

Table 8 Relative Organ Weight - Summary

Male

Test Article	Thymus		Salivary Glands		Lungs		Heart		
	x10 ⁻³ % Day 11	Day 15	% Day 11	Day 15	% Day 11	Day 15	% Day 11	Day 15	
P092 0 mg/kg/day	Mean	192.28		0.204		0.438		0.416	
	S.D.	29.90		0.017		0.022		0.026	
	n	0	5	0	5	0	5	0	5
P092 20 mg/kg/day	Mean		253.36		0.204		0.466		0.392
	S.D.		31.55		0.011		0.018		0.033
	n	0	5	0	5	0	5	0	5
P092 80 mg/kg/day	Mean		223.34		0.204		0.492 **		0.420
	S.D.		47.06		0.019		0.022		0.043
	n	0	5	0	5	0	5	0	5
P092 250 mg/kg/day	Mean	46.80		0.150		0.580		0.370	
	S.D.								
	n	1	0	1	0	1	0	1	0

Significantly different from control : **,P<0.01.

Table 8 Relative Organ Weight - Summary

Male

Test Article	Liver		Spleen		Kidneys		Adrenals		
	% Day 11	Day 15	% Day 11	Day 15	% Day 11	Day 15	x10 ⁻³ % Day 11	Day 15	
P092 0 mg/kg/day	Mean		3.110		0.270		0.848		19.64
	S.D.		0.123		0.051		0.035		0.75
	n	0	5	0	5	0	5	0	5
P092 20 mg/kg/day	Mean		3.132		0.254		0.810		21.28
	S.D.		0.076		0.042		0.073		2.00
	n	0	5	0	5	0	5	0	5
P092 80 mg/kg/day	Mean		3.456 **		0.190 *		0.842		19.70
	S.D.		0.113		0.016		0.036		2.18
	n	0	5	0	5	0	5	0	5
P092 250 mg/kg/day	Mean	4.860		0.110		0.850		34.00	
	S.D.								
	n	1	0	1	0	1	0	1	0

Significantly different from control : *,P<0.05; **,P<0.01.

Table 8 Relative Organ Weight - Summary

Male

Test Article	Dose	Testes		Prostate		Epididymides	
		% Day 11	Day 15	% Day 11	Day 15	% Day 11	Day 15
P092 0 mg/kg/day	Mean		1.028		0.156		0.206
	S.D.		0.109		0.043		0.032
	n	0	5	0	5	0	5
P092 20 mg/kg/day	Mean		1.042		0.134		0.220
	S.D.		0.079		0.026		0.025
	n	0	5	0	5	0	5
P092 80 mg/kg/day	Mean		1.236 **		0.124		0.250
	S.D.		0.086		0.013		0.038
	n	0	5	0	5	0	5
P092 250 mg/kg/day	Mean	1.230		0.080		0.170	
	S.D.						
	n	1	0	1	0	1	0

Significantly different from control : **,p<0.01.

Table 8 Relative Organ Weight - Summary

Female

Test Article	Final Body Weight		Brain		Pituitary		Thyroids		
	Dose	g	%	x10 ⁻³ %	x10 ⁻³ %				
	Day 11	Day 15	Day 11	Day 15	Day 11	Day 15	Day 11	Day 15	
P092 0 mg/kg/day	Mean		177.0		1.052		5.70		10.04
	S.D.		12.0		0.049		0.69		1.96
	n	0	5	0	5	0	5	0	5
P092 20 mg/kg/day	Mean		182.4		1.014		6.14		9.84
	S.D.		12.4		0.060		0.32		1.56
	n	0	5	0	5	0	5	0	5
P092 80 mg/kg/day	Mean		166.4		1.116		6.00		9.00
	S.D.		12.4		0.102		0.78		1.13
	n	0	5	0	5	0	5	0	5
P092 250 mg/kg/day	Mean	125.2		1.313		5.75		9.80	
	S.D.	13.4		0.170		0.76		0.91	
	n	5	0	4	0	4	0	4	0

