439-441.478-486) (388-496)] -4-mer-tetraglycyl linker (497-500) - scFc (h-CH2-CH3)-(h-CH2-CH3) (501-982) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (hinge 6-15 (501-510), CH2 R83>C (572), N84.4>G (577), V85>C (582) (511-620), CH3 E12 (636), M14 (638) (621-725), CHS>del) (501-725) -30-mer-hexakis(tetraglycyl-seryl) linker (726-755) -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (hinge 6-15 (756-765), CH2 R83>C (827), N84.4>G (832), V85>C (837) (766-875), CH3 E12 (891), M14 (893) (876-980), CHS (981-982))] (756-982)], non-glycosylated, produced in Chinese hamster ovary (CHO) cells

tarlatamab

immunoglobuline scFv-scFv-scFc, anti-[*Homo sapiens* DLL3 (delta-like ligand 3)] et anti-[*Homo sapiens* CD3E (CD3 epsilon, Leu-4)], anticorps monoclonal à chaîne unique (scFv)2-scFc, bispécifique; IG à chaîne unique scFv-scFv-scFc, anti-DLL3 et anti-CD3E (1-982) [scFv-VH-V-kappa anti-DLL3 (1-241) [VH (*Homo sapiens*IGHV4-59*01 G49>C (44) (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.7.12] (26-33.51-57.96-107) (1-118) -15-mer-tris(tétraglycyl-séryl) linker (119-133)-V-KAPPA (*Homo sapiens* IGKV3-20*01 (91.7%) -IGKJ2*01 Q120>C (234) (90.9%)) CDR-IMGT [7.3.9] (160-166.184-186.223-231) (134-241)] -6-mer-séryl-tétraglycyl-séryl linker (242-247) -scFv-VH-V-lambda anti-CD3E (248-496) [VH (*Mus musculus*IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (273-280.298-307.346-361) (248-372) -15-mer-tris(tétraglycyl-séryl) linker (373-387) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (413-421.439-441.478-486) (388-496)] -4-mer-tétraglycyl linker (497-500) - scFc (h-CH2-CH3)-(h-CH2-CH3) (501-982) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (charnière 6-15 (501-510), CH2 R83>C (572), N84.4>G (577), V85>C (582) (511-620), CH3 E12 (636), M14 (638) (621-725), CHS>del) (501-725) -30-mer-hexakis(tétraglycyl-séryl) linker (726-755) -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (charnière 6-15 (756-765), CH2 R83>C (827), N84.4>G (832), V85>C (837) (766-875), CH3 E12 (891), M14 (893) (876-980), CHS (981-982))] (756-982)], non-glycosylé, produit dans des cellules ovariennes de hamster chinois (CHO)

tarlatamab

inmunoglobulina scFv-scFv-scFc, anti-[*Homo sapiens* DLL3 (delta-like ligando 3)] y anti-[*Homo sapiens* CD3E (CD3 épsilon, Leu-4)], anticuerpo monoclonal con cadena única (scFv)2-scFc, biespecífica; IG con cadena única scFv-scFv-scFc, anti-DLL3 y anti-CD3E (1-982) [scFv-VH-V-kappa anti-DLL3 (1-241) [VH (*Homo sapiens* IGHV4-59*01 G49>C (44) (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.7.12] (26-33.51-57.96-107) (1-118) -15-mer-tris(tetraglicil-seril) linker (119-133)-V-KAPPA (*Homo sapiens* IGKV3-20*01 (91.7%) -IGKJ2*01 Q120>C (234) (90.9%)) CDR-IMGT [7.3.9] (160-166.184-186.223-231) (134-241)] -6-mer-seril-tetraglicil-seril linker (242-247) -scFv-VH-V-lambda anti-CD3E (248-496) [VH (*Mus musculus*IGHV10-1*02 (91.9%) -(IGHD) -IGHJ3*01 (86.7%)/*Homo sapiens* IGHV3-73*01 (87%) -(IGHD) -IGHJ5*01 (100%)) CDR-IMGT [8.10.16] (273-280.298-307.346-361) (248-372) -15-mer-tris(tetraglicil-seril) linker (373-387) -V-LAMBDA (*Homo sapiens* IGLV7-43*01 (85.1%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.9] (413-421.439-441.478-486) (388-496)] -4-mer-tetraglicil linker (497-500) - scFc (h-CH2-CH3)-(h-CH2-CH3) (501-982) [*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (bisagra 6-15 (501-510), CH2 R83>C (572), N84.4>G (577), V85>C (582) (511-620), CH3 E12 (636), M14 (638) (621-725), CHS>del) (501-725) -30-mer-hexakis(tetraglicil-seril) linker (726-755) -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (bisagra 6-15 (756-765), CH2 R83>C (827), N84.4>G (832), V85>C (837) (766-875), CH3 E12 (891), M14 (893) (876-980), CHS (981-982)) (756-982)]], no glicosilado, producido en las células ováricas de hámster chino (CHO)

Heavy chain / Chaîne lourde / Cadena pesada				
QVQLQESGPG	LVPKSETLSL	TCTVSGGSIS	SYYSWIRQP	PGKCLEWIGY 5C
VYYSGTTNMY	PSLKSRTVIS	VDTSKNQFSL	KLSSVTAAC	AVYYCASIAV 100
TGFYFDYWGQ	GTLVTVSSGG	GGSGGGSGGG	GGSEIVLTQS	PGLTSLSPGE 150
RVTLSCRASQ	RVNNNYLAWY	QQRPGQAPRL	LIYGASSRAT	GIPDRFSGSG 200
SGTDFTLTIS	RLEPEDFAVY	YQQYDRSFL	TFGCGTKLEI	KSGGGGSEVQ 250
LVESGGGLVQ	PGGSLKLSCA	ASGFTNKYA	MNWVRQAPGK	GLEWVARIRS 300
KYNNYATYYA	DSVKDRFTIS	RDSKNTAYL	QMNNLKTEDT	AVYYCVRHGN 350
FGNSYISYYA	YWGQGTIVTV	SSGGGGSGGG	GSGGGGSGTV	VTQEPSTLVS 400
PGGTIVLTGG	SSTGAVTSGN	YPNWVQKFG	QAPRGLIGGT	KFLAPGTPAR 450
FSGSLLGKKA	ALTLSGVQPE	DBAEYYCVLV	YSNRWVFGGG	TKLTVLGGGG 500
DKTHTCPGCP	APPELLGGPSV	FLFPPKPKDT	LMISRTPEVT	CVVVDVSHSD 550
PEVKFNWYVD	GVEVHNARTK	PCEEYQGSTY	RCVSVLTVLH	QDWLNGKEYK 60C
CKVSNKALPA	PIEKTISKAK	GQPREPQVYT	LPFSREEMTK	NQVSLTCLVK 65C
GFYPSDIAVE	WESNGQPENN	YKTTTPVVLDS	DGSFFLYSKL	TVDKSRWQQG 700
NVFSCSVMEH	ALHNHYTQKS	LSLSPGCGGS	GGGGSGGGGS	GGGGSGGGGS 750
GGGGSKDTHT	CPPCPAPELL	GGPSVFLFPP	KPKDTLMISR	TPEVTCVVVD 800
VSHEDPEVKF	NWYVDGVEVH	NARTKPCREQ	YGSTYRCVSV	LTVLHQDLNL 850
GKEYKCKVSN	KALPAPIERT	ISKAKGQPRE	PQVYTLPPSR	EEMTKNCVSL 900
TCLVKGFPYS	DIAVEWESNG	QPENNYKTTT	PVLDSGGSFF	LYSKLTVDXS 950
RWQQGNVFSC	SVMHEALHNH	YTKSLSLSP	GK	982
Post-translational modifications				
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro				
Intra-V (C23-C104)	22-95	156-222	269-345	409-477
Intra-C (C23-C104)	541-601	647-705	796-856	902-960
Intra-CH2 (C83-C85)	572-582	827-837		
Inter-VH-VL (C49-C120)	44-234			
Inter-h11 (h11-h11)	506-761			
Inter-h14 (h14-h14)	509-764			
No N-glycosylation sites / pas de sites de N-glycosylation / ningún posición de N-glicosilación				
CH2 N84.4>G:				
577, 832				
Aglycosylated				
C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal				
CHS K2:				
982				

tepegfilgrastim #
tepegfilgrastim

human granulocyte colony-stimulating factor (G-CSF, pluripoiétin) short isoform, variant (A¹>M), produced in *Escherichia coli*, pegylated with one *N*-{[α-methylpoly(oxyethylene)-ω-oxy]acetyl}-*N*-[α-methylpoly(oxyethylene)-ω-yl]glycyl group at N² of M¹ (85-95%) or at N⁶ of K¹⁷ (3-13%) or at N⁶ of K²⁴, K³⁵ and K⁴¹ (total not more than 5%);

[1-L-methionine]human granulocyte colony-stimulating factor (G-CSF, pluripoiétin) short isoform, produced in *Escherichia coli*, substituted with one *N*-{[α-methylpoly(oxyethylene)-ω-oxy]acetyl}-*N*-[α-methylpoly(oxyethylene)-ω-yl]glycyl group at N² of Met¹ (85-95%) or at N⁶ of Lys¹⁷ (3-13 %) or one other lysine residue (total not more than 5%)

tepegfilgrastim

isoforme courte du facteur de stimulation des colonies de granulocytes humain (G-CSF, pluripoiétine), variant (A¹>M), produite dans *Escherichia coli*, pégylé avec un groupe *N*-{[α-méthylpoly(oxyéthylène)-ω-oxy]acétyle}-*N*-[α-méthylpoly(oxyéthylène)-ω-yl]glycyle en N² de M¹ (85-95%) ou en N⁶ de K¹⁷ (3-13%) ou en N⁶ de K²⁴, K³⁵ et K⁴¹ (pas plus de 5% au total); [1-L-méthionine]isoforme courte du facteur de stimulation des colonies de granulocytes humain (G-CSF, pluripoiétine), produite dans *Escherichia coli*, substituée par un groupe *N*-{[α-méthylpoly(oxyéthylène)-ω-oxy]acétyle}-*N*-[α-méthylpoly(oxyéthylène)-ω-yl]glycyle en N² du Mét¹ (85-95 %) ou en N⁶ du Lys¹⁷ (3-13 %) ou d'un autre résidu de lysine (pas plus de 5 % au total)

tepegfilgrastim

factor humano de estimulación de colonias de granulocitos (G-CSF, pluripoyetina) isoforma corta, variante (A¹>M), producido en *Escherichia coli*, pegilado con un grupo *N*-{[α-metilpoli(oxietileno)-ω-oxi]acetil}-*N*-[α-metilpoli(oxietileno)-ω-il]glicilo en N² de M¹ (85-95%) o en N⁶ de K¹⁷ (3-13%) o en N⁶ de K²⁴, K³⁵ y K⁴¹ (total no más del 5%); [1-L-metionina]isoforma corta del factor humano de estimulación de colonias de granulocitos (G-CSF, pluripoyetina), producida en *Escherichia coli*, sustituido con un grupo *N*-{[α-metilpoli(oxietileno)-ω-oxi]acetil}-*N*-[α-metilpoli(oxietileno)-ω-il]glicilo en N² de la Met¹ (85-95 %) o en N⁶ de la Lis¹⁷ (3-13 %) o un otro residuo de lisina (total no más del 5 %)

C₈₅₃H₁₃₅₂N₂₂₄O₂₄₇S₉[(C₂H₄O)_{2n}] (n ≈ 400-500)

Sequence / Séquence / Secuencia

MEPLGPASSL PQSTLLKCLE QVRKIQGDGA ALQEKLCATY KLCHPPEELVL 50
LGHSLGIFWA PLSSCFSSQAL QLACGLSQLH SGLFLYQQLL QALEGISPEL 100
GPTLOTLLQC VADFATTINQ QMEELGHMAFA LQPTQGAMPA FASAFQRRAG 150
GVLVASHLQS FLEVSYRVLK HLAQF 175

Mutation / Mutation / Mutación

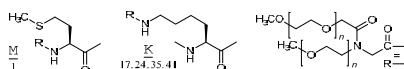
A¹>M

Post-translational modifications

Disulfide bridge location / Position du pont disulfure / Posición del puente disulfuro
37-43, 65-75

Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación
none / aucune / ninguna

Chemical modification sites / Sites de modification chimique / Posiciones de modificación química
major / principal / mayormente : M1, K17;
minor / secondaire / inferior : K24, K35, K41



tenvumestrocelum

tenvumestrocel

Allogeneic human mesenchymal stromal cells derived from umbilical cord blood (hUCB-MSCs). The cells are isolated by density-gradient centrifugation and expanded by sub-culturing of the adherent cell population. Cells (on average > 95%) express mesenchymal stromal cell related antigens (CD105 and CD73). The cells are also positive for the CD29 antigen (integrin beta-1), the CD44 antigen (cell adhesion glycoprotein), the CD13 and CD90 antigens, and the HLA-ABC histocompatibility antigen. Cells do not express (on average < 1%) hematopoietic antigens (CD45, CD34 and CD14), histocompatibility antigen (HLA-DR), endothelial cell antigen (CD31), osteoclast antigen (CD51/61), and CD64.

tenvumestrocel

Cellules stromales mésenchymateuses humaines allogéniques dérivées du sang de cordon ombilical (hUCB-MSCs). Les cellules sont isolées par centrifugation à gradient de densité et amplifiées par sous-culture de la population des cellules adhérentes. Les cellules (en moyenne > 95%) expriment des antigènes liés aux cellules stromales mésenchymateuses (CD105 et CD73). Les cellules sont également positives pour l'antigène CD29 (intégrine beta-1), l'antigène CD44 (glycoprotéine d'adhésion cellulaire), les antigènes CD13 et CD90, et l'antigène d'histocompatibilité HLA-ABC. Les cellules n'expriment pas (en moyenne < 1%) les antigènes hématopoïétiques (CD45, CD34 et CD14), l'antigène d'histocompatibilité (HLA-DR), l'antigène des cellules endothéliales (CD31), l'antigène des ostéoclastes (CD51/61), ni l'antigène CD64.

tenvumestrocel

Células estromales mesenquimales humanas alogénicas derivadas de sangre de cordón umbilical (hUCB-MSCs). Las células se aíslan por centrifugación en un gradiente de densidad y se expanden mediante subcultivo de las poblaciones adherentes. Las células (de media >95%) expresan los antígenos relacionados con las células estromales mesenquimales (CD105 y CD73). Las células son también positivas para los antígenos CD29 (integrina beta-1), CD44 (receptor de matriz extracelular), CD13 y CD90, y para el antígeno de histocompatibilidad HLA-ABC. Las células no expresan (de media <1%) antígenos hematopoyéticos (CD45, CD34 y CD14), antígeno de histocomaptibilidad (HLA-DR), antígeno de células endoteliales (CD31), antígeno de osteoclastos (CD51/61), ni CD64.

teplinovivintum

teplinovivint

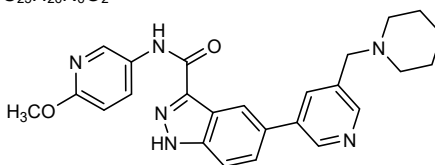
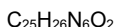
N-(6-methoxy-pyridin-3-yl)-5-{5-[(piperidin-1-yl)methyl]pyridin-3-yl}-1*H*-indazole-3-carboxamide

téplinovivint

N-(6-méthoxy-pyridin-3-yl)-5-{5-[(pipéridin-1-yl)méthyl]pyridin-3-yl}-1*H*-indazole-3-carboxamide

teplinovivint

N-(6-metoxipiridin-3-il)-5-{5-[(piperidin-1-il)metil]piridin-3-il}-1*H*-indazol-3-carboxamida

**terevalefimum**

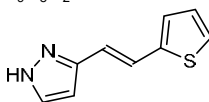
terevalefim

3-[(1*E*)-2-(thiophen-2-yl)ethen-1-yl]-1*H*-pyrazole

térévaléfim

3-[(1*E*)-2-(thiophén-2-yl)éthén-1-yl]-1*H*-pyrazole

terevalefim

3-[(1*E*)-2-(tiofen-2-il)eten-1-il]-1*H*-pirazol**tilpisertibum**

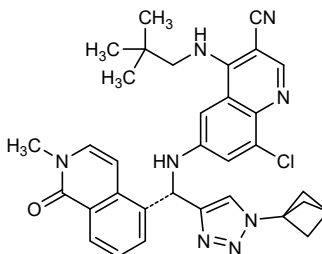
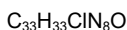
tilpisertib

6-[[[(*S*)-[1-(bicyclo[1.1.1]pentan-1-yl)-1*H*-1,2,3-triazol-4-yl](2-methyl-1-oxo-1,2-dihydroisoquinolin-5-yl)methyl]amino]-8-chloro-4-[(2,2-dimethylpropyl)amino]quinoline-3-carbonitrile

tilpisertib

6-[[[(*S*)-[1-(bicyclo[1.1.1]pentan-1-yl)-1*H*-1,2,3-triazol-4-yl](2-méthyl-1-oxo-1,2-dihydroisoquinoléin-5-yl)méthyl]amino]-8-chloro-4-[(2,2-diméthylpropyl)amino]quinoléine-3-carbonitrile

tilpisertib

6-[[[(*S*)-[1-(biciclo[1.1.1]pentan-1-il)-1*H*-1,2,3-triazol-4-il](2-metil-1-oxo-1,2-dihidroisoquinolein-5-il)metil]amino]-8-cloro-4-[(2,2-dimetilpropil)amino]quinoleína-3-carbonitrilo**tusamitamabum #**

tusamitamab

immunoglobulin G1-kappa, anti-[*Homo sapiens* CEACAM5 (carcinoembryonic antigen-related cell adhesion molecule 5, CEA, CD66e)], monoclonal antibody;

	gamma1 heavy chain (1-449) [VH (<i>Mus musculus</i> IGHV5-12-1*01 (86.6%) -(IGHD) -IGHJ3*01 (92.9%)/<i>Homo sapiens</i> IGHV3-23*01 (77.3%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -<i>Homo sapiens</i> IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), hinge 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS K>del (449) (121-449)], (223-214')-disulfide with kappa light chain (1'-214') [V-KAPPA (<i>Mus musculus</i> IGKV12-44*01 (87.4%) -IGKJ4*01 (100%)/<i>Homo sapiens</i> IGKV1-39*01 (82.1%) -IGKJ2*02 (90.9%)) CDR-IMGT [6.3.9](27-32.50-52.89-97) (1'-107') -<i>Homo sapiens</i> IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (229-229":232-232")-bisdisulfide, produced in Chinese hamster ovary (CHO)-derived cell line, glycoform alfa
tusamitamab	immunoglobuline G1-kappa, anti-[<i>Homo sapiens</i> CEACAM5 (molécule d'adhésion cellulaire 5 apparentée à l'antigène carcinoembryonnaire, CEA, CD66e)], anticorps monoclonal; chaîne lourde gamma1 (1-449) [VH (<i>Mus musculus</i> IGHV5-12-1*01 (86.6%) -(IGHD) -IGHJ3*01 (92.9%)/<i>Homo sapiens</i> IGHV3-23*01 (77.3%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -<i>Homo sapiens</i> IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), charnière 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS K>del (449) (121-449)], (223-214')-disulfure avec la chaîne légère kappa (1'-214') [V-KAPPA (<i>Mus musculus</i> IGKV12-44*01 (87.4%) -IGKJ4*01 (100%)/<i>Homo sapiens</i> IGKV1-39*01 (82.1%) -IGKJ2*02 (90.9%)) CDR-IMGT [6.3.9](27-32.50-52.89-97) (1'-107') -<i>Homo sapiens</i> IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (229-229":232-232")-bisdisulfure, produite dans une lignée cellulaire dérivée des cellules ovariennes de hamster chinois (CHO), glycoforme alfa
tusamitamab	inmunoglobulina G1-kappa, anti-[<i>Homo sapiens</i> CEACAM5 (molécula de adhesión celular 5 relacionada con el antígeno carcinoembrionario, CEA, CD66e)], anticuerpo monoclonal; cadena pesada gamma1 (1-449) [VH (<i>Mus musculus</i> IGHV5-12-1*01 (86.6%) -(IGHD) -IGHJ3*01 (92.9%)/<i>Homo sapiens</i> IGHV3-23*01 (77.3%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -<i>Homo sapiens</i> IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), bisagra 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS K>del (449) (121-449)], (223-214')-disulfuro con la cadena ligera kappa (1'-214') [V-KAPPA (<i>Mus musculus</i> IGKV12-44*01 (87.4%) -IGKJ4*01 (100%)/<i>Homo sapiens</i> IGKV1-39*01 (82.1%) -IGKJ2*02 (90.9%)) CDR-IMGT [6.3.9](27-32.50-52.89-97) (1'-107') -<i>Homo sapiens</i> IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (229-229":232-232")-bisdisulfuro, producido en una línea celular derivada de las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
EYQLQESGPG LVKPGGSLSL SCASGTFVF SYDMSVVPQT PERGLEWVAY 50
ISSGGGITYA FSTYKGRFTV SRDRAWNTLY LQMSLTSED TAVYYCAHY 100
FGSSGFFAYW CQCTLVTVSS ASTKGPVSVE LAPSSKSTSG GTAALGCLVK 150
DYFPEPTVVS WNSGALTSGV HTFFAVLQSS GLYSLSSVVT VPSSSLGTQT 200
YICNVNHKES NTKVDKKVLP KSCDKTHTCP PCPAPPELLGG PSVFLFPPKE 250
KDTLAISRTP EVTCVVVDVE HEDPEVKKNW YVDGVEVHKA KTKPREEQYN 300
STYRVVSVLT VLNQDGLNGK EYKCKVSNKA LPAPIEKTIK KAKGQPREPQ 350
VYTLPPSRDE LTKNQVSLTC LVKGFYPSDI AVEVESNGQP ENNYKTPPV 400
LDSGGSFFLY SKLTVDKSRW QQGNVFSCSV MHEALHNHYT QKSLSLSPG 449

Light chain / Chaîne légère / Cadena ligera
DIQHTQSFAS LSASVGRVIT ITCRAENIF SYLAWYQQNP GKSPKLLVYN 50
TRTLAEGVES RFSGSGSGTD FSLTISSLQF EDFATYYCQH HYGTPFTFGS 100
GTKLEIKRTV AAPSVFIFFP SDEQLKSGTA SVVCLLNIFY PREAKVQKKV 150
DNALQSGNSQ ESVTEQDSKD STYSLSSLTIT LSKADYEKKH VIACEVTHQG 200
LSSPVTKSNF RGEK 214

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22°-96° 147°-203° 264°-324° 370°-428°
22°-96° 147°-203° 264°-324° 370°-428°
Intra-L (C23-C104) 23°-88° 134°-194°
23°-88° 134°-194°
Inter-H-L (h 5-CL 126) 223°-214° 223°-214°
Inter-H-H (h 11, h 14) 229°-229° 232°-232°

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84,4:
300, 300°
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires
complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

tusamitamabum ravtansinum #
tusamitamab ravtansine

immunoglobulin G1-kappa, anti-[*Homo sapiens* CEACAM5 (carcinoembryonic antigen-related cell adhesion molecule 5, CEA, CD66e)], monoclonal antibody, conjugated to maytansinoid DM4; gamma1 heavy chain (1-449) [VH (*Mus musculus* IGHV5-12-1*01 (86.6%) -(IGHD) -IGHJ3*01 (92.9%)]/*Homo sapiens* IGHV3-23*01 (77.3%) -(IGHD) -IGHJ4*01 (92.9%)] CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%), G1m17,1 (CH1 K120 (217) (121-218), hinge 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS K>del (449) (121-449)], (223-214')-disulfide with kappa light chain (1'-214') [V-KAPPA (*Mus musculus* IGKV12-44*01 (87.4%) -IGKJ4*01 (100%)]/*Homo sapiens* IGKV1-39*01 (82.1%) -IGKJ2*02 (90.9%)] CDR-IMGT [6.3.9](27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (229-229":232-232")-bisdisulfide, produced in a Chinese hamster ovary (CHO) cell line derived from CHO-K1SV, glycoform alfa; conjugated, on an average of 3 to 4 lysyl, to maytansinoid DM4 [N2'- deacetyl-N2'-(4-mercapto-4-methyl-1-oxopentyl)- maytansine] via the reducible SPDB linker [N-succinimidyl 4-(2-pyridyldithio)butanoate] For the ravtansine part, please refer to the document "INN for pharmaceutical substances: Names for radicals, groups and others"

tusamitamab ravtansine

immunoglobuline G1-kappa, anti-[*Homo sapiens* CEACAM5 (molécule d'adhésion cellulaire 5 apparentée à l'antigène carcinoembryonnaire, CEA, CD66e)], anticorps monoclonal, conjugué au maytansinoïde DM4; chaîne lourde gamma1 (1-449) [VH (*Mus musculus* IGHV5-12-1*01 (86.6%) -(IGHD) -IGHJ3*01 (92.9%)]/*Homo sapiens* IGHV3-23*01 (77.3%) -(IGHD) -IGHJ4*01 (92.9%)] CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%), G1m17,1 (CH1 K120 (217) (121-218), charnière 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS K>del (449) (121-449)], (223-214')-disulfure avec la chaîne légère kappa (1'-214') [V-KAPPA (*Mus musculus* IGKV12-44*01 (87.4%) -IGKJ4*01 (100%)]/*Homo sapiens* IGKV1-39*01 (82.1%) -IGKJ2*02 (90.9%)] CDR-IMGT [6.3.9](27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')];

tusamitamab ravtansina

inmunoglobulina G1-kappa, anti-[*Homo sapiens* CEACAM5 (molécula de adhesión celular 5 con el antígeno carcinoembrionario, CEA, CD66e)], anticuerpo monoclonal, conjugado con maitansinoide DM4;
cadena pesada gamma1 (1-449) [VH (*Mus musculus* IGHV5-12*01 (86.6%) -(IGHD) -IGHJ3*01 (92.9%)/*Homo sapiens* IGHV3-23*01 (77.3%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%), G1m17,1 (CH1 K120 (217) (121-218), bisagra 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS K>del (449) (121-449)], (223-214')-disulfuro con la cadena ligera kappa (1'-214') [V-KAPPA (*Mus musculus* IGKV12-44*01 (87.4%) -IGKJ4*01 (100%)/*Homo sapiens* IGKV1-39*01 (82.1%) -IGKJ2*02 (90.9%)) CDR-IMGT [6.3.9](27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (229-229":232-232")-bisdisulfuro, producido en las células ováricas de hámsters chinos (CHO) línea celular derivada de CHO-K1SV, forma glicosilada alfa; conjugado, en 3 a 4 residuos lisil por término medio, con maitansinoide DM4 [N2'-deacetil-N2'-(4-mercapto-4-metil-1-oxopentil)- maitansina] mediante un conector SPDB reducible [4-(2- pirididitio)butanoato de N-succinimidilo] Para la fracción ravtansina, se ruega referirse al documento "INN for pharmaceutical substances: Names for radicals, groups and others"

Heavy chain / Chaîne lourde / Cadena pesada

```
EVQLQESGPG LVKPGGSLSL SCAASGFVFS SYDMSWVRQT PERGLENVAY 50
ISSGGGITYA PSTYKGRFTV SRDNARKNTLY LQMSNLTSSE TAVYICAAHY 100
FGSSGPFAYV GQCTLVITYSS ASTKGPSVFP LAPSSKSTSG GTAALGCLYK 150
DYFPEPTVYS WNSGALTSGV HTFPAVLQSS GLYSLSVVT VPSSSLGTQT 200
YICNVNHKPS NTKVDKKVPE KSCDKHTTCE PCFAPPELLG PSVFLFPPKP 250
KDTLMISRTPEVTCVVVDVS HEDPEVKYNN YVDGVEVHNA KTKPREEQYM 300
STYRVVSVLT VLHQDWLGKG EYKCKVSNKA LPAPIETIS KAKGQPRFPQ 350
VYTLPPSRDE LTKNQVSLTC LVKGFYPSDI AVEWESNGQP ENNYKTIPTP 400
LDSGGSFFLY SKLTVDKSRW QQGNVFVCSV MHEALHHHT QKSLSLSPG 449
```

Light chain / Chaîne légère / Cadena ligera

```
DIQMTQSPAS LSASVGDRTV ITCRASENI F SYLAWYQKPK GKSPKLLVM 50
TRTLAGVPS RFSGSGSGTD FSLTISLQEP EDFATYYCOH HYGTPTFGS 100
GTKLEIKRTV AAPSVFIFPP SDEQLKSGTA SVVCLLNMFY PREAKVQKVK 150
DNALQSGNSQ ESVTEQDSKD STYSLSSLT LT LSKADYEKHK VYACEVTHQG 200
LSSPVTKSFN RGEK 214
```

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 147-203 264-324 370-428
22"-96" 147"-203" 264"-324" 370"-428"
Intra-L (C23-C104) 23"-88" 134"-194"
23"-88" 134"-194"

Inter-H-L (h 5-CL 126) 223-214' 223"-214"

Inter-H-H (h 11, h14) 229-229" 232-232"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

300, 300"

Fucosylated complex; bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarijos complejos fucosilados.

For the ravtansine part, please refer to the document "INN for pharmaceutical substances:

Names for radicals, groups and others"

Pour la partie ravtansine, veuillez-vous référer au document "INN for pharmaceutical substances:

Names for radicals, groups and others"

Para la fracción ravtansina, se ruega referirse al documento "INN for pharmaceutical substances:

Names for radicals, groups and others"

upifitamab rilsodotinum

upifitamab rilsodotin

immunoglobulin G1-kappa, anti-[*Homo sapiens* SLC34A2 (solute carrier family 34 sodium phosphate member 2, sodium/phosphate cotransporter 2B, NaPi2b, NaPi3b, NAPI-3B)], monoclonal antibody, conjugated, via a thioether bond, to 3 to 5 flexible polymers PHF-BA-EG2-MI-AF-HPA-Ala, each comprising maleimide (MI) bioconjugation linkers and 3 to 4 auristatin F- hydroxypropylamide-L-alanine (AF-HPA-Ala), with a drug ratio of 12:1 to 15:1;

	gamma1 heavy chain (1-449) [VH (<i>Homo sapiens</i>IGHV1-46*01 (81.6%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.12] (26-33.51-58.97-108) (1-119) -glycinyI (120) -<i>Homo sapiens</i>IGHG1*03 (100%), G1m3, nG1m1 (CH1 R120 (217) (121-218), hinge 1-15 (219-233), CH2 (234-343), CH3 E12 (359), M14 (361) (344-448), CHS K>del (449)) (121-449)], (223-215')-disulfide with kappa light chain (1'-215') [V-KAPPA (<i>Mus musculus</i>IGKV10-94*01 (84.2%) -IGKJ5*01 (91.7%)/<i>Homo sapiens</i>IGKV1-33*01 (83.2%) -IGKJ2*01 (90.9%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -arginyI (108') -<i>Homo sapiens</i>IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dimer (229-229":232-232")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa; conjugated, via a thioether bond, from 3 to 5 flexible biodegradable polymers, poly(1-hydroxymethylethylene hydroxymethyl-formal) (PHF)-BA-EG2-MI-AF-HPA-Ala, each comprising maleimide (MI) conjugation linkers and 3 to 4 cytotoxic auristatin F- hydroxypropylamide-L-alanine (AF-HPA-Ala), with a drug ratio of 12:1 to 15:1
upifitamab rilsodotine	immunoglobuline G1-kappa, anti-[<i>Homo sapiens</i> SLC34A2 (membre 2 de la famille 34 sodium phosphate de transporteurs de solutés, cotransporteur 2B de sodium/phosphate, NaPi2b, NaPi3b, NAPI-3B)], anticorps monoclonal, conjugué, via une liaison thiéther, à 3 à 5 polymères flexibles PHF-BA-EG2-MI-AF-HPA-Ala, comprenant chacun des linkers de bioconjugaison maléimide (MI) et 3 à 4 auristatine F- hydroxypropylamide-alanine (AF-HPA-Ala), dans un rapport produit actif de 12:1 à 15:1; chaîne lourde gamma1 (1-449) [VH (<i>Homo sapiens</i>IGHV1-46*01 (81.6%) -(IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.12] (26-33.51-58.97-108) (1-119) -glycinyI (120) -<i>Homo sapiens</i>IGHG1*03 (100%), G1m3, nG1m1 (CH1 R120 (217) (121-218), charnière 1-15 (219-233), CH2 (234-343), CH3 E12 (359), M14 (361) (344-448), CHS K>del (449)) (121-449)], (223-215')-disulfure avec la chaîne légère kappa (1'-215') [V-KAPPA (<i>Mus musculus</i>IGKV10-94*01 (84.2%) -IGKJ5*01 (91.7%)/<i>Homo sapiens</i>IGKV1-33*01 (83.2%) -IGKJ2*01 (90.9%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -arginyI (108') -<i>Homo sapiens</i>IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dimère (229-229":232-232")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa; conjugué, via une liaison thiéther, de 3 à 5 polymères flexibles biodégradables poly(1-hydroxyméthyléthylène hydroxyméthyl-formal) (PHF)-BA-EG2-MI-AF-HPA-Ala, comprenant chacun des linkers de conjugaison maléimide (MI) et 3 à 4 auristatine F- hydroxypropylamide-L-alanine (AF-HPA-Ala) cytotoxiques, dans un rapport produit actif de 12:1 à 15:1
upifitamab rilsodotina	inmunoglobulina G1-kappa, anti-[<i>Homo sapiens</i> SLC34A2 (miembro 2 de la familia 34 sodio fosfato de transportadores de solutos, cotransportador 2B de sodio/fosfato, NaPi2b, NaPi3b, NAPI-3B)], anticuerpo monoclonal, conjugado, a través de un enlace tioéter, de 3 a 5 polímeros flexibles PHF-BA-EG2-MI-AF-HPA-Ala, que comprende cada uno de los enlaces de bioconjugación maleimida (MI) y 3 a 4 auristatina F- hidroxipropilamida-L-alanina (AF-HPA-Ala), en una relación de principio activo de 12:1 a 15:1;

cadena pesada gamma1 (1-449) [VH (*Homo sapiens* IGHV1-46*01 (81.6%) - (IGHD) -IGHJ4*01 (92.9%)) CDR-IMGT [8.8.12] (26-33.51-58.97-108) (1-119) - glicinil (120) -*Homo sapiens* IGHG1*03 (100%), G1m3, nG1m1 (CH1 R120 (217) (121-218), bisagra 1-15 (219-233), CH2 (234-343), CH3 E12 (359), M14 (361) (344-448), CHS K>del (449)) (121-449)], (223-215')-disulfuro con la cadena ligera kappa (1'-215') [V-KAPPA (*Mus musculus* IGKV10-94*01 (84.2%) -IGKJ5*01 (91.7%)/*Homo sapiens* IGKV1-33*01 (83.2%) -IGKJ2*01 (90.9%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -arginil (108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; dímero (229-229":232-232")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa; conjugado, a través de un enlace tioéter, a 3 a 5 polímeros flexibles biodegradables poli(1-hidroximetil-etileno hidroximetil-formal) (PHF)-BA-EG2-MI-AF-HPA-Ala, que comprende cada uno de los enlaces de conjugación maleimida (MI) y 3 a 4 auristatina F- hidroxipropilamida-L-alanina (AF-HPA-Ala) citotóxicos, en una relación de principio activo de 12:1 a 15:1

Heavy chain / Chaîne lourde / Cadena pesada

QVQLVQSGAE VVKPGASVKM SCKASGYFT GYNIHVKQA FGGLEWIGA 5C
IYPNGDTSY KQKFRGRATL TADTSTSTVY MELSSLRSEC SAVVYCARG 100
TARATFAWGS QGTLTVTVSSG ASTKGFSVEP LAPSSKSTSG GTALGGLVK 150
DYFEPFVTS NNSGLTSSV KTFPAVLQSS GLYSLSGVVT VPSSLSGTQT 20C
YICNVNKKPS NIKVDKRVLP KSCDKHTKCP PCPAPELLGG PSVELFPKP 25C
KOTLMISRTPT EVTCVVVDVS HEDPEVKFNK YVDGVEVHNA KTKPREEQYN 300
STYRVVSULT VLHQDWLNGK EYKCKVSNKA LPAPIEKTIS KAKGQPREPQ 350
VYTLPPSREE MTKNQVSLTC LVKGIFYPSDI AVENESNGQP ENNYKITPPV 400
LSDGGSFFLY SKLTVDKSRW QGQNVFSCSV MHEALNNHYT QKSLSLSPG 449

Light chain / Chaîne légère / Cadena ligera

DIQMTQSPSS LSAISVGDRTV ITCSAQDIG NFLNYYQQPK GKTVKVLIYY 5C
TSSLYSGVPS RFSGSGSGTD YTLTISLQP EDFATYYCQ YSKLPLTFGQ 100
GTKLELKRRT VAAPSVFIFP FDSQGLKST ASVVCLLNWF YPREAKVQWK 150
VDNALQSGHS QESVEEQGSK DSTYLSSTL TLSKADYEKR KYACEVTRQ 20C
GLSSPVTKSF NRGEK 215

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-11 (C23-C104) 22-96 147-203 264-324 370-428

22-96* 147-203* 264-324* 370-428*

Intra-I. (C23-C104) 23-88* 135-195*

23-88** 135-195**

Inter-11-I. (h 5-C1, 126)* 223-215' 223"-215"

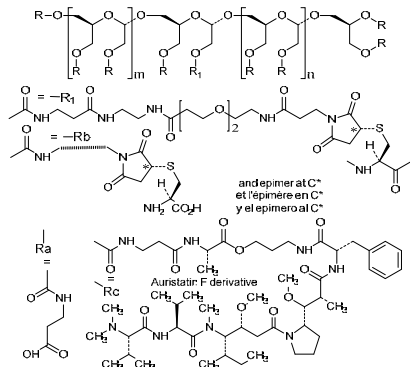
Inter-11-II. (h 11-I, h 14)* 229-229' 232-232"

*Two or three of the inter-chain disulfide bridges are not present, 3 to 5 cysteines being conjugated each via a thioether bond to one maleimide linker of one flexible polymer comprising 3 to 4 drug units.

