

Table 1.

-continued Clinical observation

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg	Finding Part	Day of experiment																					
		7----->			8----->			9----->			10----->			11----->			12----->						
		1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	1	2	3				
01 control 0	normal	-	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
		Total	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
02 2-NIBE 20	normal	-	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
		Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
	Scratched wound Neck	-	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
		+	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5		
03 2-NIBE 100	normal	-	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
		Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5

1: Before dosing    2: 30 to 60 minutes after dosing    3: 3 to 4 hours after dosing  
 2-NIBE: 2-Naphthylisobutyl ether

Table 1. -continued Clinical observation

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg	Finding Part	Day of experiment																		
		7----->			8----->			9----->			10----->			11----->			12----->			
		1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	
04 2-NIBE 500	normal	-	6	6	7	7	6	7	6	3	7	4	4	6	6	5	7	7	4	7
	Total		6	6	7	7	6	7	6	3	7	4	4	6	6	5	7	7	4	7
	Soiled fur Anogenital region	-	7	7	7	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		Total		8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Reddish tear	-	8	8	8	8	8	8	8	8	7	8	8	8	8	8	8	8	8	7
		Total		8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Smudge of perinasal area	-	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		Total		8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Salivation	-	8	7	8	8	6	7	8	3	7	8	4	6	8	6	7	8	4	7
		Total		8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Loose stool	-	7	8	8	7	8	8	6	8	8	4	8	8	6	7	8	7	8	8
		Total		8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Watery diarrhea	-	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		Total		8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Mucous feces	-	7	8	8	7	8	8	6	8	8	5	8	8	8	8	8	7	8	8
		Total		8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8

1: Before dosing    2: 30 to 60 minutes after dosing    3: 3 to 4 hours after dosing  
 2-NIBE: 2-Naphthylisobutyl ether

Table 1.

-continued Clinical observation

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg	Finding Part	Day of experiment																		
		13----->			14----->			15----->			16----->			17----->			18----->			
		1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	
01 control 0	normal	-	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
		Total	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
02 2-NIBE 20	normal	-	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
		Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
	Scratched wound Neck	-	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
		+	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5		
03 2-NIBE 100	normal	-	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
		Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5

1: Before dosing      2: 30 to 60 minutes after dosing      3: 3 to 4 hours after dosing  
 2-NIBE: 2-Naphthylisobutyl ether

Table 1. -continued Clinical observation

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg	Finding Part	Day of experiment																		
		13----->			14----->			15----->			16----->			17----->			18----->			
		1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	
04 2-NIBE 500	normal	-	4	5	5	2	5	5	4	4	5	3	4	6	0	5	6	1	5	4
		Total	4	5	5	2	5	5	4	4	5	3	4	6	0	5	6	1	5	4
	Soiled fur Anogenital region	-	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		+	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Reddish tear	-	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		+	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Smudge of perinasal area	-	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		+	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	
Salivation	-	8	5	5	8	5	5	8	4	5	8	4	6	8	5	6	8	5	4	
	+	0	3	3	0	3	3	0	4	3	0	4	2	0	3	2	0	3	4	
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	
Loose stool	-	4	8	8	4	8	8	4	8	8	4	8	8	1	8	8	1	8	8	
	+	4	0	0	4	0	0	4	0	0	4	0	0	7	0	0	7	0	0	
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	
Watery diarrhea	-	8	8	8	8	8	8	8	8	8	7	8	8	7	8	8	8	8	8	
	+	0	0	0	0	0	0	0	0	0	1	0	0	1	0	0	0	0	0	
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	
Mucous feces	-	6	8	8	4	8	8	5	8	8	5	8	8	1	8	8	2	8	8	
	+	2	0	0	4	0	0	3	0	0	3	0	0	7	0	0	6	0	0	
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	

1: Before dosing    2: 30 to 60 minutes after dosing    3: 3 to 4 hours after dosing  
 2-NIBE: 2-Naphthylisobutyl ether

Table 1.