Table 8 Relative Organ Weight - Summary

Female

Test Article	Thymus		Salivary Glands		Lungs		Heart	
		x10 ⁻³ %		%		%		%
Dose	Day 11	Day 15	Day 11	Day 15	Day 11	Day 15	Day 11	Day 15
P092	Mean	297.58		0.210		0.522		0.404
0 mg/kg/day	S.D.	28.98		0.016		0.034		0.011
	n	0	5	0	5	0	5	0
P092	Mean	280.78		0.210		0.520		0.432
20 mg/kg/day	S.D.	91.14		0.012		0.022		0.015
	n	0	5	0	5	0	5	0
P092	Mean	251.80		0.212		0.546		0.396
80 mg/kg/day	S.D.	64.92		0.025		0.027		0.031
	n	0	5	0	5	0	5	0
P092	Mean	30.78		0.130		0.648		0.395
250 mg/kg/day	S.D.	8.21		0.016		0.048		0.044
	n	4	0	4	0	4	0	4

Table 8 Relative Organ Weight - Summary

Female

Test Article	Liver		Spleen		Kidneys		Adrenals	
	% Day 11	Day 15	% Day 11	Day 15	% Day 11	Day 15	x10 ⁻³ % Day 11	Day 15
P092 0 mg/kg/day	Mean	3.108		0.220		0.838		31.16
	S.D.	0.101		0.023		0.045		2.04
	n	0	5	0	5	0	5	0
P092 20 mg/kg/day	Mean	3.300		0.226		0.872		31.72
	S.D.	0.073		0.029		0.068		3.39
	n	0	5	0	5	0	5	0
P092 80 mg/kg/day	Mean	3.610 **		0.224		0.910		36.34 *
	S.D.	0.202		0.036		0.067		3.78
	n	0	5	0	5	0	5	0
P092 250 mg/kg/day	Mean	4.603		0.083		0.835		51.33
	S.D.	0.563		0.033		0.077		11.77
	n	4	0	4	0	4	0	4

Significantly different from control : *,P<0.05; **,P<0.01.

Table 8 Relative Organ Weight - Summary

Female

Test Article	Ovaries		Uterus		
	Dose	x10 ⁻³ % Day 11	Day 15	% Day 11	Day 15
P092 0 mg/kg/day	Mean		45.60		0.292
	S.D.		6.34		0.163
	n	0	5	0	5
P092 20 mg/kg/day	Mean		48.90		0.262
	S.D.		3.75		0.138
	n	0	5	0	5
P092 80 mg/kg/day	Mean		52.94		0.356
	S.D.		4.48		0.196
	n	0	5	0	5
P092 250 mg/kg/day	Mean	35.18		0.135	
	S.D.	2.98		0.021	
	n	4	0	4	0

Table 9 Necropsy Findings - Summary

Scheduled sacrifice (Day 11, 15)

Organ Findings	Sex	Male				Female			
	Test Article	P092							
	Dose	0	20	80	250	0	20	80	250
	Dose Unit	mg/kg/day							
	Number of Animals	5	5	5	1	5	5	5	4
Number of Animals Examined	<5>	<5>	<5>	<1>	<5>	<5>	<5>	<4>	
Thymus									
Small		0	0	0	1	0	0	0	4
Spleen									
Small		0	0	0	1	0	0	0	3
Stomach									
Dilatation		0	0	0	0	0	0	0	1
Small intestine, duodenum									
Dilatation		0	0	0	0	0	0	0	1
Small intestine, jejunum									
Dilatation		0	0	0	0	0	0	0	1
Small intestine, ileum									
Dilatation		0	0	0	0	0	0	0	1
Large intestine, cecum									
Dilatation		0	0	0	0	0	0	0	1
Large intestine, colon									
Dilatation		0	0	0	0	0	0	0	1
Liver									
Nodule		0	0	0	0	0	0	0	1

Table 9 Necropsy Findings - Summary

Scheduled sacrifice (Day 11, 15)