*Deux ou trois des ponts disulfures inter-chaînes ne sont pas présents, 3 à 5 cystéines étant chacune conjuguée via une liaison thioéther à un linker-maleimide d'un polymère flexible comprenant 3 à 4 unités de principe actif.

*Faltan dos o tres puentes disulfuro inter-catenarios, 3 a 5 cisteína está conjugada a conectores de un polímero flexible que contiene 3 a 4 unidades de principio activo.

Modified residues / résidus modifiés / restos modificados:



N-terminal glutaminylation cyclization to pyroglutamylation (pL, 5-oxoprolin)

L.VIIQI:

I, I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H C12 N84, 4;

300, 300"

Fucosylated complex bi-antennary C1HO-type glycans / glycanes de type C1HO

bi-antennaires complexes fucosylés / glycanes de type C1HO biantennaires complexes fucosilados.

ursodoxicoltaurinum

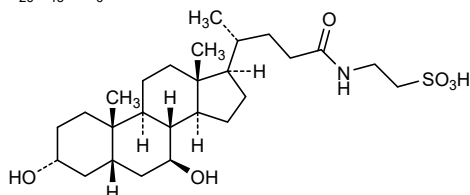
ursodoxicoltaurine

2-(3 α ,7 β -dihydroxy-5 β -cholan-24-amido)ethane-1-sulfonic acid

ursodoxicoltaurine

acide 2-(3 α ,7 β -dihydroxy-5 β -cholan-24-amido)éthane-1-sulfonique

ursodoxicoltaurina

ácido 2-(3 α ,7 β -dihidroxi-5 β -colan-24-amido)etano-1-sulfónicoC₂₆H₄₅NO₆S**vamifeportum**

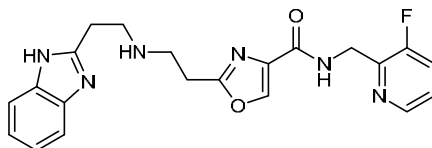
vamifeport

2-(2-[[2-(1*H*-benzimidazol-2-yl)ethyl]amino]ethyl)-*N*-[(3-fluoropyridin-2-yl)methyl]-1,3-oxazole-4-carboxamide

vamiféport

2-(2-[[2-(1*H*-benzimidazol-2-yl)éthyl]amino]éthyl)-*N*-[(3-fluoropyridin-2-yl)méthyl]-1,3-oxazole-4-carboxamide

vamifeport

2-(2-[[2-(1*H*-benzimidazol-2-il)etil]amino]etil)-*N*-[(3-fluoropiridin-2-il)metil]-1,3-oxazol-4-carboxamidaC₂₁H₂₁FN₆O₂**venadaparibum**

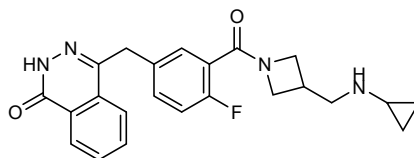
venadaparib

3⁴-fluoro-7-aza-1(1)-phthalazina-5(1,3)-azetidina-3(1,3)-benzena-8(1)-cyclopropanaoctaphane-1⁴(1³*H*),4-dione

vénadaparib

3⁴-fluoro-7-aza-1(1)-phthalazina-5(1,3)-azétidina-3(1,3)-benzèna-8(1)-cyclopropanaoctaphane-1⁴(1³*H*),4-dione

venadaparib

3⁴-fluoro-7-aza-1(1)-ftalazina-5(1,3)-azetidina-3(1,3)-bencena-8(1)-ciclopropanaoctafano-1⁴(1³*H*),4-dionaC₂₃H₂₃FN₄O₂

vevorisertibum

vevorisertib

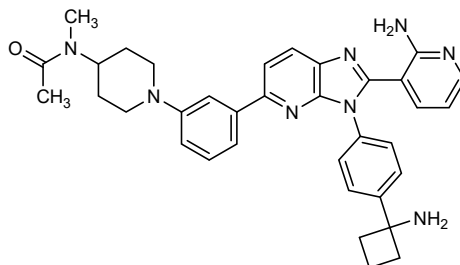
N-[1-(3-{3-[4-(1-aminocyclobutyl)phenyl]-2-(2-aminopyridin-3-yl)-3*H*-imidazo[4,5-*b*]pyridin-5-yl}phenyl)piperidin-4-yl]-*N*-methylacetamide

vévorisertib

N-[1-(3-{3-[4-(1-aminocyclobutyl)phényl]-2-(2-aminopyridin-3-yl)-3*H*-imidazo[4,5-*b*]pyridin-5-yl}phényl)pipéridin-4-yl]-*N*-méthylacétamide

vevorisertib

N-[1-(3-{3-[4-(1-aminociclobutil)fenil]-2-(2-aminopiridin-3-il)-3*H*-imidazo[4,5-*b*]piridin-5-il}fenil)piperidin-4-il]-*N*-metilacetamida

C₃₅H₃₈N₈O**vibapapogenum autoleucelum #**

vibapapogene autoleucel

Autologous peripheral blood mononuclear cells (PBMCs) expressing human papillomavirus types 16 and 18 antigens. Autologous peripheral blood mononuclear cells (PBMCs), specifically B lymphocytes and monocytes that are T-cell depleted, transfected *ex vivo* with a non-replicating adenoviral vector that expresses a codon-optimised fusion of tumour-associated antigens (TAAs) E6 and E7 derived from human papillomavirus (HPV) types 16 and 18 (HPV-16/18 E6/E7). The antigen coding region is preceded by a tPA signal sequence and transcription is under the control of the cytomegalovirus (CMV)-tetracycline 10 promoter and the SV40 poly(A) termination sequence. The vector genome is flanked by inverted terminal repeats and contains an encapsidation signal and a K35 knob.

vibapapogène autoleucel

Cellules mononucléaires du sang périphérique autologues exprimant les antigènes de papillomavirus humains de type 16 et 18. Cellules mononucléaires du sang périphérique autologues, spécifiquement les lymphocytes B et les monocytes qui sont déplétées en lymphocytes T, transfectées *ex vivo* avec une forme non répliquative du vecteur adénoviral exprimant un codon optimisé, fusion des antigènes associés à des tumeurs E6 et E7 dérivés du virus humain du papillome (HPV) types 16 et 18 (HPV-16/18 E6/E7). La région codant l'antigène est précédée d'une séquence signal tPA et la transcription est sous le contrôle du promoteur tétracycline 10 du cytomégalo virus et de la séquence de terminaison poly(A) SV40. Le génome du vecteur est flanqué de répétitions inverses et contient un signal d'encapsidation et un domaine globulaire distal (knob) K35.

vibapapogén autoleucel

Células monucleares de sangre periférica (PBMCs) autólogas que expresan antígenos del papilomavirus humano de tipo 16 y 18.
Células monucleares de sangre periférica (PBMCs) autólogas, específicamente linfocitos B y monocitos que están deplecionadas de células T, transfectadas *ex vivo* con un vector adenoviral no replicativo que expresa una fusión de los antígenos asociados a tumor (TAAs) E6 y E7 derivados del papilomavirus humano (HPV) de tipo 16 y 18 (HPV-16/18 E6/E7), con codones optimizados. La región codificadora de antígeno está precedida por una secuencia señal tPA y la transcripción está bajo el control del promotor de la tetraciclina 10 del citomegalovirus (CMV) y la secuencia poly(A) de terminación del SV50. El genoma del vector está flanqueado por repeticiones invertidas terminales y contiene una señal de encapsidación y un dominio globular distal (knob) K35.

vibosameranum #

vibosameran

Messenger RNA encoding tyrosinase.
Messenger RNA (mRNA), 5'-capped, encoding codon-optimised tyrosinase (monophenol monooxygenase, tumour rejection antigen AB), expressed as a fusion protein comprising tyrosinase, P2P16 tetanus toxoid-derived helper epitopes and a major histocompatibility complex (MHC) class I transmembrane and cytoplasmic domain, connected by two glycine/serine-rich (GS) linker peptides, flanked by 5' and 3' untranslated regions and a 3' poly(A) tail.

vibosamérán

ARN messenger codant pour la tyrosinase.
ARN messenger (ARNm), protégé en 5', codant pour la tyrosinase (monophénol monooxygénase, antigène de rejet de tumeur AB), avec des codons optimisés, exprimé sous la forme d'une protéine de fusion comprenant la tyrosinase, les épitopes auxiliaires P2P16 dérivés de l'anatoxine tétanique et un domaine transmembranaire et cytoplasmique du complexe majeur d'histocompatibilité (CMH) de classe I, reliés par deux peptides de liaison riches en glycine/sérine (GS), flanqués de régions non traduites en 5' et 3' et d'une queue poly(A) en 3'.

vibosamerán

ARN mensajero que codifica para tirosinasa.
ARN mensajero (ARNm), protegido en 5' que codifica para tirosinasa (monofenol monooxigenasa, antígeno de rechazo tumoral AB), con codones optimizados, expresada como una proteína de fusión que consta de la tirosinasa, los epítomos auxiliares P2P16 derivados del toxoide tetánico y un dominio citoplásmico y transmembrana del complejo principal de histocompatibilidad (MHC) de clase I, conectados mediante dos péptidos de enlace ricos en glicina/serina (GS), flanqueado por las regiones 5' y 3' no traducidas y una cola poly(A) en 3'.

vidutolimodum

vidutolimod

immunostimulatory CG-rich palindromic oligodeoxyribonucleotide, flanked by 3'-dG₁₀ and 5'-dG₁₀ decanucleotides:

deca[2'-deoxyguanylyl-(3'→5')]-2'-deoxyguanylyl-(3'→5')-2'-deoxyadenylyl-(3'→5')-2'-deoxycytidylyl-(3'→5')-2'-deoxyguanylyl-(3'→5')-2'-deoxyadenylyl-(3'→5')-thymidylyl-(3'→5')-2'-deoxycytidylyl-(3'→5')-2'-deoxyguanylyl-(3'→5')-thymidylyl-(3'→5')-2'-deoxycytidylyl-(3'→5')-nona[2'-deoxyguanylyl-(3'→5')]-2'-deoxyguanosine

vidutolimod

oligodésoxyribonucléotide palindromique immunostimulateur riche en CG, flanqué de décanucléotides 3'-dG₁₀ et 5'-dG₁₀: déca[2'-désoxyguanylyl-(3'→5')]-2'-désoxyguanylyl-(3'→5')-2'-désoxyadénylyl-(3'→5')-2'-désoxycytidylyl-(3'→5')-2'-désoxyguanylyl-(3'→5')-2'-désoxyadénylyl-(3'→5')-thymidylyl-(3'→5')-2'-désoxycytidylyl-(3'→5')-2'-désoxyguanylyl-(3'→5')-thymidylyl-(3'→5')-2'-désoxycytidylyl-(3'→5')-nona[2'-désoxyguanylyl-(3'→5')]-2'-désoxyguanosine

vidutolimod

oligodesoxirribonucleótido palindrómico inmunoestimulador rico en CG, flanqueado por decanucleótidos 3'-dG₁₀ y 5'-dG₁₀:

deca[2'-desoxiguanilil-(3'→5')]-2'-desoxiguanilil-(3'→5')-2'-desoxiadenilil-(3'→5')-2'-desoxicitidilil-(3'→5')-2'-desoxiguanilil-(3'→5')-2'-desoxiadenilil-(3'→5')-timidilil-(3'→5')-2'-desoxicitidilil-(3'→5')-2'-desoxiguanilil-(3'→5')-timidilil-(3'→5')-2'-desoxicitidilil-(3'→5')-nona[2'-desoxiguanilil-(3'→5')]-2'-desoxiguanosina

C₂₉₇H₃₆₃N₁₃₈O₁₇₈P₂₉

Sequence / Séquence / Secuencia

(3'→5') d(GGGGGGGGGG GACGATCGTC GGGGGGGGGG)**vimseltinibum**

vimseltinib

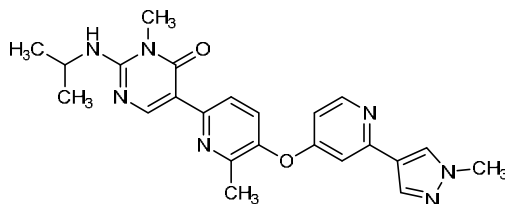
3-methyl-5-(6-methyl-5-[[2-(1-methyl-1*H*-pyrazol-4-yl)pyridin-4-yl]oxy]pyridin-2-yl)-2-[(propan-2-yl)amino]pyrimidin-4(3*H*)-one

vimseltinib

3-méthyl-5-(6-méthyl-5-[[2-(1-méthyl-1*H*-pyrazol-4-yl)pyridin-4-yl]oxy]pyridin-2-yl)-2-[(propan-2-yl)amino]pyrimidin-4(3*H*)-one

vimseltinib

3-metil-5-(6-metil-5-[[2-(1-metil-1*H*-pirazol-4-il)piridin-4-il]oxi]piridin-2-il)-2-[(propan-2-il)amino]pirimidin-4(3*H*)-ona

C₂₃H₂₅N₇O₂

vixarelimabum

vixarelimab

immunoglobulin G4-G1-lambda2, anti-[*Homo sapiens* OSMR (oncostatin M receptor, OSMRB, OSMRbeta)], *Homo sapiens* monoclonal antibody; gamma4-gamma1 heavy chain *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV1-8*01 (95.9%) -(IGHD) -IGHJ6*01 (95.0%)) CDR-IMGT [8.8.19] (26-33.51-58.97-115) (1-126)-*Homo sapiens* IGHG4*01 CH1-h-CH2, G4v5 h P10, G4v36 CH2 Q84.4 (CH1 (127-224), hinge 1-12 S10>P (234) (225-236), CH2 N84.4>Q (303) (237-346)) -IGHG1*01 CH3-CHS, G1m1 (CH3 D12 (362), L14 (364) (347-451), CHS K>del (452)) (127-452)], (140-215')-disulfide with lambda2 light chain *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-44*01 (90.8%) -IGLJ2*01 (100%)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dimer (232-232":235-235")-bisdisulfide, non-glycosylated, produced in Chinese hamster ovary (CHO)-DG44i cell line

vixarélimab

immunoglobuline G4-G1-lambda2, anti-[*Homo sapiens* OSMR (récepteur de l'oncostatine M, OSMRB, OSMRbeta)], anticorps monoclonal *Homo sapiens*; chaîne lourde gamma4-gamma1 *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV1-8*01 (95.9%) -(IGHD) -IGHJ6*01 (95.0%)) CDR-IMGT [8.8.19] (26-33.51-58.97-115) (1-126)-*Homo sapiens* IGHG4*01 CH1-h-CH2, G4v5 h P10, G4v36 CH2 Q84.4 (CH1 (127-224), charnière 1-12 S10>P (234) (225-236), CH2 N84.4>Q (303) (237-346)) (127-346) -IGHG1*01 CH3-CHS, G1m1 (CH3 D12 (362), L14 (364) (347-451), CHS K>del (452)) (347-452)], (140-215')-disulfure avec la chaîne légère lambda2 *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-44*01 (90.8%) -IGLJ2*01 (100%)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dimère (232-232":235-235")-bisdisulfure, non-glycosylé, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-DG44i

vixarelimab

inmunoglobulina G4-G1-lambda2, anti-[*Homo sapiens* OSMR (receptor de la oncostatina M, OSMRB, OSMRbeta)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma4-gamma1 *Homo sapiens* (1-452) [VH (*Homo sapiens* IGHV1-8*01 (95.9%) -(IGHD) -IGHJ6*01 (95.0%)) CDR-IMGT [8.8.19] (26-33.51-58.97-115) (1-126)-*Homo sapiens* IGHG4*01 CH1-h-CH2, G4v5 h P10, G4v36 CH2 Q84.4 (CH1 (127-224), bisagra 1-12 S10>P (234) (225-236), CH2 N84.4>Q (303) (237-346)) (127-346) -IGHG1*01 CH3-CHS, G1m1 (CH3 D12 (362), L14 (364) (347-451), CHS K>del (452)) (347-452)], (140-215')-disulfuro con la cadena ligera lambda2 *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-44*01 (90.8%) -IGLJ2*01 (100%)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dímero (232-232":235-235")-bisdisulfuro, no glicosilado, producido en las células ováricas de hámster chino (CHO) línea celular CHO-DG44i

Heavy chain / Chaîne lourde / Cadena pesada
QVQLVQSGAE VKKPGASVKV SCKASGYTFT SYEINWVRQA TGQGLEWMGW 50
MNPNSGYTGY AQKFQGRVTM TRDTSISTAY MEMSSLRSED TAVYYCARDI 100
VAANTDYFFY YGMDVWGQGT TVTVSSASTK GPSVFLAPC SRSTSESTAA 150
LGCLVKDYFP EPVTVSWNSG ALTSGVHTFP AVLQSSGLYS LSSVTVPS 200
SLGKTYTCN VDHKPSNTKV DKRVESKYGP PCPPCPAPEF LGGPSVFLFP 250
PKPKDTLMIS RTPEVTCVVV DVSQEDPEVQ FNWYDVGEV HNAKTKPREE 300
QFQSTYRVVS VLTVLHQDWL NGKEYKCKVS NKGLPSSIEK TISKAKGQPR 350
EPQVYTLPPS RDELTKQVS LTCLVKGFYP SDIAVEWESN GQPENNYKTT 400
PPVLDSDGSF FLYSKLTVDK SRWQQGNVFS CSVMHEALHN HYTKSLSLS 450
PG 452

Light chain / Chaîne légère / Cadena ligera
QSVLTQPPSA SGTGQQRVTI SCSSGNSNIG SNTVNWYHQL PGTAPKLLIY 50
NINKRPSGVP DRFGSGSKGS SASLAISGLQ SEDEADYYCS TWDDSLDGVV 100
FGGKTLTVL GQPKAAPSVT LFPPSSEELQ ANKATLVCLI SDFYPGAVTV 150
AWKADSSPVK AGVETTPSK QSNNKYAASS YLSLTPEQWK SHRSYSCQYT 200
HEGSTVEKTV APTCEC 216

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 153-209 267-327 373-431
22"-96" 153"-209" 267"-327" 373"-431"
Intra-L (C23-C104) 22"-89" 138"-197"
22"-89" 138"-197"
Inter-H-L (CH1 10-CL 126) 140-215' 140"-215"
Inter-H-H (h 8, h 11) 232-232" 235-235"

N-terminal glutaminy cyclization to pyroglutamyl (pE, 5-oxoprolyl)
H VH Q1:
1, 1"
L VL Q1:
1', 1"

No N-glycosylation sites / pas de sites de N-glycosylation / ningùn posició de N-glicosilació
H CH2 N84,4-Q (303)
Aglycosylated

vocacapsaicinum
vocacapsaicin

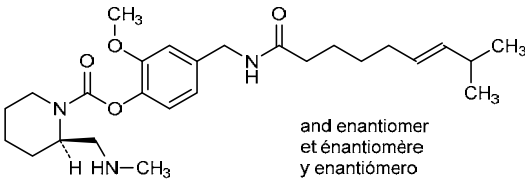
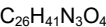
rac-2-methoxy-4-[[[(6*E*)-8-methylnon-6-enamido]methyl]phenyl (2*R*)-2-[(methylamino)methyl]piperidine-1-carboxylate

vocacapsaïcine

rac-(2*R*)-2-[(méthylamino)méthyl]pipéridine-1-carboxylate de 2-méthoxy-4-[[[(6*E*)-8-méthylnon-6-énamido]méthyl]phényle

vocacapsaïcina

rac-(2*R*)-2-[(metilamino)metil]piperidina-1-carboxilato de 4-[[[(6*E*)-8-metilnon-6-enamido]metil]-2-metoxifenilo



vodobatinibum
vodobatinib

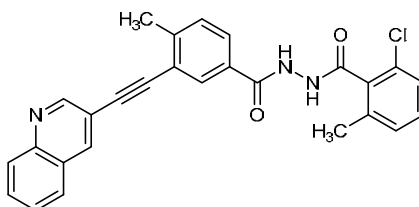
2-chloro-6-methyl-*N'*-(4-methyl-3-[(quinolin-3-yl)ethynyl]benzoyl)benzohydrazide

vodobatinib

2-chloro-6-méthyl-*N'*-(4-méthyl-3-[(quinoléin-3-yl)éthynyl]benzoyl)benzohydrazide

vodobatinib

2-cloro-6-metil-*N'*-(4-metil-3-[(quinolein-3-il)etinil]benzoil)benzohidrazida



vudalimabum

vudalimab

immunoglobulin half-IG G1-kappa/scFv-h-CH2-CH3, anti-[*Homo sapiens* CTLA4 (cytotoxic T-lymphocyte-associated protein 4, CD152)] and anti-[*Homo sapiens* PDCD1 (programmed cell death 1, PD-1, PD1, CD279)], monoclonal antibody, bispecific; gamma1 heavy chain anti-CTLA4 (1-447) [VH (*Homo sapiens* IGHV3-48*03 (90.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107)(1-118) -*Homo sapiens* IGHG1*03v, G1m3>G1m17, nG1m1 (CH1 N114>D (209) R120>K (215) (119-216), hinge 1-15 (217-231), CH2 E1.4,L1.3>P (234), L1.2>V (235), G1.1>A (236), S29>K (267), Q84.2>E (295) (232-340), CH3 E12 (356), M14 (358), L24>D (368), K26>S (370), N44>D (384), Q97>E (418), N100>D (421), M107>L (428), N114>S (434) (341-445), CHS (446-447)) (119-447)], (221-215')-disulfide with kappa light chain anti-CTLA4, *Homo sapiens* (1'-215') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (99.0%) -IGKJ1*01 (100%)) CDR-IMGT [7.3.9] (27-33.51-53.90-98) (1'-108') -*Homo sapiens* IGKC*01 (100%), Km3 A45.1 (154), V101 (192) (109'-215')]; IG scFv-h-CH2-CH3 single chain, anti-PDCD1 (1"-480") [scFv V-kappa-VH anti-PDCD1 (1"-249") [V-KAPPA (*Homo sapiens* IGKV3-15*01 (80.0%) -IGHJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1"-107") -20-mer tetrakis(glycyl-lysyl-prolyl-glycyl-seryl) linker (108"-127") -VH (*Mus musculus* IGHV6-6*02 (87.0%) -(IGHD) -IGHJ1*03 (86.7%)/*Homo sapiens* IGHV3-15*07 (81.8%) -(IGHD) -IGHJ2*01 (93.3%)) CDR-IMGT [8.10.13] (153-160.178-187.226-238) (128"-249")] -*Homo sapiens* IGHG1*03 h-CH2-CH3, nG1m1 (250"-480") [hinge 1-15 C5>S (254) (250-264) -CH2 E1.4,L1.3>P (267), L1.2>V (268), G1.1>A (269), S29>K (300) (265-373), CH3 E12 (389), E13>Q (390), M14 (391), S20>K (397), M107>L (461), N114>S (467) (374-478), CHS (479-480)]]; dimer (227-260":230-263")-bisdisulfide, produced in Chinese hamster ovary (CHO)-S cell line, glycoform alfa

vudalimab

immunoglobuline demi-IG G1-kappa/scFv-h-CH2-CH3, anti-[*Homo sapiens* CTLA4 (protéine 4 associée aux lymphocytes T cytotoxiques, CD152)] et anti-[*Homo sapiens* PDCD1 (protéine 1 de mort cellulaire programmée, PD-1, PD1, CD279)], anticorps monoclonal, bispécifique;

chaîne lourde gamma1 anti-CTLA4, (1-447) [VH (*Homo sapiens*IGHV3-48*03 (90.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens*IGHG1*03v, G1m3>G1m17, nG1m1 (CH1 N114>D (209) R120>K (215) (119-216), charnière 1-15 (217-231), CH2 E1.4,L1.3>P (234), L1.2>V (235), G1.1>A (236), S29>K (267), Q84.2>E (295) (232-340), CH3 E12 (356), M14 (358), L24>D (368), K26>S (370), N44>D (384), Q97>E (418), N100>D (421), M107>L (428), N114>S (434) (341-445), CHS (446-447)) (119-447)], (221-215')-disulfure avec la chaîne légère kappa anti-CTLA4,*Homo sapiens* (1'-215') [V-KAPPA (*Homo sapiens*IGKV3-20*01 (99.0%) -IGKJ1*01 (100%)) CDR-IMGT [7.3.9] (27-33.51-53.90-98) (1'-108') -*Homo sapiens*IGKC*01 (100%), Km3 A45.1 (154), V101 (192) (109'-215')]; IG scFv-h-CH2-CH3 chaîne unique, anti-PDCD1 (1"-480") [scFv V-kappa-VH anti-PDCD1 (1"-249") [V-KAPPA (*Homo sapiens*IGKV3-15*01 (80.0%) -IGHJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1"-107") -20-mer tétrakis(glycyl-lysyl-prolyl-glycyl-séryl) linker (108"-127") -VH (*Mus musculus*IGHV6-6*02 (87.0%) -(IGHD) -IGHJ1*03 (86.7%)/*Homo sapiens*IGHV3-15*07 (81.8%) -(IGHD) -IGHJ2*01 (93.3%)) CDR-IMGT [8.10.13] (153-160.178-187.226-238) (128"-249")]] -*Homo sapiens*IGHG1*03 h-CH2-CH3, nG1m1 (250"-480") [charnière 1-15 C5>S (254) (250-264) -CH2 E1.4,L1.3>P (267), L1.2>V (268), G1.1>A (269), S29>K (300) (265-373),CH3 E12 (389), E13>Q (390), M14 (391), S20>K (397), M107>L (461), N114>S (467) (374-478), CHS (479-480)]]; dimère (227-260":230-263")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-S, glycoforme alfa

vudalimab

immunoglobulina demi-IG G1-kappa/scFv-h-CH2-CH3, anti-[*Homo sapiens*CTLA4 (proteína 4 asociada a los linfocitos T citotóxicos, CD152)] y anti-[*Homo sapiens*PDCD1 (proteína 1 de muerte celular programada, PD-1, PD1, CD279)], anticuerpo monoclonal, biespecífico; cadena pesada gamma1 anti-CTLA4 (1-447) [VH (*Homo sapiens*IGHV3-48*03 (90.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.11] (26-33.51-58.97-107) (1-118) -*Homo sapiens*IGHG1*03v, G1m3>G1m17, nG1m1 (CH1 N114>D (209) R120>K (215) (119-216), bisagra 1-15 (217-231), CH2 E1.4,L1.3>P (234), L1.2>V (235), G1.1>A (236), S29>K (267), Q84.2>E (295) (232-340), CH3 E12 (356), M14 (358), L24>D (368), K26>S (370), N44>D (384), Q97>E (418), N100>D (421), M107>L (428), N114>S (434) (341-445), CHS (446-447)) (119-447)], (221-215')-disulfuro con la cadena ligera kappa anti-CTLA4,*Homo sapiens* (1'-215') [V-KAPPA (*Homo sapiens*IGKV3-20*01 (99.0%) -IGKJ1*01 (100%)) CDR-IMGT [7.3.9] (27-33.51-53.90-98) (1'-108') -*Homo sapiens*IGKC*01 (100%), Km3 A45.1 (154), V101 (192) (109'-215')]; IG scFv-h-CH2-CH3 cadena única, anti-PDCD1 (1"-480") [scFv V-kappa-VH anti-PDCD1 (1"-249") [V-KAPPA (*Homo sapiens*IGKV3-15*01 (80.0%) -IGHJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1"-107") -20-mer tetrakis(glicil-lisil-proil-glicil-seril) linker (108"-127") -VH (*Mus musculus*IGHV6-6*02 (87.0%) -(IGHD) -IGHJ1*03 (86.7%)/*Homo sapiens*IGHV3-15*07 (81.8%) -(IGHD) -IGHJ2*01 (93.3%)) CDR-IMGT [8.10.13] (153-160.178-187.226-238) (128"-249")]] -*Homo sapiens*IGHG1*03 h-CH2-CH3, nG1m1 (250"-480") [bisagra 1-15 C5>S (254) (250-264) -CH2 E1.4,L1.3>P (267), L1.2>V (268), G1.1>A (269), S29>K (300) (265-373),CH3 E12 (389), E13>Q (390), M14 (391), S20>K (397), M107>L (461), N114>S (467) (374-478), CHS (479-480)]]; dímero (227-260":230-263")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-S, forma glicosilada alfa

1. Heavy chain / Chaîne lourde / Cadena pesada (anti-CTLA4)	
EVQLVESGGG LVKPGGSLRL SCAASGFTFS SVTHHWVRQA	PGKGLEWVSF 50
ISYDGNKKYY ADSVKGRFTI SRDNAKNSLY LQMNSLRAED	TAVYYCARTG 100
WLGPFDYWGQ GTLVTVSSAS TKGPSVFPLA PSSKSTSGGT	AALGCLVKDY 150
FPFEPVTVSWN SGALTSGVHT FPAVLQSSGL YSLSSVTVTP	SSSLGTQTYI 200
CNVNHKPSDT VKDKKVEPKS CDKTHTCPPC PAPPVAGPSV	FLFPPKPKDT 250
LMISRTPEVT CVVVDVKHED PEVKFNWYVD GVEVHNAKT	PREEEYNSTY 300
RVVSVLTVLH QDWLNGKEYK CKVSNKALPA PIEKTIISKAK	GQPREPQVYT 350
LPSPREEMTK NQVSLTCDVS GFYPSDIAVE WESDGPENN	YKTTTPVLDS 400
DGSFFLYSKL TVDKSRWEQG DVFSCSVLHE ALHSHYTQKS	LSLSPGK 447
2. Light chain / Chaîne légère / Cadena ligera (anti-CTLA4)	
EIVLTQSPGT LSLSPGERAT LSCRASQSVS SSVLAWYQKQ	PGQAPRLLIY 50
GAFSRATGIP DRFSGSGSGT DFTLTISRLE PEDFAVYYCQ	QYGSSPWTFG 100
QGTKEVIKRT VAAPSVFIFP PSDEQLKSGT ASVVCLLNMF	YPREAKVQWK 150
VDNALQSGNS QESVTEQDSK DSTYLSSTL TLISKADYEKH	KVYACEVTHQ 200
GLSSPVTKSF NRGEC	215
3. Chain / Chaîne / Cadena IG scFv-h-CH2-CH3 (anti-PDCC1)	
EIVLTQSPAT LSASGERVT LTCRASQSVG NDVAWYQKQ	GQAPRLLIY 50
ASHRYTGVPD RFTSGSGYGE FTLTISSTSG EDFAVYYCQ	PFSSPRTFGG 100
GTKVEIKGKP GSGKPGSGKP GSGKPGSEVQ LVESGGGLVK	PGGSLRLSCV 150
ASGFTFSNYW MNWVRQAPGK GLEWVAEIRL YSNMYATHYA	ESVKGRTTIS 200
RDDSKSTLYL QMNNLKTEDT GVYYCTRYG NYGGYFDWVG	RGTLVTVSSE 250
PKSSDKTHTC PPCAPPVAG PSVFLFPKP KDITLMISRT	EVTGVVVDVK 300
HEDPEVKFNW YVDGVEVHNA KTKPREEQYN STYRVSVLT	VLHQDWLNGK 350
EYKCKVSNKA LPAPIEKTIS KAKGQPREPQ VYTLPPSREQ	MTKNQVKLTC 400
LVKGYPYSDI AVEWESNGQP ENNYKTTTPV LDSDGGSFLY	SKLTVDKSRW 450
QQGMVFCSSV LHEALSHYT QKSLSLSPGK	480
Post-translational modifications	
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro	
Intra-H (C23-C104)	22-96 145-201 261-321 367-425
Intra-L (C23-C104)	23*-89* 135*-195*
Intra-chain 3	23*-88* 149*-225* 294*-354* 400*-458*
Inter-H-L (h 5-CL 126)	221-215*
Inter-H-chain 3 (h 11, h 14)	227-260* 230-263"
N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación	
H CH2 N84,4:	
297, 330"	
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.	
C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal	
H CHS K2:	
447, 480"	

zalsenertantum tetraxetanum

zalsenertant tetraxetan

18⁴,18⁷,18¹⁰-tris(carboxymethyl)-4⁵-(2,6-dimethoxyphenyl)-7,11,15-trimethyl-3,6,16-trioxo-5²-(propan-2-yl)-2,7,11,15-tetraaza-18(1)-(1,4,7,10-tetraazacyclododecana)-4(3,1)-pyrazola-1(2)-adamantana-5(1,4)-benzenaoctadecaphane-1²-carboxylic acid

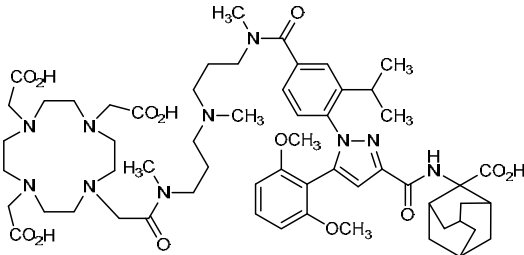
zalsénertant tétraxetan

acide 18⁴,18⁷,18¹⁰-tris(carboxyméthyl)-4⁵-(2,6-diméthoxyphényl)-7,11,15-triméthyl-3,6,16-trioxo-5²-(propan-2-yl)-2,7,11,15-tétraaza-18(1)-(1,4,7,10-tétraazacyclododécana)-4(3,1)-pyrazola-1(2)-adamantana-5(1,4)-benzénoctadécaphane-1²-carboxylique

zalsenertant tetraxetán

ácido 18⁴,18⁷,18¹⁰-tris(carboximetil)-4⁵-(2,6-dimetoxifenil)-7,11,15-trimetil-3,6,16-trioxo-5²-(propan-2-il)-2,7,11,15-tetraaza-18(1)-(1,4,7,10-tetraazaciclododecana)-4(3,1)-pirazola-1(2)-adamantana-5(1,4)-bencenaoctadecafano-1²-carboxílico

