

TABLE OF CONTENTS

II. ANNUAL REPORT

I. THE PHARMACEUTICALS AND MEDICAL DEVICES AGENCY

1. HISTORY AND PURPOSE OF THE AGENCY	1
2. SUMMARY OF OPERATIONS	
(1) Adverse Health Effect Relief Services	3
(2) Reviews and Related Operations	3
(3) Post-marketing Safety Operations	5
(4) Research & Development Promotion	5
(transferred to the National Institute of Biomedical Innovation in April 2005)	

II. OPERATION RESULTS/ACHIEVEMENT OF FY 2004

PART 1. IMPROVEMENT IN OVERALL OPERATIONS AND QUALITY IN SERVICE OF AGENCY

(1) Efficient and Flexibleactive Operations Management	
1. Operation through target management	7
2. Enhancement of operations management and top management	7
3. Establishment of advisory councils	9
4. Development of effective operations management system	10
5. Standardization of operating procedure	11
6. Promotion of information through database	11
(2) Cost Reduction by Increased Efficiency of Operations	
1. Reduction of general management expenses	12
2. Reduction of program expenses	12
3. Collection and management of contributions	15
(3) Improvement of Services to Public	
1. General consultation service	17
2. Responses to complaints from companies regarding reviews and post-marketing safety operations	19
3. Website posting	19
4. Reporting on f financial standing	19
5. Auditing and related matters	20

(4) Personnel Issues	
1. Discussion of a personnel evaluation system	20
2. Systematic implementation of training	20
3. Appropriate personnel placements	22
4. Securing human resources through open recruitment	22
5. Appropriate human resources management based on work regulations	24
(5) Ensuring Security	
1. Office entrance/Exit controls	25
2. Security of information system	25

PART 2. IMPROVEMENT IN OPERATION OF EACH DEPARTMENT OF THE AGENCY, AND IN ITS QUALITY SERVICE

1. ADVERSE HEALTH EFFECT RELIEF SERVICES	27
(1) Expeditious Processing of Relief Applications	27
(2) Unified Management of Information through the Database	29
(3) Promotion of Appropriate Communication of Information through Cross-functional Collaboration	30
(4) Surveys on Actual Situations of Adverse Health Effects from Pharmaceuticals	30
(5) Expansion of the Consultation Service	30
(6) Expansion and Review of Information Dissemination of information Regarding the Relief Services	
1. Disclosure of cases of benefit payment on website	32
2. Improvement of pamphlets and other communication tools	32
(7) Proactive Public Relations Activities	33
(8) Appropriate Conduct of Relief Services for SMON (subacute myelo-optico-neuropathy) Patients and the those patients infected with HIV-positive and AIDS Patients Infected by from Blood Preparations/Products	33

2. REVIEWS AND RELATED OPERATIONS/ POST-MARKETING SAFETY OPERATIONS

(1) Faster Access to Innovative Pharmaceuticals and Medical Devices	
1. Ensuring the benefits of pharmaceuticals and medical devices for the public and healthcare Professionals	
a. Clinical trial consultations and reviews	35
b. Grasping the needs of public and healthcare professionals	39
2. Measures for efficient and prompt reviews	

a. Approval Reviews for new pharmaceuticals	40
b. Review of new medical devices	42
c. Compliance review of application materials	44
d. Approval review for generic drugs, over-the-counter (OTC) drugs and quasi-drugs	46
3. Reinforcement of clinical trial consultation system	
a. Establishment of priority consultation system	48
b. Acceleration of clinical trial consultations for pharmaceuticals	48
4. Promotion of international harmonization	
a. Approach to international harmonization in ICH and others	49
b. Introduction of total review time	51
(2) Improvement in Reliability of Operations	
1. Planned recruitment of staff with high levels of expertise and systematic provision of opportunities for training	51
2. Development of GMP review system	52
3. Effective use of external experts	53
4. Establishment of information support system	53
5. Strengthening of partnership with foreign regulatory authorities	54
6. Evaluation of such advanced technologies as biotechnology and genomics/ cooperation in developing the national guidelines	55
7. Promotion of appropriate clinical trials	55
8. Timely provision of information including review reports	56
(3) Reinforcement of Post-Marketing Safety Operations (Reinforcement of information management and emergency management system)	
1. Basic direction of safety measures	58
2. Introduction of new method (Study to introduce data mining technique)	60
3. Establishment of sentinel medical institution network	62
4. Study on system for information collection and review on medical device malfunctions	64
5. Proper examination of reports on ADRs and medical device malfunction	65
6. Computerization of reports on ADRs and medical device malfunctions	66
7. Establishment of post-marketing safety system through information feedback	
a. Feedback to companies	66
b. Feedback to health professionals	67

c. Information provision to general consumers and patients	68
d. Improvement of the contents and quality of disseminating information	68

III. SUPPORTING INFORMATION

PART 1. OPERATIONS RELEVANT TO ADVERSE HEALTH EFFECT RELIEF SERVICES

1. ADVERSE DRUG REACTION RELIEF SERVICE

(1) Number of Applications for Benefits and Judged Cases	
(2) Contributions	71
(3) Liability Reserve	72
(4) Consultation Service	73
(5) Health and Welfare Services	73

2. BIOLOGICAL PRODUCT-DEPRIVED INFECTIOUS DISEASE RELIEF SERVICE

(1) Benefits Payment for Infectious Disease Relief	74
(2) Contributions	75

3. RELIEF SERVICES RELATED TO SMON

75

4. RELIEF SERVICE RELATED TO AIDS

(1) The Agency Conducted the Following Three Services for HIV-positive and AIDS Patients Infected from Blood Preparations/Products	76
--	----

