

Table 12-1 Incidence of AEs – Evaluation for severity (continued)
<Korea>

N=19

Severity	Mild		Moderate		Severe		Total	
	Number of subjects (events)	Incidence (%)	Number of subjects (events)	Incidence (%)	Number of subjects (events)	Incidence (%)	Number of subjects (events)	Incidence (%)
Adverse Event SOC PT	3 (4)	16	0 (0)	0	0 (0)	0	3 (4)	16
Nervous system disorders	3 (4)	16	0 (0)	0	0 (0)	0	3 (4)	16
Dizziness	1 (1)	5	0 (0)	0	0 (0)	0	1 (1)	5
Headache	1 (1)	5	0 (0)	0	0 (0)	0	1 (1)	5
Somnolence	1 (1)	5	0 (0)	0	0 (0)	0	1 (1)	5
Syncope	1 (1)	5	0 (0)	0	0 (0)	0	1 (1)	5

Incidence = the percentage of subjects

<Caucasian>

N=20

Severity	Mild		Moderate		Severe		Total	
	Number of subjects (events)	Incidence (%)	Number of subjects (events)	Incidence (%)	Number of subjects (events)	Incidence (%)	Number of subjects (events)	Incidence (%)
Adverse Event SOC PT	2(2)	10	1(1)	5			3(3)	15
Skin	1(1)	5					1(1)	5
Cold Sore	1(1)	5					1(1)	5
Nervous system disorders	1(1)	5	1(1)	5			2(2)	10
Headache	1(1)	5	1(1)	5			2(2)	10

Incidence = the percentage of subjects

Source: Appendix 8.1

* Coded using MedDRA (Ver. 12.1 or more)

12.2.3 Analysis of Adverse Events

A total of 14 AEs (6 in 5 Japanese subjects, 1 in a Chinese subject, 4 in 3 Korean subjects, 3 in 3 Caucasian subjects) were reported over the course of the study.

Four (29%) of 14 AEs were considered to be probably related to the study drug. Of the remaining 10 AEs, 2, 7 and 1 AE were considered to be unknown, probably not related and definitely not related, respectively. Thirteen (93%) of 14 AEs were rated as mild in severity. One incident of headache was rated as moderate in severity, and was considered probably related to the study drug, as the onset time was approximately 9 hours after administration of the study drug. AEs were generally short lasting and resolved without concomitant medication or other intervention.

All AEs were:

[Japanese]

- Malaise (3 incidents, reported by 2 subjects)
- Urticaria (1 incident, reported by a subject)
- Electrocardiogram PR prolongation (1 incident, reported by a subject)
- Epistaxis (1 incident, reported by a subject)

[Chinese]

- C-reactive protein increased (1 incident, reported by a subject)

[Korean]

- Dizziness (1 incident, reported by a subject)
- Headache (1 incident, reported by a subject)
- Somnolence (1 incident, reported by a subject)
- Syncope (1 incident, reported by a subject)

[Caucasian]

- Headache (2 incidents, reported by 2 subjects)
- Cold Sore (1 incident, reported by a subject)

No AEs required concomitant medication or other intervention. One AE (C-reactive protein increased in a Chinese subject) was recovering as of 9 days after administration, and all the other AEs resolved without sequelae by the end of the study.

12.2.4 Deaths, Discontinuations Due to Adverse Events, and Serious Adverse Events

There were no deaths, discontinuations due to adverse events, or serious adverse events during this study in each study site.

12.3 Clinical Laboratory Evaluation

Laboratory values (hematology, blood biochemistry and urinalysis) outside the normal reference range are flagged H (high) or L (low) as appropriate. Abnormal laboratory values by subject are listed in Appendix 8.2. Normal / abnormal rating shift in abnormal laboratory values are tabulated in Table 14.1. Summary statistics for laboratory values and shift tables for urinalysis parameters are presented in Tables 14.2 and 14.3, respectively. Follow-up/additional test by subject are listed in Appendix 8.6.

Although many subjects showed minor abnormalities in various laboratory values before and/or 48 hours after administration of the study drug, none of these except one (bilirubin in a Caucasian subject) was considered to be clinically significant. There were no apparent trends in laboratory parameters following administration of moxifloxacin.

12.4 Other Safety Assessments

12.4.1 Renal Function

Summary statistics for creatinine clearance are shown in Table 14.4. Individual creatinine clearance values before administration of the study drug are listed in Appendix 8.7.

Creatinine clearance values in each study site were as follows: 70 to <90 mL/min (1 in China, 4 in the US), 90 to <110 (6 in Japan, 4 in China, 3 in Korea, 9 in the US), 110 to <130 (8 in Japan, 10 in China, 10 in Korea, 6 in the US), 130 to <150 (4 in Japan, 5 in China, 3 in Korea, 1 in the US), 150 to <170 (2 in Japan, 3 in Korea).

12.4.2 Vital Signs

Summary statistics for vital signs are shown in Table 14.5.

There were no clinically relevant abnormalities in vital signs, and no apparent trends following dose administration.

12.4.3 12-Lead ECG

Individual 12-lead ECG abnormalities at screening, before administration, at 3, 24 and 48 hours after administration of the study drug are listed in Appendix 8.3.

Although most subjects had minor abnormalities at some time during the study, none were considered to be clinically significant.