C₅₈H₈₄N₁₀O₁₃



zanidatamabum zovodotinum #

zanidatamab zovodotin

immunoglobulin half-IG G1-kappa/scFv-h-CH2-CH3, anti-[*Homo sapiens* ERBB2 (epidermal growth factor receptor 2, receptor tyrosine protein kinase erbB-2, EGFR2, HER2, HER-2, p185c-erbB2, NEU, CD340)], humanized monoclonal antibody, biparatopic (targeting two different non-overlapping epitopes on ERBB2, on extracellular domains 2 (ECD2) and 4 (ECD4), respectively), conjugated to auristatin E; gamma1 heavy chain, anti-ERBB2 extracellular domain 2 (ECD2), humanized (1-449) [VH humanized (*Homo sapiens* IGHV3-66*01 (78.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT[8.8.12] (27-34.52-59.98-109) (1-120) -*Homo sapiens* IGHG1*01 G1m17,1 (CH1 K120 (217) (121-218), hinge 1-15 (219-233), CH2 (234-343), CH3 T6>V (353), L7>Y (354), D12 (359), L14 (361), F85.1>A (408), Y86>V (410) (344-448), CHS K2>del (449)) (121-449)], (223- 215')-disulfide with kappa light chain, anti ERBB2 ECD2, humanized (1'-215') [V-KAPPA humanized (*Homo sapiens* IGKV1-16*01 (84.2%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (28-33.51-53.90-98) (1'-108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]; IG scFv-h-CH2-CH3 single chain, anti-ERBB2 extracellular domain 4 (ECD4), humanized (1"-481") [scFv V-kappa-VH anti-ERBB2 ECD4 (1'-248') [V-KAPPA humanized (*Homo sapiens* IGKV1-39*01 (86.3%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (28-33.51-53.90-98) (1'-108') -20-mer pentakis(diglycyl-seryl-glycyl) linker (109"-128") -VH humanized (*Homo sapiens* IGHV3-66*01 (81.6%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (154-161.179-186.225-237) (129"-248")]] -dialanyl linker (249"-250") -*Homo sapiens* IGHG1*01 h-CH2-CH3, G1m1 (251"-481") [hinge 1-15, C5>S (255) (251-265), CH2 (266-375), CH3 T6>V (385), D12 (391), L14 (393), T22>L (401), K79>L (427), T81>W (429) (376-480), CHS K2>del (481)]]; dimer (229-261":232-264")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa, conjugated, on an average of 2 to 3 cysteinyl, to monomethylauristatin E (MMAE), via a cleavable maleimidocaproyl-valyl-citrullinyl-p-aminobenzylcarbonyl (mc-val-cit-PABC) type linker

zanidatamab zovodotine

immunoglobuline G1-kappa, anti-[*Homo sapiens* ERBB2 (récepteur 2 du facteur de croissance épidermique, récepteur tyrosine-protéine kinase erbB-2, EGFR2, HER2, HER-2, p185c-erbB2, NEU, CD340)], anticorps monoclonal humanisé, biparatopique (ciblant deux épitopes différents non chevauchants sur ERBB2, respectivement sur les domaines extracellulaires 2 (ECD2) et 4 (ECD4)), conjugué à l'auristatine E; chaîne lourde gamma1 anti-ERBB2 domaine extracellulaire 2 (ECD2), humanisée (1-449) [VH humanisé (*Homo sapiens* IGHV3-66*01 (78.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.12] (27-34.52-59.98-109) (1-120) -*Homo sapiens* IGHG1*01 G1m17,1 (CH1 K120 (217) (121-218), charnière 1-15 (219-233), CH2 (234- 343), CH3 T6>V (353), L7>Y (354), D12 (359), L14 (361), F85.1>A (408), Y86>V (410) (344-448), CHS K2>del (449)) (121-449)], (223-215')-disulfure avec la chaîne légère kappa, anti ERBB2 ECD2, humanisée (1'-215') [V-KAPPA humanisé (*Homo sapiens* IGKV1-16*01 (84.2%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (28-33.51-53.90-98) (1'-108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')];

IG scFv-h-CH2-CH3 chaîne unique, anti-ERBB2 domaine extracellulaire 4 (ECD4), humanisée (1"-481") [scFv V-kappa-VH anti-ERBB2 ECD4 (1'-248') [V-KAPPA humanisé (*Homo sapiens* IGKV1-39*01 (86.3%) - IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (28-33.51-53.90-98) (1'-108') -20-mer pentakis(diglycyl-seryl-glycyl) linker (109"- 128") -VH humanisé (*Homo sapiens* IGHV3-66*01 (81.6%) -(IGHD) - IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (154-161.179-186.225-237) (129"-248")]] -dialanyl linker (249"-250") -*Homo sapiens* IGHG1*01 h-CH2-CH3, G1m1 (251"-481") [charnière 1-15 C5>S (255) (251- 265), CH2 (266-375), CH3 T6>V (385), D12 (391), L14 (393), T22>L (401), K79>L (427), T81>W (429) (376-480), CHS K2>del (481)]]; dimère (229-261":232- 264")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa; conjugué, sur 2 à 3 cystéinyl en moyenne, au monométhylauristatine E (MMAE), via un linker clivable de type maléimidécaproyl-valyl-citrullinyl-*p*-aminobenzylcarbonyl (mc-val-cit-PABC).

zanidatamab zovodotina

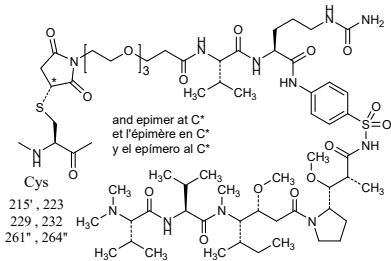
inmunoglobulina G1-kappa, anti-[*Homo sapiens* ERBB2 (receptor 2 del factor de crecimiento epidérmico, receptor tirosina-proteína kinasa erbB-2, EGFR2, HER2, HER-2, p185c-erbB2, NEU, CD340)], anticuerpo monoclonal humanizado, biparatópico (dirigiendo dos epítomos diferentes no superpuestos sobre ERBB2, respectivamente sobre los dominios extracelulares 2 (ECD2) y 4 (ECD4)), conjugado con la auristatina E; cadena pesada gamma1 anti-ERBB2 dominio extracelular 2 (ECD2), humanizada (1-449) [VH humanizado (*Homo sapiens* IGHV3-66*01 (78.8%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.12] (27-34.52-59.98-109) (1-120) -*Homo sapiens* IGHG1*01 G1m17.1 (CH1 K120 (217) (121-218), bisagra 1-15 (219-233), CH2 (234- 343), CH3 T6>V (353), L7>Y (354), D12 (359), L14 (361), F85.1>A (408), Y86>V (410) (344-448), CHS K2>del (449)) (121-449)], (223-215')-disulfuro con la cadena ligera kappa, anti ERBB2 ECD2, humanizada (1'-215') [V-KAPPA humanizado (*Homo sapiens* IGKV1-16*01 (84.2%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (28-33.51-53.90-98) (1'-108') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (154), V101 (192) (109'-215')]]; IG scFv-h-CH2-CH3 cadena única, anti-ERBB2 dominio extracelular 4 (ECD4), humanizada (1"-481") [scFv V-kappa-VH anti-ERBB2 ECD4 (1'-248') [V-KAPPA humanizado (*Homo sapiens* IGKV1-39*01 (86.3%) - IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (28-33.51-53.90-98) (1'-108') -20-mer pentakis(diglicil-seril-glicil) linker (109"- 128") -VH humanizado (*Homo sapiens* IGHV3-66*01 (81.6%) -(IGHD) - IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (154-161.179-186.225-237) (129"-248")]] -dialanyl linker (249"-250") -*Homo sapiens* IGHG1*01 h-CH2-CH3, G1m1 (251"-481") [bisagra 1-15 C5>S (255) (251- 265), CH2 (266-375), CH3 T6>V (385), D12 (391), L14 (393), T22>L (401), K79>L (427), T81>W (429) (376-480), CHS K2>del (481)]]; dímero (229-261":232- 264")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa; conjugado, sobre 2 a 3 restos cisteinil en término medio, con monometilauristatina E (MMAE), mediante un conector escindible de tipo maleimidécaproyl-valil-citrullinil-*p*-aminobenzilcarbonil (mc-val-cit-PABC).

1. Heavy chain / Chaîne lourde / Cadena pesada (anti-ERBB2 ECD2)					
GEVQLVESGG	GLVQPGSLR	LSCAASGFTF	ADYTMWVRQ	APGKLEWVG	50
DVNPNSGGSI	YNQRFGRFT	FSVDRSKNTL	YLQMNSLRAE	DTAAYVCARN	100
LGPSFYFDYW	QGGLTLTVSS	ASTKGPSVFP	LAPSSKSTSG	GTAALGCLVK	150
DYFPEPVTVS	WNSGALTSGV	HTFPAVLQSS	GLYSLSSVVT	VPSSSLGTQT	200
YICNVNPKPS	NTKVDKKEVP	KSCDKHTTCP	PCPAPELLGG	PSVFLFPPKP	250
KDTLMISRTL	EVTCVVVDVS	HEDPEVKFNW	YVDGVEVHNA	KTKPREEQYN	300
STYRVVSVLT	VLHQDWLNGK	EYKCKVSNKA	LPAPTEKTI	KAKGQPREPQ	350
VYVYPPSRDE	LTKNQVSLTC	LVKGFYPSDI	AVEWESNGQP	ENNYKTTTPV	400
LDSDGSFALV	SKLTVDKSRW	QQGNVFSQSV	MHEALHNHYT	QKSLSLSPG	449
2. Light chain / Chaîne légère / Cadena ligera (anti-ERBB2 ECD2)					
GDIQMTQSPS	SLASVSGDRV	TIITCKASQDV	SIGVAMWYQK	PGKAPKLLIY	50
SASYRYTGVP	SRFSGSGSGT	DFTLTISSLQ	PEDFATYYCQ	QYYIYPATFG	100
QGTVKEIKRT	VAAPSVFIFP	PSDEQLKSGT	ASVCLLNNF	YPREAKVQWK	150
VDNALQSGNS	QESVTEQDSK	DSTYLSSTLT	TLSKADYEKH	KYACEVTHQ	200
GLSSPVTKSF	NRGEC				215
3. Chain / Chaîne / Cadena: IG scFv-h-CH2-CH3 (anti-ERBB2 ECD4)					
GDIQMTQSPS	SLASVSGDRV	TIITCKASQDV	NTAVAMWYQK	PGKAPKLLIY	50
SASFLYSGVP	SRFSGSGSGT	DFTLTISSLQ	PEDFATYYCQ	QHYTTPPTFG	100
QGTKVEIKGG	SGGSGSGSGS	GSGSGSGSGV	QLVESGGGLV	QPGGSLRLSC	150
AASGFNIKDT	YIHWVRQAPG	KGLEWARIY	PTNGYTRYAD	SVKGRFTISA	200
DTSKNTAYLQ	MNSLRAEDTA	VYYCSRWGGD	GFYAMDYWGQ	GLTVTVSSAA	250
EPKSSDKTHT	CPPCPAPELL	GGPSVFLFPP	KPKDTLMISR	TPEVTCVVD	300
VSHEDPEVKF	NWYVDGVEVH	NAKTKPREEQ	YNSTYRVVSV	LTVLHQDWLN	350
GKEYKCKVSN	KALPAPIEKT	ISKAKGQPRE	PQVYVLPSPR	DELTKNQVSL	400
LCLVKGFYPS	DIAVEWESNG	QPENNYLTWP	PVLSDSGSFF	LYSKLTVDKS	450
RWQQGNVFSC	SVMHEALHNH	YTKQSLSLSP	G		481

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 23-97 147-203 264-324 370-428
Intra-L (C23-C104) 24'-89' 135'-195'
Intra-chain 3 24"-89" 150"-224" 296"-356" 402"-460"
Inter-H-L (h 5-CL 126)* 223-215'
Inter-H-chain 3 (h 11, h 14)* 229-261" 232-264"
*At least two of the three inter-chain disulfide bridges are not present, an average of 2 to 3 cysteinyl being conjugated each via a thioether bond to a drug linker.
*Au moins deux des trois ponts disulfures inter-chaînes ne sont pas présents, 2 à 3 cystéinyl en moyenne étant chacun conjugué via une liaison thioéther à un linker-principe actif.
*Al menos dos de los tres puentes disulfuro inter-catenarios no están presentes, una media de 2 a 3 cisteinil está conjugada a conectores de principio activo.

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
300, 332"
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantennarios complejos fucosilados.

Modified residues / résidus modifiés / restos modificados:



zatolmilastum
zatolmilast

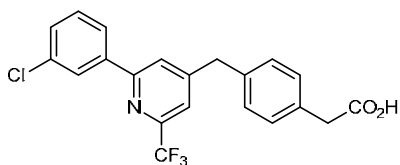
(4-[[2-(3-chlorophenyl)-6-(trifluoromethyl)pyridin-4-yl)methyl]phenyl)acetic acid

zatolmilast

acide (4-[[2-(3-chlorophényl)-6-(trifluorométhyl)pyridin-4-yl]méthyl]phényl)acétique

zatolmilast

ácido (4-[[2-(3-clorofenil)-6-(trifluorometil)piridin-4-il]metil]fenil)acético

C₂₁H₁₅ClF₃NO₂

zelavespibum

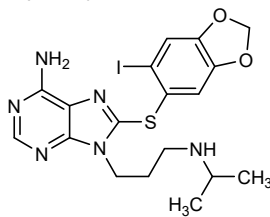
zelavespib

8-[(6-iodo-2*H*-1,3-benzodioxol-5-yl)sulfanyl]-9-{3-[(propan-2-yl)amino]propyl}-9*H*-purin-6-amine

zélavespib

8-[(6-iodo-2*H*-1,3-benzodioxol-5-yl)sulfanyl]-9-{3-[(propan-2-yl)amino]propyl}-9*H*-purin-6-amine

zelavespib

8-[(6-iodo-2*H*-1,3-benzodioxol-5-il)sulfanyl]-9-{3-[(propan-2-il)amino]propil}-9*H*-purin-6-aminaC₁₈H₂₁IN₆O₂S

zimberelimabum #

zimberelimab

immunoglobulin G4-lambda, anti-[*Homo sapiens* PDCD1 (programmed cell death 1, PD-1, PD1, CD279)], *Homo sapiens* monoclonal antibody; gamma4 heavy chain *Homo sapiens* (1-450) [VH (*Homo sapiens* IGHV4-39*01 (87.9%) -(IGHD) - IGHJ4*01 (100%)) CDR-IMGT [10.7.15] (26-35.53-59.98-112) (1-123)-*Homo sapiens* IGHG4*01 (CH1 (124-221), hinge 1-12 S10>P (231) (222-233), CH2 (234-343), CH3 (344-448), CHS (449-450) (124-450)), (137-215')-disulfide with lambda light chain *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV2-14*01 (94.9%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.10] (26-34.52-54.91-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dimer (229-229'':232-232'')-bisdisulfide, produced in Chinese hamster ovary (CHO)-K1 cell line, glycoform alfa

zimberélimab

immunoglobuline G4-lambda, anti-[*Homo sapiens* PDCD1 (protéine 1 de mort cellulaire programmée, PD-1, PD1, CD279)], anticorps monoclonal *Homo sapiens*; chaîne lourde gamma4 *Homo sapiens* (1-450) [VH (*Homo sapiens* IGHV4-39*01 (87.9%) -(IGHD) - IGHJ4*01 (100%)) CDR-IMGT [10.7.15] (26-35.53-59.98-112) (1-123)-*Homo sapiens* IGHG4*01 (CH1 (124-221), charnière 1-12 S10>P (231) (222-233), CH2 (234-343), CH3 (344-448), CHS (449-450) (124-450)), (137-215')-disulfure avec la chaîne légère

zimberelimab

lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV2-14*01 (94.9%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.10] (26-34.52-54.91-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dimère (229-229":232-232")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-K1, glycoforme alfa

immunoglobulina G4-lambda, anti-[*Homo sapiens* PDCD1 (proteína 1 de muerte celular programada PD-1, PD1, CD279), anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma4 *Homo sapiens* (1-450) [VH (*Homo sapiens* IGHV4-39*01 (87.9%) -IGHD) -IGHJ4*01 (100%)) CDR-IMGT [10.7.15] (26-35.53-59.98-112) (1-123)-*Homo sapiens* IGHG4*01 (CH1 (124-221), bisagra 1-12 S10>P (231) (222-233), CH2 (234-343), CH3 (344-448), CHS (449-450) (124-450)], (137-215')-disulfuro con la cadena ligera lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV2-14*01 (94.9%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.10] (26-34.52-54.91-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dímero (229-229":232-232")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-K1, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada

QLQLQESGPG	LVKPSSETLTL	TCTVSADSI	STTYVWVIR	QPPGKGLEWI	50
GSISYSGSTY	YNPSLKSRT	VSVDTSKNQF	SLKLNSVAAT	DTALYYCARH	100
LGYNTRYLPF	DYWGQGT	LVT VSSASTKGPS	VFPLAPCSRS	TSESTAALGC	150
LVKDYPPEPV	TVSWNSGALT	SGVHTFPAVL	QSSGLYSLSS	VVTPSSSLG	200
TKTYTCNVDH	KPSNTKVDKR	VESKYGPPCP	PCPAPEFLGG	PSVFLFPPKP	250
KDTLMISRT	EVTCVVVDVS	QEDPEVQFNW	YVDGVEVHNA	KTKPREEQFN	300
STYRVVSVLT	VLHQDWLNGK	EYKCKVSNKG	LPSSIEKTIS	KAKGQPREPQ	350
VYTLPPSQEE	MTKNQVSLTC	LVKGFYPSDI	AVEWESNGQP	ENNYKTTTPPV	400
LDSGGSFFLY	SRLTVDKSRW	QEGNVFSCSV	MHEALHNYHT	QKSLSLSLGK	450

Light chain / Chaîne légère / Cadena ligera

QSALTQPASV	SGSPGQSITI	SCTGTSSDVG	FYNYVSWYQQ	HPGKAPELMI	50
YDVSNRPSGV	SDRFSGSKSG	NTASLTISGL	QAEDEADYYC	SSYTSISTWV	100
FGGGTKLTVL	GQPKAAPSVT	LFPPSSEELQ	ANKATLVCLT	SDFYPGAVTV	150
AWKADSSPVK	AGVETTTPSK	QSNKNYAASS	YLSLTPEQWK	SHRSYSCQVT	200
HEGSTVEKTV	APTECS				216

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104)	22"-97"	150"-206"	264"-324"	370"-428"
	22"-97"	150"-206"	264"-324"	370"-428"

Intra-L (C23-C104)	22"-90"	138"-197"
	22"-90"	138"-197"

Inter-H-L (CH1 10-CL 126)	137"-215"	137"-215"
---------------------------	-----------	-----------

Inter-H-H (h 8, h 11)	229"-229"	232"-232"
-----------------------	-----------	-----------

N-terminal glutaminyl cyclization to pyroglutamyl (pE, 5-oxoprolyl)

L VH Q1:

I, I"

L VL Q1:

I', I'''

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

300, 300"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires

complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

450, 450"

Please find below the names from proposed INN List 124 – COVID-19 (special edition) that have reached the status of recommended INN. Veuillez trouver ci-dessous les noms de la Liste 124 des DCI proposées – COVID-19 (édition spéciale) ayant atteint le statut de DCI recommandées. Por favor, abajo se encuentran los nombres de la Lista 124 de DCI propuestas – COVID-19 (edición especial) que han alcanzado el estatus de DCI recomendadas.

Latin, English, French, Spanish: <i>Recommended INN</i>	<i>Chemical name or description; Molecular formula; Graphic formula</i>
<i>DCI Recommandée</i>	<i>Nom chimique ou description; Formule brute; Formule développée</i>
<i>DCI Recomendada</i>	<i>Nombre químico o descripción; Fórmula molecular; Fórmula desarrollada</i>

abdavomeranum #

abdavomeran

messenger RNA (mRNA), 5'-capped, encoding the codon-optimised receptor binding domain (RBD) of the SARS-CoV-2 (severe acute respiratory syndrome coronavirus 2, GenBank: MN908947.3) spike (S) glycoprotein, expressed as a fusion protein 5' with an extended S glycoprotein signal peptide and connected 3' by a glycine/serine (GS)-rich linker peptide with a T4 fibrin trimerization domain, flanked by 5' and 3' untranslated regions and a 3' poly(A) tail; contains N1-methylpseudouridine instead of uridine (*all-U>m¹Ψ*).

abdavoméran

ARN messager (ARNm), protégé en 5', codant pour le domaine de liaison au récepteur (RBD) de la glycoprotéine spike (S) du SARS-CoV-2 (coronavirus 2 du syndrome respiratoire aigu sévère, GenBank: MN908947.3) avec des codons optimisés, exprimé comme une protéine de fusion 5'-avec un peptide signal de la glycoprotéine S allongé et lié en 3' par un peptide de liaison riche en glycine / sérine (GS) avec un domaine de trimérisation de la fibrine de T4, flanqué par des régions 5' et 3' non traduites et une queue poly(A); contient de la N1-méthylpseudouridine au lieu de l'uridine (*tout-U>m¹Ψ*).

abdavomerán

RNA mensajero (mRNA), protegido en 5', que codifica para el dominio de unión al receptor (RBD) de la glicoproteína de la espícula (S) de SRAS-CoV-2 (coronavirus 2 del síndrome respiratorio agudo severo, GenBank: MN908947.3) con codones optimizados, expresado como una proteína de fusión en 5' con un péptido señal de glicoproteína S extendido y conectado en 3' mediante un péptido enlazador (linker) rico en glicina/serina (GS) con un dominio fibrina de trimerización de T4, flanqueado por regiones 5' y 3' no traducidas y una cola poly(A); contiene N1-metilpseudouridina en lugar de uridina (*all-U>m¹Ψ*).

alunacedasum alfa #

alunacedase alfa

human angiotensin-converting enzyme 2 extracellular soluble domain (18-740, 1-723 in the current sequence), dimer, glycosylated, produced in Chinese hamster ovary (CHO) cells; human angiotensin-converting enzyme 2 (ACE2, angiotensin-converting enzyme homolog, ACEH, ACE-related carboxypeptidase, metalloprotease MPROT15, EC:3.4.17.23), soluble extracellular (18-740)-domain of the proprotein, non-covalent dimer, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

alunacédase alfa domaine extracellulaire soluble de l'enzyme de conversion de l'angiotensine de type 2 humaine (18-740, 1-723 dans la séquence actuelle), dimère, glycosylé, produit dans des cellules ovariennes de hamster chinois (CHO);enzyme de conversion de l'angiotensine de type 2 humaine (ECA2, enzyme de conversion homologue de l'angiotensine, HECA, carboxypeptidase liée à l'ECA, métalloprotéase MPROT15, EC: 3.4.17.23), domaine protéique extracellulaire soluble (18-740), dimère non covalent, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

alunacedasa alfa enzima humana convertidora de la angiotensina 2 dominio extracelular soluble (18-740, 1-723 en la secuencia actual) dímero, glicosilado, producido en células ováricas de hámster Chino (CHO); enzima humana convertidora de la angiotensina 2 (ECA2, homólogo enzima covertidora de angiotensina, HECA, ECA-carboxipeptidasa relacionada, metaloproteasa MPROT15, EC:3.4.17.23), soluble extracelular (18-740)-dominio de la proproteína, dímero no covalente, producido en células ováricas de hámster Chino (CHO), glicofoma alfa

Sequence / Séquence / Secuencia
QSTIEEQART FLDKFNHEAE DLFYQSSLAS WNYNTNITEE NVQNMNNAGD 50
KWSAFLKEQS TLAQMYPLQE IQNLTVKLQL QALQQNGSSV LSEDKSKRLN 100
TILNTMSTIY STGKVCNPDN PQECLLLEPG LNEIMANSLD YNERLWAWES 150
WRSEVGKQLR FLYEYVVLK NEMARANHYE DYGDYWRGDY EVNGVDGYDY 200
SPGQLIEDVE HTFEEIKPLY EHLHAYVRK LMNAYFSYIS FIGCLPAHL 250
GDMWGRFWIN LYSLTVPFGQ KPNIDVTDAM VDQAWDAQRI FKEAEKFEVS 300
VGLPNMTQGF WENSMLTDFG NVQKAVCHPT AWDLGKGDFR ILMCTKVIMD 350
DFLTAHHEMG HIQYDMAYAA QPFLLRNGAN EGFHEAVGEI MSLSAATEKH 400
LKSIGLLSPD EQEDNETEIN FLLKQALTIV GTLPFTYMLE KWRWMVFKGE 450
IPKQWMMKW WEMKREIVGV VEPVEHDETY CDPASLFHVS NDYSFIRYIT 500
RTLYQFQFQE ALQCAAKHEG FLHKCDISNS TEAGQKLFNM LRLGKSEWPT 550
LALENVVGAK NMNVRPLNY FEPLFTWLDK QNKNSFVGWS TDWSPYADQS 600
IKVRISLKSA LGDKAYEWNH NEMYLFRSSV AYAMRQYFLK VKNQMLFGE 650
EDVRVANLKP RISFNFFVTA EKNVSDIIPR TEVEKAIRMS RSPINDAERL 700
NDNSLEFLGI QPTLGPFNQF FVS 723

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
116-124, 327-344, 513-525 116'-124', 327'-344', 513'-525'
Cys-SH: 244, 481, 244', 481'

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
N36, N73, N86, N305, N415, N529, N36', N73', N86', N305', N415', N529' (97-100% each)
N673, N673' (4% each)

O-glycosylation sites / Sites de O-glycosylation / Posiciones de O-glicosilación
S704, S704' or T713, T713' or S723, S723'

Other modifications: Q1, Q1'>Glp (pyroGlu);
S723, S723'; partially clipped (~4%)

atibuclimabum #
atibuclimab immunoglobulin G4-kappa, anti-[*Homo sapiens* CD14, membrane (mCD14) and soluble (sCD14)], chimeric monoclonal antibody; gamma4 heavy chain chimeric (1-442) [VH *Mus musculus* (Musmus IGHV3-2*02 (95.9%) -(IGHD) -Musmus IGHJ3*01 (92.9%) A128>S (115)) CDR-IMGT [9.7.8] (26-34.52-58.97-104) (1-115) -*Homo sapiens* IGHG4*01 (100%), CH1 (116-213),hinge 1-12 (214-225), CH2 (226-335), CH3 (336-440), CHS (441-442)] (116-442)], (129-218')-disulfide with kappa light chain chimeric (1'-218') [V-KAPPA *Mus musculus* (Musmus IGKV3-5*01 (97.0%) -IGKJ2*01 (100%)) CDR-IMGT [10.3.9] (27-36.54-56.93-101) (1'-111') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (157), V101 (195) (112'-218')]; dimer (221-221":224-224")-bisdisulfide, produced in a Chinese hamster ovary (CHO)-DG44 cell line, glycoform alfa

atibuclimab immunoglobuline G4-kappa anti-[*Homo sapiens* CD14, membrane (mCD14) et soluble (sCD14)], anticorps monoclonal chimérique; chaîne lourde gamma4 chimérique (1-442) [VH *Mus musculus* (Musmus IGHV3-2*02 (95.9%) -(IGHD) -Musmus IGHJ3*01 (92.9%) A128>S (115)) CDR-IMGT [9.7.8] (26-34.52-58.97-104) (1-115) - *Homo sapiens* IGHG4*01 (100%), CH1 (116-213), charnière 1-12 (214-225), CH2 (226-335), CH3 (336-440), CHS (441-442)) (116-442)], (129-218')-disulfure avec la chaîne légère kappa chimérique (1'-218') [V-KAPPA *Mus musculus* (Musmus IGKV3-5*01 (97.0%) - IGKJ2*01 (100%)) CDR-IMGT [10.3.9] (27-36.54-56.93-101) (1'-111') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (157), V101 (195) (112'-218'')]; dimère (221-221":224-224")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-DG44, glycoforme alfa

atibuclimab inmunoglobulina G4-kappa anti-[*Homo sapiens* CD14, membrana (mCD14) y soluble (sCD14)], anticuerpo monoclonal quimérico; cadena pesada gamma4 quimérico (1-442) [VH *Mus musculus* (Musmus IGHV3-2*02 (95.9%) -(IGHD) -Musmus IGHJ3*01 (92.9%) A128>S (115)) CDR-IMGT [9.7.8] (26-34.52-58.97-104) (1-115) - *Homo sapiens* IGHG4*01 (100%), CH1 (116-213), bisagra 1-12 (214-225), CH2 (226-335), CH3 (336-440), CHS (441-442)) (116-442)], (129-218')-disulfuro con la cadena ligera kappa quimérica (1'-218') [V-KAPPA *Mus musculus* (Musmus IGKV3-5*01 (97.0%) - IGKJ2*01 (100%)) CDR-IMGT [10.3.9] (27-36.54-56.93-101) (1'-111') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (157), V101 (195) (112'-218'')]; dímero (221-221":224-224")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-DG44, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada			
DVLQQLSGPG	LVKPSQSLSL	TCTVTGYSIT	SDSAWNWIRQ
YISYSGSTSY	NPSLKSIRSI	TRDTSKNQFF	LQLNSVTTED
RFAYWGQGT	VTVSGASTKG	PSVFPLAPCS	RSTSESTAAL
PVTVSNWSGA	LTSGVHTFPA	VLQSSGLYSL	SSVVTVPSSS
DHKPSNTKVD	KRVESKYGPP	CPSCPAPEFL	GGPSVFLFPP
TPEVTQVVVD	VSQEDPEVQF	NWYVDGVEVH	NAKTKPREEQ
LTVLHQDMLN	GKEYKCKVSN	KGLPSSIEKT	ISKAKQPRE
EEMTKNQVSL	TCLVKGFPYS	DIAEWESNG	QPENNYKTP
LYSLRTVDKS	RWQEGNVFSC	SVMHEALHNN	YTKSLSLSL
		GK	442
Light chain / Chaîne légère / Cadena ligera			
DIVLTQSPAS	LAVSLGQRAT	ISCRASESVD	SYVNSFLHWY
LYRASNLQS	GIPARFSGSG	SRTDFTLTIN	PVEADDVATY
TFGGGKLEI	KRTVAAPSVF	IFPPSDEQLK	SGTASVVCLL
QWKVDNALQS	GNSQESVTEQ	DSKDSTYSLS	STLTLSKADY
THQGLSSPVT	KSFNRGEC		218
Post-translational modifications			
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro			
Intra-H (C23-C104)	22-96	142-198	256-316
	22"-96"	142"-198"	256"-316"
Intra-L (C23-C104)	23"-92"	138"-198"	
	23"-92"	138"-198"	
Inter-H-L (CH1 10-CL 126)	129-218"	129"-218"	
Inter-H-H (h 8, h 11)	221-221"	224-224"	
N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación			
H CH2 N84.4:			
292, 292"			
Fucosylated complex bi-antennary CHO-type glycans (G0F predominant) / glycanes de type CHO bi-antennaires complexes fucosylés (G0F prédominant) / glicanos de tipo CHO biantenarios complejos fucosilados (G0F predominante).			
C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal			
H CHS K2:			
442, 442"			

bamlanivimabum #

bamlanivimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) spike (S) protein], *Homo sapiens* monoclonal antibody; gamma1 heavy chain *Homo sapiens* (1-455) [VH (*Homo sapiens* IGHV1-69*04 (99.0%) -(IGHD) -IGHJ6*01 (90.9%) T123>A (120)) CDR-IMGT [8.8.18] (26-33.51-58.97-114) (1-125) -*Homo sapiens* IGHG1*03 (100%) G1m3, nG1m1 (CH1 R120 (222) (126-223), hinge 1-15 (224-238), CH2 (239-348), CH3 E12 (364), M14 (366) (349-453), CHS (454-455)) (126-454)], (228-214')-disulfide with kappa light chain *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-39*01 (97.9%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (234-234'':237-237'')-bisdisulfide, produced in a Chinese hamster ovary (CHO) cell line derived from CHO-K1SV, lacking the glutamine synthetase (GS) gene, glycoform alfa

bamlanivimab

immunoglobuline G1-kappa anti-[*Homo sapiens* protéine spike (S) du coronavirus 2 du syndrome respiratoire aigu sévère (SARS-CoV-2)], anticorps monoclonal *Homo sapiens*; chaîne lourde gamma1 *Homo sapiens* (1-455) [VH (*Homo sapiens* IGHV1-69*04 (99%) -(IGHD) -IGHJ6*01 (90.9%) T123>A (120)) CDR-IMGT [8.8.18] (26-33.51-58.97-114) (1-125) -*Homo sapiens* IGHG1*03 (100%) G1m3, nG1m1 (CH1 R120 (222) (126-223), charnière 1-15 (224-238), CH2 (239-348), CH3 E12 (364), M14 (366) (349-453), CHS (454-455)) (126-454)], (228-214')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-39*01 (97.9%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (234-234'':237-237'')-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) dérivant de la lignée cellulaire CHO-K1SV, ne présentant pas le gène de la glutamine synthétase, glycoforme alfa

bamlanivimab

inmunoglobulina G1-kappa anti-[*Homo sapiens* proteína espícula (S) del coronavirus 2 del síndrome respiratorio agudo severo (SRAS-CoV-2)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma1 *Homo sapiens* (1-455) [VH (*Homo sapiens* IGHV1-69*04 (99%) -(IGHD) -IGHJ6*01 (90.9%) T123>A (120)) CDR-IMGT [8.8.18] (26-33.51-58.97-114) (1-125) -*Homo sapiens* IGHG1*03 (100%) G1m3, nG1m1 (CH1 R120 (222) (126-223), bisagra 1-15 (224-238), CH2 (239-348), CH3 E12 (364), M14 (366) (349-453), CHS (454-455)) (126-454)], (228-214')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-39*01 (97.9%) -IGKJ1*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (234-234'':237-237'')-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular derivada de CHO-K1SV, en ausencia del gen glutamina sintetasa (GS), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada	
QVQLVQSGAE VKKPGSSVKY SCKASGGTFS NYAISWVRQA PGQGLEWMGR	50
IIPILGIANY AQKFGQGRVTI TADKSTSTAY MELSSLRSED TAVYYCARGY	100
YEARHYYYYY AMDVWGQCTA VTWSASTKG FSVFPLAPSS KSTSGGTAAAL	150
GCLVKDYEPF PVTVSWNSGA LTSGVHTFFA VLQSSGLYSL SSVTVVPSSS	200
LGTOTYICNV NHKPSNTKVD KRVEPKSCDK THTCPFCPAF ELLGGPSVFL	250
FFPKPDTLM ISRTPEVTCV VVDVSHEDPE VKENWYVDGV EVHNAKTKER	300
EEQYNSTYRV VSVLTIVLHQD WLNGKEYCKK VSNKALPAPI ERTISKAKGQ	350
PREPOVYTLF PSREEMTKNQ VSLTCLVKG F YPSDIAVENE SNGQFENNYK	400
TTPFVLDSGG SFFLYSKLTV DKSRWQQGNV FSCSVMHREAL HNHVTQKSL	450
LSFGK	455
Light chain / Chaîne légère / Cadena ligera	
DIQMTQSFSS LSASVGDRTV ITCRASQGIS SYLSWYQKP GKAPKLLIYA	50
ASSLQSGVPS RFSGSGSGTD FTLTITSLQP EDFATYYCQQ SYSTPRFTFGQ	100
GTRVEIKRTV AAPSVFIFFP SDEQLKSGTA SVVCLLNIFY PREAKVQWKV	150
DNALQSGNSQ ESVTEQDSKD STYLSLSTLT LSKADYEKHK VYACEVTHQG	200
LSSPVTKSFN RGEK	214
Post-translational modifications	
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro	
Intra-H (C23-C104)	22°-96° 152°-208° 269°-329° 375°-433°
	22°-96° 152°-208° 269°-329° 375°-433°
Intra-L (C23-C104)	23°-88° 134°-194°
	23°-88° 134°-194°
Inter-H-L (h 5-CL 126)	228°-214° 228°-214°
Inter-H-H (h 11, h 14)	234°-234° 237°-237°
N-terminal glutamine cyclization to pyroglutamyI (pE, S-oxopropyl)	
H V H Q I:	
I, I"	
N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación	
H CH2 N84.4:	
305, 305"	
Fucosylated complex bi-antennary CHO-type glycans (G0F predominant) / glycanes de type CHO bi-antennaires complexes fucosylés (G0F predominant) / glicanos de tipo CHO biantenarios complejos fucosilados (G0F predominante).	
C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal	
H CHS K2:	
455, 455"	

casirivimabum #
casirivimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) spike (S) protein, receptor binding domain (RBD)], *Homo sapiens* monoclonal antibody; gamma1 heavy chain *Homo sapiens* (1-450) [VH (*Homo sapiens* IGHV3-11*01 (96.9%) -(IGHD) - IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), hinge 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-214')-disulfide with kappa light chain *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-33*01 (96.8%) - IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107") -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214''); dimer (229-229":232-232")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

casirivimab

immunoglobuline G1-kappa anti-[*Homo sapiens* protéine spike (S) du coronavirus 2 du syndrome respiratoire aigu sévère (SARS-CoV-2), domaine de liaison au récepteur (RBD)], anticorps monoclonal *Homo sapiens*;

casirivimab

chaîne lourde gamma1 *Homo sapiens* (1-450) [VH (*Homo sapiens* IGHV3-11*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), charnière 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-214')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-33*01 (96.8%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (229-229":232-232")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

inmunoglobulina G1-kappa anti-[*Homo sapiens* proteína espícula (S) del coronavirus 2 del síndrome respiratorio agudo severo (SRAS-CoV-2), dominio de unión al receptor (RBD)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma1 *Homo sapiens* (1-450) [VH (*Homo sapiens* IGHV3-11*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), bisagra 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-214')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-33*01 (96.8%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (229-229":232-232")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada

QVQLVESGGG LVKPGGSLRL SCAASGFTFS DYYMSWIRQA PGKGLEWVS 50
ITYSGSTIYY ADSVKGRFTI SRDNAKSLY LQMSLRAED TAVYYCARD 100
GTTMVPFDYW GQGTLVTVSS ASTKGPSVFP LAPSSKSTSG GTAALGLCVK 150
DYFPEPVTVS WNSGALTSGV HTFPAVLQSS GLYSLSVVV VPSSSLGTQT 200
YICNVNHKPS NTKVDKKEP KSCDKTHTCP PCPAPELLGG PSVFLFPPKP 250
KDTLMISRTP EVTCVVDVVS HEDPEVKFNW YVDGVEVHNA KTKPREEQYN 300
STYRVVSVLT VLHQDWLNGK EYCKYKSNKA LPAPTEKITS KAKGQPREPQ 350
VYTLPPSRDE LTKNQVSLTC LVKGFYPSDI AVEVESNGQP ENNYKTTTPV 400
LDSGGSFFLY SKLTVDKSRW QQGNVFSVSV MHEALHNHYT QKSLSLSPGK 450

Light chain / Chaîne légère / Cadena ligera

DIQMTQSPSS LSASVGDRTV ITCQASQDIT NYLWVYQQK GKAPKLLIYA 50
ASNLETGVPS RFGSGSGSDT FTFTISGLQP EDIATYYCQ YDNLPLTFGG 100
GTRVEIKRTV AAPSVEIFPP SDEQLKSGTA SVVCLLNIFY PREAKVQWVK 150
DNALQSGNSQ ESVTEQDSKD STYLSSTLT LSKADYEKHK VYACEVTHQG 200
LSSPVTKSFN RGEK 214

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 147-203 264-324 370-428
22"-96" 147"-203" 264"-324" 370"-428"

Intra-L (C23-C104) 23"-88" 134"-194"
23"-88" 134"-194"

Inter-H-L (h 5-CL 126) 223-214' 223"-214"

Inter-H-H (h 11, h 14) 229-229" 232-232"

N-terminal glutamine cyclization to pyroglutamyl (pE, 5-oxoprolyl)

H VH Q1:

I, I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

300, 300"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantennarios complejos fucosilados.