-continued Clinical observation

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg	Finding Part	Day of experiment																		
		19----->			20----->			21----->			22----->			23----->			24----->			
		1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	
01 control 0	normal	-	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
		Total	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
02 2-NIBE 20	normal	-	5	5	5	5	5	5	4	4	4	4	4	4	4	4	4	4	4	4
		Total	5	5	5	5	5	5	4	4	4	4	4	4	4	4	4	4	4	4
	Scratched wound Neck	-	5	5	5	5	5	5	4	4	4	4	4	4	4	4	4	4	4	4
		+	0	0	0	0	0	0	1	1	1	1	1	1	1	1	1	1	1	1
Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5		
03 2-NIBE 100	normal	-	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5		

1: Before dosing    2: 30 to 60 minutes after dosing    3: 3 to 4 hours after dosing  
 2-NIBE: 2-Naphthylisobutyl ether

Table 1. -continued Clinical observation

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg	Finding Part	Day of experiment																		
		19----->			20----->			21----->			22----->			23----->			24----->			
		1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	1	2	3	
04 2-NIBE 500	normal	-	1	5	5	3	5	7	3	3	7	5	4	7	1	4	8	4	5	8
	Total		1	5	5	3	5	7	3	3	7	5	4	7	1	4	8	4	5	8
	Soiled fur Anogenital region	-	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Reddish tear	-	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Smudge of perinasal area	-	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Salivation	-	8	5	5	8	5	7	8	3	7	8	4	7	8	4	8	8	5	8
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Loose stool	-	1	8	8	3	8	8	3	8	8	5	8	8	1	8	8	4	8	8
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Watery diarrhea	-	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8
	Mucous feces	-	2	8	8	4	8	8	3	8	8	5	8	8	4	8	8	5	8	8
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8

1: Before dosing    2: 30 to 60 minutes after dosing    3: 3 to 4 hours after dosing  
 2-NIBE: 2-Naphthylisobutyl ether

Table 1.

-continued Clinical observation

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg	Finding Part	Day of experiment													
		25----->			26----->			27----->			28----->			29	
		1	2	3	1	2	3	1	2	3	1	2	3		
01 control 0	normal	-	10	10	10	10	10	10	10	10	10	10	10	10	10
		Total	10	10	10	10	10	10	10	10	10	10	10	10	10
02 2-NIBE 20	normal	-	4	4	4	4	4	4	4	4	4	4	4	4	5
		Total	4	4	4	4	4	4	4	4	4	4	4	4	5
	Scratched wound Neck	-	4	4	4	4	4	4	4	4	4	4	4	4	5
		+	1	1	1	1	1	1	1	1	1	1	1	1	0
		Total	5	5	5	5	5	5	5	5	5	5	5	5	5
03 2-NIBE 100	normal	-	5	5	5	5	5	5	5	5	5	5	5	5	5
		Total	5	5	5	5	5	5	5	5	5	5	5	5	5

1: Before dosing    2: 30 to 60 minutes after dosing    3: 3 to 4 hours after dosing  
 2-NIBE: 2-Naphthylisobutyl ether