12.5 Safety Conclusions

Eighty eligible subjects were enrolled for this clinical pharmacokinetic study in order to investigate the pharmacokinetic profile of single oral dose of 400 mg of moxifloxacin in healthy adult male subjects among four ethnics. Of 80 subjects, one was withdrew his consent with personal reason and dropped out before administrating the study drug. All other subjects satisfied with all of the inclusion criteria and none of the exclusion criteria. Seventy-nine subjects completed the study and were evaluated for safety in each study site.

A total of 14 AEs (6 in 5 Japanese subjects, 1 in a Chinese subject, 4 in 3 Korean subjects and 3 in 3 Caucasian subjects) were observed during the study, of which it was considered that 4 events (urticaria in a Japanese subject, dizziness and headache in 2 Korean subjects, headache in a Caucasian subject) were considered to be probably related to the study drug. The most frequently reported AEs were malaise (3 incidents in 2 Japanese subjects) and headache (2 incidents in 2 Caucasian subjects and 1 incident in a Korean subject). All AEs except one incident of headache were mild in severity. AEs were generally short lasting and resolved without concomitant medication or other intervention. There were no deaths or serious AEs.

Laboratory measurements and clinical safety assessments (vital signs, physical examinations and 12-lead ECG) did not appear to show any clinically relevant abnormalities arising from the administration of moxifloxacin.

13. DISCUSSION AND OVERALL CONCLUSIONS

All treatment-related AEs except one were mild in severity, and none required concomitant medication or intervention. Laboratory and other safety assessments did not appear to show any clinically relevant abnormalities arising from the administration of moxifloxacin. Moxifloxacin showed the similar safety results in these four ethnic groups.

The data from this study indicate that moxifloxacin given in oral dose of 400 mg is safe and relatively well-tolerated by healthy male Japanese, Chinese, Korean and Caucasian subjects.

14. TABLES AND FIGURES REFERRED TO BUT NOT INCLUDED IN THE TEXT

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14.1 Summary Table of Normal/Abnormal Rating Shift in Abnormal Laboratory Values (continued)

Parameter	Japanese			Chinese			Korean			Caucasian		
	48 Hours -1 Hour		High	48 Hours -1 Hour		High	48 Hours -1 Hour		High	48 Hours -1 Hour		High
	Low	Normal	Low	Normal	Low	Normal	Low	Normal	Low	Normal	Low	Normal
Total Protein	Low	0	0	0	0	0	0	0	0	0	0	0
	Normal	0	19	0	18	0	18	0	19	0	0	20
	High	0	0	0	2	0	0	0	0	0	0	0
Albumin	Low	0	0	0	0	0	0	0	0	0	0	0
	Normal	0	20	0	20	0	20	0	19	0	0	20
	High	0	0	0	0	0	0	0	0	0	0	0
Total Cholesterol	Low	0	0	0	0	0	0	0	0	0	0	0
	Normal	1	19	0	18	0	18	0	18	1	0	16
	High	0	0	0	1	0	1	0	0	0	0	1
HDL Cholesterol	Low	0	0	0	0	0	0	0	0	0	0	0
	Normal	1	18	0	12	0	12	0	10	1	0	16
	High	0	1	0	3	0	3	0	2	5	0	0
LDL Cholesterol	Low	2	0	0	2	0	2	0	0	0	0	0
	Normal	0	15	0	10	0	10	0	13	2	0	7
	High	0	1	2	0	0	0	0	0	4	0	0
Triglyceride	Low	0	0	0	0	0	0	0	0	0	0	0
	Normal	0	20	0	14	0	14	0	17	1	0	19
	High	0	0	0	3	0	3	0	0	1	0	0
BUN	Low	0	0	0	0	0	0	0	0	0	0	0
	Normal	0	20	0	20	0	20	0	11	0	0	20
	High	0	0	0	0	0	0	0	0	0	0	0
Creatinine	Low	0	0	0	0	0	0	0	0	0	0	0
	Normal	0	20	0	20	0	20	0	19	0	0	19
	High	0	0	0	0	0	0	0	0	0	0	0
Uric Acid	Low	1	0	0	0	0	0	0	0	0	0	0
	Normal	0	19	0	17	0	17	0	15	0	0	19
	High	0	0	0	0	0	0	0	2	2	0	0