Organ	Sex	Male				Female			
	Test Article	P092							
Findings	Dose	0	20	80	250	0	20	80	250
	Dose Unit	mg/kg/day							
	Number of Animals	5	5	5	1	5	5	5	4
	Number of Animals Examined	<5>	<5>	<5>	<1>	<5>	<5>	<5>	<4>
Kidney									
Dilatation, pelvis		0	1	0	0	0	0	0	0
Testis									
Small		1	0	0	0				
Epididymis									
Small		1	0	0	0				
Seminal vesicle									
Small		0	0	0	1				
Prostate									
Small		0	0	0	1				
Skin									
Erosion		0	0	0	0	0	0	0	2
Fur loss		0	0	0	0	0	0	0	2

Table 9 Necropsy Findings - Summary

Organ Findings	Sex	Death	
		Male	Female
	Test Article	P092	P092
	Dose	250	250
	Dose Unit	mg/kg/day	mg/kg/day
	Number of Animals	4	1
	Number of Animals Examined	<4>	<1>
Thymus			
Small		3	0
Spleen			
Small		4	1
Stomach			
Dark reddish patch, glandular stomach, mucosa		3	1
Small intestine, duodenum			
Dilatation		1	0
Small intestine, jejunum			
Dilatation		1	0
Small intestine, ileum			
Dilatation		1	0
Large intestine, cecum			
Dilatation		4	1
Large intestine, colon			
Dilatation		1	0
Seminal vesicle			
Small		2	
Prostate			
Small		2	

Appendix 1 - 1 Clinical Sign P092 0 mg/kg/day Male

Animal	Day	1	2	3	4	5	6	7	8	9	10	11	12	13
Number	Findings	Time	10	20	10	20	10	20	10	20	10	20	10	20
10101	No Abnormality													
10102	No Abnormality													
10103	No Abnormality													
10104	No Abnormality													
10105	No Abnormality													

Animal	Day	14	15		
Number	Findings	Time	10	20	am
10101	No Abnormality				
10102	No Abnormality				
10103	No Abnormality				
10104	No Abnormality				
10105	No Abnormality				

+, Present; 1, Slight; 2, Moderate; 3, Severe;
 Time 10, Before dosing; Time 20, About 2 hours after dosing;

Appendix 1 - 2 Clinical Sign P092 20 mg/kg/day Male

Animal		Day	1	2	3	4	5	6	7	8	9	10	11	12	13	
Number	Findings	Time	10	20	10	20	10	20	10	20	10	20	10	20	10	20
10201	No Abnormality															
10202	No Abnormality															
10203	No Abnormality															
10204	No Abnormality															
10205	No Abnormality															

Animal		Day	14	15
Number	Findings	Time	10	20 am
10201	No Abnormality			
10202	No Abnormality			
10203	No Abnormality			
10204	No Abnormality			
10205	No Abnormality			

+, Present; 1, Slight; 2, Moderate; 3, Severe;
 Time 10, Before dosing; Time 20, About 2 hours after dosing;

Appendix 1 - 3 Clinical Sign P092 80 mg/kg/day Male

Animal	Day	1	2	3	4	5	6	7	8	9	10	11	12	13
Number	Findings	Time	10	20	10	20	10	20	10	20	10	20	10	20
10301	No Abnormality													
10302	No Abnormality													
10303	No Abnormality													
10304	No Abnormality													
10305	No Abnormality													

Animal	Day	14	15		
Number	Findings	Time	10	20	am
10301	No Abnormality				
10302	No Abnormality				
10303	No Abnormality				
10304	No Abnormality				
10305	No Abnormality				

+, Present; 1, Slight; 2, Moderate; 3, Severe;
 Time 10, Before dosing; Time 20, About 2 hours after dosing;