Table 1. -continued Clinical observation

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg	Finding Part	Day of experiment													
		25----->			26----->			27----->			28----->			29	
		1	2	3	1	2	3	1	2	3	1	2	3		
04 2-NIBE 500	normal	-	4	5	8	2	4	7	1	6	8	1	4	8	2
		Total	4	5	8	2	4	7	1	6	8	1	4	8	2
	Soiled fur	-	8	8	8	8	8	8	8	8	8	8	8	8	8
	Anogenital region	+	0	0	0	0	0	0	0	0	0	0	0	0	0
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8
	Reddish tear	-	8	8	8	8	8	8	8	8	8	8	8	8	8
		+	0	0	0	0	0	0	0	0	0	0	0	0	0
		Total	8	8	8	8	8	8	8	8	8	8	8	8	8
	Smudge of perinasal area	-	8	8	8	8	8	8	8	8	8	8	8	8	8
		+	0	0	0	0	0	0	0	0	0	0	0	0	0
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	
Salivation	-	8	5	8	8	4	7	8	6	8	8	4	8	8	
	+	0	3	0	0	4	1	0	2	0	0	4	0	0	
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	
Loose stool	-	4	8	8	2	8	8	1	8	8	1	8	8	2	
	+	4	0	0	6	0	0	7	0	0	7	0	0	6	
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	
Watery diarrhea	-	8	8	8	8	8	8	8	8	8	8	8	8	8	
	+	0	0	0	0	0	0	0	0	0	0	0	0	0	
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	
Mucous feces	-	5	8	8	3	8	8	1	8	8	4	8	8	3	
	+	3	0	0	5	0	0	7	0	0	4	0	0	5	
	Total	8	8	8	8	8	8	8	8	8	8	8	8	8	

1: Before dosing    2: 30 to 60 minutes after dosing    3: 3 to 4 hours after dosing  
 2-NIBE: 2-Naphthylisobutyl ether



Table 1.

-continued Clinical observation

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg	Finding Part	Day of experiment														
		29	30	31	32	33	34	35	36	37	38	39	40	41	42	43
01 control 0	normal	-	5	5	5	5	5	5	5	5	5	5	5	5	5	5
		Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5
04 2-NIBE 500	normal	-	1	5	5	5	5	5	5	5	5	5	5	5	5	5
		Total	1	5	5	5	5	5	5	5	5	5	5	5	5	5
	Loose stool	-	1	5	5	5	5	5	5	5	5	5	5	5	5	5
		+	4	0	0	0	0	0	0	0	0	0	0	0	0	0
	Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5	
	Mucous feces	-	1	5	5	5	5	5	5	5	5	5	5	5	5	5
+		4	0	0	0	0	0	0	0	0	0	0	0	0	0	
Total	5	5	5	5	5	5	5	5	5	5	5	5	5	5		

2-NIBE: 2-Naphthylisobutyl ether

Table 2.

## Body weight

Exp.No. 9933 (115-212)

Sex: Male

Group No. Dose mg/kg		Day of experiment										Gain 1-28	Unit:g
		1	4	8	11	15	18	22	25	28			
01 control 0	N	10	10	10	10	10	10	10	10	10	10	10	
	Mean	142	162	191	212	240	265	291	312	330	188		
	S.D.	5	6	14	13	17	16	19	21	22	21		
02 2-NIBE 20	N	5	5	5	5	5	5	5	5	5	5		
	Mean	144	163	195	217	249	272	300	319	339	195		
	S.D.	6	7	6	7	7	10	9	11	14	9		
03 2-NIBE 100	N	5	5	5	5	5	5	5	5	5	5		
	Mean	145	164	195	217	245	264	289	305	319	174		
	S.D.	6	6	7	9	12	16	21	25	27	31		
04 2-NIBE 500	N	10	10	10	10	10	10	10	10	10	10		
	Mean	143	130##	164**	180**	202**	216**	230**	243**	256**	113**		
	S.D.	6	15	14	17	22	24	31	30	34	30		

Significantly different from 01 group ## P  $\lll$  0.01 (Steel)  
 Significantly different from 01 group \*\* P  $\lll$  0.01 (Dunnett)  
 2-NIBE: 2-Naphthylisobutyl ether

Table 2. -continued Body weight

Exp.No. 9933 (115-212)

Sex: Male

Group No. Dose	mg/kg	N	Day of experiment					Gain 29-42	Unit:g
			29	32	36	39	42		
01 control	0	N	5	5	5	5	5	5	
		Mean	333	346	357	367	375	41	
		S.D.	19	20	20	22	19	7	
04 2-NIBE	500	N	5	5	5	5	5	5	
		Mean	252**	271**	293*	307*	317*	65**	
		S.D.	43	44	47	46	43	9	

Significantly different from 01 group \* P ≤ 0.05 , \*\* P ≤ 0.01 (Dunnett)  
2-NIBE: 2-Naphthylisobutyl ether

Table 2.