14.1 Summary Table of Normal/Abnormal Rating Shift in Abnormal Laboratory Values (continued)

Parameter	Japanese			Chinese			Korean			Caucasian		
	48 Hours -1 Hour	Low	Normal	High	48 Hours -1 Hour	Low	Normal	High	48 Hours -1 Hour	Low	Normal	High
AST	Low	0	0	0	Low	0	0	0	Low	0	0	0
	Normal	0	20	0	Normal	0	20	0	Normal	0	18	0
	High	0	0	0	High	0	0	0	High	0	2	0
ALT	Low	0	0	0	Low	0	0	0	Low	0	0	0
	Normal	0	20	0	Normal	0	19	1	Normal	0	19	0
	High	0	0	0	High	0	0	0	High	0	0	0
γ-GTP	Low	0	0	0	Low	0	0	0	Low	0	0	0
	Normal	0	19	0	Normal	0	20	0	Normal	0	17	0
	High	0	1	0	High	0	0	0	High	0	1	0
Total Bilirubin	Low	0	0	0	Low	0	0	0	Low	0	0	0
	Normal	0	19	1	Normal	0	16	0	Normal	0	8	0
	High	0	0	0	High	0	3	1	High	0	6	5
Direct Bilirubin	Low	0	0	0	Low	0	0	0	Low	0	0	0
	Normal	0	20	0	Normal	0	19	0	Normal	0	19	0
	High	0	0	0	High	0	0	1	High	0	0	0
ALP	Low	0	0	0	Low	0	0	0	Low	0	0	0
	Normal	1	18	0	Normal	0	20	0	Normal	0	19	0
	High	0	0	1	High	0	0	0	High	0	0	0
LDH	Low	1	0	0	Low	0	1	0	Low	0	0	0
	Normal	1	17	0	Normal	3	16	0	Normal	0	11	1
	High	0	1	0	High	0	0	0	High	0	7	0
CK	Low	1	0	0	Low	0	0	0	Low	0	0	0
	Normal	1	17	0	Normal	0	20	0	Normal	0	19	0
	High	0	1	0	High	0	0	0	High	0	1	0
Na	Low	0	0	0	Low	0	0	0	Low	0	0	0
	Normal	0	20	0	Normal	0	20	0	Normal	0	18	0
	High	0	0	0	High	0	0	0	High	0	0	0
K	Low	0	0	0	Low	0	0	0	Low	0	0	0
	Normal	0	20	0	Normal	0	20	0	Normal	0	19	0
	High	0	0	0	High	0	0	0	High	0	0	0
Cl	Low	0	0	0	Low	0	0	0	Low	0	0	0
	Normal	0	20	0	Normal	0	20	0	Normal	0	19	0
	High	0	0	0	High	0	0	0	High	0	0	0

14.1 Summary Table of Normal/Abnormal Rating Shift in Abnormal Laboratory Values (continued)

Parameter	Japanese						Chinese						Korean						Caucasian					
	48 Hours -1 Hour		Low	Normal	High	-1 Hour	48 Hours -1 Hour		Low	Normal	High	-1 Hour	48 Hours -1 Hour		Low	Normal	High	-1 Hour	48 Hours -1 Hour		Low	Normal	High	
	Low	High	Low	Normal	High		Low	Normal	High	Low	Normal		High	Low	Normal	High	Low		Normal	High	Low	Normal	High	
Blood Sugar	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	0	0	0	
	Normal	0	19	0	0	Normal	0	20	0	0	0	Normal	0	17	0	0	0	0	0	Normal	0	20	0	
	High	0	1	0	0	High	0	0	0	0	0	High	0	0	0	0	0	0	0	High	0	0	0	
CRP	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	0	0	0	
	Normal	0	19	0	0	Normal	0	19	1	0	0	Normal	0	19	0	0	0	0	0	Normal	0	18	0	
	High	0	1	0	0	High	0	0	0	0	0	High	0	0	0	0	0	0	0	High	0	2	0	
Urine Protein	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	0	0	0	
	Normal	0	20	0	0	Normal	0	19	0	0	0	Normal	0	18	0	0	0	0	0	Normal	0	20	0	
	High	0	0	0	0	High	0	1	0	0	0	High	0	0	0	0	0	0	0	High	0	0	0	
Urine Glucose	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	0	0	0	
	Normal	0	20	0	0	Normal	0	20	0	0	0	Normal	0	19	0	0	0	0	0	Normal	0	20	0	
	High	0	0	0	0	High	0	0	0	0	0	High	0	0	0	0	0	0	0	High	0	0	0	
Bilirubin	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	0	0	0	
	Normal	0	20	0	0	Normal	0	20	0	0	0	Normal	0	19	0	0	0	0	0	Normal	0	18	0	
	High	0	0	0	0	High	0	0	0	0	0	High	0	0	0	0	0	0	0	High	0	2	0	
Ketone bodies	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	0	0	0	
	Normal	0	20	0	0	Normal	0	20	0	0	0	Normal	0	18	0	0	0	0	0	Normal	0	15	1	
	High	0	0	0	0	High	0	0	0	0	0	High	0	1	0	0	0	0	0	High	0	3	1	
Occult Blood	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	0	0	0	
	Normal	0	19	0	0	Normal	0	19	0	0	0	Normal	0	18	0	0	0	0	0	Normal	0	19	0	
	High	0	1	0	0	High	0	1	0	0	0	High	0	0	0	0	0	0	0	High	0	0	1	
pH	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	1	0	0	
	Normal	0	20	0	0	Normal	0	20	0	0	0	Normal	0	19	0	0	0	0	0	Normal	0	19	0	
	High	0	0	0	0	High	0	0	0	0	0	High	0	0	0	0	0	0	0	High	0	0	0	
Urobilinogen	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	0	0	0	
	Normal	0	20	0	0	Normal	0	20	0	0	0	Normal	0	19	0	0	0	0	0	Normal	0	20	0	
	High	0	0	0	0	High	0	0	0	0	0	High	0	0	0	0	0	0	0	High	0	0	0	
Urinary Sediment	Low	0	0	0	0	Low	0	0	0	0	0	Low	0	0	0	0	0	0	0	Low	0	0	0	
	Normal	0	7	5	0	Normal	0	20	0	0	0	Normal	0	19	0	0	0	0	0	Normal	0	20	0	
	High	0	6	2	0	High	0	0	0	0	0	High	0	0	0	0	0	0	0	High	0	0	0	