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

450, 450"

cilgavimabum

cilgavimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) spike (S) protein, receptor binding domain (RBD)], *Homo sapiens* monoclonal antibody; gamma1 heavy chain *Homo sapiens* (1-461) [VH (*Homo sapiens*IGHV3-15*01 (96.0%) - (IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.10.22] (26-33.51-60.99-120) (1-131) -*Homo sapiens*IGHG1*03 G1m3, nG1m1, G1v21 CH2 Y15.1, T16, E18, G1v39 CH2 F1.3, E1.2, S116 (CH1 R120 (228) (132-229), hinge 1-15 (230-244), CH2 L1.3>F (248), L1.2>E (249), M15.1>Y (266), S16>T (268), T18>E (270), P116>S (345) (245-354), CH3 E12 (370), M14 (372) (355-459), CHS (460-461)) (132-461)], (234-219')-disulfide with kappa light chain *Homo sapiens* (1'-219') [V-KAPPA (*Homo sapiens*IGKV4-1*01 (96.0%) -IGKJ4*01 (100%)) CDR-IMGT [12.3.8] (27-38.56-58.95-102) (1'-112') -*Homo sapiens*IGKC*01 (100%) Km3 A45.1 (158), V101 (196) (113'-219')]; dimer (240-240":243-243")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

cilgavimab

immunoglobuline G1-kappa anti-[*Homo sapiens* protéine spike (S) du coronavirus 2 du syndrome respiratoire aigu sévère (SARS-CoV-2), domaine de liaison au récepteur (RBD)], anticorps monoclonal *Homo sapiens*; chaîne lourde gamma1 *Homo sapiens* (1-461) [VH (*Homo sapiens*IGHV3-15*01 (96.0%) - (IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.10.22] (26-33.51-60.99-120) (1-131) -*Homo sapiens*IGHG1*03 G1m3, nG1m1, G1v21 CH2 Y15.1, T16, E18, G1v39 CH2 F1.3, E1.2, S116 (CH1 R120 (228) (132-229), charnière 1-15 (230-244), CH2 L1.3>F (248), L1.2>E (249), M15.1>Y (266), S16>T (268), T18>E (270), P116>S (345) (245-354), CH3 E12 (370), M14 (372) (355-459), CHS (460-461)) (132-461)], (234-219')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-219') [V-KAPPA (*Homo sapiens*IGKV4-1*01 (96.0%) -IGKJ4*01 (100%)) CDR-IMGT [12.3.8] (27-38.56-58.95-102) (1'-112') -*Homo sapiens*IGKC*01 (100%) Km3 A45.1 (158), V101 (196) (113'-219')]; dimère (240-240":243-243")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

cilgavimab

inmunoglobulina G1-kappa anti-[*Homo sapiens* proteína espícula (S) del coronavirus 2 del síndrome respiratorio agudo severo (SRAS-CoV-2), dominio de unión al receptor (RBD)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma1 *Homo sapiens* (1-461) [VH (*Homo sapiens*IGHV3-15*01 (96.0%) - (IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.10.22] (26-33.51-60.99-120) (1-131) -*Homo sapiens*IGHG1*03 G1m3, nG1m1, G1v21 CH2 Y15.1, T16, E18, G1v39 CH2 F1.3, E1.2, S116 (CH1 R120 (228) (132-229), bisagra 1-15 (230-244), CH2 L1.3>F (248), L1.2>E (249), M15.1>Y (266), S16>T (268), T18>E (270), P116>S (345) (245-354), CH3 E12 (370), M14 (372) (355-459), CHS (460-461)) (132-461)], (234-219')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-219') [V-KAPPA (*Homo sapiens*IGKV4-1*01 (96.0%) -IGKJ4*01 (100%)) CDR-IMGT [12.3.8] (27-38.56-58.95-102) (1'-112') -*Homo sapiens*IGKC*01 (100%) Km3 A45.1 (158), V101 (196) (113'-219')]; dímero (240-240":243-243")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
EVQLVESGGG LVKPGGSLRL SCAASGFTFR DVWMSWVRQA PGKGLEWVGR 50
IKSKIDGGTT DYAAPVKGRF TISRDDSKNT LYLQMNSLKT EDTAVYYCTT 100
AGSYYYDTVG PGLPEGKFDY WGQGLTLVTVS SASTKGPSVF PLAPSSKSTS 150
GGTAALGCLV KDYFPEPVTV SWNSGALTSG VHTFPAVLQS SGLYSLSSVV 200
TVPSSSLGTQ TYICNVNHKP SNTKVDKRVE PKSCDKTHTC PPCPAPEFEG 250
GPSVFLFPPK PKDTLYITRE PEVTCVVVDV SHEDPEVKFN WYVDGVEVHN 300
AKTKPREEQY NSTYRVVSVL TVLHQDWLNG KEYCKVSNK ALPASIEKTI 350
SKAKGQPREP QVYTLPPSRE EMTKNQVSLT CLVKGFYPSD IAVEWESNGQ 400
PENNYKTPPP VLDSGGSFFL YSKLTVDKSR WQQGNVFSKS VMHEALHNNY 450
TQKSLSLSPG K 461

Light chain / Chaîne légère / Cadena ligera
DIVMTQSPDS LAVSLGERAT INCKSSQSVL YSSNNKNYLA WYQQKPGQPP 50
KLLMYWASTR ESGVPDRFSG SGSGAEFTLT ISSLQAEDVA IYYCQYYYST 100
LTFGGGTKEV IKRTVAAPSV FIFPPSDEQL KSGTASVCL LNNFYPREAK 150
VQWKVDNALQ SGNSQESVTE QDSKDSYSL SSSLTLISKAD YEKHKVYACE 200
VTHQGLSSPV TKSFNRGEC 219

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-98 158-214 275-335 381-439
22"-98" 158"-214" 275"-335" 381"-439"
Intra-L (C23-C104) 23'-94' 139'-199'
23"-94" 139"-199"
Inter-H-L (h 5-CL 126) 234-219' 234"-219"
Inter-H-H (h 11, h 14) 240-240" 243-243"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
311, 311*
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires
complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.
C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal
H CHS K2:
461, 461*

dazcapistatum
dazcapistat

rac-N-[(2R)-4-amino-3,4-dioxo-1-phenylbutan-2-yl]-4-(2-fluorophenyl)-2-methyl-1,3-oxazole-5-carboxamide

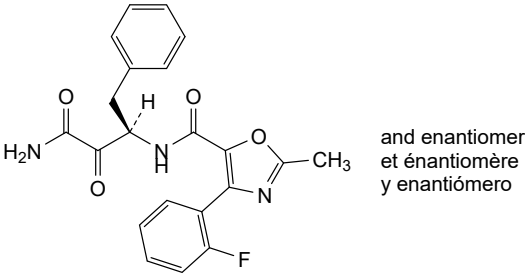
dazcapistat

rac-N-[(2R)-4-amino-3,4-dioxo-1-phénylbutan-2-yl]-4-(2-fluorophényl)-2-méthyl-1,3-oxazole-5-carboxamide

dazcapistat

rac-N-[(2R)-4-amino-3,4-dioxo-1-fenilbutan-2-il]-4-(2-fluorofenil)-2-metil-1,3-oxazol-5-carboxamida

C₂₁H₁₈FN₃O₄



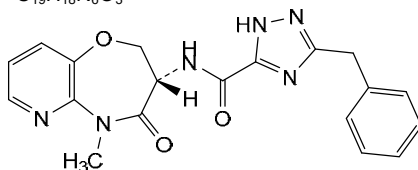
eclitasertibum
eclitasertib

5-benzyl-N-[(3S)-5-methyl-4-oxo-2,3,4,5-tetrahydropyrido[3,2-b][1,4]oxazepin-3-yl]-1H-1,2,4-triazole-3-carboxamide

éclitasertib 5-benzyl-*N*-[(3*S*)-5-méthyl-4-oxo-2,3,4,5-tétrahydropyrido[3,2-*b*][1,4]oxazépin-3-yl]-1*H*-1,2,4-triazole-3-carboxamide

eclitasertib 5-bencil-*N*-[(3*S*)-5-metil-4-oxo-2,3,4,5-tetrahidropirido[3,2-*b*][1,4]oxazepin-3-il]-1*H*-1,2,4-triazol-3-carboxamida

C₁₉H₁₈N₆O₃



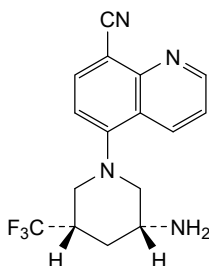
enpatoranum

enpatoran 5-[(3*R*,5*S*)-3-amino-5-(trifluorométhyl)piperidin-1-yl]quinoline-8-carbonitrile

enpatoran 5-[(3*R*,5*S*)-3-amino-5-(trifluorométhyl)pipéridin-1-yl]quinoléine-8-carbonitrile

enpatorán 5-[(3*R*,5*S*)-3-amino-5-(trifluorometil)piperidin-1-il]quinoleína-8-carbonitrilo

C₁₆H₁₅F₃N₄



etesevimabum

etesevimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) spike (S) protein, receptor binding domain (RBD)], *Homo sapiens* monoclonal antibody; gamma1 heavy chain *Homo sapiens* (1-449) [VH (*Homo sapiens* IGHV3-66*01 (96.9%) - (IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.7.13] (26-33.51-57.96-108) (1-119) -*Homo sapiens* IGHG1*03 G1m3, nG1m1, G1v14 A1.3, A1.2 (CH1 R120 (216) (120-217), hinge 1-15 (218-232), CH2 L1.3>A (236), L1.2>A (237), (233-342), CH3 E12 (358), M14 (360) (343-447), CHS (448-449)) (120-449)], (222-216')-disulfide with kappa light chain *Homo sapiens* (1'-216') [V-KAPPA (*Homo sapiens* IGKV1-39*01 (97.9%) -IGKJ2*01 (100%)) CDR-IMGT [6.3.11] (27-32.50-52.89-99) (1'-109') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (155), V101 (193) (110'-216')]; dimer (228-228'':231-231'')-bisdisulfide, produced in a Chinese hamster ovary (CHO) cell line derived from CHO-K1SV, lacking the glutamine synthetase (GS) gene, glycoform alfa

étésévimab

immunoglobuline G1-kappa anti-[*Homo sapiens* protéine spike (S) du coronavirus 2 du syndrome respiratoire aigu sévère (SARS-CoV-2), domaine de liaison au récepteur (RBD)], anticorps monoclonal *Homo sapiens*;
chaîne lourde gamma1 *Homo sapiens* (1-449) [VH (*Homo sapiens*IGHV3-66*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.7.13] (26-33.51-57.96-108) (1-119) -*Homo sapiens*IGHG1*03 G1m3, nG1m1, G1v14 A1.3, A1.2 (CH1 R120 (216) (120-217), charnière 1-15 (218-232), CH2 L1.3>A (236), L1.2>A (237), (233-342), CH3 E12 (358), M14 (360) (343-447), CHS (448-449)) (120-449)], (222-216')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-216') [V-KAPPA (*Homo sapiens*IGKV1-39*01 (97.9%) -IGKJ2*01 (100%)) CDR-IMGT [6.3.11] (27-32.50-52.89-99) (1'-109') -*Homo sapiens*IGKC*01 (100%) Km3 A45.1 (155), V101 (193) (110'-216')]; dimère (228-228":231-231")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) dérivant de la lignée cellulaire CHO-K1SV, ne présentant pas le gène de la glutamine synthétase, glycoforme alfa

etesevimab

immunoglobulina G1-kappa anti-[*Homo sapiens* proteína espícula (S) del coronavirus 2 del síndrome respiratorio agudo severo (SRAS-CoV-2), dominio de unión al receptor (RBD)], anticuerpo monoclonal *Homo sapiens*;
cadena pesada gamma1 *Homo sapiens* (1-449) [VH (*Homo sapiens*IGHV3-66*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.7.13] (26-33.51-57.96-108) (1-119) -*Homo sapiens*IGHG1*03 G1m3, nG1m1, G1v14 A1.3, A1.2 (CH1 R120 (216) (120-217), bisagra 1-15 (218-232), CH2 L1.3>A (236), L1.2>A (237), (233-342), CH3 E12 (358), M14 (360) (343-447), CHS (448-449)) (120-449)], (222-216')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-216') [V-KAPPA (*Homo sapiens*IGKV1-39*01 (97.9%) -IGKJ2*01 (100%)) CDR-IMGT [6.3.11] (27-32.50-52.89-99) (1'-109') -*Homo sapiens*IGKC*01 (100%) Km3 A45.1 (155), V101 (193) (110'-216')]; dímero (228-228":231-231")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular derivada de CHO-K1SV, en ausencia del gen Glutamina Sintetasa (GS), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
EVQLVESGGG LVQPGGSLRL SCAASGFTVS SNYMSWVRQA PGKGLEWWSV 50
IYSGGSTFYA DSVKGRFTIS RDNSMNTLFL QMNSLRAEDT AVYYCARVLP 100
MYGDLIDYWG QGTLVTYSSA STKGPSVFPPL APSSKSTSGG TAALGCLVLD 150
YFPEPTVTSW NSGALTSQGVH TFAVLQSSG LYSLSVSVTV PSSSLGTQTY 200
ICNVNHPKPSN TKVDKRVPEK SCDKTHTCPP CPAPEAAGGP SVFLFPKPK 250
DTLMISRTPE VTCVVDVDSH EDPEVKFNWY VDGVEVHNAK TKPREQVNS 300
TYRVVSVLTV LQDMLNGKE YKCKVSNKAL PAPIEKTISK AKGQPREPQV 350
YTLPPSREEM TKNQVSLTCL VKGFYPSDIA VEWESNGQPE NNYKTTTPVL 400
DSGGSFFLYS KLTVDKSRWQ QGNVFSCSVM HEALHNHYTQ KSLSLSPGK 449

Light chain / Chaîne légère / Cadena ligera
DIVMTQSPSS LSASVGDRTV ITCRASQGIS RYLNWYQQKPK GKAPKLLIYA 50
ASSLQSGVPS RFGSGSGSDT FTLTISSLQP EDFATYYCQQ SYSTPPEYTF 100
GQGTKLEIKR TVAAPSFIIF PPSDEQLKSG TASVCLLNN FYPREKVVQW 150
KQDNALQSGN SQESVTEQDS KDSTYSLSST LTLTKADYEK HKVYACEVTH 200
QGLSSPVTKS FNRGEC 216

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-95 146-202 263-323 369-427
22"-95" 146"-202" 263"-323" 369"-427"

Intra-L (C23-C104) 23'-88' 136'-196'
23'''-88''' 136'''-196'''

Inter-H-L (h 5-CL 126) 222-216' 222"-216"

Inter-H-H (h 11, h 14) 228-228" 231-231"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

299, 299"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantennarios complejos fucosilados.

C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

449, 449"

ganulamieranum #

ganulamieran

messenger RNA (mRNA), 5'-capped, encoding the codon-optimised receptor binding domain (RBD) of the SARS-CoV-2 (severe acute respiratory syndrome coronavirus 2, GenBank: MN908947.3) spike (S) glycoprotein expressed as a fusion protein, 5' with an extended S glycoprotein signal peptide, and 3' with a T4 fibrin trimerization domain and the S glycoprotein transmembrane domain (TM) connected by two glycine/serine-rich (GS) linker peptides, flanked by 5' and 3' untranslated regions and a 3' poly(A) tail; contains N1-methylpseudouridine instead of uridine (*all-U>m¹Ψ*).

ganulamieran

ARN messenger (ARNm), protégé en 5', codant pour le domaine de liaison au récepteur (RBD) de la glycoprotéine spike (S) du SARS-CoV-2 (coronavirus 2 du syndrome respiratoire aigu sévère, GenBank: MN908947.3) avec des codons optimisés, exprimés sous forme de protéine de fusion, 5' avec un peptide signal de la glycoprotéine S allongé et 3' avec un domaine de trimérisation de la fibrine de T4 et le domaine transmembranaire (TM) de la glycoprotéine S, liés au moyen de deux peptides de liaison (linker) riches en glycine / sérine (GS), flanqués de régions 5' et 3' non traduites et d'une queue poly(A); contient de la N1-méthylpseudouridine au lieu de l'uridine (*tout-U>m¹Ψ*).

ganulamieran

RNA mensajero (mRNA), protegido en 5', que codifica para el dominio de unión al receptor (RBD) de la glicoproteína de la espícula (S) de SRAS-CoV-2 (coronavirus 2 del síndrome respiratorio agudo severo, GenBank: MN908947.3) con codones optimizados, expresado como una proteína de fusión, en 5' con un péptido señal de glicoproteína S extendido y en 3' con un dominio fibrina de trimerización de T4 y el dominio transmembrana (TM) de la glicoproteína S, conectado mediante dos péptidos enlazadores (linker) ricos en glicina/sérina (GS), flanqueado por regiones 5' y 3' no traducidas y una cola poly(A); contiene N1-metilpseudouridina en lugar de uridina (*all-U>m¹Ψ*).

goflikiceptum #

goflikicept

human interleukin-1 receptor accessory protein fragment (21-358, 1-338 in the current sequence) fused via peptidyl linker ³³⁹GSGGG³⁴³ to a human immunoglobulin G1 Fc fragment (344-570, C-terminal Lys clipped) variant (S>C⁴⁷⁷, T>W⁴⁸⁹, K>A⁵³²), covalent dimer with human interleukin-1 receptor type 1 fragment (21-333, 1-313 in the current sequence) fused via peptidyl linker ³¹⁴GSGG³¹⁸ to a human immunoglobulin G1 Fc fragment (319-545, C-terminal Lys clipped) variant (Y>C⁴⁴⁷, T>S⁴⁶⁴, L>A⁴⁶⁶, F>K⁵⁰³, Y>V⁵⁰⁵), glycosylated, produced in Chinese hamster ovary (CHO) cells; fusion protein comprising the (1-338)-fragment of the human interleukin 1 receptor accessory protein (IL1RAP, IL-1RAcP, interleukin-1 receptor 3, IL1R3), a GSGGG linker (339-343), and the (S>C⁴⁷⁷, T>W⁴⁸⁹, K>A⁵³²) variant of the C-terminal 227-peptide (Fc fragment) of human immunoglobulin G1 (344-570), (349-324':352-327':477-447')-trisdissulfide with a fusion protein comprising the (21-333)-fragment of the precursor of human interleukin 1 receptor type 1 (IL1R1, IL1Rα, IL1R type I, p80, CD121a) (1-313), a GSGGG linker (314-318), and the (Y>C⁴⁴⁷, T>S⁴⁶⁴, L>A⁴⁶⁶, F>K⁵⁰³, Y>V⁵⁰⁵) variant of the C-terminal 227-peptide (Fc fragment) of human immunoglobulin G1 (319-545), produced in Chinese hamster ovary (CHO) cells, glycoform alfa

goflikicept

fragment de la protéine accessoire du récepteur de l'interleukine 1 humaine (21-358, 1-338 dans la séquence actuelle) fusionné via un peptide liant ³³⁹GSGGG³⁴³ au fragment Fc de l'immunoglobuline humaine G1 (344-570, C-terminal attaché Lys) variant (S>C⁴⁷⁷, T>W⁴⁸⁹, K>A⁵³²), dimère covalent avec fragment de récepteur de l'interleukine-1 humaine de type 1 (21-333, 1-313 dans la séquence actuelle) fusionné via un peptide liant ³¹⁴GSGG³¹⁸ à un fragment Fc d'immunoglobuline humaine G1 (319-545, attaché au variant C-terminal Lys) (Y>C⁴⁴⁷, T>S⁴⁶⁴, L>A⁴⁶⁶, F>K⁵⁰³, Y>V⁵⁰⁵), glycosylé, produit dans des cellules ovariennes de hamster chinois (CHO); une protéine de fusion comprenant le fragment (1-338) de la protéine accessoire du récepteur de l'interleukine 1 humaine (IL1RAP, IL-1RAcP, récepteur 3 de l'interleukine-1, IL1R3), un linker GSGGG (339-343), et le variant (S>C⁴⁷⁷, T>W⁴⁸⁹, K>A⁵³²) du peptide C-terminal-227 (fragment Fc) de l'immunoglobuline humaine G1 (344-570), (349-324': 352-327': 477-447')-trisdissulfure avec une protéine de fusion comprenant le fragment- (21-333) du précurseur du récepteur de l'interleukine humaine de type 1 (IL1R1, IL1Rα, IL1R type I, p80, CD121a) (1-313), une liaison GSGGG (314-318) et le variant (Y>C⁴⁴⁷, T>S⁴⁶⁴, L>A⁴⁶⁶, F>K⁵⁰³, Y>V⁵⁰⁵) du peptide C-terminal-227 (fragment Fc) de l'immunoglobuline humaine G1 (319-545), produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

goflikicept

receptor de la interleukina-1 humana fragmento de la proteína complemento (21-358, 1-338 en la secuencia actual) fusionado a través de un enlace peptídico ³³⁹GSGGG³⁴³ a la inmunoglobulina humana G1 Fc fragmento (344-570, fijado al C-terminal Lis) variante (S>C⁴⁷⁷, T>W⁴⁸⁹, K>A⁵³²), dímero covalente con receptor de interleukina-1 humana fragmento tipo 1 (21-333, 1-313 en la secuencia actual) fusionado a través del enlace peptídico ³¹⁴GSGG³¹⁸ a una inmunoglobulina humana G1 Fc fragmento (319-545, fijado al C-terminal Lis) variante (Y>C⁴⁴⁷, T>S⁴⁶⁴, L>A⁴⁶⁶, F>K⁵⁰³, Y>V⁵⁰⁵), glicosilado, producido en células ováricas de hámster Chino (CHO). una proteína de fusión que comprende el fragmento- (1-338) del receptor humano de la interleukina 1 proteína accesorio (IL1RAP, IL-1RAcP, interleukina-1 receptor 3, IL1R3), un enlace GSGGG (339-343), y la variante (S>C⁴⁷⁷, T>W⁴⁸⁹, K>A⁵³²) del terminal-C péptido-227 (Fc fragmento) de la inmunoglobulina humana G1 (344-570), (349-324':352-327':477-447')-trisdissulfuro con una proteína de fusión que comprende el fragmento-(21-333) del precursor del receptor tipo 1 de la interleukina humana (IL1R1, IL1Rα, IL1R tipo I, p80, CD121a) (1-313), un enlace GSGGG (314-318), y la (Y>C⁴⁴⁷, T>S⁴⁶⁴, L>A⁴⁶⁶, F>K⁵⁰³, Y>V⁵⁰⁵) variante del terminal-C péptido-227 (Fc fragmento) de inmunoglobulina humana G1 (319-545), producido en células ováricas de hámster Chino (CHO), glicofoma alfa

Sequence / Séquence / Secuencia		
IL1RAP-GSG ₃ -Fc		
SERCDWGLD	TMRIQVFD	EPARIKCLF EHLKFNYS AHSAGTLIW 50
YWTRQDRLE	EPINFRLPEN	RISKEKDLV FRPTLLNDGT NYTCMLRNTT 100
YCSKVAFLPE	VVQKDSCFNS	PMKLPVHKLY IEYGIQRITC PNVDGYFPSS 150
VKPTITWYMG	CYKIQNFNNV	IPEGMNL SFL IALISNNGNY TCVVTYPENG 200
RTFHLTRTLT	VKVVGSPKNA	VPPVIHSPND HVVYEKEPGE ELLIPTVYF 250
SFLMDSRNEV	WMTIDGKKPD	DITIDVTINE SISHSRTEDE TRTQILSIKK 300
VTSEDLKRSY	VCHARSAKGE	VAKAAKVQK VPAPRYTVGS GGG DKHTHTCP 350
PCPAPELLGG	PSVFLFPPKP	KDTLMISRTP EVTCVVVDVS HEDPEVKFNM 400
YVDGVEVHNA	KTKPREEQYN	STYRVVSVLT VLHQDMLNGK EYKCKVSNKA 450
LPAPIEKTIS	KAKGQPREPQ	VYTLPPCRDE LTKNQVSLW LVKGFYPSDI 500
AVEWESNGQP	ENNYKTTTPV	LDSGGSFFLY S ALTVDKSRW QQGNVFCSSV 550
MHEALHNHYT	QKSLSLSPGK	
IL1R1-GSG ₃ -Fc		
DKCKEREKI	ILVSSANEID	VRPCPLNPNE HKGTITWYKD DSKTPVSTEQ 50
ASRIHQHKEK	LWFVPAKVED	SGHYCYVRN SSYCLRIKIS AKFVENEPNL 100
CYNAQAIFKQ	KLPVAGDGGL	VCPYMEFFKN ENNELPKLQW YKDCPKLLD 150
NIHFSGVKDR	LIVMNVAEKH	RGNYTCHASY TYLGKQYPIT RVIEFITLEE 200
NKPTRPVIIS	PANETMEVDL	GSQIQLICNV TGQLSDIAYW KWNGSVIDED 250
DPVLGEDYYS	VENPANKRRS	TLITVLNISE IESRFYKHPF TCFAKNTHGI 300
DAAYIQLIYP	VTN SGGG DK	THTCPPCPAP ELLGGSPVFL FPPKPKDTLM 350
ISRTPEVTCV	VVDVSHEDPE	VKFNWYVDGV EVHNAKTKPR EEQYNSTYRV 400
VSVLTVLHQD	WLNKGEYKCK	VSNKALPAPI EKTISKAKGQ PREPQVCTLP 450
PSRDELTKNQ	VSL SC AVKGF	YPSDIAVEWE SNGQPENNYK TTPPVLDSDG 500
SF K LVSKLTV	DKSRWQQGNV	FSCVMHEAL HNHYTQKSLS LSPGK 545
Mutations / Mutations / Mutaciones		
S ⁴⁷⁷ → C , T ⁴⁸⁹ → W , K ⁵³² → A Y ⁴⁴⁷ → C , T ⁴⁶⁴ → S , I ⁴⁶⁶ → A , E ⁵⁰³ → K , Y ⁵⁰⁵ → V		
Post-translational modifications		
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro		
Inter-chain: 349-324' 352-327' 477-447'		
Intra-chain IL1RAP-GSG3-Fc : 4-102 27-94 117-161 140-192 246-312 384-444 490-548		
Intra-chain IL1R1-GSG3-Fc : 3'-84' 24'-76' 101'-144' 122'-176' 228'-292' 359'-419' 465'-523'		
Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación		
N37, N87, N91, N98, N176, N189, N279, N420; N80', N173', N213', N229', N243', N277', N313', N395'		
Other modifications: K570, K545' clipped		

imdevimabum #
imdevimab

immunoglobulin G1-lambda, anti-[*Homo sapiens* severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) spike (S) protein, receptor binding domain (RBD)], *Homo sapiens* monoclonal antibody;
gamma1 heavy chain *Homo sapiens* (1-450) [VH (*Homo sapiens* IGHV3-30*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218),hinge 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-215')-disulfide with lambda light chain *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV2-14*01 (93.9%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.10] (26-34.52-54.91-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dimer (229-229":232-232")-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

imdévimab

immunoglobuline G1-lambda anti-[*Homo sapiens* protéine spike (S) du coronavirus 2 du syndrome respiratoire aigu sévère (SARS-CoV-2), domaine de liaison au récepteur (RBD)], anticorps monoclonal *Homo sapiens*;

chaîne lourde gamma1 *Homo sapiens* (1-450) [VH (*Homo sapiens* IGHV3-30*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), charnière 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-215')-disulfure avec la chaîne légère lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV2-14*01 (93.9%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.10] (26-34.52-54.91-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dimère (229-229":232-232")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

imdevimab

immunoglobulina G1-lambda anti-[*Homo sapiens* proteína espícula (S) del coronavirus 2 del síndrome respiratorio agudo severo (SRAS-CoV-2), dominio de unión al receptor (RBD)], anticuerpo monoclonal *Homo sapiens*;

cadena pesada gamma1 *Homo sapiens* (1-450) [VH (*Homo sapiens* IGHV3-30*01 (96.9%) -(IGHD) -IGHJ4*01 (100%)) CDR-IMGT [8.8.13] (26-33.51-58.97-109) (1-120) -*Homo sapiens* IGHG1*01 (100%) G1m17,1 (CH1 K120 (217) (121-218), bisagra 1-15 (219-233), CH2 (234-343), CH3 D12 (359), L14 (361) (344-448), CHS (449-450)) (121-450)], (223-215')-disulfure con la cadena ligera lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV2-14*01 (93.9%) -IGLJ3*02 (100%)) CDR-IMGT [9.3.10] (26-34.52-54.91-100) (1'-110') -*Homo sapiens* IGLC2*01 (100%) (111'-216')]; dímero (229-229":232-232")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada

QVQLVESGGG	VVQPGRSRLR	SCAASGFTFS	NYAMYVWRQA	PGKGLEWVAV	50
ISYDGSNKYY	ADSVKGRFTI	SRDNSKNTLY	LQMNSLRTE	TAVYYCASGS	100
DYGDYLLVYW	QGGLTVTVSS	ASTKGPSVFP	LAPSSKSTSG	GTAALGCLVK	150
DYFPEPVTVS	NSGALTSGV	HTFPAVLQSS	GLYSLSSVVT	VPSSSLGTQT	200
YICNVNHNKPS	NTKVDKKVEP	KSCDKHTHTC	PCPAPELLGG	PSVFLFPPKP	250
KDTLMISRTP	EVTGVVVDVS	HEDPEVKFNW	YVDGVEVHNA	KTKPREEQYN	300
STYRVVSVLT	VLHQDWLNGK	EYKCKVSNKA	LPAPIEKTI	KAKGQPREPQ	350
VYTLPPSRDE	LTKNQVSLTC	LVKGFYPSDI	AVEMESNQGP	ENNYKTTTPV	400
LDSDGSFFLY	SKLTVDKSRW	QQGNVFSCSV	MHEALHNHYT	QKSLSLSPGK	450

Light chain / Chaîne légère / Cadena ligera

QSALTQPASV	SGSPGQSITI	SCTGTSSDVG	GYNYVSWYQQ	HPGKAPKLMI	50
YDYSKRPSGV	SNRFGSKSG	NTASLTISGL	QSEDEADYYC	NSLTISISTWV	100
FGGGTKLTVL	GQPKAAPSVT	LFPPSSEELQ	ANKATLVCLII	SDFYPGAVTV	150
AWKADSSPVK	AGVETTTPSK	QSNKYYAASS	YLSLTPEQWK	SHRSYSCQVT	200
HEGSTVEKTV	APTECS				216

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104)	22-96	147-203	264-324	370-428
	22"-96"	147"-203"	264"-324"	370"-428"

Intra-L (C23-C104)	22'-90'	138'-197'
	22'''-90'''	138'''-197'''

Inter-H-L (h 5-CL 126)	223-215"	223"-215"
------------------------	----------	-----------

Inter-H-H (h 11, h 14)	229-229"	232-232"
------------------------	----------	----------

N-terminal glutamine cyclization to pyroglutamyl (pE, 5-oxoprolyl)

H VH Q1:

I, I"

L VL Q1:

I', I'''

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

300, 300"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

C-terminal lysine clipping / Coupeure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

