

Appendix 5

Necropsy Findings

B120716

Animal: 50305

Sex: Female

Material: Thiopental, P092

Dose: 150 mg/kg

Removal Reason: Scheduled Sacrifice

Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 10401	Sex: Male
Material: Thiopental, P092	Dose: 500 mg/kg
	Removal Reason: Scheduled Sacrifice
	Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 10402	Sex: Male
Material: Thiopental, P092	Dose: 500 mg/kg
	Removal Reason: Scheduled Sacrifice
	Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 10403	Sex: Male
Material: Thiopental, P092	Dose: 500 mg/kg
	Removal Reason: Scheduled Sacrifice
	Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 10404

Sex: Male

Material: Thiopental, P092

Dose: 500 mg/kg

Removal Reason: Scheduled Sacrifice

Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 10405	Sex: Male
Material: Thiopental, P092	Dose: 500 mg/kg
	Removal Reason: Scheduled Sacrifice
	Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 50401	Sex: Female
Material: Thiopental, P092	Dose: 500 mg/kg
	Removal Reason: Scheduled Sacrifice
	Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 50402	Sex: Female
Material: Thiopental, P092	Dose: 500 mg/kg
	Removal Reason: Scheduled Sacrifice
	Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 50403

Sex: Female

Material: Thiopental, P092

Dose: 500 mg/kg

Removal Reason: Scheduled Sacrifice

Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 50404	Sex: Female
Material: Thiopental, P092	Dose: 500 mg/kg
	Removal Reason: Scheduled Sacrifice
	Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Appendix 5

Necropsy Findings

B120716

Animal: 50405

Sex: Female

Material: Thiopental, P092

Dose: 500 mg/kg

Removal Reason: Scheduled Sacrifice

Study Day (Week) of Death: 15 (3)

Gross Pathology Observations [Correlation]:

Necropsy Findings : Absent

Date	Calibration curve			Back ground (% of LLOQ)			QC sample			
	Concentration (ng/mL)		%RE	Sample	Peak area		Concentration (ng/mL)			
	Nominal	Measured			P092	IS	Nominal	No.	Measured	%RE
2012/11/28	5	5.10	2.0	Blank	0	8861	10	1	10.6	6.0
	10	10.3	3.0		(0.0)	(3.0)		2	8.81	-11.9
	20	18.7	-6.5	Zero	0	289930	50	1	50.7	1.4
	50	47.2	-5.6		(0.0)	--		2	42.1	-15.8
	100	105	5.0	LLOQ	5020	295893	800	1	771	-3.6
	200	209	4.5					2	720	-10.0
	500	486	-2.8							
	1000	1000	0.0							
2012/12/6	5	4.69	-6.2	Blank	0	8921	10	1	9.09	-9.1
	10	9.73	-2.7		(0.0)	(3.1)		2	8.74	-12.6
	20	19.9	-0.5	Zero	0	288930	50	1	44.1	-11.8
	50	53.4	6.8		(0.0)	--		2	45.7	-8.6
	100	106	6.0	LLOQ	3473	287198	800	1	757	-5.4
	200	191	-4.5					2	713	-10.9
	500	510	2.0							
	1000	990	-1.0							

RE: Relative error, LLOQ: Lower limit of quantification

%RE = (measured value - nominal value)/nominal value × 100

The value in parenthesis denotes peak area ratio (%) of back ground peak area to that in LLOQ.

--: Not applicable

<Male>

Dose (mg/kg)	Animal No.	Time (h)	Reanalysis (ng/mL)	Initial analysis (ng/mL)	RE* (%)
50	20201	4	12.1	12.6	-4.0
	20202	4	12.4	13.6	-9.2
	20203	24	8.10	10.1	-22.0
150	20301	4	33.4	38.0	-12.9
		24	13.9	14.0	-0.7
	20302	1	34.7	37.8	-8.6
		24	10.1	10.2	-1.0
	20303	24	15.6	13.9	11.5
500	20401	0.5	128	126	1.6
		8	14.5	14.2	2.1
	20402	2	58.5	58.3	0.3
		8	14.4	14.7	-2.1

<Female>

Dose (mg/kg)	Animal No.	Time (h)	Reanalysis (ng/mL)	Initial analysis (ng/mL)	RE* (%)
50	60201	1	15.4	20.4	-27.9
		4	10.2	11.9	-15.4
	60203	1	29.2	39.0	-28.7
		2	17.8	21.1	-17.0
150	60301	1	17.7	20.8	-16.1
	60302	1	28.3	30.4	-7.2
		8	9.17	10.3	-11.6
	60303	4	17.0	17.6	-3.5
500	60401	1	46.5	46.7	-0.4
		24	10.7	10.9	-1.9
	60402	1	30.4	31.2	-2.6
		24	13.5	14.0	-3.6

*: RE (%)= 100 × (Reanalysis value-Initial analysis value)/[(Reanalysis value+Initial analysis value)/2]

Day numbers relative to Start Date

Group	Sex	Animal	Clinical Sign	Site	1	1	1	1	1	1	2
					Before dosing	Time 10	Time 20	Time 30	Time 40	Time 50	a.m.
5	m	20101	No Abnormalities Detected		X	X	X	X	X	X	X
		20102	No Abnormalities Detected		X	X	X	X	X	X	X
		20103	No Abnormalities Detected		X	X	X	X	X	X	X