14.2 Summary Statistics for Laboratory Values

Hematology

Parameter	Japanese		Chinese		Korean		Caucasian			
	Unit	-1 Hour	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours
RBC	Number of subjects	20	20	20	20	19	19	20	20	20
	Mean	494.1	480.3	4.9	5.0	518.1	523.5	5.0	5.0	5.1
	SD	33.36	22.42	0.37	0.34	26.09	31.36	0.27	0.27	0.31
	Minimum	453.0	444.0	4.3	4.4	468.0	453.0	4.5	4.5	4.6
	Maximum	499.00	474.00	4.86	5.07	522.00	520.00	4.97	4.97	5.02
Reticulocyte	Number of subjects	20	20	20	20	19	19	20	20	20
	Mean	12.5	11.2	48.0	51.9	10.7	11.3	1.2	1.2	1.1
	SD	2.76	3.07	11.84	12.00	3.24	3.14	0.45	0.45	0.38
	Minimum	8.0	5.1	32.4	34.9	5.1	7.0	0.5	0.5	0.6
	Maximum	13.00	10.40	46.73	52.60	10.10	10.90	1.15	1.15	1.05
Hemoglobin	Number of subjects	20	20	20	20	19	19	20	20	20
	Mean	15.1	14.7	160.1	161.1	15.5	15.7	15.3	15.3	15.5
	SD	0.97	0.69	14.30	10.32	0.84	0.85	0.91	0.91	0.84
	Minimum	13.6	13.4	131.0	140.0	13.8	14.0	13.9	13.9	14.6
	Maximum	15.00	14.75	160.00	162.50	15.50	15.80	15.25	15.25	15.40
Hematocrit	Number of subjects	20	20	20	20	19	19	20	20	20
	Mean	43.8	42.5	43.8	45.0	46.0	46.4	44.3	44.3	44.5
	SD	2.48	1.91	2.51	2.82	2.05	2.08	2.34	2.34	2.22
	Minimum	39.7	38.9	40.5	39.6	42.0	41.8	40.0	40.0	41.6
	Maximum	43.15	42.45	43.70	45.90	46.20	46.80	44.45	44.45	44.45
Platelet	Number of subjects	20	20	20	20	19	19	20	20	20
	Mean	24.6	23.9	200.5	217.2	23.8	24.1	203.9	203.9	206.0
	SD	4.01	4.11	34.23	41.76	6.14	5.98	27.09	27.09	29.75
	Minimum	16.6	16.4	130.0	133.0	13.1	11.5	153.0	153.0	156.0
	Maximum	24.60	23.55	208.00	224.00	23.70	25.10	201.50	201.50	204.50
WBC	Number of subjects	20	20	20	20	19	19	20	20	20
	Mean	56.9	51.8	5.4	5.3	5,832.1	5,350.5	5.4	5.4	4.6
	SD	11.96	11.11	1.55	1.34	1,114.20	927.95	1.29	1.29	0.85
	Minimum	34.4	33.8	3.3	4.0	3,780.0	3,490.0	3.0	3.0	3.3
	Maximum	55.15	49.70	5.29	4.89	5,740.00	5,500.00	5.10	5.10	4.50
		77.9	81.7	10.1	9.8	7,880.0	7,750.0	8.9	8.9	5.9

14.2 Summary Statistics for Laboratory Values (continued)

Hematology

Parameter	Japanese		Chinese		Korean		Caucasian		
	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours
Neutrophil	Number of subjects	%	20	20	20	19	19	20	20
	Mean		53.8	53.8	57.1	56.9	49.2	57.3	54.9
	SD		7.16	5.87	10.06	10.52	7.22	8.65	7.10
	Minimum		43.6	39.5	43.4	43.2	35.8	43.0	41.0
	Median		52.05	54.25	58.25	56.35	51.60	56.00	56.00
	Maximum		68.3	62.1	81.8	82.9	61.3	76.0	67.0
Lymphocyte	Number of subjects	%	20	20	20	20	19	20	20
	Mean		36.1	36.1	32.2	33.5	39.3	31.0	34.2
	SD		6.94	5.42	8.86	9.05	6.89	7.32	7.11
	Minimum		20.4	27.9	13.1	13.2	23.9	16.0	23.0
	Median		36.30	37.45	30.40	34.40	38.30	32.00	34.00
	Maximum		46.8	47.3	44.7	47.5	49.3	44.0	47.0
Monocyte	Number of subjects	%	20	20	20	20	19	20	20
	Mean		5.9	5.9	5.6	4.6	8.0	8.3	8.0
	SD		1.43	1.34	1.02	1.08	1.78	1.89	1.00
	Minimum		3.8	4.3	3.3	2.2	6.0	5.0	5.0
	Median		5.60	5.85	5.55	4.70	8.10	8.00	8.00
	Maximum		9.6	10.1	7.7	6.8	13.2	13.0	9.0
Eosinophil	Number of subjects	%	20	20	20	20	19	20	20
	Mean		3.6	3.7	3.1	2.6	2.8	2.9	2.3
	SD		2.02	1.98	1.86	1.93	1.02	1.66	1.45
	Minimum		1.3	1.3	0.9	0.5	1.3	1.0	1.0
	Median		3.35	3.45	2.60	2.20	2.50	3.00	2.00
	Maximum		8.8	7.3	8.6	8.5	5.1	6.0	6.0
Basophil	Number of subjects	%	20	20	20	20	19	20	20
	Mean		0.6	0.6	0.7	0.5	0.5	0.6	0.7
	SD		0.32	0.32	0.24	0.21	0.33	0.24	0.47
	Minimum		0.2	0.2	0.2	0.1	0.2	0.0	0.0
	Median		0.55	0.50	0.70	0.50	0.30	1.00	1.00
	Maximum		1.3	1.6	1.2	1.0	1.5	1.0	1.0