-continued Body weight

Exp.No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg		Day of experiment									Gain 1-28	Unit:g
		1	4	8	11	15	18	22	25	28		
01 control 0	N	10	10	10	10	10	10	10	10	10	10	10
	Mean	116	129	142	153	161	167	178	184	191	75	
	S.D.	4	4	6	6	10	10	11	12	11	11	
02 2-NIBE 20	N	5	5	5	5	5	5	5	5	5	5	
	Mean	114	126	140	153	162	169	181	187	193	79	
	S.D.	4	6	9	10	12	11	12	15	13	11	
03 2-NIBE 100	N	5	5	5	5	5	5	5	5	5	5	
	Mean	117	130	149	160	171	178	188	193	201	84	
	S.D.	4	6	8	10	11	13	15	14	15	12	
04 2-NIBE 500	N	10	10	8	8	8	8	8	8	8	8	
	Mean	116	105**	132*	141*	154	164	173	178	188	73	
	S.D.	4	8	7	11	10	10	14	15	19	18	

Significantly different from 01 group \* P  $\leq$  0.05 , \*\* P  $\leq$  0.01 (Dunnett)

2-NIBE: 2-Naphthylisobutyl ether

Table 2. -continued Body weight

Exp.No. 9933 (115-212)

Sex: Female

Group No. Dose	mg/kg	Day of experiment					Gain 29-42	Unit:g
		29	32	36	39	42		
01 control	N	5	5	5	5	5	5	
	Mean	191	199	204	206	209	19	
	S.D.	16	16	13	12	15	4	
04 2-NIBE	N	5	5	5	5	5	5	
	Mean	191	199	206	210	215	25	
	S.D.	20	22	21	22	22	5	

2-NIBE: 2-Naphthylisobutyl ether

Table 3.

## Food consumption

Exp. No. 9933 (115-212)

Sex: Male

Group No. Dose mg/kg		Day of experiment						Unit:g/animal/day
		1 => 8	8 => 15	15 => 22	22 => 28	29 => 36	36 => 42	
01 control 0	N	10	10	10	10	5	5	
	Mean	18	19	20	19	24	23	
	S.D.	1	1	1	1	1	1	
02 2-NIBE 20	N	5	5	5	5			
	Mean	19	21#	21#	20			
	S.D.	1	1	1	1			
03 2-NIBE 100	N	5	5	5	5			
	Mean	18	20	20	19			
	S.D.	1	2	2	2			
04 2-NIBE 500	N	10	10	10	10	5	5	
	Mean	13##	19	17#	16*	29**	25	
	S.D.	3	3	4	3	3	2	

Significantly different from 01 group # P  $\lll$  0.05, ## P  $\lll$  0.01 (Steel)  
 Significantly different from 01 group \* P  $\lll$  0.05, \*\* P  $\lll$  0.01 (Dunnett)  
 2-NIBE: 2-Naphthylisobutyl ether

Table 3.

-continued Food consumption

Exp. No. 9933 (115-212)

Sex: Female

Group No. Dose mg/kg		Day of experiment						Unit:g/animal/day
		1 => 8	8 => 15	15 => 22	22 => 28	29 => 36	36 => 42	
01 control 0	N	10	10	10	10	5	5	
	Mean	14	14	14	13	18	18	
	S.D.	1	1	1	1	1	1	
02 2-NIBE 20	N	5	5	5	5			
	Mean	14	15	14	13			
	S.D.	3	1	2	1			
03 2-NIBE 100	N	5	5	5	5			
	Mean	15	15	15	13			
	S.D.	1	1	2	1			
04 2-NIBE 500	N	8	8	8	8	5	5	
	Mean	11##	14	14	14	23##	20	
	S.D.	2	2	2	2	3	4	