450, 450"

molnupiravirum

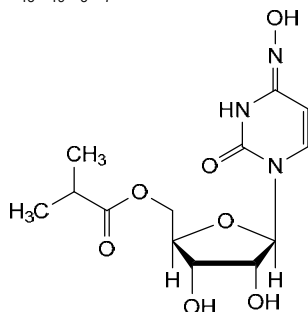
molnupiravir

*N*⁴-hydroxycytidine 5'-(2-methylpropanoate)

molnupiravir

5'-(2-méthylpropanoate) de *N*⁴-hydroxycytidine

molnupiravir

5'-(2-metilpropanoato) de *N*⁴-hidroxicitidinaC₁₃H₁₉N₃O₇**nezulcitinibum**

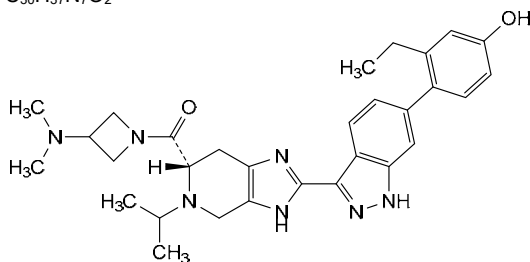
nezulcitinib

[3-(dimethylamino)azetidin-1-yl]{{(6*S*)-2-[6-(2-ethyl-4-hydroxyphenyl)-1*H*-indazol-3-yl]-5-(propan-2-yl)-4,5,6,7-tetrahydro-1*H*-imidazo[4,5-*c*]pyridin-6-yl}methanone

nézulcitinib

[3-(diméthylamino)azétidin-1-yl]{{(6*S*)-2-[6-(2-éthyl-4-hydroxyphényl)-1*H*-indazol-3-yl]-5-(propan-2-yl)-4,5,6,7-tétrahydro-1*H*-imidazo[4,5-*c*]pyridin-6-yl}méthanone

nezulcitinib

[3-(dimetilamino)azetidin-1-il]{{(6*S*)-2-[6-(2-etil-4-hidroksifenil)-1*H*-indazol-3-il]-5-(propan-2-il)-4,5,6,7-tetrahidro-1*H*-imidazo[4,5-*c*]piridin-6-il}metanonaC₃₀H₃₇N₇O₂**pemziviptadilum #**

pemziviptadil

human L-methionyl vasoactive intestinal peptide (2-29) fused to 121 repeats of different elastin-derived pentapeptides (VPGVG, VPGGG, VPGAG, VPGWP) (30-634), produced in *Escherichia coli*;
 fusion protein comprising L-methionyl (1)-vasoactive intestinal polypeptide (human VIP) (2-29) and an elastin-like artificial polymer (30-629) of 120 alternating pentapeptides of three types VPGVG, VPGGG, and VPGAG, and a C-terminal pentapeptide VPGWP (630-634), produced in *Escherichia coli*

pemziviptadil

L-méthionyl peptide vasoactif intestinal humain (2-29) fusionné à 121 répétitions de différents pentapeptides dérivés de l'élastine (VPGVG, VPGGG, VPGAG, VPGWP) (30-634), produits dans *Escherichia coli* ; protéine de fusion comprenant le polypeptide intestinal vasoactif L-méthionyl (1) (VIP humain) (2-29) et un polymère artificiel de type élastine (30-629) de 120 pentapeptides alternés de trois types VPGVG, VPGGG et VPGAG, et un pentapeptide C-terminal VPGWP (630-634), produit dans *Escherichia coli*

pemziviptadil

péptido humano L-metionil vasoactivo intestinal (2-29) fusionado a 121 repeticiones de pentapéptidos derivados de elastina diferentes (VPGVG, VPGGG, VPGAG, VPGWP) (30-634), producido en *Escherichia coli*; proteína de fusión que comprende polipéptido intestinal vasoactivo L-metionil (1) (humano VIP) (2-29) y un polímero artificial parecido a la elastina (30-629) de 120 pentapéptidos alternos de tres tipos VPGVG, VPGGG, y VPGAG, y un pentapéptido C-terminal VPGWP (630-634), producido en *Escherichia coli*

Sequence / Séquence / Secuencia

MHSDAVFTDN	YTRLRKQMAV	KKYLSILNV	PVGVPVGVG	PGGVPAGAV	50
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	100
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	150
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	200
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	250
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	300
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	350
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	400
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	450
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	500
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	550
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PVGVPVGVG	PGGVPAGAV	600
PVGVPVGVG	PVGVPGGGV	PAGVPGGGV	PWGP		634

positions 30-634: 12 × pattern AABCAAACB of 10 pentapeptides of three types
A = **VPGVG**, B = **VPGGG**, and C = **VPGAG** plus a C-terminal pentapeptide **VPGWP**

pidacmeranum #

pidacmeran

self-replicating messenger RNA (mRNA), 5'-capped, encoding *Venezuelan equine encephalitis virus* (VEEV) RNA-dependent RNA polymerase (VEEV proteins nsP1, nsP2, nsP3 and nsP4) and a full-length codon-optimised pre-fusion stabilised conformation variant (K986P and V987P) of the SARS-CoV-2 (severe acute respiratory syndrome coronavirus 2, GenBank: MN908947.3) spike (S) glycoprotein, flanked by 5' and 3' untranslated regions and a 3' poly(A) tail; replication of the RNA is under control of a VEEV subgenomic promoter.

pidacmèran

ARN messenger (ARNm) auto-répliquant, protégé en 5', codant pour l'ARN polymérase ARN-dépendante du *virus de l'encéphalite équine vénézuélienne* (protéines VEEV nsP1, nsP2, nsP3 et nsP4) et une variante complète de pré-fusion de conformation stabilisée (K986P et V987P) de la glycoprotéine spike (S) du SARS-CoV-2 (coronavirus 2 du syndrome respiratoire aigu sévère, GenBank: MN908947.3) avec des codons optimisés, flanqués de régions 5' et 3' non traduites une queue poly(A) 3'; la réplication de l'ARN est sous le contrôle du promoteur sous-génomique de VEEV.

pidacmerán

RNA mensajero auto replicativo (mRNA), protegido en 5', que codifica para la RNA polimerasa dependiente de RNA del *virus de la encefalitis equina venezolana* (proteínas VEEV nsP1, nsP2, nsP3 y nsP4) y una variante pre-fusión completa de conformación estabilizada (K986P and V987P) de la glicoproteína de la espícula (S) del SRAS-CoV-2 (coronavirus 2 del síndrome respiratorio agudo severo, GenBank: MN908947.3) con codones optimizados, flanqueada por regiones 5' y 3' no traducidas y una cola poly(A) en 3'; la replicación del ARN está bajo el control del promotor subgenómico de VEEV.

regdanvimabum #

regdanvimab

immunoglobulin G1-lambda, anti-[*Homo sapiens* severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) spike (S) protein, receptor binding domain (RBD)], humanized and *Homo sapiens* monoclonal antibody;
gamma1 heavy chain humanized (1-458) [VH (*Homo sapiens* IGHV2-70*12 (92.9%) -(IGHD) - IGHJ6*01 (100%)) CDR-IMGT [10.7.20] (26-35.53-59.98-117) (1-128) -*Homo sapiens* IGHG1*08p (100%) G1m3,1 (CH1 R120 (225) (129-226), hinge 1-15 (227-241), CH2 (242-351), CH3 D12 (367), L14 (369) (352-456), CHS (457-458)) (129-458)], (231-215')-disulfide with lambda light chain *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-51*01 (100%) - IGLJ2*01 (90.9%) K123>E (106)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC2*01 (98.1%) S45>G (156), T82>K (167) (111'-216')]; dimer (237-237":240-240")-bisdisulfide, produced in a Chinese hamster ovary (CHO) cell line derived from CHO-K1, glycoform alfa

regdanvimab

immunoglobuline G1-lambda anti-[*Homo sapiens* protéine spike (S) du coronavirus 2 du syndrome respiratoire aigu sévère (SARS-CoV-2), domaine de liaison au récepteur (RBD)], anticorps monoclonal humanisé et *Homo sapiens*;
chaîne lourde gamma1 humanisée (1-458) [VH (*Homo sapiens* IGHV2-70*12 (92.9%) -(IGHD) - IGHJ6*01 (100%)) CDR-IMGT [10.7.20] (26-35.53-59.98-117) (1-128) -*Homo sapiens* IGHG1*08p (100%) G1m3,1 (CH1 R120 (225) (129-226), charnière 1-15 (227-241), CH2 (242-351), CH3 D12 (367), L14 (369) (352-456), CHS (457-458)) (129-458)], (231-215')-disulfure avec la chaîne légère lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-51*01 (100%) -IGLJ2*01 (90.9%) K123>E (106)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC2*01 (98.1%) S45>G (156), T82>K (167) (111'-216')]; dimère (237-237":240-240")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) dérivant de la lignée cellulaire CHO-K1, glycoforme alfa

regdanvimab

inmunoglobulina G1-lambda anti-[*Homo sapiens* proteína espícula (S) del coronavirus 2 del síndrome respiratorio agudo severo (SRAS-CoV-2), dominio de unión al receptor (RBD)], anticuerpo monoclonal humanizado y *Homo sapiens*; cadena pesada gamma1 humanizada (1-458) [VH (*Homo sapiens*IGHV2-70*12 (92.9%) -(IGHD) -IGHJ6*01 (100%)) CDR-IMGT [10.7.20] (26-35.53-59.98-117) (1-128) -*Homo sapiens*IGHG1*08p (100%) G1m3,1 (CH1 R120 (225) (129-226), bisagra 1-15 (227-241), CH2 (242-351), CH3 D12 (367), L14 (369) (352-456), CHS (457-458)) (129-458)], (231-215')-disulfuro con la cadena ligera lambda *Homo sapiens* (1'-216') [V-LAMBDA (*Homo sapiens* IGLV1-51*01 (100%) -IGLJ2*01 (90.9%) K123>E (106)) CDR-IMGT [8.3.11] (26-33.51-53.90-100) (1'-110') -*Homo sapiens* IGLC2*01 (98.1%) S45>G (156), T82>K (167) (111'-216')]; dímero (237-237":240-240")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular derivada de CHO-K1, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada

```

QITLKESGPT LVKFTQTLTL TCSFSGFSLT TSGVGVGWIR QFPKALEWL 50
ALIDWDNKKY HTSLKTRLTI ISKDTSKNQV VLTMTNMDPV DTATYYCARI 100
EGFLRYNRNY YYGMDVWGO GTTVTVSSAS TKGPSVFPLA PSSKSTSGGT 150
AALGCLVRDY FEPFVTVSWN SGALTSGVHT FPAVLQSSGL YSLSSVVTVF 20C
SSSLGTQTYI CNVNHKFSNT KVDKRVFEKS CDKTHTCPPC PAPELLGGFS 25C
VELFPKPKD TLMISRTPEV TCVVVDVSHS DPEVKENWYV DGEVHNART 30C
KPREEQYNST YRVVSVLTVL HQDWLNGKEY KCKVSNKALP APIEKTISKA 35C
KGQPREPQVY TLPPSRDEL TKNQVSLTCLV KGFYPSDIAV EWESNGQPEN 400
NKYTPPVLD SDGSFFLYSK LTVDKSRWQQ GNVPSCSVMH EALHNHYTQR 450
SLSLSPGK 458

```

Light chain / Chaîne légère / Cadena ligera

```

ELVLTQPPSV SAAFGQKVTI SCSSGSSNIG NNYVSWYQQL FGTAPKLLIY 50
DNKRPSGPI DRESGSKSGT SATLGITGLQ TGDEADYCG TWDSSLSAGV 100
FGGGTELTVL GQPKAAPSVT LFPSSSEELQ ANKATLVCLI SDFYPGAVTV 150
AWKADGSFVK AGVETTKESK QSNKYAASS YLSLTPEQWK SHRSYSCQVT 20C
HEGSTVERTV ATECS 216

```

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22'-97' 155"-211" 272"-332" 378"-436"
 22"-97" 155"-211" 272"-332" 378"-436"

Intra-L (C23-C104) 22'-89' 138"-197"
 22"-89" 138"-197"

Inter-H-L (h 5-CL 126) 231-215' 231"-215"

Inter-H-H (h 11, h 14) 237-237" 240-240"

N-terminal glutamine cyclization to pyroglutanyl (pE, 5-oxoprolyl)

H VHQ1:
 1, 1"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:
 308, 308"

Fucosylated complex bi-antennary CHO-type glycans (G0F predominant) / glycanes de type CHO bi-antennaires complexes fucosylés (G0F prédominant) / glicanos de tipo CHO biantenaricos complejos fucosilados (G0F predominante).

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K.2:
 458, 458"

reluscovtogenum ralaplasmidum

reluscovtogene ralaplasmid

DNA plasmid expressing a consensus sequence* of the full-length SARS-CoV-2 (severe acute respiratory syndrome coronavirus 2) spike (S) glycoprotein, with an N-terminal IgE leader sequence, under control of the human CMV promoter (hCMV promoter) and a bovine growth hormone poly-adenylation signal (bGH poly(A)). The plasmid also contains a pUC origin of replication and a kanamycin antibiotic resistance gene.

*based upon sequences published on GISAID (Global Initiative on Sharing All Influenza Data) in early 2020.

réluscovtogène ralaplasamide

plasmide d'ADN exprimant une séquence consensus* complète de la glycoprotéine spike (S) du SARS-CoV-2 (coronavirus 2 du syndrome respiratoire aigu sévère), avec une séquence leader N-terminale d'IgE, sous contrôle du promoteur du CMV humain (promoteur hCMV) et d'un signal de polyadénylation de l'hormone de croissance bovine (bGH poly(A)). Le plasmide contient également une origine de répllication pUC et un gène de résistance à la kanamycine.

*basé sur des séquences publiées dans GISAID (Global Initiative on Sharing All Influenza Data) début 2020.

reluscovtogén ralaplasimida

plásmido de DNA que expresa una secuencia consenso* completa de la glicoproteína de la espícula (S) de SARS-CoV-2 (coronavirus 2 del síndrome respiratorio agudo severo), con una secuencia líder N-terminal de la IgE, bajo el control del promotor de CMV humano (promotor hCMV) y una señal de poliadenilación de la hormona de crecimiento bovina (bGH poly(A)). El plásmido contiene también un origen de replicación pUC y un gen de resistencia al antibiótico kanamicina.

*basada en secuencias publicadas en GISAID (Global Initiative on Sharing All Influenza Data) a principios de 2020.

sotrovimabum #
sotrovimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) spike (S) protein, receptor binding domain (RBD)], *Homo sapiens* monoclonal antibody; gamma1 heavy chain *Homo sapiens* (1-457) [VH (*Homo sapiens* IGHV1-18*01 (92.9%) -(IGHD) -IGHJ5*01 (92.9%)) CDR-IMGT [8.8.20] (26-33.51-58.97-116) (1-127) -*Homo sapiens* IGHG1*01 G1m17,1, G1v24 CH3 L107, S114(CH1 K120 (224) (128-225), hinge 1-15 (226-240), CH2 (241-350), CH3 D12 (366), L14 (368), M107>L (438), N114>S (444) (351-455), CHS (456-457)) (128-457)], (230-214')-disulfide with kappa light chain *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (96.7%) -IGKJ4*01 (100%)) CDR-IMGT [7.3.8] (27-33.51-53.90-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (236-236":239-239")-bisdisulfide, produced in a Chinese hamster ovary (CHO)-K1 cell line, glycoform alfa

sotrovimab

immunoglobuline G1-kappa anti-[*Homo sapiens* protéine spike (S) du coronavirus 2 du syndrome respiratoire aigu sévère (SARS-CoV-2), domaine de liaison au récepteur (RBD)], anticorps monoclonal *Homo sapiens*;

chaîne lourde gamma1 *Homo sapiens* (1-457) [VH (*Homo sapiens* IGHV1-18*01 (92.9%) -(IGHD) -IGHJ5*01 (92.9%)) CDR-IMGT [8.8.20] (26-33.51-58.97-116) (1-127) -*Homo sapiens* IGHG1*01 G1m17,1, G1v24 CH3 L107, S114 (CH1 K120 (224) (128-225), charnière 1-15 (226-240), CH2 (241-350), CH3 D12 (366), L14 (368), M107>L (438), N114>S (444) (351-455), CHS (456-457)) (128-457)], (230-214')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (96.7%) -IGKJ4*01 (100%)) CDR-IMGT [7.3.8] (27-33.51-53.90-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (236-236'':239-239')-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) lignée cellulaire CHO-K1, glycoforme alfa

sotrovimab

immunoglobulina G1-kappa anti-[*Homo sapiens* proteína espícula (S) del coronavirus 2 del síndrome respiratorio agudo severo (SRAS-CoV-2), dominio de unión al receptor (RBD)], anticuerpo monoclonal *Homo sapiens*;

cadena pesada gamma1 *Homo sapiens* (1-457) [VH (*Homo sapiens* IGHV1-18*01 (92.9%) -(IGHD) -IGHJ5*01 (92.9%)) CDR-IMGT [8.8.20] (26-33.51-58.97-116) (1-127) -*Homo sapiens* IGHG1*01 G1m17,1, G1v24 CH3 L107, S114 (CH1 K120 (224) (128-225), bisagra 1-15 (226-240), CH2 (241-350), CH3 D12 (366), L14 (368), M107>L (438), N114>S (444) (351-455), CHS (456-457)) (128-457)], (230-214')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-214') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (96.7%) -IGKJ4*01 (100%)) CDR-IMGT [7.3.8] (27-33.51-53.90-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (236-236'':239-239')-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular CHO-K1, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada

QVQLVQSGAE	VKKPGASVKV	SCKASGYPT	SYGISWVRQA	PGQGLEWMGW	50
ISTYQGNTRY	AQKFGQRTM	TTDTSTTTGY	MELRLRSD	TAVYYCARDY	100
TRGAWFGESL	IGGFDNWGGQ	TLTVTSSAST	KGPSVFPLAP	SSKSTSGGTA	150
ALGLVKVDYF	PEPVTVSWNS	GALTSGVHTF	PAVLQSSGLY	SLSSVTVPSY	200
SSLGTQTYIC	NVNHKPSNTK	VDKKVEPKSC	DKTHTCPPCP	APPELLGGPSV	250
FLFPPPKPDT	LMISRTPEVT	CVVVDVSHED	PEVKFNWYVD	GVEVHNAKTK	300
PREEQYNSTY	RVVSVTLVLH	QDWLNGKEYK	CKVSNKALPA	PIEKTISKAK	350
GQPREPQVYT	LPSPRDELTK	NQVSLTCLVK	GFYPDSIAVE	WESNGQPENN	400
YKTTTPPVLD	DGSFFLYSKL	TVDKSRWQQG	NVFSCSVLHE	ALHSHTQKLS	450
LSLSPGK					457

Light chain / Chaîne légère / Cadena ligera

EIVLTQSPGT	LSLSPGERAT	LSCRASQTVS	STSLAWYQQK	PGQAPRLLIY	50
GASSRATGIP	DRFSGSGSGT	DFTLTISRLE	PEDFAVYYCQ	QHDSTLTFFG	100
GTKVEIKRTV	AAPSVFIFPP	SDEQLKSGTA	SVVCLLNIFY	PREAKVQWKV	150
DNALQSGNSQ	ESVTEQDSKD	STYLSSTLT	LSKADYEKHK	VYACEVTHQG	200
LSPVTKSFN	RGEK				214

Post-translational modifications

Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro

Intra-H (C23-C104) 22-96 154-210 271-331 377-435
22"-96" 154"-210" 271"-331" 377"-435"

Intra-L (C23-C104) 23'-89' 134'-194'
23'''-89''' 134'''-194'''

Inter-H-L (h 5-CL 126) 230-214' 230"-214"

Inter-H-H (h 11, h 14) 236-236" 239-239"

N-terminal glutamine cyclization to pyroglutamyl (pE, 5-oxoprolyl)

H VH Q1:

1, 1"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación

H CH2 N84.4:

307, 307"

Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires complexes fucosylés / glicanos de tipo CHO biantenarijos complejos fucosilados.

Deamidation sites / Sites de désamidation / Posiciones de desamidación

H CH2 N98: 325, 325"

H CH3 N44: 394, 394"

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal

H CHS K2:

457, 457"

subasumstatum

subasumstat

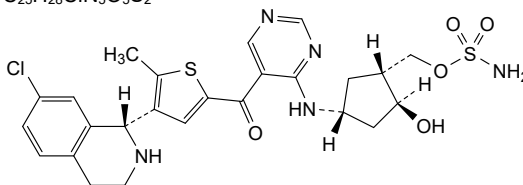
{{(1*R*,2*S*,4*R*)-4-[(5-{4-[(1*R*)-7-chloro-1,2,3,4-tetrahydroisoquinolin-1-yl]-5-methylthiophene-2-carbonyl}pyrimidin-4-yl)amino]-2-hydroxycyclopentyl)methyl sulfamate

subasumstat

sulfamate de {{(1*R*,2*S*,4*R*)-4-[(5-{4-[(1*R*)-7-chloro-1,2,3,4-tétrahydroisoquinoléin-1-yl]-5-méthylthiophène-2-carbonyl}pyrimidin-4-yl)amino]-2-hydroxycyclopentyl)méthyle

subasumstat

sulfamato de {{(1*R*,2*S*,4*R*)-4-[(5-{4-[(1*R*)-7-cloro-1,2,3,4-tetrahydroisoquinolein-1-il]-5-metiltiofeno-2-carbonil}pirimidin-4-il)amino]-2-hidroxiciclopentil}metilo

$$\text{C}_{25}\text{H}_{28}\text{ClN}_5\text{O}_5\text{S}_2$$
**tixagevimabum #**

tixagevimab

immunoglobulin G1-kappa, anti-[*Homo sapiens* severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) spike (S) protein, receptor binding domain (RBD)], *Homo sapiens* monoclonal antibody; gamma1 heavy chain *Homo sapiens* (1-453) [VH (*Homo sapiens* IGHV1-58*01 (98.0%) -(IGHD) -IGHJ3*02 (93.8%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Homo sapiens* IGHG1*03 G1m3, nG1m1, G1v21 CH2 Y15.1, T16, E18, G1v39 CH2 F1.3, E1.2, S116 (CH1 R120 (220) (124-221), hinge 1-15 (222-236), CH2 L1.3>F (240), L1.2>E (241), M15.1>Y (258), S16>T (260), T18>E (262), P116>S (337) (237-346), CH3 E12 (362), M14 (364) (347-451), CHS (452-453)) (124-453)], (226-216')-disulfide with kappa light chain *Homo sapiens* (1'-216') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (98.9%) -IGKJ1*01 (100%)) CDR-IMGT [7.3.10] (27-33.51-53.90-99) (1'-109') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (155), V101 (193) (110'-216')]; dimer (232-232':235-235')-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

tixagévimab

immunoglobuline G1-kappa anti-[*Homo sapiens* protéine spike (S) du coronavirus 2 du syndrome respiratoire aigu sévère (SARS-CoV-2), domaine de liaison au récepteur (RBD)], anticorps monoclonal *Homo sapiens*; chaîne lourde gamma1 *Homo sapiens* (1-453) [VH (*Homo sapiens* IGHV1-58*01 (98.0%) -(IGHD) -IGHJ3*02 (93.8%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Homo sapiens* IGHG1*03 G1m3, nG1m1, G1v21 CH2 Y15.1, T16, E18, G1v39 CH2 F1.3, E1.2, S116 (CH1 R120 (220) (124-221), charnière 1-15 (222-236), CH2 L1.3>F (240), L1.2>E (241), M15.1>Y (258), S16>T (260), T18>E (262), P116>S (337) (237-346), CH3 E12 (362), M14 (364) (347-451), CHS (452-453)) (124-453)], (226-216')-disulfure avec la chaîne légère kappa *Homo sapiens* (1'-216') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (98.9%) -IGKJ1*01 (100%)) CDR-IMGT [7.3.10] (27-33.51-53.90-99) (1'-109') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (155), V101 (193) (110'-216')]; dimère (232-232':235-235')-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

tixagevimab

inmunoglobulina G1-kappa anti-[*Homo sapiens* proteína espícula (S) del coronavirus 2 del síndrome respiratorio agudo severo (SRAS-CoV-2), dominio de unión al receptor (RBD)], anticuerpo monoclonal *Homo sapiens*;
cadena pesada gamma1 *Homo sapiens* (1-453) [VH (*Homo sapiens* IGHV1-58*01 (98.0%) -(IGHD) -IGHJ3*02 (93.8%)) CDR-IMGT [8.8.16] (26-33.51-58.97-112) (1-123) -*Homo sapiens* IGHG1*03 G1m3, nG1m1, G1v21 CH2 Y15.1, T16, E18, G1v39 CH2 F1.3, E1.2, S116 (CH1 R120 (220) (124-221), bisagra 1-15 (222-236), CH2 L1.3>F (240), L1.2>E (241), M15.1>Y (258), S16>T (260), T18>E (262), P116>S (337) (237-346), CH3 E12 (362), M14 (364) (347-451), CHS (452-453)) (124-453)], (226-216')-disulfuro con la cadena ligera kappa *Homo sapiens* (1'-216') [V-KAPPA (*Homo sapiens* IGKV3-20*01 (98.9%) -IGKJ1*01 (100%)) CDR-IMGT [7.3.10] (27-33.51-53.90-99) (1'-109') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (155), V101 (193) (110'-216')]; dímero (232-232":235-235")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO), forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
QMQLVQSGPE VKKPGTSVKV SCKASGFTFM SSAVQWVRQA RGQRLEWIGW 50
IVIGSGNTNY AQKFQERVIT TRDMSTSTAY MELSSLRSED TAVYYCAAPY 100
CSSISGNDGF DIWGQGTMTV VSSASTKGPS VFPLAPSSKS TSGGTAALGC 150
LVKDYFPEPV TVSWNSGALT SGVHTFPAVL QSSGLYSLSS VVTVPSSSLG 200
TQTYICNVNH KPSNTKVDKR VEPKSCDKTH TCPPCPAPEF EGGPSVFLFP 250
PKPKDLYIT REPEVTCVVV DVSHEDPEVK FNMYYVDGVEV HNAKTPREE 300
QYNSTYRVVS VLTVLHQDWL NGKEYKCKVS NKALPASIEK TISKAKGQPR 350
EPQVYTLPPS REEMTKNQVS LTCLVKGFYP SDIAVEWESN GQPENNYKTT 400
PPVLDSGGSF FLYSKLTVDK SRWQGGNVFS CSMVHEALHN HYTKLSLSLS 450
PGK 453

Light chain / Chaîne légère / Cadena ligera
EVLVTQSPGT LSLSPGERAT LSCRASQSVS SSYLAWYQK PGQAPRLLIY 50
GASSRATGIP DRFSGSGSGT DFTLTISRLE PEDFAVYYCQ HYGSSRGWTF 100
GQGTKEIKR TVAAPSVEIF PPSDEQLKSG TASVVCLLNN FYPREKVQW 150
KVDNALQSGN SQESVTEQDS KDSTYLSST LTLSKADYEK HKVYACEVTH 200
QGLSSPVTKS FNRGEC 216

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22'-96" 101'-106" 150'-206" 267'-327" 373'-431"
22'-96" 101'-106" 150'-206" 267'-327" 373'-431"
Intra-H (CDR3) 101'-106" 101'-106"
101'-106"
Intra-L (C23-C104) 23'-89" 136'-196"
23'-89" 136'-196"
Inter-H-L (h 5-CL 126) 226'-216" 226'-216"
Inter-H-H (h 11, h 14) 232'-232" 235'-235"

N-terminal glutamine cyclization to pyroglutamyl (pE, 5-oxoprolyl)
H VH Q1:
I, I"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
303, 303"
Fucosylated complex bi-antennary CHO-type glycans / glycanes de type CHO bi-antennaires
complexes fucosylés / glicanos de tipo CHO biantenarios complejos fucosilados.

C-terminal lysine clipping / Coupure de la lysine C-terminale / Recorte de lisina C-terminal
H CHS K2:
453, 453"

tozinameranum #
tozinameran

messenger RNA (mRNA), 5'-capped, encoding a full-length, codon-optimised pre-fusion stabilised conformation variant (K986P and V987P) of the SARS-CoV-2 (severe acute respiratory syndrome coronavirus 2, GenBank: MN908947.3) spike (S) glycoprotein, flanked by 5' and 3' untranslated regions and a 3' poly(A) tail; contains N1-methylpseudouridine instead of uridine (*all-U>m¹Ψ*).

tozinaméran	ARN messenger (ARNm), protégé en 5', codant pour un variant de pré-fusion stabilisé en conformation (K986P et V987P) de la glycoprotéine spike (S) du SARS-CoV-2 (coronavirus 2 du syndrome respiratoire aigu sévère, GenBank: MN908947.3) complet, avec des codons optimisés, flanqués de régions 5' et 3' non traduites et d'une queue poly(A) 3'; contient de la N1-méthylpseudouridine au lieu de l'uridine (<i>tout-U>m¹Ψ</i>).
tozinamerán	RNA mensajero (mRNA), protegido en 5', que codifica para una variante pre-fusión de conformación estabilizada (K986P and V987P) de la glicoproteína de la espícula (S) del SRAS-CoV-2 (coronavirus 2 del síndrome respiratorio agudo severo, GenBank: MN908947.3) completa, con codones optimizados, flanqueada por regiones 5' y 3' no traducidas y una cola poly(A) en 3'; contiene N1-metilpseudouridina en lugar de uridina (<i>all-U>m¹Ψ</i>).
zansecimabum # zansecimab	immunoglobulin G4-kappa, anti-[<i>Homo sapiens</i> ANGPT2 (angiotensinogen 2)], humanized monoclonal antibody; gamma4 heavy chain humanized (1-447) [VH (<i>Homo sapiens</i> IGHV1-18*01 (86.7%) -(IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) - <i>Homo sapiens</i> IGHG4*01, G4v5 h P10,G4v4 CH2 A1.3, A1.2 (CH1 (122-219), hinge 1-12 S10>P (229) (220-231), CH2 F1.3>A (235), L1.2>A (236) (232-341), CH3 (342-446), CHS K>del (447)) (122-447)], (135-214')-disulfide with kappa light chain humanized (1'-214') [V-KAPPA (<i>Homo sapiens</i> IGKV1-12*01 (83.2%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') - <i>Homo sapiens</i> IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimer (227-227":230-230")-bisdisulfide, produced in a Chinese hamster ovary (CHO) cell line derived from CHO-GSKO, glycoform alfa
zansécimab	immunoglobuline G4-kappa anti-[<i>Homo sapiens</i> ANGPT2 (angiotensinogen 2)], anticorps monoclonal humanisé; chaîne lourde gamma4 humanisée (1-447) [VH (<i>Homo sapiens</i> IGHV1-18*01 (86.7%) -(IGHD) -IGHJ4*01 (93.3%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) - <i>Homo sapiens</i> IGHG4*01 G4v5 h P10,G4v4 CH2 A1.3, A1.2 (CH1 (122-219), charnière 1-12 S10>P (229) (220-231), CH2 F1.3>A (235), L1.2>A (236) (232-341), CH3 (342-446), CHS K>del (447)) (122-447)], (135-214')-disulfure avec la chaîne légère kappa humanisée (1'-214') [V-KAPPA (<i>Homo sapiens</i> IGKV1-12*01 (83.2%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') - <i>Homo sapiens</i> IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dimère (227-227":230-230")-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO) dérivant de la lignée cellulaire CHO-GSKO, glycoforme alfa
zansecimab	inmunoglobulina G4-kappa anti-[<i>Homo sapiens</i> ANGPT2 (angiotensinogen 2)], anticuerpo monoclonal humanizado;

cadena pesada gamma4 humanizada (1-447) [VH (*Homo sapiens*IGHV1-18*01 (86.7%) -(IGHD) - IGHJ4*01 (93.3%)) CDR-IMGT [8.8.14] (26-33.51-58.97-110) (1-121) -*Homo sapiens* IGHG4*01 G4v5 h P10,G4v4 CH2 A1.3, A1.2 (CH1 (122-219), bisagra 1-12 S10>P (229) (220-231), CH2 F1.3>A (235), L1.2>A (236) (232-341), CH3 (342-446), CHS K>del (447)) (122-447)], (135-214')-disulfuro con la cadena ligera kappa humanizada (1'-214') [V-KAPPA (*Homo sapiens* IGKV1-12*01 (83.2%) -IGKJ4*01 (100%)) CDR-IMGT [6.3.9] (27-32.50-52.89-97) (1'-107') -*Homo sapiens* IGKC*01 (100%) Km3 A45.1 (153), V101 (191) (108'-214')]; dímero (227-227":230-230")-bisdisulfuro, producido en las células ováricas de hámster chino (CHO) línea celular derivada de CHO-GSKO, forma glicosilada alfa

Heavy chain / Chaîne lourde / Cadena pesada
QVQLVQSGAE VKKPGASVKV SCKASGYST DYNMVMVRQA PGQGLEWMGY 50
IDPYNGGTGY NQKFEGRVTM TDDTSTSTAY MELRSLRSD TAVVYCARTR 100
DRYDVWYFDV WQGGLTLVTVS SASTKGPSVF PLAPCSRSTS ESTAALGCLV 150
KDYFPEPVTV SWNSGALTSG VHTFPAVLQS SGLYSLSSV TVPSSSLGK 200
TYTCNVDHKP SNTKVDKRVE SKYGPCCPPC PAPEAAGGPS VLFPPKPKD 250
TLMISRTPEV TCVVVDVSQE DPEVQFNWYV DGEVHNAKT KPREEQFNST 300
YRVVSVLTVL HQDWLNGKEY KCKVSNKGLP SSIEKTIKA KGQPREPQVY 350
TLPPSQEEMT KNQVSLTCLV KGFYPSDIAV EWESNGQPEN NYKTTTPVLD 400
SDGSFFLYSR LTVDKSRWQE GNVFSCSVMH EALHNHYTQK SLSLSLG 447

Light chain / Chaîne légère / Cadena ligera
DIQMTQSPSS VSASVGDRVT ITCKASQDQVY IAVAWYQQK GKAPKLLIYW 50
ASTRDGTGVP RFGSGSGGTD FTLTISSLQ EDFAITYYCHQ YSSYPPTFGG 100
GTKVEIKRTV AAPSVFIFPP SDEQLKSGTA SVVCLLNIFY PREAKVQWKV 150
DNALQSGNSQ ESVTEQDSKD STYLSSTLT LSKADYEKHK VYACEVTHQG 200
LSSPVTKSFN RGE 214

Post-translational modifications
Disulfide bridges location / Position des ponts disulfure / Posiciones de los puentes disulfuro
Intra-H (C23-C104) 22-96 148-204 262-322 368-426
22"-96" 148"-204" 262"-322" 368"-426"
Intra-L (C23-C104) 23'-88' 134'-194'
23"-88" 134"-194"
Inter-H-L (CH1 10-CL 126) 135-214' 135"-214"
Inter-H-H (h 8, h 11) 227-227" 230-230"

N-terminal glutamine cyclization to pyroglutamyl (pE, 5-oxopropyl)
H VH Q1:
1, 1"

N-glycosylation sites / Sites de N-glycosylation / Posiciones de N-glicosilación
H CH2 N84.4:
298, 298"
Fucosylated complex bi-antennary CHO-type glycans (G0F predominant) / glycanes de type CHO bi-antennaires complexes fucosylés (G0F prédominant) / glicanos de tipo CHO biantenarijos complejos fucosilados (G0F predominante).

zorecimeranum #
zorecimeran

messenger RNA (mRNA), 5'-capped, encoding a full-length codon-optimised pre-fusion stabilized conformation variant (K986P and V987P) of the SARS-CoV-2 (severe acute respiratory syndrome coronavirus 2; NC_045512.2) spike (S) glycoprotein, flanked by 5' and 3' untranslated regions; the 3' untranslated region comprises a part of the human alpha globin gene 3' UTR sequence element, a synthetic poly(A) tail and a poly(C)-rich sequence followed by a histone stem loop sequence as a 3' tail.

zoréciméran

ARN messenger (ARNm), protégé en 5', codant pour un variant de pré-fusion stabilisé en conformation (K986P et V987P) de la glycoprotéine spike (S) du SARS-CoV-2 (coronavirus 2 du syndrome respiratoire aigu sévère), NC_045512.2) complet, avec des codons optimisés, flanqués de régions 5' et 3' non traduites. La région 3' non traduite comprend une partie de l'élément de séquence 3' UTR du gène de l'alpha globine humaine, une queue poly(A) synthétique et une séquence riche en poly(C) suivie d'une séquence d'histones stem-loop comme extrémité en 3'.

zorecimerán

RNA mensajero (mRNA), protegido en 5', que codifica para una variante pre-fusión de conformación estabilizada (K986P and V987P) de la glicoproteína de la espícula (S) del SRAS-Cov-2 (coronavirus 2 del síndrome respiratorio agudo severo, NC_045512.2) completa, con codones optimizados, flanqueada por regiones 5' y 3' no traducidas. La región 3' no traducida comprende una parte del elemento secuencia 3' UTR del gen de la alfa globina humana, una cola poly(A) sintética y una secuencia rica en poly(C) seguida de una secuencia de histona stem-loop cola 3'.