14.2 Summary Statistics for Laboratory Values (continued)

Blood Biochemistry

Parameter	Japanese			Chinese			Korean			Caucasian		
	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours
Total Protein	Number of subjects	20	20	20	20	20	g/dL	19	19	19	20	20
	Mean	7.1	6.9	77.4	76.5	76.5	g/dL	7.1	7.1	7.2	6.9	7.1
	SD	0.45	0.32	3.52	3.43	3.43	g/dL	0.39	0.39	0.50	0.40	0.42
	Minimum	6.0	6.2	67.2	67.4	67.4	g/dL	6.2	6.2	6.2	6.3	6.4
	Maximum	7.8	7.5	83.0	81.0	81.0	g/dL	7.8	7.8	8.0	7.6	8.0
Albumin	Number of subjects	20	20	20	20	20	g/dL	19	19	19	20	20
	Mean	4.6	4.5	45.8	45.1	45.1	g/dL	4.4	4.4	4.5	4.5	4.6
	SD	0.24	0.17	2.63	3.09	3.09	g/dL	0.23	0.23	0.31	0.26	0.25
	Minimum	4.2	4.2	40.7	38.4	38.4	g/dL	4.1	4.1	4.0	4.1	4.2
	Maximum	4.9	4.8	46.10	45.55	45.55	g/dL	4.40	4.40	4.50	4.50	4.60
Total Cholesterol	Number of subjects	20	20	20	20	20	mg/dL	19	19	19	20	20
	Mean	178.1	169.1	4.4	4.4	4.4	mg/dL	174.8	174.8	183.9	170.0	175.2
	SD	31.28	30.30	0.53	0.59	0.59	mg/dL	35.39	35.39	35.21	26.88	26.26
	Minimum	119.0	108.0	3.6	3.5	3.5	mg/dL	115.0	115.0	123.0	97.0	101.0
	Maximum	230.0	223.0	4.31	4.37	4.37	mg/dL	170.00	170.00	181.00	170.50	178.00
HDL Cholesterol	Number of subjects	20	20	20	20	20	mg/dL	19	19	19	20	20
	Mean	60.5	55.0	1.2	1.2	1.2	mg/dL	51.2	51.2	52.1	51.9	47.7
	SD	11.84	9.82	0.25	0.26	0.26	mg/dL	10.79	10.79	9.16	12.44	8.96
	Minimum	36.0	33.0	0.7	0.7	0.7	mg/dL	28.0	28.0	30.0	35.0	32.0
	Maximum	91.0	73.0	1.19	1.16	1.16	mg/dL	50.00	50.00	53.00	53.50	47.00
LDL Cholesterol	Number of subjects	20	20	20	20	20	mg/dL	78.0	78.0	71.0	90.0	70.0
	Mean	105.1	102.6	2.7	2.6	2.6	mg/dL	105.4	105.4	115.6	99.3	108.9
	SD	25.88	25.64	0.55	0.55	0.55	mg/dL	32.02	32.02	31.32	23.80	23.42
	Minimum	62.0	58.0	1.8	1.9	1.9	mg/dL	61.0	61.0	75.0	49.0	50.0
	Maximum	153.0	151.0	2.74	2.57	2.57	mg/dL	102.00	102.00	112.00	99.50	112.50
Triglyceride	Number of subjects	20	20	20	20	20	mg/dL	165.0	165.0	173.0	143.0	155.0
	Mean	87.4	86.4	1.3	1.3	1.3	mg/dL	100.9	100.9	110.8	94.5	93.0
	SD	35.36	29.67	0.38	0.36	0.36	mg/dL	48.40	48.40	54.93	26.42	23.04
	Minimum	40.0	53.0	0.7	0.6	0.6	mg/dL	45.0	45.0	54.0	35.0	35.0
	Maximum	161.0	148.0	1.24	1.27	1.27	mg/dL	89.00	89.00	81.00	96.50	94.50