Significantly different from 01 group ## P  $\leq$  0.01 (Steel)  
 2-NIBE: 2-Naphthylisobutyl ether

Table 4. Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Male

Signs	Dose level (mg/kg)	Administration period (Week)					Recovery period (Week)			
		B. G.	1	2	3	4	1	2		
No. of animals	0	10	10	10	10	10	5	5		
	20	5	5	5	5	5	-	-		
	100	5	5	5	5	5	-	-		
	500	10	10	10	10	10	5	5		
REMOVAL FROM CAGE Ease of removal	Very easy	0	8	8	10	8	8	5	5	
		20	4	4	5	5	3	-	-	
		100	5	3	5	4	4	-	-	
		500	5	6	9	10	9	5	5	
	Easy	0	2	2	0	2	2	0	0	
		20	1	1	0	0	2	-	-	
		100	0	2	0	1	1	-	-	
		500	5	4	1	0	1	0	0	
	Vocalization	None	0	10	10	10	10	10	5	5
			20	5	5	5	5	5	-	-
			100	5	5	5	5	5	-	-
			500	10	10	10	10	10	5	5

B. G. : Before grouping



Table 4. -continued Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Male

Signs	Dose level (mg/kg)	B. G.	Administration period (Week)				Recovery period (Week)		
			1	2	3	4	1	2	
HANDLING OBSERVATIONS									
Muscle tone	Normal	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Subnormal temperature	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Piloerection	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Staining hair	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Unkempt hair	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Skin color	Normal	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Lacrimation	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5

B. G. : Before grouping

Table 4. -continued Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Male

Signs	Dose level (mg/kg)	B. G.	Administration period (Week)				Recovery period (Week)		
			1	2	3	4	1	2	
Exophthalmos	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Pupillary size	Normal	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Salivation	Absent	0	10	10	10	10	10	5	5
		20	5	5	4	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	8	3	2	5	5
	Slight	0	0	0	0	0	0	0	0
		20	0	0	1	0	0	-	-
		100	0	0	0	0	0	-	-
		500	0	0	2	7	6	0	0
	Marked	0	0	0	0	0	0	0	0
		20	0	0	0	0	0	-	-
		100	0	0	0	0	0	-	-
		500	0	0	0	0	2	0	0

B. G. : Before grouping

Table 4. -continued Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Male

Signs	Dose level (mg/kg)	B. G.	Administration period (Week)				Recovery period (Week)		
			1	2	3	4	1	2	
OBSERVATIONS IN ARENA									
Posture	Normal	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Motor activity	Normal	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Respiration	Normal	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Lid closure	Wide open	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Gait	Normal	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	4	5	5	5	-	-
		500	10	10	10	10	10	5	5
	Indifference immobile	0	0	0	0	0	0	0	0
		20	0	0	0	0	0	-	-
		100	0	1	0	0	0	-	-
		500	0	0	0	0	0	0	0
Tremor	None	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5

B. G. : Before grouping

Table 4. -continued Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Male

Signs	Dose level (mg/kg)	B. G.	Administration period (Week)				Recovery period (Week)		
			1	2	3	4	1	2	
Twitch	None	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Convulsion	None	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Stereotypic behavior	None	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5
Abnormal behavior	None	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	10	10	10	10	5	5

B. G. : Before grouping

Table 4. -continued Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Male

Signs	Dose level (mg/kg)	Administration period (Week)				Recovery period (Week)			
		B. G.	1	2	3	4	1	2	
<b>SENSORIMOTOR FUNCTION</b>									
Approach contact	No reaction	0				0		0	
		20				0		-	
		100				0		-	
		500				1		0	
	Normal a)	0				10		5	
		20				5		-	
		100				5		-	
		500				9		5	
	Touch response	No reaction	0				0		0
			20				0		-
			100				0		-
			500				1		0
Moderate reaction		0				10		5	
		20				5		-	
		100				5		-	
		500				8		5	
Freezing		0				0		0	
		20				0		-	
		100				0		-	
		500				1		0	
Pinna response	Normal	0				10		5	
		20				5		-	
		100				5		-	
		500				10		5	