Severity Codes: X = Present

Group 5 - 0 mg/kg Group 6 - 50 mg/kg Group 7 - 150 mg/kg Group 8 - 500 mg/kg

Time 10 : About 0.5 hours after dosing; Time 20 : About 1 hours after dosing; Time 30 : 2 hours after dosing; Time 40 : About 4 hours after dosing;

Day numbers relative to Start Date

Group	Sex	Animal	Clinical Sign	Site	1	1	1	1	1	1	2
					Before dosing	Time 10	Time 20	Time 30	Time 40	Time 50	a.m.
6	m	20201	No Abnormalities Detected		X	X	X	X	X	X	X
		20202	No Abnormalities Detected		X	X	X	X	X	X	X
		20203	No Abnormalities Detected		X	X	X	X	X	X	X

Severity Codes: X = Present

Group 5 - 0 mg/kg Group 6 - 50 mg/kg Group 7 - 150 mg/kg Group 8 - 500 mg/kg

Time 10 : About 0.5 hours after dosing; Time 20 : About 1 hours after dosing; Time 30 : 2 hours after dosing; Time 40 : About 4 hours after dosing;

Day numbers relative to Start Date

Group	Sex	Animal	Clinical Sign	Site	Day numbers relative to Start Date						
					1 Before dosing	1 Time 10	1 Time 20	1 Time 30	1 Time 40	1 Time 50	2 a.m.
7	m	20301	No Abnormalities Detected		X	X	X	X	.	X	X
			Diarrhea		X	.	.
		20302	No Abnormalities Detected		X	X	X	X	X	.	X
			Diarrhea		X	.
		20303	No Abnormalities Detected		X	X	X	X	X	X	X

Severity Codes: X = Present

Group 5 - 0 mg/kg Group 6 - 50 mg/kg Group 7 - 150 mg/kg Group 8 - 500 mg/kg

Time 10 : About 0.5 hours after dosing; Time 20 : About 1 hours after dosing; Time 30 : 2 hours after dosing; Time 40 : About 4 hours after dosing;

Day numbers relative to Start Date

Group	Sex	Animal	Clinical Sign	Site	1	1	1	1	1	1	2	
					Before dosing	Time 10	Time 20	Time 30	Time 40	Time 50	a.m.	
8	m	20401	No Abnormalities Detected		X	X	X	.	X	.	X	
			Diarrhea		.	.	.	X	.	X	.	
		20402	No Abnormalities Detected		X	X	X	X
			Diarrhea		.	.	.	X	X	X	X	.
		20403	No Abnormalities Detected		X	X	X	X	X	X	.	X
			Diarrhea		X	.

Severity Codes: X = Present

Group 5 - 0 mg/kg Group 6 - 50 mg/kg Group 7 - 150 mg/kg Group 8 - 500 mg/kg

Time 10 : About 0.5 hours after dosing; Time 20 : About 1 hours after dosing; Time 30 : 2 hours after dosing; Time 40 : About 4 hours after dosing;

Day numbers relative to Start Date

Group	Sex	Animal	Clinical Sign	Site	1	1	1	1	1	1	2
					Before dosing	Time 10	Time 20	Time 30	Time 40	Time 50	a.m.
5	f	60101	No Abnormalities Detected		X	X	X	X	X	X	X
		60102	No Abnormalities Detected		X	X	X	X	X	X	X
		60103	No Abnormalities Detected		X	X	X	X	X	X	X

Severity Codes: X = Present

Group 5 - 0 mg/kg Group 6 - 50 mg/kg Group 7 - 150 mg/kg Group 8 - 500 mg/kg

Time 10 : About 0.5 hours after dosing; Time 20 : About 1 hours after dosing; Time 30 : 2 hours after dosing; Time 40 : About 4 hours after dosing;

Day numbers relative to Start Date

Group	Sex	Animal	Clinical Sign	Site	1	1	1	1	1	1	2
					Before dosing	Time 10	Time 20	Time 30	Time 40	Time 50	a.m.
6	f	60201	No Abnormalities Detected		X	X	X	X	X	X	X
		60202	No Abnormalities Detected		X	X	X	X	X	X	X
		60203	No Abnormalities Detected		X	X	X	X	X	X	X

Severity Codes: X = Present

Group 5 - 0 mg/kg Group 6 - 50 mg/kg Group 7 - 150 mg/kg Group 8 - 500 mg/kg

Time 10 : About 0.5 hours after dosing; Time 20 : About 1 hours after dosing; Time 30 : 2 hours after dosing; Time 40 : About 4 hours after dosing;

Day numbers relative to Start Date

Group	Sex	Animal	Clinical Sign	Site	1	1	1	1	1	1	2
					Before dosing	Time 10	Time 20	Time 30	Time 40	Time 50	a.m.
7	f	60301	No Abnormalities Detected		X	X	X	X	X	.	X
			Diarrhea		X	.
		60302	No Abnormalities Detected		X	X	X	X	X	.	X
			Diarrhea		X	.
		60303	No Abnormalities Detected		X	X	X	X	X	X	X

Severity Codes: X = Present

Group 5 - 0 mg/kg Group 6 - 50 mg/kg Group 7 - 150 mg/kg Group 8 - 500 mg/kg

Time 10 : About 0.5 hours after dosing; Time 20 : About 1 hours after dosing; Time 30 : 2 hours after dosing; Time 40 : About 4 hours after dosing;