14.2 Summary Statistics for Laboratory Values (continued)

Blood Biochemistry

Parameter	Japanese			Chinese			Korean			Caucasian		
	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours
BUN	Number of subjects	20	20	20	20	20	mg/dL	19	19	mg/dL	20	20
	Mean	12.9	11.3	4.8	4.1	10.0	11.3	10.0	10.0	12.5	13.9	
	SD	2.76	2.62	0.73	0.61	1.53	1.86	1.53	7.0	2.44	1.77	
	Minimum	9.0	8.0	3.4	3.1	4.19	9.0	7.0	11.00	8.0	11.0	
	Maximum	13.00	11.00	4.72	4.19	5.2	11.00	10.00	13.0	12.00	14.00	
Creatinine	Number of subjects	20	20	20	20	20	mg/dL	19	19	mg/dL	20	20
	Mean	0.8	0.8	79.8	84.8	84.8	0.9	1.0	1.0	1.0	1.0	
	SD	0.10	0.10	7.82	6.65	7.00	0.08	0.09	0.08	0.10	0.11	
	Minimum	0.7	0.7	70.0	72.0	72.0	0.8	0.8	0.8	0.8	0.8	
	Maximum	1.0	1.1	95.0	97.0	97.0	1.1	1.2	1.2	1.1	1.2	
Uric Acid	Number of subjects	20	20	20	20	20	mg/dL	19	19	mg/dL	20	20
	Mean	5.9	5.8	340.9	356.5	356.5	6.4	6.1	6.1	6.2	6.0	
	SD	0.80	0.77	42.57	47.39	47.39	0.94	0.88	0.88	0.98	0.87	
	Minimum	3.5	3.5	271.0	277.0	277.0	4.7	5.0	5.0	4.6	4.5	
	Maximum	7.5	7.0	415.0	436.0	436.0	8.2	8.3	8.3	6.05	6.05	
AST	Number of subjects	20	20	20	20	20	IU/L	19	19	IU/L	20	20
	Mean	18.9	17.2	19.4	20.3	20.3	21.2	18.4	18.4	22.1	21.2	
	SD	4.72	4.09	3.55	4.58	4.58	6.29	3.10	3.10	6.82	4.07	
	Minimum	13.0	12.0	13.0	15.0	15.0	15.0	13.0	13.0	15.0	16.0	
	Maximum	19.00	16.00	19.00	19.50	19.50	20.00	17.00	17.00	20.00	20.50	
ALT	Number of subjects	20	20	20	20	20	IU/L	19	19	IU/L	20	20
	Mean	18.7	16.0	16.4	18.9	18.9	17.5	17.6	17.6	20.0	19.1	
	SD	7.86	6.64	6.89	8.17	8.17	6.70	7.09	7.09	9.00	7.19	
	Minimum	7.0	8.0	7.0	8.0	8.0	10.0	10.0	10.0	11.0	11.0	
	Maximum	19.00	15.00	15.00	17.00	17.00	15.00	15.00	15.00	17.50	17.00	
γ-GTP	Number of subjects	20	20	20	20	20	IU/L	19	19	IU/L	20	20
	Mean	23.0	20.9	17.7	17.5	17.5	24.0	23.3	23.3	20.2	19.9	
	SD	10.11	8.49	4.46	4.58	4.58	13.99	12.04	12.04	14.36	12.81	
	Minimum	13.0	12.0	12.0	11.0	11.0	9.0	11.0	11.0	4.0	4.0	
	Maximum	21.00	18.50	16.50	16.50	16.50	18.00	19.00	19.00	17.50	16.00	
		55.0	45.0	25.0	24.0	24.0	57.0	49.0	49.0	72.0	65.0	

14.2 Summary Statistics for Laboratory Values (continued)

Blood Biochemistry

Parameter	Japanese				Chinese				Korean				Caucasian			
	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	Unit	-1 Hour	48 Hours	
Total Bilirubin	Number of subjects	20	20	20	20	20	mg/dL	20	20	19	19	mg/dL	20	20	20	
	Mean	1.0	1.0	1.0	15.6	13.2		15.6	13.2	1.2	1.2		1.2	1.2	0.8	
	SD	0.24	0.26	0.26	6.49	6.51		6.49	6.51	0.26	0.24		0.26	0.24	0.44	
	Minimum	0.6	0.7	0.7	7.9	7.5		7.9	7.5	0.7	0.8		0.7	0.8	0.3	
	Maximum	1.05	1.00	1.00	15.35	12.30		15.35	12.30	1.30	1.20		1.30	1.20	0.80	
Direct Bilirubin	Number of subjects	20	20	20	20	20	mg/dL	20	20	19	19	mg/dL	20	20	20	
	Mean	0.2	0.2	0.2	2.7	2.5		2.7	2.5	0.3	0.3		0.3	0.1		
	SD	0.06	0.07	0.07	1.88	2.03		1.88	2.03	0.08	0.06		0.08	0.06		
	Minimum	0.1	0.1	0.1	0.3	0.3		0.3	0.3	0.2	0.2		0.2	0.1		
	Maximum	0.20	0.20	0.20	2.73	2.14		2.73	2.14	0.30	0.30		0.30	0.11		
ALP	Number of subjects	20	20	20	20	20	IU/L	20	20	19	19	IU/L	20	20	20	
	Mean	216.2	204.3	204.3	72.1	71.6		72.1	71.6	58.7	56.7		58.7	57.9		
	SD	58.43	55.98	55.98	21.87	19.51		21.87	19.51	15.07	12.87		15.07	10.74		
	Minimum	139.0	128.0	128.0	56.0	53.0		56.0	53.0	40.0	37.0		40.0	36.0		
	Maximum	223.50	209.50	209.50	69.00	65.00		69.00	65.00	55.00	54.00		55.00	56.00		
LDH	Number of subjects	20	20	20	20	20	IU/L	20	20	19	19	IU/L	20	20	20	
	Mean	142.8	130.1	130.1	123.2	112.3		123.2	112.3	220.6	169.6		220.6	146.5		
	SD	26.88	22.03	22.03	17.59	13.02		17.59	13.02	46.79	33.14		46.79	23.07		
	Minimum	90.0	91.0	91.0	95.0	86.0		95.0	86.0	150.0	135.0		150.0	115.0		
	Maximum	140.50	132.00	132.00	121.00	112.00		121.00	112.00	217.00	165.00		217.00	139.50		
CK	Number of subjects	20	20	20	20	20	IU/L	20	20	19	19	IU/L	20	20	20	
	Mean	106.1	80.3	80.3	69.9	70.3		69.9	70.3	95.0	80.1		95.0	66.6		
	SD	49.76	22.87	22.87	27.29	41.15		27.29	41.15	22.53	19.92		22.53	18.65		
	Minimum	54.0	46.0	46.0	38.0	26.0		38.0	26.0	62.0	53.0		62.0	36.0		
	Maximum	90.50	72.50	72.50	64.00	60.00		64.00	60.00	92.00	73.00		92.00	65.50		
Maximum	277.0	129.0	129.0	171.0	195.0		171.0	195.0	142.0	126.0		142.0	114.0			