Electronic structure available on Mednet: <http://mednet.who.int/>
Structure électronique disponible sur Mednet: <http://mednet.who.int/>
Estructura electrónica disponible en Mednet: <http://mednet.who.int/>
* <http://www.who.int/medicines/services/inn/publication/en/>

AMENDMENTS TO PREVIOUS LISTS
MODIFICATIONS APPORTÉES AUX LISTES ANTÉRIEURES
MODIFICACIONES A LAS LISTAS ANTERIORES

Recommended International Nonproprietary Names (Rec. INN): List 76
Dénominations communes internationales recommandées (DCI Rec.): Liste 76
Denominaciones Comunes Internacionales Recomendadas (DCI Rec.): Lista 76
(WHO Drug Information, Vol. 30, No. 3, 2016)

p.483 **blontuvetmabum #**

blontuvetmab
 blontuvetmab
 blontuvetmab

replace the description by the following one
remplacer la description par la suivante
sustitúyase la descripción por la siguiente

immunoglobulin G2_V-kappa-C-lambda, anti-[**Canis lupus familiaris** MS4A1 (membrane-spanning 4-domains subfamily A member 1, CD20)], caninized monoclonal antibody; gamma2 heavy chain chimeric (1-448) [*Mus musculus* VH (*Mus musculus* IGHV1-15*01 -(IGHD) -IGHJ1*03) [8.8.6] (1-113) -*Canis lupus familiaris* IGHG2*02 (CH1 T26>Q (131) (114-211), hinge (212-229), CH2 (230-339), CH3 (340-446), CHS (447-448)) (114-448)], (128-218')-disulfide with V-kappa-C-lambda light chain chimeric (1'-219') [*Mus musculus* V-KAPPA (*Mus musculus* IGKV8-30*01 -IGKJ5*01) [12.3.9] (1'-112') -*Canis lupus familiaris* IGL1CS1*01 V45.3>I (162) (114'-219')]; dimer (225-225'':228-228'')-bisdisulfide

immunoglobuline G2_V-kappa-C-lambda, anti-[**Canis lupus familiaris** MS4A1 (membre 1 de la sous-famille A à 4 domaines transmembranaires, CD20)], anticorps monoclonal caninisé; chaîne lourde gamma2 chimérique (1-448) [*Mus musculus* VH (*Mus musculus* IGHV1-15*01 -(IGHD) -IGHJ1*03) [8.8.6] (1-113) -*Canis lupus familiaris* IGHG2*02 (CH1 T26>Q (131) (114-211), charnière (212-229), CH2 (230-339), CH3 (340-446), CHS (447-448)) (114-448)], (128-218')-disulfure avec la chaîne légère V-kappa-C-lambda chimérique (1'-219') [*Mus musculus* V-KAPPA (*Mus musculus* IGKV8-30*01 -IGKJ5*01) [12.3.9] (1'-112') -*Canis lupus familiaris* IGL1CS1*01 V45.3>I (162) (114'-219')]; dimère (225-225'':228-228'')-bisdisulfure

immunoglobulina G2_V-kappa-C-lambda, anti-[**Canis lupus familiaris** MS4A1 (miembro 1 de la subfamilia A con 4 dominios transmembranarios, CD20)], anticuerpo monoclonal caninizado; cadena pesada gamma2 quimérica (1-448) [*Mus musculus* VH (*Mus musculus* IGHV1-15*01 -(IGHD) -IGHJ1*03) [8.8.6] (1-113) -*Canis lupus familiaris* IGHG2*02 (CH1 T26>Q (131) (114-211), bisagra (212-229), CH2 (230-339), CH3 (340-446), CHS (447-448)) (114-448)], (128-218')-disulfuro con la cadena ligera V-kappa-C-lambda quimérica (1'-219') [*Mus musculus* V-KAPPA (*Mus musculus* IGKV8-30*01 -IGKJ5*01) [12.3.9] (1'-112') -*Canis lupus familiaris* IGL1CS1*01 V45.3>I (162) (114'-219')]; dímero (225-225'':228-228'')-bisdisulfuro

p.534 **tamtuvetmabum #**
 -535 tamtuvetmab
 tamtuvetmab
 tamtuvetmab

replace the description by the following one
remplacer la description par la suivante
sustitúyase la descripción por la siguiente

immunoglobulin G2_V-kappa-C-lambda, anti-[**Canis lupus familiaris** CD52], caninized monoclonal antibody;
 gamma2 heavy chain chimeric (1-456) [chimeric VH (*Rattus norvegicus* IGHV7S6*01 (97.00%) -(IGHD) - *Canis lupus familiaris* IGHJ-E2RCC8) [8.10.12] (1-121) -*Canis lupus familiaris* IGHG2*02 (CH1 (122-219), hinge (220-237), CH2 (238-347), CH3 (348-454), CHS (455-456))(122-456)], (136-212')-disulfide with V-kappa-C-lambda light chain chimeric (1'-213') [*Rattus norvegicus* V-KAPPA (*Rattus norvegicus* IGKV22S7 (93.70%) -IGKJ1*01) [6.3.9] (1'-107') - *Canis lupus familiaris* IGLC1S1*01 V45.3>I (156) (108'-213')]; dimer (233-233":236-236")-bisdisulfide

immunoglobuline G2_V-kappa-C-lambda, anti-[**Canis lupus familiaris** CD52], anticorps monoclonal caninisé;
 chaîne lourde gamma2 chimérique (1-456) [VH chimérique (*Rattus norvegicus* IGHV7S6*01 (97.00%) -(IGHD) -*Canis lupus familiaris* IGHJ-E2RCC8) [8.10.12] (1-121) -*Canis lupus familiaris* IGHG2*02 (CH1 (122-219), charnière (220-237), CH2 (238-347), CH3 (348-454), CHS (455-456)) (122-456)], (136-212')-disulfure avec la chaîne légère V-kappa-C-lambda chimérique (1'-213') [*Rattus norvegicus* V-KAPPA (*Rattus norvegicus* IGKV22S7 (93.70%) -IGKJ1*01) [6.3.9] (1'-107') -*Canis lupus familiaris* IGLC1S1*01 V45.3>I (156) (108'-213')]; dimère (233-233":236-236")-bisdisulfure

inmunoglobulina G2_V-kappa-C-lambda, anti-[**Canis lupus familiaris** CD52], anticuerpo monoclonal caninizado;
 cadena pesada gamma2 quimérica (1-456) [VH quimérico (*Rattus norvegicus* IGHV7S6*01 (97.00%) -(IGHD) -*Canis lupus familiaris* IGHJ-E2RCC8) [8.10.12] (1-121) -*Canis lupus familiaris* IGHG2*02 (CH1 (122-219), bisagra (220-237), CH2 (238-347), CH3 (348-454), CHS (455-456))(122-456)], (136-212')-disulfuro con la cadena ligera V-kappa-C-lambda quimérica (1'-213') [*Rattus norvegicus* V-KAPPA (*Rattus norvegicus* IGKV22S7 (93.70%) -IGKJ1*01) [6.3.9] (1'-107') -*Canis lupus familiaris* IGLC1S1*01 V45.3>I (156) (108'-213')]; dímero (233-233":236-236")-bisdisulfuro

Recommended International Nonproprietary Names (Rec. INN): List 77**Dénominations communes internationales recommandées (DCI Rec.): Liste 77****Denominaciones Comunes Internacionales Recomendadas (DCI Rec.): Lista 77****(WHO Drug Information, Vol. 31, No. 1, 2017)****p.118 ranevetmabum #**

-119 ranevetmab *replace the description by the following one*
 ranevetmab *remplacer la description par la suivante*
 ranevetmab *sustitúyase la descripción por la siguiente*

immunoglobulin G1-kappa, anti-[**Canis lupus familiaris** NGF (nerve growth factor, nerve growth factor beta polypeptide, NGFB, beta-NGF)], caninized monoclonal antibody;
 gamma1 heavy chain (1-453) [caninized VH (*Rattus norvegicus*IGHV5S13*01 (71.40%) -(IGHD)-IGHJ4*01) [8.7.16] (1-122) -*Canis lupus familiaris*IGHG1*01 (CH1 (123-219), hinge (220-233), CH2 (234-343), CH3 (344-451), CHS (452-453)) (123-453)], (137-213')-disulfide with kappa light chain (1'-217') [caninized V-KAPPA (*Rattus norvegicus*IGKV12S34*01 (76.80%) -IGKJ2-3*01) [6.3.9] (1'-107') -*Canis lupus familiaris*IGKC*01 (108'-213') -4-mer (214'-217')]; dimer (224-224'':226-226'':232-232'')-trisdifulfide

immunoglobuline G2-kappa, anti-[**Canis lupus familiaris** NGF (facteur de croissance du nerf, facteur de croissance du nerf polypeptide bêta, NGFB, bêta-NGF)], anticorps monoclonal caninisé;
 chaîne lourde gamma2 (1-453) [VH caninisé (*Rattus norvegicus*IGHV5S13*01 (71.40%) -(IGHD)-IGHJ4*01) [8.7.16] (1-122) -*Canis lupus familiaris*IGHG1*01 (CH1 (123-219), charnière (220-233), CH2 (234-343), CH3 (344-451), CHS (452-453)) (123-453)], (137-213')-disulfure avec la chaîne légère kappa (1'-217') [V-KAPPA caninisé (*Rattus norvegicus*IGKV12S34*01 (76.80%) -IGKJ2-3*01) [6.3.9] (1'-107') -*Canis lupus familiaris*IGKC*01 (108'-213') -4-mer (214'-217')]; dimère (224-224'':226-226'':232-232'')-trisdisulfure

inmunoglobulina G2-kappa, anti-[**Canis lupus familiaris** NGF (factor de crecimiento de los nervios, factor de crecimiento de nervios polipéptido beta, NGFB, beta-NGF)], anticuerpo monoclonal caninizado;
 cadena pesada gamma2 (1-453) [VH caninizado (*Rattus norvegicus*IGHV5S13*01 (71.40%) -(IGHD)-IGHJ4*01) [8.7.16] (1-122) -*Canis lupus familiaris*IGHG1*01 (CH1 (123-219), bisagra (220-233), CH2 (234-343), CH3 (344-451), CHS (452-453)) (123-453)], (137-213')-disulfuro con la cadena ligera kappa (1'-217') [V-KAPPA caninizado (*Rattus norvegicus*IGKV12S34*01 (76.80%) -IGKJ2-3*01) [6.3.9] (1'-107') -*Canis lupus familiaris*IGKC*01 (108'-213') -4-mer (214'-217')]; dímero (224-224'':226-226'':232-232'')-trisdisulfuro

Recommended International Nonproprietary Names (Rec. INN): List 79**Dénominations communes internationales recommandées (DCI Rec.): Liste 79****Denominaciones Comunes Internacionales Recomendadas (DCI Rec.): Lista 79****(WHO Drug Information, Vol. 32, No. 1, 2018)**

p.170

suprimáse
 toglatato de tigilanol

insertese
 tiglatato de tigilanol

Recommended International Nonproprietary Names (Rec. INN): List 81**Dénominations communes internationales recommandées (DCI Rec.): Liste 81****Denominaciones Comunes Internacionales Recomendadas (DCI Rec.): Lista 81***(WHO Drug Information, Vol. 31, No. 1, 2017)***olenasufligenum relduparvovecum #**

olenasufligene relduparvovec *A revised sequence has been uploaded to the INN MedNet, which includes three nucleotide insertions within the CAG promoter compared to the previous version, increasing the total length of the vector to 4075bp. The table of features was revised accordingly.*

olénasufligène relduparvovec *Une séquence révisée a été téléchargée sur MedNet DCI, et comprend trois insertions de nucléotides dans le promoteur CAG par rapport à la version précédente, augmentant la longueur totale du vecteur à 4075 pb. Le tableau des caractéristiques a été révisé en conséquence.*

olenasufligén relduparvovec *Se cargó una secuencia revisada en MedNet DCI e incluye tres inserciones de nucleótidos dentro del promotor CAG en comparación con la versión anterior, aumentando la longitud total del vector a 4075 pb. La tabla de características se revisó en consecuencia.*

Recommended International Nonproprietary Names (Rec. INN): List 83**Dénominations communes internationales recommandées (DCI Rec.): Liste 83****Denominaciones Comunes Internacionales Recomendadas (DCI Rec.): Lista 83***(WHO Drug Information, Vol. 34, No. 1, 2020)***p.41 efbemalenograstimum alfa #**

-42 efbemalenograstim alfa *replace the description and the structure by the following ones*

efbémalénograstim alfa *remplacer la description et la structure par les suivantes*

efbemalenograstim alfa *sustitúyase la descripción y la estructura por las siguientes*

human granulocyte colony-stimulating factor (G-CSF) fragment fused via a peptidyl linker to a human immunoglobulin G2 Fc fragment variant, dimer:

[human granulocyte colony-stimulating factor (G-CSF, pluripoiétin) short **V³⁶, S³⁷, E³⁸>del**] isoform (1-174)]-[GSG₃S(G₄S)₂ linker (175-190)]-[human immunoglobulin G2 Fc fragment (223 C-terminal residues) (*Homo sapiens*IGHG2*01 (natural S³⁴⁴>A variant); hinge (191-197), CH2 (P²⁹⁷>S) (198-306), CH3 (307-411), CHS (412-413) (191-413))] fusion protein, dimer (193-193':196-196')-bisdisulfide, produced in Chinese hamster ovary (CHO) cells, glycoform alfa

fragment du facteur de stimulation des colonies de granulocytes humain (G-CSF) lié par un peptide à un variant du fragment Fc de l'immunoglobuline humaine G2, dimère:

[facteur humain de stimulation des colonies de granulocytes (G-CSF, pluripoiétine) **V³⁶, S³⁷, E³⁸>del**] isoforme courte (1-174)]-[GSG₃S(G₄S)₂ linker (175-190)]-[fragment Fc de l'immunoglobuline humaine G2 (résidus 223 C-terminaux) (*Homo sapiens*IGHG2*01 (variant naturel S³⁴⁴>A); charnière (191-197), CH2 (P²⁹⁷>S) (198-306), CH3 (307-411), CHS (412-413) (191-413))] protéine de fusion, dimère (193-193':196-196')-bisdisulfure, produit par des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

fragmento del factor de estimulación de las colonias de granulocitos humano (G-CSF) que se une por un péptido a una variante del fragment Fc de la inmunoglobulina humana G2, dímero:

[factor humano de estimulación de las colonias de granulocitos (G-CSF, pluripoyetina) [V³⁶, S³⁷, E³⁸-del] isoforma corta (1-174)]-[GS₃S(G₄S)₂ conector (175-190)]-[fragmento Fc de la inmunoglobulina humana G2 (residuo 223 C-terminal) (*Homo sapiens* IGHG2*01 (variante natural S³⁴⁴>A); bisagra (191-197), CH2 (P²⁹⁷>S) (198-306), CH3 (307-411), CHS (412-413) (191-413))] proteína de fusión, dímero (193-193':196-196')-bisdisulfuro, producido en las células ováricas de hamsters chinos (CHO), glicoforma alfa

Sequence / Séquence / Secuencia:

```
TPLGPASSLP QSFLLKCLEQ VRKIQGDGAA LQEKLCATYK LCHPEELVLL 50
GHSLGIPWAP LSSCPSQALQ LAGCLSQLHS GLFLYQGLLQ ALEGISPGLG 100
PTLDTLQLDV ADFATTIWQQ MEELGMAPAL QPTQGAMPAP ASAFQRRAGG 150
VLVASHLQSF LEVSYRVLRH LAQPGSGGGG GGGGSGGGG VECPPCPAPP 200
VAGPSVFLFP PKPKDTLMIS RTPEVTCVVV DVSHEDEPEVQ FNWYVDGVEV 250
HNAKTKPRE QFNSTFRVVS VLTVVHQDWL NGKEYKCKVS NKGLPASIEK 300
TISKTKGQPR EPQVYTLPPS REEMTKNQVS LTCLVKGFYP SDIAVEWESN 350
GQPENNYKTT PPM L DSDGSF FLYSKLTVDK SRWQQGNVFS CSMVHEALHN 400
HYTQKSLSL PGK 413
```

Disulfide bridge location / Position de la pont disulfure / Posición del puente disulfuro:

intra-G-CSF: 36-42, 64-74, 36'-42', 64'-74'
intra-IgG2 Fc: 227-287, 333-391, 227'-287', 333'-391'
inter-IgG2 Fc: 193-193', 196-196'

Glycosylation sites / Sites de glycosylation / Posiciones de glicosilación

N-linked glycans: Asn263, Asn263'

O-linked glycans: Thr133, Thr133'

p.298
-299

narsoplimab #

narsoplimab
narsoplimab
narsoplimab

replace the description by the following one
remplacer la description par la suivante
sustitúyase la descripción por la siguiente

immunoglobulin G4-lambda, anti-[*Homo sapiens* MASP2 (mannan-binding lectin serine peptidase 2, mannan-binding lectin serine protease 2, mannan-binding lectin serine protease 1 pseudogene 1, MASP1P1)], *Homo sapiens* monoclonal antibody; **gamma4** heavy chain *Homo sapiens* (1-445) [VH (*Homo sapiens* IGHV2-26*01 (94.0%) -(IGHD) -IGHJ4*01 (100%)) [10.7.10] (1-118) -*Homo sapiens* IGHG4*01 (CH1 (119-216), hinge 1-12 S10>P (226) (217-228), CH2 (229-338), CH3 (339-443), CHS (444-445)) (119-445)], (132-211')-disulfide with lambda light chain *Homo sapiens* (1'-212') [V-LAMBDA (*Homo sapiens* IGLV3-1*01 (93.5%) -IGLJ2*01 (100%)) [6.3.9] (1'-106') -*Homo sapiens* IGLC2*01 (100%) (107'-212')]; dimer (224-224'':227-227'')-bisdisulfide, produced in Chinese hamster ovary (CHO)-S cell line, glycoform alfa

immunoglobuline G4-lambda, anti-[*Homo sapiens* MASP2 (sérine peptidase 2 de la lectine liant le mannane, sérine protéase 2 de la lectine liant le mannane, pseudogène 1 de la protéase 1 de la lectine liant le mannane, MASP1P1)], anticorps monoclonal *Homo sapiens*;
chaîne lourde **gamma4** *Homo sapiens* (1-445) [VH (*Homo sapiens* IGHV2-26*01 (94.0%) -(IGHD) -IGHJ4*01 (100%)) [10.7.10] (1-118) -*Homo sapiens* IGHG4*01 (CH1 (119-216), charnière 1-12 S10>P (226) (217-228), CH2 (229-338), CH3 (339-443), CHS (444-445)) (119-445)], (132-211')-disulfure avec la chaîne légère lambda *Homo sapiens* (1'-212') [V-LAMBDA (*Homo sapiens* IGLV3-1*01 (93.5%) -IGLJ2*01 (100%)) [6.3.9] (1'-106') -*Homo sapiens* IGLC2*01 (100%) (107'-212')]; dimère (224-224'':227-227'')-bisdisulfure, produit dans des cellules ovariennes de hamsters chinois (CHO) lignée cellulaire S, glycoforme alfa

inmunoglobulina G4-lambda, anti-[*Homo sapiens* MASP2 (serina peptidasa 2 de la lectina que se une al manano, serina proteasa 2 de la lectina unida al manano, pseudogen 1 de la proteasa 1 de la lectina unida al manano, MASP1P1)], anticuerpo monoclonal *Homo sapiens*; cadena pesada gamma4 *Homo sapiens* (1-445) [VH (*Homo sapiens*IGHV2-26*01 (94.0%) -(IGHD) -IGHJ4*01 (100%)) [10.7.10] (1-118) - *Homo sapiens*IGHG4*01 (CH1 (119-216), bisagra 1-12 S10>P (226) (217-228), CH2 (229-338), CH3 (339-443), CHS (444-445)) (119-445)], (132-211')-disulfuro con la cadena ligera lambda *Homo sapiens* (1'-212') [V-LAMBDA (*Homo sapiens*IGLV3-1*01 (93.5%) -IGLJ2*01 (100%)) [6.3.9] (1'-106') -*Homo sapiens*IGLC2*01 (100%) (107'-212')]; dímero (224-224":227-227")-bisdisulfuro, producido en las células ováricas de hamsters chinos (CHO) línea celular S, glicoforma alfa

Proposed International Nonproprietary Names (Prop. INN): List 122

Dénominations communes internationales proposées (DCI Prop.): Liste 122

Denominaciones Comunes Internacionales Propuestas (DCI Prop.): Lista 122

(WHO Drug Information, Vol. 33, No. 2, 2019)

p.831 eflepedocokinum alfa#

-832 eflepédocokine alfa *remplacer la description par la suivante*
 eflepedocokina alfa *sustitúyase la descripción por la siguiente*

interleukine 22 humaine (IL22, cytokine Zcyto18, facteur inductible dérivé des lymphocytes T lié à IL10, IL-TIF) (1-146), fusionnée via un peptide liant GSG₃S(G₄S)₂ (147-162) au fragment Fc C-terminal de l'immunoglobuline G2 humaine (163-385), P²⁶⁹>S-mutant **S³¹⁶>A-variant**, dimère (165-165':168-168')-bisdisulfure, produit dans des cellules ovariennes de hamster chinois (CHO), glycoforme alfa

interleukina 22 humana (IL22, citoquina Zcyto18, factor inducible derivado de los linfocitos T relacionado con IL10, IL-TIF) (1-146), fusionada a través de un péptido que se une a GSG₃S(G₄S)₂ (147-162) con el fragmento Fc C-terminal de la inmunoglobulina G2 humana (163-385), P²⁶⁹>S-mutante **S³¹⁶>A-variante**, dímero (165-165':168-168')-bisdisulfuro, producido en las células ováricas de hamster chino (CHO), glicoforma alfa

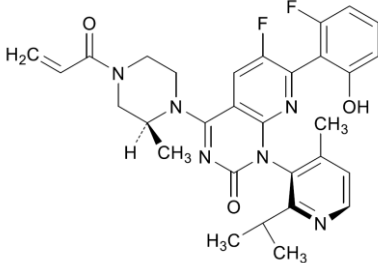
Procedure and Guiding Principles / Procédure et Directives / Procedimientos y principios generales

The text of the *Procedures for the Selection of Recommended International Nonproprietary Names for Pharmaceutical Substances* and *General Principles for Guidance in Devising International Nonproprietary Names for Pharmaceutical Substances* will be reproduced in proposed INN lists only.

Les textes de la *Procédure à suivre en vue du choix de dénominations communes internationales recommandées pour les substances pharmaceutiques* et des *Directives générales pour la formation de dénominations communes internationales applicables aux substances pharmaceutiques* seront publiés seulement dans les listes des DCI proposées.

El texto de los *Procedimientos de selección de denominaciones comunes internacionales recomendadas para las sustancias farmacéuticas* y de los *Principios generales de orientación para formar denominaciones comunes internacionales para sustancias farmacéuticas* aparece solamente en las listas de DCI propuestas.

毒薬・劇薬等の指定審査資料のまとめ

化学名・別名	(1 <i>M</i>)-6-Fluoro-7-(2-fluoro-6-hydroxyphenyl)-1-[4-methyl-2-(propan-2-yl)pyridin-3-yl]-4-[(2 <i>S</i>)-2-methyl-4-(prop-2-enoyl)piperazin-1-yl]pyrido[2,3- <i>d</i>]pyrimidin-2(1 <i>H</i>)-one [別名：Sotorasib・（ソトラシブ）]
構造式	
効能・効果	がん化学療法後に増悪したKRAS G12C変異陽性の切除不能な進行・再発の非小細胞肺癌
用法・用量	通常、成人にはソトラシブとして960mgを1日1回経口投与する。なお、患者の状態により適宜減量する。
劇薬等の指定	
市販名及び有効成分・分量	[アムジェン株式会社] 製剤：ルマケラス錠120mg（1錠中ソトラシブ120mg（力価）含有）
毒性	GLP条件下で実施したラットを用いた28日間反復投与毒性試験において、sotorasibの30、100及び200 mg/kgを投与したときの忍容性は良好であり、使用した動物の10%に重篤な毒性が発現する投与量（STD10）は200 mg/kgを超えていた。ラットで認められたsotorasibに関連した主な変化は軽微から軽度であり、腎尿細管上皮の変性／壊死、脾臓重量の増加、白血球の増加、並びに網状赤血球数及び赤血球指数の変化を伴う循環赤血球（RBC）量の減少（ヘモグロビン、赤血球数及びヘマトクリット）と、病理組織学的検査における腎臓髄質外帯外側での腎尿細管上皮の変性／壊死であった。尿細管における変化は28日間の休薬期間終了後に部分的に回復したことから、回復あるいは修復が持続していることが確認された。しかしながら、少数の尿細管の周辺に間質性繊維増殖がみられた。ラットにおけるsotorasibの投与に関連した脾臓重量の増加は、28日間の休薬期間終了時には認められず、血液学的パラメータの変化はすべて対照群と同程度であった。ラットを用いた3カ月間反復投与及び2カ月間回復性試験では、sotorasibのより高い用量（60、180及び750 mg/kg）まで評価しており、ヒトの臨床曝露量を網羅する、より高い全身曝露が得られた。腎尿細管変性／壊死の発現頻度及び重症度（軽微から顕著）は、28日間反復投与毒性試験における腎臓の変化と比較して高かった。これは、試験期間が長いこと及びsotorasibの全身曝露量が高いことに起因したものと考えられた。休薬期間終了時、すべての用量において、sotorasibの投与に関連した腎臓の変化は部分的に回復した。ラットを用いた3カ月間反復投与毒性試験におけるSTD10は180 mg/kgであった。 GLP条件下で実施したイヌを用いた28日間反復投与毒性試験において、sotorasibの10、30及び300 mg/kgを投与したときの忍容性は良好であった。重篤な毒性が発現しない最大投与量（HNSTD）は300 mg/kgを超えていた。イヌにおけるsotorasibの投与に関連した主な変化は、網状赤血球数の減少を伴う軽微から軽度の赤血球系パラメータの減少であった。GLP条件下で実施したイヌを用いた3カ月間反復投与毒性試験では、sotorasibのより高い全身曝露を得るために、高い用量（200及び1000 mg/kg/dayを100及び500 mg/kgの用量で1日2回投与）を用いて評価した。しかしながら、sotorasibの1000 mg/kg/dayを投与しても、ラットを用いた3カ月間反復投与毒性試験と比較して高い曝露は得られなかった。sotorasibの投与に関連した所見

	として、胆嚢内の異物、軽微から軽度の血液学的検査及び血液生化学的検査パラメータの変化、肝臓、下垂体又は甲状腺の病理組織学的変化がみられたが、これらは重篤な毒性ではなく、肝細胞酵素誘導に対する適応性反応又は二次的反応と判断した。イヌにおいて腎毒性は認められなかった。本試験におけるHNSTDは1000 mg/kg/dayであった。
副作用	国際共同第I/II相試験である20170543試験の第I相及び第II相パートにおいて本剤960 mgを1回以上投与された非小細胞肺癌患者190例（日本人13例を含む）中128例（67.4%）に副作用が認められ、主な副作用（発現率5%以上）は、下痢（27.9%）、悪心、アラニンアミノトランスフェラーゼ増加、アスパラギン酸アミノトランスフェラーゼ増加（各16.3%）、疲労（11.1%）、血中アルカリホスファターゼ増加（7.9%）、嘔吐（7.4%）及び腹痛（5.3%）であった。
会社	アムジェン株式会社 製剤：製造販売

ソトラシブ 添付資料一覧

アムジェン株式会社

第3部 品質に関する文書

3.2 データ又は報告書

3.2.S 原薬

資料番号	資料表題	著者	実施期間	試験実施場所	評価・参考
3.2.S	原薬	Amgen	—	海外	評価

3.2.P 製剤

資料番号	資料表題	著者	実施期間	試験実施場所	評価・参考
3.2.P	製剤	Amgen	—	海外	評価

3.2.A その他

資料番号	資料表題	著者	実施期間	試験実施場所	評価・参考
3.2.A	その他	Amgen	—	海外	評価

第3部 品質に関する文書
3.3 参考文献

資料番号	著者・表題・出典
3.3-1	
3.3-2	
3.3-3	
3.3-4	
3.3-5	
3.3-6	

第4部 非臨床試験報告書

4.2 試験報告書

4.2.1 薬理試験

4.2.1.1 効力を裏付ける試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.1.1-1	Study R20150198 - Biochemical Characterization of AMG 510 and Metabolites in KRAS ^{G12C/C118A} and KRAS ^{C118A} Nucleotide Exchange Assays	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-2	Study R20150199 - Cellular Characterization of AMG 510	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-3	Study R20190078 - Effects of AMG 510 on KRAS Signaling in NCI-H358 and MIA PaCa-2 Cells In Vitro	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-4	Study 153894 - In Vitro Combination Effects of AMG 510 with MAPK Pathway, PI3K Pathway, and Other Targeted Inhibitors	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-5	Study R20150188 - AMG 510 Inhibits Phospho-ERK1/2 Signaling in <i>KRAS p.G12C</i> Pancreatic MIA PaCa-2 T2 Tumors in Female Athymic Nude Mice	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-6	Study R20190129 - AMG 510 Inhibits ERK1/2 Phosphorylation Signaling and Results in Covalent Occupancy of KRAS ^{G12C} in <i>KRAS p.G12C</i> NSCLC NCI-H358 Tumors in Female Athymic Nude Mice	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-7	Study R20150192 - Effect of AMG 510 on Phospho-ERK1/2 Signaling at Early Timepoints (0.25 to 2 hours) in <i>KRAS p.G12C</i> Pancreatic MIA PaCa-2 T2 Tumors in Female Athymic Nude Mice	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-8	Study R20150189 - The Effect of Once-daily Dosing of AMG 510 on Tumor Growth in the <i>KRAS p.G12C</i> Pancreatic MIA PaCa-2 T2 Xenograft Model in Female Athymic Nude Mice	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-9	Study R20150190 - The Effect of Once-daily Dosing of AMG 510 on Tumor Growth in the <i>KRAS p.G12C</i> NSCLC H358 Xenograft Model in Female Athymic Nude Mice	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-10	Study R20150191 - The Effect of Once-daily Dosing of AMG 510 on Tumor Growth in the <i>KRAS p.G12V</i> Colorectal Carcinoma SW480-1AC Xenograft Model in Female Athymic Nude Mice	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-11	Study R20190130 - AMG 510 Inhibited ERK 1/2 Phosphorylation in a Time and Dose Dependent Manner in Human KRAS <i>p.G12C</i> CRC Patient-derived Xenograft Tumors in Female NOD/SCID IL2rg (NSG) Mices	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-12	Study R20190131 - Once Daily Dosing of AMG 510 Inhibited Tumor Growth in a Human <i>KRAS p.G12C</i> CRC Patient-derived Xenograft Model in Female NSG Mice	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-13	Study R20180033 - The Effect of AMG 510 in Combination with MEK Inhibitor 1009089 (PD-0325901) in the <i>KRAS p.G12C</i> NSCLC H358 Xenograft Model in Female Athymic Nude Mice	Amgen	2017/1/1 - 2017/1/1	海外	評価
4.2.1.1-14	Study 153358 - Effect of AMG 510 in Combination with Afatinib on Tumor Growth in the <i>KRAS p.G12C</i> NSCLC NCI-H358 Xenograft Model in Female Athymic Nude Mice	Amgen	2017/1/1 - 2017/1/1	海外	評価

4.2.1.1-15	Study 153397 - The Effect of AMG 510 in Combination with RMC-4550 (SHP2i), on Tumor Growth in the <i>KRAS p.G12C</i> NSCLC NCI-H358 Xenograft Model in Female Athymic Nude Mice	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.1-16	Study R20180032 - The Effect of AMG 510 in Combination with Carboplatin on Tumor Growth in the <i>KRAS p.G12C</i> NSCLC H358 Xenograft Model in Female Athymic Nude Mice	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.1-17	Study R20180002 - AMG 510 Inhibits ERK1/2 Phosphorylation in Murine Colorectal CT-26 <i>KRAS p.G12C</i> -H10 Tumors in Female Balb/c Mice	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.1-18	Study R20190110 - AMG 510 Inhibits ERK1/2 Phosphorylation in Murine Colorectal CT-26 <i>KRAS p.G12C</i> -H10 Tumors in Female Balb/c Mice	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.1-19	Study R20190128 - The Effect of Once-daily Dosing of AMG 510 on Tumor Growth in the Parental CT-26 <i>KRAS p.G12D</i> Colorectal Carcinoma Syngeneic Model in Female Balb/c Mice	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.1-20	Study R20180034 - The Effect of AMG 510 in Combination with anti-PD-1 antibody on Tumor Growth in the Murine Colorectal CT-26 <i>KRAS p.G12C</i> -H10 Syngeneic Model in Female Balb/c Mice	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.1-21	Study R20180035 - The Effect of AMG 510 on Tumor Infiltrating Lymphocytes in CT-26 <i>KRAS p.G12C</i> -H10 Colorectal Tumors in Female Balb/c Mice	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.1-22	Study R20190105 - The Effect of AMG 510 and/or Anti-PD-1 on CT-26 <i>KRAS p.G12C</i> -H10 Colorectal Tumors by Immunohistochemistry (IHC) in the Female BALB/c Mouse	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.1-23	Study R20180036 - The Effect of AMG 510 on Gene Expression and T-Cell Priming in CT-26 <i>KRAS p.G12C</i> -H10 Colorectal Tumors in Female Balb/c Mice	Amgen	20■■■■ - 20■■■■	海外	評価

4.2.1.2 副次的薬理試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.1.2-1	Study 125831 - mRNA Expression Assessment of KRAS	Amgen	20■■■■	海外	評価
4.2.1.2-2	Study 124452 - <i>In Vitro</i> Pharmacology Study of 3365648 and 3365626 – Early Panel	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.2-3	Study 124453 - <i>In Vitro</i> Pharmacology Late Panel Study of 3365648 and 3365626	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.2-4	Study R20150219 - Cysteine Proteome Profiling of AMG 510 in NCI-H358	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.2-5	Study 124807 - <i>In Vitro</i> Pharmacology Study of 3368167 Early Panel	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.2-6	Study 124787 - KINOMEScan™ Profiling Service Primary Screen Report	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.2-7	Study 124808 - <i>In Vitro</i> Pharmacology Study of 3368167 Late Panel	Amgen	20■■■■ - 20■■■■	海外	評価

4.2.1.2-8	Study 153413 - <i>In Vitro</i> Pharmacology New Early Panel Study of 3375854 and 3413829	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.2-9	Study 153414 - <i>In Vitro</i> Pharmacology New Late Panel Study of 3375854 and 3413829	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.2-10	Study 153415 - KINOMEScan™ Profiling Service Primary Screen Report	Amgen	20■■■■ - 20■■■■	海外	評価

4.2.1.3 安全性薬理試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.1.3-1	Study 150431 - Effect of AMG 510 on Cloned hERG Potassium Channels Expressed in Human Embryonic Kidney Cells	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.3-2	Study 150458 - AMG 510: Single-Dose Oral Gavage Cardiovascular Safety Pharmacology Evaluation in Male Telemetry-Instrumented Conscious Dogs	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.1.3-3	Study 124803 - AMG3368167: Acute <i>In Vitro</i> Effect on hERG recorded From HEK Cells	Amgen	20■■■■ - 20■■■■	海外	参考
4.2.1.3-4	Study 153419 - hERG ³ H-Dofetilide Filter Binding Assay – Study of 3375854#1 and 3413829#1	Amgen	20■■■■ - 20■■■■	海外	参考