Appendix 1 - 5 Clinical Sign P092 0 mg/kg/day Female

Animal	Day	1	2	3	4	5	6	7	8	9	10	11	12	13						
Number	Findings	Time	10	20	10	20	10	20	10	20	10	20	10	20	10	20	10	20	10	20
50101	No Abnormality																			
50102	No Abnormality																			
50103	No Abnormality																			
50104	No Abnormality																			
50105	No Abnormality																			

Animal	Day	14	15		
Number	Findings	Time	10	20	am
50101	No Abnormality				
50102	No Abnormality				
50103	No Abnormality				
50104	No Abnormality				
50105	No Abnormality				

+ , Present; 1 , Slight; 2 , Moderate; 3 , Severe;
Time 10 , Before dosing; Time 20 , About 2 hours after dosing;

Appendix 1 - 6 Clinical Sign P092 20 mg/kg/day Female

Animal	Day	1	2	3	4	5	6	7	8	9	10	11	12	13
Number	Findings	Time	10	20	10	20	10	20	10	20	10	20	10	20
50201	No Abnormality													
50202	No Abnormality													
50203	No Abnormality													
50204	No Abnormality													
50205	No Abnormality													

Animal	Day	14	15		
Number	Findings	Time	10	20	am
50201	No Abnormality				
50202	No Abnormality				
50203	No Abnormality				
50204	No Abnormality				
50205	No Abnormality				

+, Present; 1, Slight; 2, Moderate; 3, Severe;
 Time 10, Before dosing; Time 20, About 2 hours after dosing;

Appendix 1 - 7 Clinical Sign P092 80 mg/kg/day Female

Animal	Day	1	2	3	4	5	6	7	8	9	10	11	12	13
Number	Findings	Time	10	20	10	20	10	20	10	20	10	20	10	20
50301	No Abnormality													
50302	No Abnormality													
50303	No Abnormality													
50304	No Abnormality													
50305	No Abnormality													

Animal	Day	14	15		
Number	Findings	Time	10	20	am
50301	No Abnormality				
50302	No Abnormality				
50303	No Abnormality				
50304	No Abnormality				
50305	No Abnormality				

+ , Present; 1 , Slight; 2 , Moderate; 3 , Severe;
 Time 10 , Before dosing; Time 20 , About 2 hours after dosing;

Appendix 1 - 8 Clinical Sign P092 250 mg/kg/day Female

Animal Number	Findings	Day Time	1		2		3		4		5		6		7		8		9		10		11		12		13		
			10	20	10	20	10	20	10	20	10	20	10	20	10	20	10	20	10	20	10	20	10	20	10	20	10	20	
50401	No Abnormality																												
50402	Death																			+									
50403	No Abnormality																												
50404	Emaciation																			+	+	+	+	+					
	Loss of fur	08																		+	+	+	+	+					
	Erosion	08																				+	+	+					
50405	Emaciation																					+	+	+					
	Loss of fur	07																					+	+	+				
	Loss of fur	08																		+	+	+	+	+					
	Erosion	07																							+				

Animal Number	Findings	Day Time	14		15	
			10	20	am	
50401	Death					
50402	Death					
50403	Death					
50404	Emaciation					
	Loss of fur	08				
	Erosion	08				
50405	Emaciation					
	Loss of fur	07				
	Loss of fur	08				
	Erosion	07				

+, Present; 1, Slight; 2, Moderate; 3, Severe;

Time 10, Before dosing; Time 20, About 2 hours after dosing;

07, Chest; 08, Abdomen;

Appendix 2 - 1 Body Weight P092 0 mg/kg/day Male Unit : g

Animal Number	Day	1	4	8	11	14
10101		186	217	252	288	308
10102		174	198	217	236	253
10103		175	202	235	261	279
10104		164	185	217	242	260
10105		183	212	245	272	289

Appendix 2 - 2 Body Weight P092 20 mg/kg/day Male Unit : g

Animal Number	Day	1	4	8	11	14
10201		186	216	245	280	293
10202		172	198	224	256	272
10203		178	205	239	261	282
10204		170	191	224	250	267
10205		177	201	230	256	275