14.2 Summary Statistics for Laboratory Values (continued)

Blood Biochemistry

Parameter	Japanese		Chinese		Korean		Caucasian	
	-1 Hour	48 Hours	-1 Hour	48 Hours	-1 Hour	48 Hours	-1 Hour	48 Hours
Na	Unit							
	Number of subjects							
	Mean	141.4	141.4	140.5	140.3	139.6	140.7	138.9
	SD	0.88	1.04	1.40	1.17	2.06	0.99	1.73
	Minimum	140.0	139.0	138.2	138.3	133.0	139.0	135.0
K	Unit							
	Number of subjects							
	Mean	4.2	4.1	4.0	4.1	4.1	4.0	4.5
	SD	0.28	0.26	0.31	0.30	0.20	0.21	0.31
	Minimum	3.6	3.6	3.5	3.7	3.7	3.5	3.9
Cl	Unit							
	Number of subjects							
	Mean	103.3	104.4	105.1	103.9	105.3	104.7	100.5
	SD	1.62	1.50	1.76	1.32	2.11	1.97	1.99
	Minimum	101.0	102.0	102.2	101.7	100.0	101.0	96.0
Blood Sugar	Unit							
	Number of subjects							
	Mean	96.1	92.0	5.3	5.1	77.5	83.0	86.0
	SD	6.55	3.76	0.34	0.26	5.18	4.74	6.55
	Minimum	85.0	84.0	4.7	4.6	69.0	72.0	75.0
CRP	Unit							
	Number of subjects							
	Mean	0.3	0.2	0.5	0.9	0.0	0.1	1.5
	SD	0.29	0.14	0.63	2.39	0.08	0.07	1.66
	Minimum	0.1	0.1	0.1	0.1	0.0	0.0	0.1
Maximum	Median	0.10	0.20	0.30	0.23	0.00	0.00	0.70
	Maximum	0.6	0.3	2.4	11.0	0.3	0.2	6.0

14.2 Summary Statistics for Laboratory Values (continued)

Urinalysis

Parameter	Japanese		Chinese		Korean		Caucasian	
	-1 Hour	48 Hours	-1 Hour	48 Hours	-1 Hour	48 Hours	-1 Hour	48 Hours
pH								
Number of subjects	20	20	20	20	19	19	20	20
Mean	6.4	6.1	5.8	5.7	5.7	5.8	6.1	6.3
SD	0.32	0.28	0.44	0.47	0.31	0.34	0.21	0.38
Minimum	6.0	6.0	5.0	5.0	5.5	5.5	6.0	6.0
Median	6.50	6.00	5.50	5.50	5.50	6.00	6.00	6.00
Maximum	7.0	7.0	7.0	6.5	6.5	6.5	6.5	7.0