4.2.2 薬物動態試験

4.2.2.1 分析法及びバリデーション報告書

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.2.1-1	Study 150506 - Determination of AMG 510 in K ₂ EDTA Sprague Dawley Rat Plasma Using Plasma Using LC-MS/MS Following Protein Precipitation	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.2.1-2	Study 152694 - Validation of an LC-MS/MS Assay for AMG 510 in Rabbit Plasma with K ₂ EDTA	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.2.1-3	Study 150505 - Determination of AMG 510 in K ₂ EDTA Beagle Dog Plasma Using LC-MS/MS Following Protein Precipitation	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.2.1-4	Study 151603 - Validation of an LC-MS/MS Assay for AMG 510 in Rat Plasma with K ₂ EDTA (and Addendum 1)	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.2.1-5	Study 151604 - Validation of an LC-MS/MS Assay for AMG 510 in Dog Plasma with K ₂ EDTA (and Addendum 1)	Amgen	20■■■■ - 20■■■■	海外	評価

4.2.2.2 吸収

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.2.2-1	Study 150529 - Preclinical Pharmacokinetics of AMG 510	Amgen	20■■■■	海外	評価

4.2.2.2-2	Study 125554 - Pharmacokinetics of AMG 510 Following Oral Solution, Suspension, or Tablet Dose Administration to Male Beagle Dogs	Amgen	20■■■	海外	評価
-----------	---	-------	-------	----	----

4.2.2.3 分布

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.2.3-1	Study 150563 - <i>In Vitro</i> Permeability Assessment of AMG 510 and Its Metabolite (AMG3368167) in MDCK Transwell Assay	Amgen	20■■■	海外	評価
4.2.2.3-2	Study 150530 - Mouse, Rat, Dog, Monkey, and Human <i>In Vitro</i> Plasma Protein Binding of AMG 510 and AMG3368167	Amgen	20■■■	海外	評価
4.2.2.3-3	Study 150532 - <i>In Vitro</i> Assessment of Blood-to-Plasma Partitioning of AMG 510 in Mouse, Rat, Dog, Monkey, and Human	Amgen	20■■■	海外	評価
4.2.2.3-4	Study 152495 - Pharmacokinetics, Distribution, Metabolism, and Excretion of ¹⁴ C-AMG510 Following Oral Administration to Rats	Amgen	20■■■ - 20■■■	海外	評価

4.2.2.4 代謝

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.2.4-1	Study 150531 - Characterization of <i>In Vitro</i> Metabolites of AMG 510 Formed in Mouse, Rat, Dog, Monkey and Human Hepatic Microsomes and Hepatocytes	Amgen	20■■■	海外	評価

4.2.2.5 排泄

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.2.5-1	Study 124686 - Metabolism and Excretion of AMG 510 Following a Single Intravenous or Oral Dose to Male Sprague-Dawley Rats	Amgen	20■■■	海外	評価
4.2.2.5-2	Study 153304 - Pharmacokinetics, Absorption, Metabolism, and Excretion of ¹⁴ C-AMG 510 Following Oral Administration to Dogs	Amgen	20■■■ - 20■■■	海外	評価

4.2.3 毒性試験

4.2.3.2 反復投与毒性試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.3.2-1	Study 124431 - AMG3365648: 7-Day Oral Toxicology Study in the Sprague Dawley Rat	Amgen	20■■■ - 20■■■	海外	参考

4.2.3.2-2	Study 150428 - AMG 510: 28-Day Oral Toxicology Study in the Sprague Dawley Rat with a 28-Day Recovery Period	Amgen	20■■■ - 20■■■	海外	評価
4.2.3.2-3	Study 150432 - AMG 510: A 3-Month Oral Toxicology Study in the Sprague Dawley Rat with a 2-Month Recovery Period	Amgen	20■■■ - 20■■■	海外	評価
4.2.3.2-4	Study 124273 - AMG3365648: Exploratory 7-Day Oral Toxicology Study in the Female Beagle Dog	Amgen	20■■■ - 20■■■	海外	参考
4.2.3.2-5	Study 151550 - AMG 510: Exploratory 7-Day Oral Toxicology Study in the Beagle Dog (and Amendment 1)	Amgen	20■■■	海外	参考
4.2.3.2-6	Study 150429 - AMG 510: 28-Day Oral Toxicology Study in the Beagle Dog	Amgen	20■■■ - 20■■■	海外	評価
4.2.3.2-7	Study 150433 - AMG 510: A 3-Month BID Oral Toxicology Study in the Beagle Dog	Amgen	20■■■ - 20■■■	海外	評価

4.2.3.3 遺伝毒性試験

4.2.3.3.1 In vitro試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.3.3.1-1	Study 124824 - Bacterial Reverse Mutation Assay in 24-well Plates (Micro Ames) with AMG3365648 #6	Amgen	20■■■ - 20■■■	海外	参考
4.2.3.3.1-2	Study 124825 - <i>In Vitro</i> Micro HPBL Micronucleus Assay with AMG3365648 #6	Amgen	20■■■ - 20■■■	海外	参考
4.2.3.3.1-3	Study 150436 - AMG 510: Bacterial Reverse Mutation Test in <i>Salmonella typhimurium</i> and <i>Escherichia coli</i>	Amgen	20■■■ - 20■■■	海外	評価

4.2.3.3.2 In vivo試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.3.3.2-1	Study 150435 - AMG 510 - Combined In Vivo Mammalian Erythrocyte Micronucleus Test and Alkaline Comet Assay in Rat	Amgen	20■■■ - 20■■■	海外	評価

4.2.3.5 生殖発生毒性試験

4.2.3.5.2 胚・胎児発生に関する試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.3.5.2-1	Study 150434 - AMG 510: Embryo-Fetal Development Toxicology Study in the Sprague Dawley Rat	Amgen	20■■■ - 20■■■	海外	評価
4.2.3.5.2-2	Study 150808 - AMG 510: Maternal Tolerability Study in the Pregnant New Zealand White Rabbit	Amgen	20■■■ - 20■■■	海外	参考
4.2.3.5.2-3	Study 150810 - AMG 510: Embryo-Fetal Development Study in the New Zealand White Rabbit	Amgen	20■■■ - 20■■■	海外	評価

4.2.3.7 その他の毒性試験

4.2.3.7.3 メカニズム試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.3.7.3-1	Study 153127 - AMG 510: Exploratory 7-Day Oral Toxicology Study in the Male Sprague Dawley Rat	Amgen	2019/01/01 - 2019/01/01	海外	参考
4.2.3.7.3-2	Study 153409 - AMG 510 and AMG3368167: Cytochrome P450 and UDP Glucuronosyltransferase Induction in Cultured Beagle Dog Hepatocytes	Amgen	2019/01/01 - 2019/01/01	海外	参考

4.2.3.7.5 代謝物の毒性試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.3.7.5-1	Study 125020 - Bacterial Reverse Mutation Assay in 24-well Plates (Micro Ames) with AMG3368167#2	Amgen	2019/01/01 - 2019/01/01	海外	参考
4.2.3.7.5-2	Study 125324 - <i>In Vitro</i> Micro HPBL Micronucleus Assay with AMG3368167#4	Amgen	2019/01/01 - 2019/01/01	海外	参考
4.2.3.7.5-3	Study 153305 - In Silico Mutagenicity Predictions of AMG3413829 (M18)	Amgen	2019/01/01 - 2019/01/01	海外	参考
4.2.3.7.5-4	Study 153306 - In Silico Mutagenicity Prediction of AMG3375854 (M10)	Amgen	2019/01/01 - 2019/01/01	海外	参考

4.2.3.7.6 不純物の毒性試験

資料番号	資料表題	著者	実施期間 (年/月/日)	試験実施場所	評価・参考
4.2.3.7.6-1	Study 152458 - Mixture of AMG 510 and Impurities: A 28-Day Oral Toxicology Study in the Sprague Dawley Rat	Amgen	2019/01/01 - 2019/01/01	海外	評価
4.2.3.7.6-2	Study 153451 - Mixture of AMG 510 and Impurities: A 28-Day Oral Toxicology Study in the Sprague Dawley Rat	Amgen	2019/01/01 - 2019/01/01	海外	評価
4.2.3.7.6-3	Study 125621 - Bacterial Reverse Mutation Assay in 24-well Plates (Micro Ames) with AMG3379613	Amgen	2019/01/01 - 2019/01/01	海外	参考
4.2.3.7.6-4	Study 153049 - AMG3396902: GLP Bacterial Reverse Mutation Test in <i>Salmonella typhimurium</i> and <i>Escherichia coli</i>	Amgen	2019/01/01 - 2019/01/01	海外	評価
4.2.3.7.6-5	Study 153050 - AMG3414168: GLP Bacterial Reverse Mutation Test in <i>Salmonella typhimurium</i> and <i>Escherichia coli</i>	Amgen	2019/01/01 - 2019/01/01	海外	評価
4.2.3.7.6-6	Study 153051 - AMG3412607 GLP Bacterial Reverse Mutation Test in <i>Salmonella typhimurium</i> and <i>Escherichia coli</i>	Amgen	2019/01/01 - 2019/01/01	海外	評価
4.2.3.7.6-7	Study 153052 - AMG3368167: GLP Bacterial Reverse Mutation Test in <i>Salmonella typhimurium</i> and <i>Escherichia coli</i>	Amgen	2019/01/01 - 2019/01/01	海外	評価
4.2.3.7.6-8	Study 153053 - AMG3379613: GLP Bacterial Reverse Mutation Test in <i>Salmonella typhimurium</i> and <i>Escherichia coli</i>	Amgen	2019/01/01 - 2019/01/01	海外	評価
4.2.3.7.6-9	Study 153054 - AMG3379620: GLP Bacterial Reverse Mutation Test in <i>Salmonella typhimurium</i> and <i>Escherichia coli</i>	Amgen	2019/01/01 - 2019/01/01	海外	評価

4.2.3.7.6-10	Study 153494 - AMG3379613 Lot No. MH135223-001 and Lot No. MH144641-001: Bacterial Reverse Mutation Test in 24-well Plates (Micro Ames) in <i>Salmonella typhimurium</i> and <i>Escherichia coli</i>	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.3.7.6-11	Study 154030 - AMG3422551: Bacterial Reverse Mutation Test in 24-well Plates (Micro Ames) in <i>Salmonella Typhimurium</i> and <i>Escherichia Coli</i>	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.3.7.6-12	Study 154031 - AMG3422536: Bacterial Reverse Mutation Test in 24-well Plates (Micro Ames) in <i>Salmonella Typhimurium</i> and <i>Escherichia Coli</i>	Amgen	20■■■■ - 20■■■■	海外	評価
4.2.3.7.6-13	Study 154032 - AMG3422547: Bacterial Reverse Mutation Test in 24-well Plates (Micro Ames) in <i>Salmonella Typhimurium</i> and <i>Escherichia Coli</i>	Amgen	20■■■■ - 20■■■■	海外	評価

4.2.3.7.7 その他の試験

資料番号	資料表題	著者	実施期間 (年／月／日)	試験実施場所	評価・参考
4.2.3.7.7-1	Study 124666 - Screening Assay to Assess the Phototoxic Potential of AMG3365648	Amgen	20■■■■ - 20■■■■	海外	参考

第4部 非臨床試験報告書

4.3 参考文献

資料番号	著者・表題・出典
4.3-1	AACR Project GENIE Consortium. AACR project GENIE: powering precision medicine through an international consortium. <i>Cancer Discovery</i> . 2017;7:818-831.
4.3-2	Anders MW. Glutathione-dependent bioactivation of haloalkanes and haloalkenes. <i>Drug metabolism reviews</i> . 2004a;36:583-594.
4.3-3	Anders MW. Formation and toxicity of anesthetic degradation products. <i>Annual review of pharmacology and toxicology</i> . 2004b;45:147-176.
4.3-4	Baillie TA. Approaches to Mitigate the Risk of Serious Adverse Reactions in Covalent Drug Design. <i>Expert Opinion on Drug Discovery</i> . 2020:1-44.
4.3-5	Barbacid M. Ras genes. <i>Annual Review of Biochemistry</i> . 1987;56:779-827.
4.3-6	Barnett LMA, Cummings BS. Nephrotoxicity and Renal Pathophysiology: A Contemporary Perspective. <i>Toxicological Sciences</i> . 2018;164:379-390.
4.3-7	Barrow P, Schmitt G. Regulatory Approaches to Nonclinical Reproductive Toxicity Testing of Anti-Cancer Drugs. <i>Anti-cancer agents in medicinal chemistry</i> . 2017;17:1171-1183.
4.3-8	Benjamin SA, Stephens LC, Hamilton BF, Saunders WJ, Lee AC, Saunders GM, et al. Associations between lymphocytic thyroiditis, hypothyroidism, and thyroid neoplasia in beagles. <i>Veterinary pathology</i> . 1996;33:486-494.
4.3-9	Biernacka A, Tsongalis PD, Peterson JD, de Abreu FB, Black CC, Gutmann EJ, et al. The potential utility of re-mining results of somatic mutation testing: KRAS status in lung adenocarcinoma. <i>Cancer genetics</i> . 2016;209:195-198.
4.3-10	Canon J, Rex K, Saiki AY, Mohr C, Cooke K, Bagal D, et al. The clinical KRAS(G12C) inhibitor AMG 510 drives anti-tumour immunity. <i>Nature</i> . 2019;575:217-223.
4.3-11	Chang EH, Gonda MA, Ellis RW, Scolnick EM, Lowy DR. Human genome contains four genes homologous to transforming genes of Harvey and Kirsten murine sarcoma viruses. <i>Proceedings of the National Academy of Sciences</i> . 1982;79:4848-4852.
4.3-12	Cooper AJ, Krasnikov BF, Niatetskaya ZV, Pinto JT, Callery PS, Villar MT, et al. Cysteine S-conjugate beta-lyases: important roles in the metabolism of naturally occurring sulfur and selenium-containing compounds, xenobiotics and anticancer agents. <i>Amino acids</i> . 2011;41:7-27.
4.3-13	Cooper AJL, Hanigan MH. Metabolism of Glutathione S-Conjugates: Multiple Pathways. <i>Comprehensive Toxicology</i> . 2018;363-406.
4.3-14	Cully M, Downward J. SnapShot: Ras Signaling. <i>Cell</i> . 2008;133:1292-1292.e1291.
4.3-15	Daminet S, Ferguson DC. Influence of drugs on thyroid function in dogs. <i>Journal of veterinary internal medicine</i> . 2003;17:463-472.
4.3-16	Ebert PJ, Cheung J, Yang Y, McNamara E, Hong R, Moskalenko M, et al. MAP kinase inhibition promotes T cell and anti-tumor activity in combination with PD-L1 checkpoint blockade. <i>Immunity</i> . 2016;44:609-621.
4.3-17	European Medicines Agency (EMA). Guideline on the Investigation of Drug Interactions. 2013.
4.3-18	Gainor JF, Varghese AM, Ou S-HI, Kabraji S, Awad MM, Katayama R, et al. ALK Rearrangements Are Mutually Exclusive with Mutations in EGFR or KRAS: An Analysis of 1,683 Patients with Non-Small Cell Lung Cancer. <i>Clin Cancer Res</i> . 2013;19:4273-4281.
4.3-19	Green T, Odum J, Nash JA, Foster JR. Perchloroethylene-induced rat kidney tumors: an investigation of the mechanisms involved and their relevance to humans. <i>Toxicol Appl Pharmacol</i> . 1990;103:77-89.
4.3-20	Gul Altuntas T, Kharasch ED. Biotransformation of L-cysteine S-conjugates and N-acetyl-L-cysteine S-conjugates of the sevoflurane degradation product fluoromethyl-2,2-difluoro-1-(trifluoromethyl)vinyl ether (compound A) in human kidney in vitro: interindividual variability in N-acetylation, N-deacetylation, and beta-lyase-catalyzed metabolism. <i>Drug metabolism and disposition the biological fate of chemicals</i> . 2002;30:148-154.

資料番号	著者・表題・出典
4.3-21	Hall A, Marshall CJ, Spurr NK, Weiss RA. Identification of transforming gene in two human sarcoma cell lines as a new member of the ras gene family located on chromosome 1. <i>Nature</i> . 1983;303:396-400.
4.3-22	Harvey JJ. An unidentified virus which causes the rapid production of tumours in mice. <i>Nature</i> . 1964;204:1104-1105.
4.3-23	Hobbs GA, Der CJ, Rossman KL. RAS isoforms and mutations in cancer at a glance. <i>Journal of Cell Science</i> . 2016;129:1287-1292.
4.3-24	Horvath JJ, Witmer CM, Witz G. Nephrotoxicity of the 1:1 acrolein-glutathione adduct in the rat. <i>Toxicol Appl Pharmacol</i> . 1992;117:200-207.
4.3-25	Houle CD, Finch GL, Mauthe RJ, Potter MD, Walisser JA, Gardner IB, et al. Effects of lersivirine on canine and rodent thyroid function. <i>Toxicologic pathology</i> . 2014;42:897-912.
4.3-26	ICH Harmonized Tripartite Guideline Q3A(R2): Impurities in new drug substances. 2006.
4.3-27	ICH Harmonized Tripartite Guideline Q3B(R2): Impurities in new drug products. 2006.
4.3-28	ICH Harmonized Tripartite Guideline S2(R1): Guidance on genotoxicity testing and data interpretation for pharmaceuticals intended for human use. 2011.
4.3-29	ICH Harmonized Tripartite Guideline S5(R3): Detection of reproductive and developmental toxicity for human pharmaceuticals. 2020.
4.3-30	ICH Harmonized Tripartite Guideline S9: Nonclinical evaluation for anticancer pharmaceuticals 2009.
4.3-31	ICH Harmonized Tripartite Guideline S9: Nonclinical evaluation for anticancer pharmaceuticals: Questions and Answers 2018.
4.3-32	Iyer RA, Anders MW. Cysteine conjugate beta-lyase-dependent biotransformation of the cysteine S-conjugates of the sevoflurane degradation product compound A in human, nonhuman primate, and rat kidney cytosol and mitochondria. <i>Anesthesiology</i> . 1996;85:1454-1461.
4.3-33	Janes MR, Zhang J, Li LS, Hansen R, Peters U, Guo X, et al. Targeting KRAS Mutant Cancers with a Covalent G12C-Specific Inhibitor. <i>Cell</i> . 2018;172:578-589.
4.3-34	Jannin A, Penel N, Ladsous M, Vantyghem MC, Do Cao C. Tyrosine kinase inhibitors and immune checkpoint inhibitors-induced thyroid disorders. <i>Critical reviews in oncology/hematology</i> . 2019;141:23-35.
4.3-35	Jousma E, Rizvi TA, Wu J, Janhofer D, Dombi E, Dunn RS, et al. Preclinical assessments of the MEK inhibitor PD-0325901 in a mouse model of Neurofibromatosis type 1. <i>Pediatric blood & cancer</i> . 2015;62:1709-1716.
4.3-36	Karnoub AE, Weinberg RA. Ras oncogenes: split personalities. <i>Nature Reviews Molecular Cell Biology</i> . 2008; 9:517-531.
4.3-37	Keller D, Brennan R, Leach K. Clinical and Nonclinical Adverse Effects of Kinase Inhibitors. In: <i>L Urbán, Patel VF, Vaz RJ, eds. Antitargets and Drug Safety. Wiley-VCH Verlag GmbH & Co. KGaA</i> ; 2015:365-400.
4.3-38	Kirsten WH, Mayer LA. Morphologic responses to a murine erythroblastosis virus. <i>Journal of the National Cancer Institute</i> . 1967;39:311-335.
4.3-39	Lash LH, Nelson RM, Van Dyke RA, Anders MW. Purification and characterization of human kidney cytosolic cysteine conjugate beta-lyase activity. <i>Drug metabolism and disposition</i> . 1990;18:50-54.
4.3-40	Laurberg P. Paradoxically subnormal serum T ₄ and T ₃ in dogs after prolonged excessive TSH stimulation of the thyroid caused by post-cAMP refractoriness of thyroid hormone secretion. <i>Metabolism</i> . 1989;38:265-270.
4.3-41	Liu M. Relative Response Factor of Impurities in AMG 510. Amgen Memorandum MEMO-407042. 2020.
4.3-42	Lock EA and Antoine DJ. 14.03 - Renal Xenobiotic Metabolism. In: CA McQueen, ed. <i>Comprehensive Toxicology (Third Edition)</i> . Oxford: Elsevier; 2018:30-55.
4.3-43	Maziasz T, Kadambi VJ, Silverman L, Fedyk E, Alden CL. Predictive toxicology approaches for small molecule oncology drugs. <i>Toxicologic pathology</i> . 2010;38:148-164.

資料番号	著者・表題・出典
4.3-44	Mccarthy RI, Lock EA, Hawksworth GM. Cytosolic C-S lyase activity in human kidney samples-relevance for the nephrotoxicity of halogenated alkenes in man. <i>Toxicology and industrial health</i> . 1994;10:103-112.
4.3-45	Mcclain RM. Mechanistic considerations for the relevance of animal data on thyroid neoplasia to human risk assessment. <i>Mutation research</i> . 1995;333:131-142.
4.3-46	McCormick F. K-Ras protein as a drug target. <i>J Mol Med</i> . 2016;94:253-258.
4.3-47	Mcdonald ER, 3rd, De Weck A, Schlabach MR, Billy E, Mavrakis KJ, Hoffman GR, et al. Project DRIVE: A Compendium of Cancer Dependencies and Synthetic Lethal Relationships Uncovered by Large-Scale, Deep RNAi Screening. <i>Cell</i> . 2017;170:577-592.
4.3-48	Mesich ML, Mayhew PD, Paek M, Holt DE, Brown DC. Gall bladder mucocoeles and their association with endocrinopathies in dogs: a retrospective case-control study. <i>The Journal of small animal practice</i> . 2009;50:630-635.
4.3-49	Milburn MV, Tong L, deVos AM, Brünger A, Yamaizumi Z, Nishimura S, et al. Molecular switch for signal transduction: structural differences between active and inactive forms of protooncogenic ras proteins. <i>Science</i> . 1990;247:939-945.
4.3-50	Miller RP, Tadagavadi RK, Ramesh G, Reeves WB. Mechanisms of Cisplatin nephrotoxicity. <i>Toxins</i> . 2010;2:2490-2518.
4.3-51	Miyahisa I, Sameshima T, Hixon MS. Rapid Determination of the Specificity Constant of Irreversible Inhibitors (k_{inact}/K_i) by Means of an Endpoint Competition Assay. <i>Angewandte Chemie</i> . 2015;127:14305-14308.
4.3-52	Mutlib AE, Gerson RJ, Meunier PC, Haley PJ, Chen H, Gan LS, et al. The species-dependent metabolism of efavirenz produces a nephrotoxic glutathione conjugate in rats. <i>Toxicol Appl Pharmacol</i> . 2000;169:102-113.
4.3-53	NCCN Clinical Practice Guidelines in Oncology (NCCN Guidelines [®]) Non-Small Cell Lung Cancer Version 3.2018 - February 21, 2018.
4.3-54	Neumann J, Zeindl-Eberhart E, Kirchner T, Jung A. Frequency and type of KRAS mutations in routine diagnostic analysis of metastatic colorectal cancer. <i>Pathology – Research and Practice</i> . 2009;205:858-862.
4.3-55	Nichols RJ, Haderk F, Stahlhut C, Schulze CJ, Hemmati G, Wildes D, et al. RAS nucleotide cycling underlies the SHP2 phosphatase dependence of mutant BRAF-, NF1-and RAS-driven cancers. <i>Nature cell biology</i> . 2018;20:1064-1073.
4.3-56	Nitzsche D. Effect of maternal feed restriction on prenatal development in rats and rabbits – A review of published data. <i>Regulatory Toxicology and Pharmacology</i> . 2017;90:95-103.
4.3-57	Ostrem JM, Peters U, Sos ML, Wells JA, Shokat KM. K-Ras (G12C) inhibitors allosterically control GTP affinity and effector interactions. <i>Nature</i> . 2013;503:548-551.
4.3-58	Ostrem JM, Shokat KM. Direct small-molecule inhibitors of KRAS: from structural insights to mechanism-based design. <i>Nature Reviews Drug Discovery</i> . 2016;15:771-785.
4.3-59	Pai EF, Kabsch W, Krengel U, Holmes KC, John J, Wittinghofer A. Structure of the guanine-nucleotide-binding domain of the Ha-ras oncogene product p21 in the triphosphate conformation. <i>Nature</i> . 1989;341:209-214.
4.3-60	Patricelli MP, Janes MR, Li LS, Hansen R, Peters U, Kessler LV, et al. Selective Inhibition of Oncogenic KRAS Output with Small Molecules Targeting the Inactive State. <i>Cancer Discov</i> . 2016;6:316-329.
4.3-61	Perazella MA. Pharmacology behind Common Drug Nephrotoxicities. <i>Clinical journal of the American Society of Nephrology</i> . 2018;13:1897-1908.
4.3-62	Prior IA, Lewis PD, Mattos C. A comprehensive survey of Ras mutations in cancer. <i>Cancer research</i> . 2012;72:2457-2467.
4.3-63	Richardson TA, Klaassen CD. Disruption of thyroid hormone homeostasis in Ugt1a-deficient Gunn rats by microsomal enzyme inducers is not due to enhanced thyroxine glucuronidation. <i>Toxicol Appl Pharmacol</i> . 2010;248:38-44.
4.3-64	Rood JJM, Jamalpoor A, Van Hoppe S, Van Haren MJ, Wasmann RE, Janssen MJ, et al. Extrahepatic metabolism of ibrutinib. <i>Investigational New Drugs</i> . 2021;39:1-14.

資料番号	著者・表題・出典
4.3-65	Scheffler M, Ihle MA, Hein R, Merkelbach-Bruse S, Scheel AH, Siemanowski J, et al. K-ras Mutation Subtypes in NSCLC and Associated Co-occurring Mutations in Other Oncogenic Pathways. <i>Journal of Thoracic Oncol</i> . 2019;14:606-616.
4.3-66	Scheffzek K, Ahmadian MR, Kabsch W, Wiesmüller L, Lautwein A, Schmitz F, et al. The Ras-RasGAP complex: structural basis for GTPase activation and its loss in oncogenic Ras mutants. <i>Science (New York, NY)</i> . 1997;277:333-338.
4.3-67	Simanshu DK, Nissley DV, McCormick F. RAS proteins and their regulators in human disease. <i>Cell</i> . 2017;170:17-33.
4.3-68	Taparowsky E, Shimizu K, Goldfarb M, Wigler M. Structure and activation of the human N-ras gene. <i>Cell</i> . 1983;34:581-586.
4.3-69	Testa B, Clement B. Chapter 24 - Biotransformation Reactions and their Enzymes. <i>The Practice of Medicinal Chemistry (Fourth Edition)</i> . Academic Press; 2015:561-584.
4.3-70	Townsend DM, Hanigan MH. Inhibition of gamma-glutamyl transpeptidase or cysteine S-conjugate beta-lyase activity blocks the nephrotoxicity of cisplatin in mice. <i>The Journal of pharmacology and experimental therapeutics</i> . 2002;300:142-148.
4.3-71	Tsuchida S. Glutathione Transferases. <i>Encyclopedia of Cancer (Second Edition)</i> . Academic Press; 2002;297-307.
4.3-72	United States Food and Drug Administration (FDA). In Vitro Drug Interaction Studies – Cytochrome P450 Enzyme- and Transporter-Mediated Drug Interactions Guidance for Industry. 2020.
4.3-73	Vernazza P, Wang C, Pozniak A, Weil E, Pulik P, Cooper DA, et al. Efficacy and safety of lersivirine (UK-453,061) versus efavirenz in antiretroviral treatment-naïve HIV-1-infected patients: week 48 primary analysis results from an ongoing, multicenter, randomized, double-blind, phase IIb trial. <i>Journal of acquired immune deficiency syndromes</i> . 2013;62:171-179.
4.3-74	Vetter IR, Wittinghofer A. The guanine nucleotide-binding switch in three dimensions. <i>Science</i> . 2001;294:1299-1304.
4.3-75	Zabka TS, Fielden MR, Garrido R, Tao J, Fretland AJ, Fretland JL, et al. Characterization of xenobiotic-induced hepatocellular enzyme induction in rats: anticipated thyroid effects and unique pituitary gland findings. <i>Toxicologic pathology</i> . 2011;39:664-677.
4.3-76	Zhang L, Hanigan MH. Role of cysteine S-conjugate beta-lyase in the metabolism of cisplatin. <i>The Journal of pharmacology and experimental therapeutics</i> . 2003;306:988-994.
4.3-77	Addeo A, Banna GL and Friedlaender A. KRAS G12C Mutations in NSCLC: From Target to Resistance. <i>Cancers</i> . 2021;13:2541.
4.3-78	Li S, Liu S, Deng J, Akbay EA, Hai J, Ambrogio C, et al. Assessing Therapeutic Efficacy of MEK Inhibition in a KRASG12C-Driven Mouse Model of Lung Cancer. <i>Clin Cancer Res</i> . 2018;24:4854-4864.

資料番号	著者・表題・出典
4.3-79	Muñoz-Maldonado C, Zimmer Y and Medová M. A Comparative Analysis of Individual RAS Mutations in Cancer Biology. <i>Front Oncol.</i> 2019;9: 1088.
4.3-80	Newlaczyl AU, Coulson JM and Prior IA. Quantification of spatiotemporal patterns of Ras isoform expression during development. <i>Sci Rep.</i> 2017;7:41297.
4.3-81	Román M, Baraibar I, López I, Nadal E, Rolfo C, Vicen S, et al. KRAS oncogene in non-small cell lung cancer: clinical perspectives on the treatment of an old target. <i>Mol Cancer.</i> 2018; 17: 33.
4.3-82	Temko D, Tomlinson IPM, Severini S, Schuster-Böckler B and Graham BA. The effects of mutational processes and selection on driver mutations across cancer types. <i>Nature Com.</i> 2018;9:1857.
4.3-83	Thein K, Banks K, Saam J, Raymond VM, Roszik J, Meric-Bernstam F, et al. The prevalence of KRAS ^{G12C} mutations utilizing circulating tumor DNA (ctDNA) in 80,911 patients with cancer. <i>J Clin Oncol.</i> 2020;38: Issue15 suppl. 3547.

第5部 臨床試験報告書

5.2 臨床試験一覧表

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.2	Tabular Listing of All Clinical Studies	Amgen	—	—	—	—

5.3 試験報告書及び関連情報

5.3.1 生物薬剤学試験報告書

5.3.1.1 バイオアベイラビリティ(BA)試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.1.1-1	Study 20190316 - An Open-label, Randomized, 2-period, 2-treatment Crossover Study to Assess the Effect of Food on the Pharmacokinetics of AMG 510 in Healthy Subjects	Amgen	20■■■■—20■■■	海外	参考	有

5.3.1.2 比較BA試験及び生物学的同等性(BE)試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.1.2-1	Study 20190500 - An Open-label, Randomized, 2-Period, 2-Treatment Crossover Study to Evaluate the Relative Bioavailability of AMG 510 Administered as Tablets and as a Water Dispersion in Healthy Subjects	Amgen	20■■■■—20■■■	海外	参考	有

5.3.1.4 生物学的及び理化学的分析法検討報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.1.4-1	Study 150636 - Determination of AMG 510 in K ₂ EDTA Human Plasma Using LC-MS/MS Following Protein Precipitation	Amgen	20■■■■—20■■■	海外	参考	無
5.3.1.4-2	Study 151842 Amendment 2 - Determination of AMG 510 in K ₂ EDTA Human Plasma Using LC-MS/MS Following Protein Precipitation (Revised Range)	Amgen	20■■■■—20■■■	海外	参考	無
5.3.1.4-3	Study 153277 - Determination of AMG 510 in Human Urine Using LC-MS/MS Following Sample Dilution	Amgen	20■■■■—20■■■	海外	参考	無
5.3.1.4-4.1	Study 150840 - Quantitation of AMG 510 in Human Plasma via HPLC with MS/MS Detection	Amgen	20■■■	海外	参考	無
5.3.1.4-4.2	Study 150840 Addendum 1 - Quantitation of AMG 510 in Human Plasma via HPLC with MS/MS Detection	Amgen	20■■■	海外	参考	無
5.3.1.4-4.3	Study 150840 Addendum 2 - Quantitation of AMG 510 in Human Plasma via HPLC with MS/MS Detection	Amgen	20■■■	海外	参考	無

5.3.1.4-4.4	Study 150840 Addendum 3 - Quantitation of AMG 510 in Human Plasma via HPLC with MS/MS Detection	Amgen	20■■■	海外	参考	無
5.3.1.4-4.5	Study 150840 Addendum 4 - Quantitation of AMG 510 in Human Plasma via HPLC with MS/MS Detection	Amgen	20■■■	海外	参考	無
5.3.1.4-5	Study 150840 Digoxin - Quantitation of Digoxin in Human Plasma via HPLC with MS/MS Detection	Amgen	20■■■	海外	参考	無
5.3.1.4-6	Study 150840 - Analytical Method Validation Report for Metformin in Human EDTA K ₃ Plasma	Amgen	20■■■	海外	参考	無
5.3.1.4-7	Study 150840 - Validation of a High Performance Liquid Chromatographic Method Using Tandem Mass Spectrometry Detection for the Determination of Metformin (10 to 2500 ng/mL) in Human Urine	Amgen	20■■■	海外	参考	無
5.3.1.4-8	Study 150840 - Quantitation of Midazolam and 1-Hydroxymidazolam in Human Plasma via HPLC with MS/MS Detection	Amgen	20■■■	海外	参考	無
5.3.1.4-9	Study 150840 Itraconazole - Quantitation of Itraconazole and 2-Hydroxyitraconazole in Human Plasma via HPLC with MS/MS Detection	Amgen	20■■■	海外	参考	無
5.3.1.4-10.1	Study 153140 - Quantitation of AMG 510 in Human Urine via LC-MS/MS	Amgen	20■■■	海外	参考	無
5.3.1.4-10.2	Study 153140 Addendum 1 - Quantitation of AMG 510 in Human Urine via LC-MS/MS	Amgen	20■■■	海外	参考	無
5.3.1.4-11.1	Study 153141 - Quantitation of AMG 510 Metabolites M10 (AMG3375854), M18 (AMG3413829), and M24 (AMG3368167) in Human Plasma via LC-MS/MS	Amgen	20■■■	海外	参考	無
5.3.1.4-11.2	Study 153141 Addendum 1 - Quantitation of AMG 510 Metabolites M10 (AMG3375854), M18 (AMG3413829), and M24 (AMG3368167) in Human Plasma via LC-MS/MS	Amgen	20■■■	海外	参考	無
5.3.1.4-11.3	Study 153141 Addendum 2 - Quantitation of AMG 510 Metabolites M10 (AMG3375854), M18 (AMG3413829), and M24 (AMG3368167) in Human Plasma via LC-MS/MS	Amgen	20■■■	海外	参考	無
5.3.1.4-12	bacsr-20190317 - Bioanalytical Report - Study 20190317 - A Phase I, Open-Label Study to Evaluate the Drug-Drug Interactions Between Metformin and AMG 510 in Healthy Subjects	Amgen	20■■■	海外	参考	無
5.3.1.4-13	bacsr-20190318 - Bioanalytical Report - Study 20190318 - Quantitation of Itraconazole in Human Plasma via HPLC with MS/MS Detection	Amgen	20■■■	海外	参考	無

5.3.2 ヒト生体試料を用いた薬物動態関連の試験報告書

5.3.2.1 血漿蛋白結合試験報告書

資料番号	資料表題	著者	試験実施期間 (年/月/日)	試験実施場所 (国内/海外)	評価・参考	申請電子データ 提出の有無
5.3.2.1-1	Study 153486 - Human In Vitro Plasma Protein Binding of AMG3375854 and AMG3413829	Amgen	20■■■	海外	参考	無