B. G. : Before grouping

a) Rat slowly approaches and sniffs at object or turns away

Table 4. -continued Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Male

Signs	Dose level (mg/kg)	Administration period (Week)				Recovery period (Week)		
		B. G.	1	2	3	4	1	2
Pain response(Tail pinch)	Slowly turns	0				1	0	
		20				0	-	
		100				0	-	
		500				0	0	
	Normal	0				9	5	
		20				5	-	
		100				5	-	
		500				10	5	
Pupillary reflex	Normal	0				10	5	
		20				5	-	
		100				5	-	
		500				10	5	
Air righting reflex	Normal	0				10	5	
		20				5	-	
		100				5	-	
		500				10	5	

B. G. : Before grouping

Table 4. -continued Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Female

Signs	Dose level (mg/kg)	B. G.	Administration period (Week)				Recovery period (Week)		
			1	2	3	4	1	2	
No. of animals	0	10	10	10	10	10	5	5	
	20	5	5	5	5	5	-	-	
	100	5	5	5	5	5	-	-	
	500	10	9	8	8	8	5	5	
REMOVAL FROM CAGE									
Ease of removal	Very easy	0	8	10	7	10	10	5	5
		20	3	5	3	5	5	-	-
		100	4	5	4	5	5	-	-
		500	9	7	6	7	8	5	5
	Easy	0	2	0	3	0	0	0	0
		20	2	0	2	0	0	-	-
		100	1	0	1	0	0	-	-
		500	1	1	2	1	0	0	0
	Moderately difficult	0	0	0	0	0	0	0	0
		20	0	0	0	0	0	-	-
		100	0	0	0	0	0	-	-
		500	0	1	0	0	0	0	0
Vocalization	None	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	9	8	8	8	5	5

B. G. : Before grouping

Table 4. -continued Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Female

Signs	Dose level (mg/kg)	B. G.	Administration period (Week)				Recovery period (Week)		
			1	2	3	4	1	2	
<b>HANDLING OBSERVATIONS</b>									
Muscle tone	Normal	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	9	8	8	8	5	5
Subnormal temperature	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	9	8	8	8	5	5
Piloerection	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	9	8	8	8	5	5
Staining hair	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	7	8	8	8	5	5
	Very soiled	0	0	0	0	0	0	0	0
		20	0	0	0	0	0	-	-
		100	0	0	0	0	0	-	-
		500	0	2	0	0	0	0	0
Unkempt hair	Absent	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	9	8	8	8	5	5
Skin color	Normal	0	10	10	10	10	10	5	5
		20	5	5	5	5	5	-	-
		100	5	5	5	5	5	-	-
		500	10	9	8	8	8	5	5

B. G. : Before grouping



Table 4. -continued Detailed clinical observations and sensory reactivity to stimuli of different types

Exp. No. 9933 (115-212)

Sex : Female

Signs	Dose level (mg/kg)	Administration period (Week)					Recovery period (Week)	
		B. G.	1	2	3	4	1	2
Lacrimation	Absent	0	10	10	10	10	5	5
		20	5	5	5	5	-	-
		100	5	5	5	5	-	-
		500	10	9	8	8	5	5
Exophthalmos	Absent	0	10	10	10	10	5	5
		20	5	5	5	5	-	-
		100	5	5	5	5	-	-
		500	10	9	8	8	5	5
Pupillary size	Normal	0	10	10	10	10	5	5
		20	5	5	5	5	-	-
		100	5	5	5	5	-	-
		500	10	9	8	8	5	5
Salivation	Absent	0	10	10	10	10	5	5
		20	5	5	5	5	-	-
		100	5	5	5	5	-	-
		500	10	8	5	5	6	5
	Slight	0	0	0	0	0	0	0
		20	0	0	0	0	0	-
		100	0	0	0	0	0	-
		500	0	1	3	3	2	0

B. G. : Before grouping