14.3 Shift Tables for Urinalysis Parameters

Parameter	Japanese						Chinese						Korean						Caucasian						
	48 Hours		-	+	++	+++	48 Hours		-	+	++	+++	48 Hours		-	+	++	+++	48 Hours		-	+	++	+++	
	Negative	(+)					Negative	(+)					Negative	(+)					Negative	(+)					Negative
Urine Protein	-1 Hour	14	6	0	0	0	0	-1 Hour	19	0	0	0	-1 Hour	18	1	0	0	0	-1 Hour	17	3	0	0	0	0
	Negative							Negative					Negative						Negative						
	(+)	0	0	0	0	0	0	(+)	1	0	0	0	(+)	0	0	0	0	0	(+)	0	0	0	0	0	0
	+	0	0	0	0	0	0	+	0	0	0	0	+	0	0	0	0	0	+	0	0	0	0	0	0
Urine Glucose	-1 Hour	20	0	0	0	0	0	-1 Hour	20	0	0	0	-1 Hour	19	0	0	0	0	-1 Hour	20	0	0	0	0	0
	Negative							Negative					Negative						Negative						
	(+)	0	0	0	0	0	0	(+)	0	0	0	0	(+)	0	0	0	0	0	(+)	0	0	0	0	0	0
	+	0	0	0	0	0	0	+	0	0	0	0	+	0	0	0	0	0	+	0	0	0	0	0	0
Bilirubin	-1 Hour	20	0	0	0	0	0	-1 Hour	20	0	0	0	-1 Hour	19	0	0	0	0	-1 Hour	18	0	0	0	0	0
	Negative							Negative					Negative						Negative						
	(+)	0	0	0	0	0	0	(+)	0	0	0	0	(+)	0	0	0	0	0	(+)	0	0	0	0	0	0
	+	0	0	0	0	0	0	+	0	0	0	0	+	0	0	0	0	0	+	0	0	0	0	0	0
Ketone bodies	-1 Hour	20	0	0	0	0	0	-1 Hour	20	0	0	0	-1 Hour	18	0	0	0	0	-1 Hour	15	1	0	0	0	0
	Negative							Negative					Negative						Negative						
	(+)	0	0	0	0	0	0	(+)	0	0	0	0	(+)	0	0	0	0	0	(+)	0	0	0	0	0	0
	+	0	0	0	0	0	0	+	0	0	0	0	+	0	0	0	0	0	+	0	0	0	0	0	0
Occult Blood	-1 Hour	19	0	0	0	0	0	-1 Hour	19	0	0	0	-1 Hour	18	1	0	0	0	-1 Hour	19	0	0	0	0	0
	Negative							Negative					Negative						Negative						
	(+)	1	0	0	0	0	0	(+)	1	0	0	0	(+)	0	0	0	0	0	(+)	0	0	0	0	0	0
	+	0	0	0	0	0	0	+	0	0	0	0	+	0	0	0	0	0	+	0	0	0	0	0	0
Urobilinogen	-1 Hour	0	0	0	0	0	0	-1 Hour	20	0	0	0	-1 Hour	0	0	0	0	0	-1 Hour	20	0	0	0	0	0
	Negative							Negative					Negative						Negative						
	(+)	0	20	0	0	0	0	(+)	0	0	0	0	(+)	1	18	0	0	0	(+)	0	0	0	0	0	0
	+	0	0	0	0	0	0	+	0	0	0	0	+	0	0	0	0	0	+	0	0	0	0	0	0
Urinary Sediment	-1 Hour	9	0	5	0	0	0	-1 Hour	20	0	0	0	-1 Hour	19	0	0	0	0	-1 Hour	20	0	0	0	0	0
	Negative							Negative					Negative						Negative						
	(+)	0	0	0	0	0	0	(+)	0	0	0	0	(+)	0	0	0	0	0	(+)	0	0	0	0	0	0
	+	4	0	2	0	0	0	+	0	0	0	0	+	0	0	0	0	0	+	0	0	0	0	0	0

14.4 Summary Statistics for Creatinine clearance

Race	Parameter	Number of Subjects	Creatinine clearance [mL/min] at -1 Hour				
			Mean	SD	Minimum	Median	Maximum
Japanese	Creatinine clearance	20	122.1	19.26	96.0	120.00	169.0
Chinese	Creatinine clearance	20	118.7	17.49	84.0	123.00	144.0
Korean	Creatinine clearance	19	125.8	20.62	93.0	122.00	169.0
Caucasian	Creatinine clearance	20	101.5	14.36	77.0	103.00	130.0

14.5 Summary Statistics for Vital Signs

Race	Parameter	Scheduled Time	Number of Subjects	Mean	SD	Minimum	Median	Maximum
Japanese	SBP [mmHg]	-1 Hour	20	103.7	8.27	88.0	102.50	119.0
		3 Hours	20	97.3	9.49	77.0	97.50	113.0
		24 Hours	20	101.8	9.30	82.0	100.50	120.0
		48 Hours	20	101.2	10.40	85.0	100.50	119.0
	DBP [mmHg]	-1 Hour	20	61.4	8.56	49.0	60.00	76.0
		3 Hours	20	54.4	7.30	41.0	53.00	67.0
		24 Hours	20	58.5	6.86	47.0	57.50	74.0
		48 Hours	20	58.6	8.22	46.0	58.50	77.0
	Pulse rate [bpm]	-1 Hour	20	65.1	4.96	57.0	64.00	74.0
		3 Hours	20	67.7	7.13	57.0	66.50	81.0
		24 Hours	20	68.0	7.15	51.0	69.50	81.0
		48 Hours	20	66.2	7.66	53.0	66.00	82.0
	Body temperature [°C]	-1 Hour	20	36.2	0.35	35.3	36.20	36.6
		3 Hours	20	36.5	0.29	35.7	36.55	36.9
		24 Hours	20	36.3	0.38	35.4	36.30	36.8
		48 Hours	20	36.4	0.20	36.0	36.40	36.8
Chinese	SBP [mmHg]	-1 Hour	20	116.5	9.71	100.0	119.00	134.0
		3 Hours	20	113.3	8.97	100.0	110.00	130.0
		24 Hours	20	110.9	11.19	94.0	110.00	138.0
		48 Hours	20	114.7	11.23	96.0	115.00	136.0
	DBP [mmHg]	-1 Hour	20	74.7	10.31	60.0	79.00	88.0
		3 Hours	20	75.1	8.98	60.0	78.00	86.0
		24 Hours	20	73.0	8.84	60.0	72.00	86.0
		48 Hours	20	77.1	7.38	60.0	78.00	88.0
	Pulse rate [bpm]	-1 Hour	20	62.5	6.42	52.0	62.00	78.0
		3 Hours	20	58.6	5.35	52.0	57.00	68.0
		24 Hours	20	59.3	4.17	52.0	60.00	68.0
		48 Hours	20	64.3	7.71	50.0	62.00	80.0
	Body temperature [°C]	-1 Hour	20	36.0	0.20	35.8	36.00	36.4
		3 Hours	20	36.1	0.32	35.8	36.00	37.0
		24 Hours	20	36.0	0.22	35.8	36.00	36.5
		48 Hours	20	36.2	0.24	35.9	36.10	36.7