5.3.2.2 肝代謝及び薬物相互作用試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.2.2-1	Study 150537 - Identification of the Major Enzymes Involved in the In Vitro Metabolism of AMG 510	Amgen	20■■■	海外	参考	無
5.3.2.2-2	Study 154076 - In Vitro Characterization of the Glutathione Conjugation of AMG 510	Amgen	20■■■	海外	参考	無
5.3.2.2-3	Study 150535 - Evaluation of the Inhibitory Potential of AMG 510 and AMG 3368167 Against Eight Principal Human Cytochrome P450 Isozymes Using Human Hepatic Microsomes	Amgen	20■■■	海外	参考	無
5.3.2.2-4	Study 153423 - Determination of the Inhibition of CYP1A2, CYP2B6, CYP2C8, CYP2C9, CYP2C19, CYP2D6, and CYP3A by AMG 3375854 and AMG 3413829 in Human Liver Microsomes	Amgen	20■■■	海外	参考	無
5.3.2.2-5	Study 150536 - Evaluation of AMG 510 and AMG3368167 as Inducers of CYP1A2, CYP2B6, CYP2C8, CYP2C9, CYP2C19, and CYP3A4 in Primary Human Hepatocytes	Amgen	20■■■	海外	参考	無
5.3.2.2-6	Study 153424 - An In Vitro Evaluation of the Effect of AMG3375854 and AMG3413829 on mRNA and Enzymatic Activity Levels of Cytochrome P450 Genes in Cultured Human Hepatocytes	Amgen	20■■■	海外	参考	無
5.3.2.2-7	Study 153303 - Peak Area Ratios for AMG 510 and Identified Human Metabolites in Matrix-normalized Rat, Dog, and Human Plasma Samples	Amgen	20■■■	海外	参考	無
5.3.2.2-8	Study 150539 - Assessment of AMG 510 and its Metabolite AMG3368167 as Inhibitors of Human BCRP, P-gp, MATE1, MATE2-K, OAT1, OAT3, OCT1, OCT2, OATP1B1, and OATP1B3 Mediated Transport	Amgen	20■■■-20■■■	海外	参考	無
5.3.2.2-9	Study 153425 - Assessment of AMG 3375854 and AMG 3413829 as inhibitors of human OAT1, OAT3, OCT1, OCT2, OATP1B1, OATP1B3, MATE1, MATE2-K, BCRP and P-gp mediated transport	Amgen	20■■■-20■■■	海外	参考	無

5.3.2.3 他のヒト生体試料を用いた試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.2.3-1	Study 150540 - Assessment of AMG 510 and its Metabolite AMG3368167 as a Substrate of Human BCRP and P-gp Mediated Transport	Amgen	20■■■-20■■■	海外	参考	無
5.3.2.3-2	Study 154075 - Nonspecific Binding of AMG 510 and Three of Its Metabolites to Human Liver Microsomes	Amgen	20■■■	海外	参考	無

5.3.3 臨床薬物動態（PK）試験報告書

5.3.3.1 健康被験者におけるPK及び初期忍容性試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.3.1-1.1	Study 20190321 - A Phase I, Open-label Study of the Absorption, Metabolism, and Excretion of [¹⁴ C]-AMG 510 Following a Single Oral Dose in Healthy Male Subjects	Amgen	20██/██/██－20██/██/██	海外	参考	無
5.3.3.1-1.2	Study 20190321 addendum - A Phase I, Open-label Study of the Absorption, Metabolism, and Excretion of [¹⁴ C]-AMG 510 Following a Single Oral Dose in Healthy Male Subjects	Amgen	20██/██/██－20██/██/██	海外	参考	無

5.3.3.4 外因性要因を検討したPK試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.3.4-1	Study 20190315 - An Open-label, Drug-drug Interaction Study to Evaluate the Effect of AMG 510 on the Pharmacokinetics of Digoxin, a P-glycoprotein Substrate, in Healthy Subjects	Amgen	20██/██/██－20██/██/██	海外	参考	有
5.3.3.4-2.1	Study 20190317 - A Phase I, Open-label Study to Evaluate the Drug-drug Interactions Between Metformin and AMG 510 in Healthy Subjects	Amgen	20██/██/██－20██/██/██	海外	参考	有
5.3.3.4-2.2	Study 20190317 errata - A Phase I, Open-label Study to Evaluate the Drug-drug Interactions Between Metformin and AMG 510 in Healthy Subjects	Amgen	20██/██/██－20██/██/██	海外	参考	無
5.3.3.4-3	Study 20190318 - An Open-label Study to Evaluate the Drug-drug Interaction Effect of Itraconazole, a CYP3A4 Inhibitor, on the Pharmacokinetics of AMG 510 in Healthy Subjects	Amgen	20██/██/██－20██/██/██	海外	参考	有
5.3.3.4-4	Study 20190319 - An Open-label Study to Evaluate the Drug-drug Interaction Effect of Rifampin, a CYP3A4 Inducer, on the Pharmacokinetics of AMG 510 in Healthy Subjects	Amgen	20██/██/██－20██/██/██	海外	評価	有
5.3.3.4-5	Study 20190320 - An Open-label Study to Evaluate the Drug-drug Interaction Effect of Omeprazole, a Proton-pump Inhibitor, on the Pharmacokinetics of AMG 510 in Healthy Subjects	Amgen	20██/██/██－20██/██/██	海外	評価	有
5.3.3.4-6.1	Study 20200199 - An Open-label Study to Evaluate the Impact of Acid-reducing Agents on the Pharmacokinetics of AMG 510 in Healthy Subjects under Fed Conditions	Amgen	20██/██/██－20██/██/██	海外	評価	有
5.3.3.4-6.2	Study 20200199 errata - An Open-label Study to Evaluate the Impact of Acid-reducing Agents on the Pharmacokinetics of AMG 510 in Healthy Subjects under Fed Conditions	Amgen	20██/██/██－20██/██/██	海外	評価	無
5.3.3.4-7	Study 153468 - Assessment of Clinical Drug-Drug Interaction of AMG 510 Using Physiologically Based Pharmacokinetic Modeling	Amgen	20██/██/██	－	参考	有

5.3.3.5 ポピュレーションPK試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.3.5-1	Study 152921 - Population Pharmacokinetic Evaluation of AMG 510 Monotherapy in Healthy and Advanced Solid Tumor Subjects With a Specific KRAS Mutation From Phase 1 and Phase 2 Studies	Amgen	20██/██/██	－	参考	有

5.3.4 臨床薬力学（PD）試験報告書

5.3.4.2 患者におけるPD試験及びPK/PD試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.4.2-1	Study 152922 - Exposure-Response of Sotorasib in Subjects With Advanced Solid Tumors With a Specific KRAS Mutation	Amgen	20■■■■	-	参考	有

5.3.5 有効性及び安全性試験報告書

5.3.5.1 申請する適応症に関する比較対照試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.5.1-1	Ongoing Study Safety Report: Study 20190009 - A Phase 3 Multicenter, Randomized, Open Label, Active-controlled, Study of AMG 510 Versus Docetaxel for the Treatment of Previously Treated Locally Advanced and Unresectable or Metastatic NSCLC Subjects With Mutated <i>KRAS p.G12C</i>	Amgen	2020/6/4－実施中 Cutoff: 2020/9/1	国内／海外	参考	無

5.3.5.2 非対照試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.5.2-1.1	Study 20170543 Phase-1 Portion interim analysis - A Phase 1/2, Open-label Study Evaluating the Safety, Tolerability, Pharmacokinetics, Pharmacodynamics, and Efficacy of AMG 510 Monotherapy in Subjects With Advanced Solid Tumors With <i>KRAS p.G12C</i> Mutation and AMG 510 Combination Therapy in Subjects With Advanced NSCLC With <i>KRAS p.G12C</i> Mutation (CodeBreak 100)	Amgen	2018/8/27－実施中 Cutoff: 2020/7/6	国内／海外	評価	有
5.3.5.2-1.2	Study 20170543 errata - Phase-1 Portion interim analysis - A Phase 1/2, Open-label Study Evaluating the Safety, Tolerability, Pharmacokinetics, Pharmacodynamics, and Efficacy of AMG 510 Monotherapy in Subjects With Advanced Solid Tumors With <i>KRAS p.G12C</i> Mutation and AMG 510 Combination Therapy in Subjects With Advanced NSCLC With <i>KRAS p.G12C</i> Mutation (CodeBreak 100)	Amgen	2018/8/27－実施中 Cutoff: 2020/7/6	国内／海外	評価	無
5.3.5.2-1.3	Study 20170543 errata2 - Phase-1 Portion interim analysis - A Phase 1/2, Open-label Study Evaluating the Safety, Tolerability, Pharmacokinetics, Pharmacodynamics, and Efficacy of AMG 510 Monotherapy in Subjects With Advanced Solid Tumors With <i>KRAS p.G12C</i> Mutation and AMG 510 Combination Therapy in Subjects With Advanced NSCLC With <i>KRAS p.G12C</i> Mutation (CodeBreak 100)	Amgen	2018/8/27－実施中 Cutoff: 2020/7/6	国内／海外	評価	無
5.3.5.2-2.1	Study 20170543 Phase-2 Portion primary analysis- A Phase 1/2, Open-label Study Evaluating the Safety, Tolerability, Pharmacokinetics, Pharmacodynamics, and Efficacy of AMG 510 Monotherapy in Subjects With Advanced Solid Tumors With <i>KRAS p.G12C</i> Mutation and AMG 510 Combination Therapy in Subjects With Advanced NSCLC With <i>KRAS p.G12C</i> Mutation (CodeBreak 100)	Amgen	2019/8■■－実施中 Cutoff: 2020/9/1	国内／海外	評価	有

5.3.5.2-2.2	Study 20170543 errata - Phase-2 Portion primary analysis- A Phase 1/2, Open-label Study Evaluating the Safety, Tolerability, Pharmacokinetics, Pharmacodynamics, and Efficacy of AMG 510 Monotherapy in Subjects With Advanced Solid Tumors With <i>KRAS p.G12C</i> Mutation and AMG 510 Combination Therapy in Subjects With Advanced NSCLC With <i>KRAS p.G12C</i> Mutation (CodeBreak 100)	Amgen	2019/8 ■■■ー実施中 Cutoff: 2020/9/1	国内／海外	評価	無
5.3.5.2-3	Study 20170543 PRO Data - A Phase 1/2, Open-label Study Evaluating the Safety, Tolerability, Pharmacokinetics, Pharmacodynamics, and Efficacy of AMG 510 Monotherapy in Subjects With Advanced Solid Tumors With <i>KRAS p.G12C</i> Mutation and AMG 510 Combination Therapy in Subjects With Advanced NSCLC With <i>KRAS p.G12C</i> Mutation (CodeBreak 100)	Amgen	2019/8 ■■■ー実施中 Cutoff: 2020/9/1	国内／海外	評価	無
5.3.5.2-4	Study 20170543 Phase-2 Portion supplemental - A Phase 1/2, Open-label Study Evaluating the Safety, Tolerability, Pharmacokinetics, Pharmacodynamics, and Efficacy of AMG 510 Monotherapy in Subjects With Advanced Solid Tumors With <i>KRAS p.G12C</i> Mutation and AMG 510 Combination Therapy in Subjects With Advanced NSCLC With <i>KRAS p.G12C</i> Mutation (CodeBreak 100)	Amgen	2019/8 ■■■ー実施中 Cutoff: 2020/12/1	国内／海外	評価	無

5.3.5.3 複数の試験成績を併せて解析した報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.5.3-1	Integrated Summary of Safety ISS TFLs Supplemental Statistical Analysis Plan	Amgen	—	—	—	無
5.3.5.3-2	Japan NDA analysis JNDA TFLs Supplemental Statistical Analysis Plan	Amgen	—	—	—	無

5.3.5.4 その他の臨床試験報告書

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.5.4-1	Observational Research Study 20180277 - A Retrospective Cohort Study of Patient Characteristics and Treatment Outcomes Among Patients With <i>KRAS p.G12C</i> Mutation-positive Advanced Non-small Cell Lung Cancer in the AACR Project GENIE Database	Amgen	20■■■■ (Date of report)	海外	参考	無
5.3.5.4-2	Observational Research Study 20190037 - Descriptive Analysis of Patients With <i>KRAS G12C</i> Mutant Solid Tumors in the Syapse Database	Amgen	20■■■■ (Date of report)	海外	参考	無
5.3.5.4-3	Observational Research Study 20200097 - A Retrospective Cohort Study of Patient Characteristics and Treatment Outcomes Among Patients With <i>KRAS p.G12C</i> Mutation-positive Advanced Non-small Cell Lung Cancer in the Flatiron Health-Foundation Medicine Clinico-Genomics Database	Amgen	20■■■■ (Date of report)	海外	参考	無
5.3.5.4-4	Observational Research Study 20200132 - A Retrospective Cohort Study of Patient Characteristics and Treatment Outcomes Among Patients With Advanced Non-small Cell Lung Cancer in the Flatiron Health-Foundation Medicine Clinico-Genomics Database	Amgen	20■■■■ (Date of report)	海外	参考	無

5.3.5.4-5	Observational Research Study 20190344 - <i>KRAS</i> ^{G12C} Mutation in Non-Small Cell Lung Cancer (NSCLC): Epidemiology and Clinical Outcomes: Systematic Literature Review	Amgen	20■■■■ (Date of report)	海外	参考	無
5.3.5.4-6	Observational Research Study 20200090 - Qualitative Interviews in Patients With <i>KRAS</i> ^{p.G12C} -Mutated Non-Small Cell Lung Cancer	Amgen	20■■■■ (Date of report)	海外	参考	無
5.3.5.4-7.1	Ongoing Study Safety Report: Study 20190135 - A Phase 1b Study Evaluating the Safety, Tolerability, Pharmacokinetics, and Efficacy of AMG 510 (pINN Sotorasib) in Combination With Trametinib in Subjects With Advanced Solid Tumors With <i>KRAS</i> ^{p.G12C} Mutation (CodeBreak 101 Subprotocol A)	Amgen	2019/12■■■ - 実施中 Cutoff : 2020/9/1	海外	参考	無
5.3.5.4-7.2	Ongoing Study Safety Report: Study 20190135 - A Phase 1b Study Evaluating the Safety, Tolerability, Pharmacokinetics, and Efficacy of AMG 510 (pINN Sotorasib) in Combination With RMC-4630 in Subjects With Advanced Solid Tumors With <i>KRAS</i> ^{p.G12C} Mutation (CodeBreak 101 Subprotocol C)	Amgen	20■■■■ - 実施中 Cutoff : 2020/9/1	海外	参考	無
5.3.5.4-7.3	Ongoing Study Safety Report: Study 20190135 - A Phase 1b Study Evaluating the Safety, Tolerability, Pharmacokinetics, and Efficacy of AMG 510 in Combination With Afatinib in Subjects With Advanced Non-small Cell Lung Cancer (NSCLC) With <i>KRAS</i> ^{p.G12C} Mutation (CodeBreak 101 Subprotocol D)	Amgen	20■■■■ - 実施中 Cutoff : 2020/9/1	海外	参考	無
5.3.5.4-7.4	Ongoing Study Safety Report: Study 20190135 - A Phase 1b Study Evaluating the Safety, Tolerability, Pharmacokinetics, and Efficacy of AMG 510 in Combination With Atezolizumab in Subjects With Advanced Non-small Cell Lung Cancer (NSCLC) With <i>KRAS</i> ^{p.G12C} Mutation (CodeBreak 101 Subprotocol E)	Amgen	20■■■■ - 実施中 Cutoff : 2020/9/1	海外	参考	無
5.3.5.4-7.5	Ongoing Study Safety Report: Study 20190135 - A Phase 1b Study Evaluating the Safety, Tolerability, and Efficacy of AMG 510 in Combination With Panitumumab and in Combination with Panitumumab and FOLFIRI in Subjects With Advanced Solid Tumors With <i>KRAS</i> ^{p.G12C} Mutation (CodeBreak 101 Subprotocol H)	Amgen	20■■■■ - 実施中 Cutoff : 2020/9/1	国内／海外	参考	無
5.3.5.4-8	Ongoing Study Safety Report: Study 20190147 - A Phase 1, Open-label Study Evaluating the Safety, Tolerability, Pharmacokinetics, and Efficacy of AMG 510 in Subjects of Chinese Descent With Advanced/Metastatic Solid Tumors With <i>KRAS</i> ^{p.G12C} Mutation (CodeBreak105)	Amgen	2020/4/28 - 実施中 Cutoff : 2020/9/1	海外	参考	無
5.3.5.4-9	Observational Research Study 20190325 - Patient Characteristics and Clinical Outcomes by the Presence of and Variants of <i>KRAS</i> Mutation in Japanese Advanced Non-small Cell Lung Cancer (NSCLC) Patients: LC-SCRUM-Japan	Amgen	20■■■■ (Date of report)	国内	参考	無

5.3.7 患者データ一覧表及び症例記録

資料番号	資料表題	著者	試験実施期間 (年／月／日)	試験実施場所 (国内／海外)	評価・参考	申請電子データ 提出の有無
5.3.7	患者データ一覧表及び症例記録 JNDA Listings	Amgen	—	—	—	無

第5部 臨床試験報告書

5.4 参考文献

資料番号	著者・表題・出典
5.4-1	American Association for Cancer Research (AACR) Project GENIE Consortium. AACR Project GENIE: Powering precision medicine through an international consortium. <i>Cancer Discov.</i> 2017;7(8):818-831.
5.4-2	Arbour KC, Rizvi H, Plodkowski AJ, et al. Clinical characteristics and anti-PD-(L)1 treatment outcomes of KRAS-G12C mutant lung cancer compared to other molecular subtypes of KRAS-mutant lung cancer. <i>J Clin Oncol.</i> 2020;38(15 Suppl):9596.
5.4-3	Barbacid M. <i>ras</i> GENES. <i>Annu Rev Biochem.</i> 1987;56:779-827.
5.4-4	Barlesi F, Mazieres J, Merlio JP, et al. Routine molecular profiling of patients with advanced non-small-cell lung cancer: results of a 1-year nationwide programme of the French Cooperative Thoracic Intergroup (IFCT). <i>Lancet.</i> 2016;387:1415-1426.
5.4-5	Bezjak A, Tu D, Seymour L, et al. Symptom improvement in lung cancer patients treated with erlotinib: quality of life analysis of the National Cancer Institute of Canada Clinical Trials Group Study BR.21. <i>J Clin Oncol.</i> 2006;24(24):3831-3837.
5.4-6	Biernacka A, Tsongalis PD, Peterson JD, et al. The potential utility of re-mining results of somatic mutation testing: <i>KRAS</i> status in lung adenocarcinoma. <i>Cancer Genet.</i> 2016;209:195-198.
5.4-7	Blumenthal GM, Karuri SW, Zhang H, et al. Overall response rate, progression-free survival, and overall survival with targeted and standard therapies in advanced non-small-cell lung cancer: US Food and Drug Administration trial-level and patient-level analyses. <i>J Clin Oncol.</i> 2015;33(9):1008-1014.
5.4-8	Borghaei H, Paz-Ares L, Horn L, et al. Nivolumab versus docetaxel in advanced nonsquamous non-small-cell lung cancer. <i>N Engl J Med.</i> 2015;373(17):1627-1639.
5.4-9	Brahmer J, Reckamp KL, Baas P, et al. Nivolumab versus docetaxel in advanced squamous-cell non-small-cell lung cancer. <i>N Engl J Med.</i> 2015;373(2):123-135.
5.4-10	Brunelli L, Caiola E, Marabese M, Broggin M and Pastorelli R. Capturing the metabolomic diversity of KRAS mutants in non-small-cell lung cancer cells. <i>Oncotarget.</i> 2014;5:4722-4731.
5.4-11	Burns TF, Borghaei H, Ramalingam SS, Mok TS and Peters S. Targeting <i>KRAS</i> -mutant non-small-cell lung cancer: one mutation at a time, with a focus on <i>KRAS G12C</i> mutations. <i>J Clin Oncol.</i> 2020;38:4208-4218.
5.4-12	Canon J, Rex K, Saiki AY, et al. The clinical KRAS(G12C) inhibitor AMG 510 drives anti-tumour immunity. <i>Nature.</i> 2019;575:217-223.
5.4-13	Clarke JM, Wang X, Ready NE. Surrogate clinical endpoints to predict overall survival in non-small cell lung cancer trials - are we in a new era? <i>Transl Lung Cancer Res.</i> 2015;Dec;4(6):804-808.
5.4-14	Clopper CJ and Pearson ES. The use of confidence or fiducial limits illustrated in the case of the binomial. <i>Biometrika.</i> 1934;26(4):404-413.
5.4-15	Cocks K, King MT, Velikova G, de Castro Jr G, Martyn St-James M, Fayers PM, Brown JM. Evidence-based guidelines for interpreting change scores for the European Organisation for the Research and Treatment of Cancer Quality of Life Questionnaire Core 30. <i>Eur J Cancer.</i> 2012;48:1713-1721.
5.4-16	Cooley ME, Short TH, Moriarty HJ. Symptom prevalence, distress, and change over time in adults receiving treatment for lung cancer. <i>Psychooncology.</i> 2003;12:694-708.
5.4-17	Couey MA, Bell RB, Patel AA, et al. Delayed immune-related events (DIRE) after discontinuation of immunotherapy: diagnostic hazard of autoimmunity at a distance. <i>J Immunother Cancer.</i> 2019;7:165.
5.4-18	Cui W, Franchini F, Alexander M, et al. Assessing the significance of KRAS G12C mutation: Clinicopathologic features, treatments, and survival outcomes in a real-world KRAS mutant non-small cell lung cancer cohort. <i>J Clin Oncol.</i> 2020a;38(15 Suppl):e19324.
5.4-19	Cui W, Franchini F, Alexander M, et al. Real world outcomes in KRAS G12C mutation positive non-small cell lung cancer. <i>Lung Cancer.</i> 2020b;146:310-317.
5.4-20	Cully M and Downward J. SnapShot: Ras Signaling. <i>Cell.</i> 2008;133:1292.
5.4-21	Drilon A, Laetsch TW, Kummar S, et al. Efficacy of larotrectinib in <i>TRK</i> fusion-positive cancers in adults and children. <i>N Engl J Med.</i> 2018;378(8):731-739.

資料番号	著者・表題・出典
5.4-22	Eisenhauer EA, Therasse P, Bogaerts J, et al. New response evaluation criteria in solid tumours: revised RECIST guideline (version 1.1). <i>Eur J Cancer</i> . 2009;45:228-247.
5.4-23	Ettinger DS, Wood DE, Aisner DL, et al. NCCN Guidelines Insights: Non–small cell lung cancer, Version 5. 2017. <i>J Natl Compr Canc Netw</i> . 2017;15(4):504-535.
5.4-24	Ettinger DS, Wood DE, Aggarwal C, et al. NCCN Guidelines Insights: Non–small Cell Lung Cancer, Version 1. 2020. <i>J Natl Compr Canc Netw</i> . 2019;17(12):1464-1472.
5.4-25	Fernández-Medarde A and, Santos E. Ras in cancer and developmental diseases. <i>Genes Cancer</i> . 2011;2(3):344-358.
5.4-26	Fossella FV, DeVore R, Kerr RN, et al; TAX 320 Non-Small Cell Lung Cancer Study Group. Randomized phase III trial of docetaxel versus vinorelbine or ifosfamide in patients with advanced non–small-cell lung cancer previously treated with platinum-containing chemotherapy regimens. <i>J Clin Oncol</i> . 2000;18(12):2354-2362.
5.4-27	Fossella FV, Lynch T, Shepherd FA. Second line chemotherapy for NSCLC: establishing a gold standard. <i>Lung Cancer</i> . 2002;38:s5-s12.
5.4-28	Gainor JF, Varghese AM, Ou SH, et al. <i>ALK</i> rearrangements are mutually exclusive with mutations in <i>EGFR</i> or <i>KRAS</i> : an analysis of 1,683 patients with non–small cell lung cancer. <i>Clin Cancer Res</i> . 2013;19(15):4273-4281.
5.4-29	Garon EB, Ciuleanu TE, Arrieta O, et al. Ramucirumab plus docetaxel versus placebo plus docetaxel for second-line treatment of stage IV non-small-cell lung cancer after disease progression on platinum-based therapy (REVEL): a multicentre, double-blind, randomised phase 3 trial. <i>Lancet</i> . 2014;384:665-673.
5.4-30	Garrido P, Olmedo ME, Gómez A, et al. Treating <i>KRAS</i> -mutant NSCLC: latest evidence and clinical consequences. <i>Ther Adv Med Oncol</i> . 2017;9(9):589-597.
5.4-31	Gridelli C, de Castro Carpeno J, Dingemans AM, et al. Safety and efficacy of bevacizumab plus standard-of-care treatment beyond disease progression in patients with advanced non-small cell lung cancer: The AvaALL randomized clinical trial. <i>JAMA Oncol</i> . 2018;4(12):e183486.
5.4-32	Hames ML, Chen H, Jams W, et al. Correlation between <i>KRAS</i> mutation status and response to chemotherapy in patients with advanced non-small cell lung cancer. <i>Lung Cancer</i> . 2016;92:29-34.
5.4-33	Hanna N, Shepherd FA, Fossella FV, et al. Randomized phase III trial of pemetrexed versus docetaxel in patients with non–small-cell lung cancer previously treated with chemotherapy. <i>J Clin Oncol</i> . 2004;22(9):1589-1597.
5.4-34	Herbst RS, O'Neill VJ, Fehrenbacher L, et al. Phase II study of efficacy and safety of bevacizumab in combination with chemotherapy or erlotinib compared with chemotherapy alone for treatment of recurrent or refractory non–small-cell lung cancer. <i>J Clin Oncol</i> . 2007;25(30):4743-4750.
5.4-35	Herbst RS, Baas P, Kin DW, et al. Pembrolizumab versus docetaxel for previously treated, PD-L1-positive, advanced non-small-cell lung cancer (KEYNOTE-010): a randomised controlled trial. <i>Lancet</i> . 2016;387:1540-1550.
5.4-36	Hierro C, Matos I, Martin-Liberal J, et al. Agnostic-histology approval of new drugs in oncology: Are we already there? <i>Clin Cancer Res</i> . 2019;25(11):3210-3219.
5.4-37	Janes MR, Zhang J, Li LS, et al. Targeting <i>KRAS</i> mutant cancers with a covalent G12C-specific inhibitor. <i>Cell</i> . 2018;172:578-589.
5.4-38	Johnson ML, Sima CS, Chaff J, et al. Association of <i>KRAS</i> and <i>EGFR</i> mutations with survival in patients with advanced lung adenocarcinomas. <i>Cancer</i> . 2013;119:356-362.
5.4-39	Jones RP, Sutton PA, Evans JP, et al. Specific mutations in <i>KRAS</i> codon 12 are associated with worse overall survival in patients with advanced and recurrent colorectal cancer. <i>Br J Cancer</i> . 2017;116:923-929.
5.4-40	Liao YC, Liao WY, Shun SC, YU CJ, Yang PC, Lai YH. Symptoms, psychological distress, and supportive care needs in lung cancer patients. <i>Support Care Cancer</i> . 2011;19:1743-1751.
5.4-41	Liu SY, Sun H, Zhou JY, et al. Clinical characteristics and prognostic value of the <i>KRAS G12C</i> mutation in Chinese non-small cell lung cancer patients. <i>Biomark Res</i> . 2020;8:22.
5.4-42	Maringwa JT, Quinten C, King M, et al; EORTC PROBE project and the Lung Cancer Group. Minimal important differences for interpreting health-related quality of life scores from the EORTC QLQ-C30 in lung cancer patients participating in randomized controlled trials. <i>Support Care Cancer</i> . 2011;19:1753-1760.

資料番号	著者・表題・出典
5.4-43	Martorell PM, Huerta M, Compañ Quilis A, et al. Coexistence of <i>EGFR</i> , <i>KRAS</i> , <i>BRAF</i> , and <i>PIK3CA</i> mutations and <i>ALK</i> rearrangement in a comprehensive cohort of 326 consecutive spanish nonsquamous NSCLC patients. Clin Lung Cancer. 2017;Nov;18(6):e395-402.
5.4-44	McCormick F. K-Ras protein as a drug target. J Mol Med. 2016;94:253-258.
5.4-45	McCormick F. Progress in targeting RAS with small molecule drugs. Biochem J. 2019;476 365-374.
5.4-46	McDonald ER, de Weck A, Schlabach MR, et al. Project DRIVE: A compendium of cancer dependencies and synthetic lethal relationships uncovered by large-scale deep RNAi screening. Cell. 2017;170:577-592.
5.4-47	Nolte S, Liegl G, Petersen MA, et al; EORTC Quality of Life Group. General population normative data for the EORTC QLQ-C30 health-related quality of life questionnaire based on 15,386 persons across 13 European countries, Canada and the Unites States. Eur J Cancer. 2019;107:153-163.
5.4-48	Ostrem JM, Peters U, Sos ML, Wells JA, Shokat KM. K-Ras (G12C) inhibitors allosterically control GTP affinity and effector interactions. Nature. 2013;503:548-551.
5.4-49	Ostrem JM and Shokat KM. Direct small-molecule inhibitors of KRAS; from structural insights to mechanism-based design. Nature Rev Drug Discov. 2016;15:771-785.
5.4-50	Oxnard GR, Wilcox KH, Gonen M, et al. Response rate as a regulatory end point in single-arm studies of advanced solid tumors. JAMA Oncol. 2016;2(6):772-779.
5.4-51	Park S, Kim JY, Lee SH, et al. KRAS G12C mutation as a poor prognostic marker of pemetrexed treatment in non-small cell lung cancer. Korean J Intern Med. 2017;32:514-522.
5.4-52	Patricelli MP, Janes MR, Li LS, et al. Selective inhibition of oncogenic KRAS output with small molecules targeting the inactive state. Cancer Discov. 2016;6(3):316-329.
5.4-53	Pignatti F, Jonsson B, Blumenthal G, Justice R. Assessment of benefits and risks in development of targeted therapies for cancer - The view of regulatory authorities. Mol Oncol. 2015;9:1034-1041.
5.4-54	Planchard D, Popat S, Kerr K, et al; ESMO Guidelines Committee. Metastatic non-small cell lung cancer: ESMO Clinical Practice Guidelines for diagnosis, treatment and follow-up. Ann Oncol. 2018;29 (Supplement 4):iv192-iv237.
5.4-55	Prior IA, Lewis PD, Mattos C. A comprehensive survey of Ras mutations in cancer. Cancer Res. 2012;72 (10):2457-2467.
5.4-56	Rittmeyer A, Barlesi F, Waterkamp D, et al. Atezolizumab versus docetaxel in patients with previously treated non-small-cell lung cancer (OAK): a phase 3, open-label, multicentre randomised controlled trial. Lancet. 2017;389:255-265.
5.4-57	Román M, Baraibar I, López I, et al. KRAS oncogene in non-small cell lung cancer: clinical perspectives on the treatment of an old target. Mol Cancer. 2018;17:33.
5.4-58	Scheffler M, Ihle MA, Hein R, et al. K-ras mutation subtypes in NSCLC and associated co-occurring mutations in other oncogenic pathways. J Thorac Oncol. 2019;14(4):606-616.
5.4-59	Scheffzek K, Ahmadian MR, Kabsch W, Wiesmüller L, Lautwein A, Schmitz F, et al. The Ras-RasGAP complex: Structural basis for GTPase activation and its loss in oncogenic Ras mutants. Science. 1997;277:333-338.
5.4-60	Scott NW, Fayes PM, Aaronson NK, et al; EORTC Quality of Life Group. EORTC QLQ-C30 Reference Values: This manual presents reference data for the QLQ-C30 based upon data provided by EORTC Quality of Life Group members and other users of the QLQ-C30. July 2008.
5.4-61	Shepherd FA, Dancey J, Ramlau R, et al. Prospective randomized trial of docetaxel versus best supportive care in patients with non-small-cell lung cancer previously treated with platinum-based chemotherapy. J Clin Oncol. 2000;18(10):2095-2103.
5.4-62	Shepherd FA, Rodrigues Pereira JR, Ciuleanu T, et al. Erlotinib in previously treated non-small-cell lung cancer. N Engl J Med. 2005;353(2):123-132.
5.4-63	Simanshu DK, Nissley DV, McCormick F. RAS proteins and their regulators in human disease. Cell. 2017;170:17-33.

資料番号	著者・表題・出典
5.4-64	Soria JC, Felip E, Cobo M, et al. Afatinib versus erlotinib as second-line treatment of patients with advanced squamous cell carcinoma of the lung (LUX-Lung 8): an open-label randomised controlled phase 3 trial. <i>Lancet Oncol.</i> 2015;16:897-907.
5.4-65	Svaton M, Fiala O, Pesek M, et al. The prognostic role of <i>KRAS</i> mutation in patients with advanced NSCLC treated with second- or third-line chemotherapy. <i>Anticancer Res.</i> 2016;36:1077-1082.
5.4-66	Tamiya Y, Zenke Y, Matsumoto S, et al. Therapeutic impact of mutation subtypes and concomitant <i>STK11</i> mutations in <i>KRAS</i> -mutated non-small cell lung cancer (NSCLC): A result of nationwide genomic screening project (LC-SCRUM-Japan). <i>J Clin Oncol.</i> 2020;38(15 suppl):9589.
5.4-67	Thein K, Banks K, Saam J, et al. The prevalence of <i>KRAS</i> ^{G12C} mutations utilizing circulating tumor DNA (ctDNA) in 80,911 patients with cancer. <i>J Clin Oncol.</i> 2020;38(15 suppl):3547.
5.4-68	Tishelman C, Degner LF, Rudman A, et al. Symptoms in patients with lung carcinoma: Distinguishing distress from intensity. <i>Cancer.</i> 2005;104:2013-2021.
5.4-69	Tishelman C, Petersson LM, Degner LF, Sprangers MAG. Symptom prevalence, intensity, and distress in patients with inoperable lung cancer in relation to time of death. <i>J Clin Oncol.</i> 2007;25(34):5381-5389.
5.4-70	Van Cutsem E, Cervantes A, Nordlinger B, Arnold D; ESMO Guidelines Working Group. Metastatic colorectal cancer: ESMO Clinical Practice Guidelines for diagnosis, treatment and follow-up. <i>Ann Oncol.</i> 2014;25 (Supplement 3):iii1–iii9.
5.4-71	Vokes EE, Ready N, Felip E, et al. Nivolumab versus docetaxel in previously treated advanced non-small-cell lung cancer (CheckMate 017 and CheckMate 057): 3-year update and outcomes in patients with liver metastases. <i>Ann Oncol.</i> 2018;29:959-965.
5.4-72	Wiesweg M, Kasper S, Worm K, et al. Impact of <i>RAS</i> mutation subtype on clinical outcome - a cross-entity comparison of patients with advanced non-small cell lung cancer and colorectal cancer. <i>Oncogene.</i> 2019;38(16):2953-2966.
5.4-73	Xie C, Ying L, Lan-Lan L, et al. Identification of a new potent inhibitor targeting <i>KRAS</i> in non-small cell lung cancer cells. <i>Front Pharmacol.</i> 2017;8:823.
5.4-74	澤端章好, 浅村尚生, 呉屋朝幸, 他. 2002年の肺癌治療例の全国集計に関する報告. <i>日呼外会誌.</i> 2010;24:110-24.

添付すべき資料がない項目リスト

第3部 品質に関する文書

3.2.R 各極の要求資料

第4部 非臨床試験報告書

4.2.1.4 薬力学的薬物相互作用

4.2.2.6 薬物動態学的薬物相互作用

4.2.2.7 その他の薬物動態試験

4.2.3.1 単回投与毒性試験

4.2.3.4 がん原性試験

4.2.3.5.1 受胎能及び着床までの初期胚発生に関する試験

4.2.3.5.3 出生前及び出生後の発生並びに母体の機能に関する試験

4.2.3.5.4 新生児を用いた試験

4.2.3.6 局所刺激性試験

4.2.3.7.1 抗原性試験

4.2.3.7.2 免疫毒性試験

4.2.3.7.4 依存性試験

第5部 臨床試験報告書

5.3.1.3 In Vitro-In Vivoの関連を検討した試験報告書

5.3.3.2 患者におけるPK及び初期忍容性試験報告書

5.3.3.3 内因性要因を検討したPK試験報告書

5.3.4.1 健康被験者におけるPD試験及びPK/PD試験報告書

5.3.6 市販後の使用経験に関する報告書