PART 2. REVIEW AND RELATED OPERATIONS AND POST-MARKETING OPERATIONS

1. FACE-TO-FACE CONSULTATION

(1) Existing Services	77
(2) New Services	77
1. Simple consultation services concerning generic drugs, over-the-counter (OTC) drugs and quasi drugs	77
2. CT Consultations and pre-approval consultations services for medical devices and in vitro diagnostics	77
3. Simple consultation services regarding medical devices and in vitro diagnostic drugs	78
4. Services of preliminary interview about medical devices and in vitro diagnostics	

	78
5. Review on designation of prioritized face-to-face advice	78
2. OPERATION ON APPROVAL REVIEW FOR PHARMACEUTICALS ETC.	
(1) Approval Review for Pharmaceuticals and Quasi Drugs	78
1. Review for new drugs approval	79
2. Review for the approval of prioritized applications	79
3. Disclosure of review progress to applicants	80
4. Approval review for in vitro diagnostics	81
(2) Re-examination and Re-evaluation of Pharmaceuticals	81
3. REVIEW FOR MEDICAL DEVICES APPROVAL	
(1) Approval Review for Medical Devices	81
1. Approval review for new medical devices	81
2. Approval review for improved medical devices	82
3. Approval review for generic medical devices	82
4. Approval review for prioritized medical devices	82
5. Disclosure about review progress to applicants	82
4. CONFORMITY AUDIT	
(1) Conformity Audit for Application Materials for New Drug Approval	83
(2) Conformity Audit for Generic Drugs	83
(3) Conformity Audit of Application Materials for Re-examination	83
(4) Conformity Audit of Application Materials for Re-evaluation	83
(5) On-site Conformity Review of Pharmaceuticals Safety Tests (GLP Review)	84
(6) On-site Conformity Review of Drug Clinical Study (GCP Review)	
1. GCP review for new drugs	84
2. GCP review for generic drugs	84
(7) On-site GPMS* (Good post-marketing surveillance practice) review	84
5. OTHER SERVICES RELATED TO REVIEWS	
(1) Investigation of Notified CT Protocol	85
(2) Investigation on Reported Adverse Health Effects (AHEs) that Happened in Clinical Trials	85
(3) Services Related to Prior Assessment on Pharmaceuticals / Medical Devices that Utilize Cells and Tissues, Drugs for Gene Therapy and Cartagena Related Issues etc.	86

(4) Review and Confirmation of Exporting License Application	87
--	----

6. ASSURANCE OF COMPLIANCE AND STANDARD AND QUALITY CONTROL

(1) On-site GMP Audit of Pharmaceuticals etc.	87
(2) Making Draft Standards	88

7. POST-MARKETING SAFETY OPERATIONS

(1) Collection of ADR Reports	88
(2) Investigation/review of ADR reports and others	90
(3) Providing Safety Information	92
(4) Consultation Services for Consumers	93
(5) Collection of Safety Measures Contributions	95

Incorporated Administrative Agency – Pharmaceutical and Medical Devices Agency (PMDA)

Midterm Targets	96
-----------------	----

Incorporated Administrative Agency – Pharmaceutical and Medical Devices Agency (PMDA)

Midterm Plan	103
--------------	-----

IV. STATISTICAL TABLES

1. Adverse Health Effect Relief Service

Table 1 Number of Cases on Adverse Health Effect Relief Benefits 1980-2004	120
--	-----

Table 2 Numbers of claims and benefit amounts in Adverse Health Effect Relief Services	121
--	-----

Table 3 Numbers of claims and benefit amounts in Adverse Health Effect Relief Services in each prefecture	122
---	-----

Table 4 Transition in the breakdown of different diseases (symptoms) due to adverse reactions	124
---	-----

Table 5 Transition in the number of pharmaceuticals causing adverse reactions by different categories of drug actions	125
---	-----

Table 6 Adverse Drug Reaction Funds from Manufactures	126
---	-----

Table 7 Number of Consultations for Relief Benefits	127
---	-----

Table 8 Biological Product-derived Infectious Disease Relief Service	128
--	-----

Relief Service for SMON Patients

Table 9 Payment of Healthcare Allowances and Nursing Expenses	129
---	-----

Relief Service for HIV-positive and AIDS Patients

Table 10	Number of Claim Cases and Payment Amounts related to Research and Study Projects	130
Table 11	Number of Claim Cases and Payment Amounts related to Healthcare Support Service	131
Table 12	Number of Claims and Benefit Amounts by Types in Relief Services for HIV-positive and AIDS Patients	132

Relief Service of HIV-positive and AIDS Patients

Table 13	Number of Consultation Cases for Relief Services	133
Table 14	Number of Applications and Approvals for Drugs (FY1997 – FY2004)	134
Table 15	Number of Applications and Approvals for Medical Devices (FY1997 – FY2004)	135
Table 16	Review Status for generic drugs, etc. in FY 2004	136
Table 17	Clinical Trial Consultation Achievements	137
Table 18	A list of approved items in 2004 (new drugs)	139
Table 19	A list of approved items in 2004 (new medical devices)	149
Table 20	Number of Reports on ADRs and Medical Device Malfunctions	151
Table 21	Measures for safety strategies and revision to "precautions on use" related to pharmaceuticals implemented by the Ministry of Health, Labour and Welfare in 2004 – 2004 Designations	152
Table 22	Revision of “Precautions on Use” related to medical devices: 2004 Designations and notification on self-assessment 2004 designations	156
Table 23	Safety information of pharmaceuticals and devices in FY 2004 (No.200-211)	158
Table 24	User Fee Lists	160