プロピオノニトリルのマウスを用いた吸入による2週間毒性試験報告書

試験番号:0447

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(DEAD AND MORIBUND ANIMALS)

TABLE 1
SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES OF MALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE

	0pp	om	6	3.3ppm	ı	1:	2.5ppn	า	:	25ppm			50ppm		1	00ppn	า
Week-Day on Study	Av.Wt.	No.of Surviv. 5>	Av.Wt.	% of cont. < 5>	No.of Surviv.	Av.Wt.	% of cont. < 5>	No.of Surviv.	Av.Wt.	% of cont. < 5>	No.of Surviv.	Av.Wt.	% of cont. < 5>	No.of Surviv.	Av.Wt.	% of cont. < 5>	No.of Surviv.
0-0	23.3 (5)	5/5	23.3 (5)	100	5/5	23.3 (5)	100	5/5	23.4 (5)	100	5/5	23.2 (5)	100	5/5	23.2 (5)	100	5/5
1-2	23.9 (5)	5/5	24.1 (5)	101	5/5	23.5 (5)	98	5/5	23.8 (5)	100	5/5	23.9 (5)	100	5/5	22.0 ( 2)	92	2/5
1-4	23.9 (5)	5/5	24.6 (5)	103	5/5	23.6 (5)	99	5/5	24.0 (5)	100	5/5	24.1 (5)	101	5/5	23.2 ( 2)	97	2/5
1-7	24.4 (5)	5/5	25.3 (5)	104	5/5	23.8 (5)	98	5/5	24.2 (5)	99	5/5	24.1 (5)	99	5/5	23.9 (2)	98	2/5
2-3	24.8 (5)	5/5	25.2 (5)	102	5/5	24.5 (5)	99	5/5	24.8 (5)	100	5/5	24.8 (5)	100	5/5	24.9 (2)	100	2/5
2-7	25.4 (5)	5/5	25.9 (5)	102	5/5	24.7 (5)	97	5/5	25.3 (5)	100	5/5	24.6 (5)	97	5/5	25.0 (2)	98	2/5

TABLE 2 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES OF FEMALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE

	lq0	pm	(	6.3ppm	ı	1	2.5ppn	n	2	25ppm	1		50ppm		10	00ppm	1
Week-Day on Study	Av.Wt.	No.of Surviv. 5>	Av.Wt.	% of cont. < 5>	No.of Surviv.	Av.Wt.	% of cont. < 5>	No.of Surviv.	Av.Wt.	% of cont. < 5>	No.of Surviv.	Av.Wt.	% of cont. < 5>	No.of Surviv.	Av.Wt.	% of cont. < 5>	No.of Surviv.
0-0	18.3 (5)	5/5	18.3 ( 5)	100	5/5	18.3 ( 5)	100	5/5	18.3 ( 5)	100	5/5	18.3 ( 5)	100	5/5	18.3 ( 5)	100	5/ 5
1-2	18.3 (5)	5/5	18.3 (5)	100	5/5	17.9 (5)	98	5/5	18.6 (5)	102	5/5	17.8 (5)	97	5/5	- ( -)	_	0/5
1-4	18.5 (5)	5/5	18.9 (5)	102	5/5	18.6 (5)	101	5/5	19.4 (5)	105	5/5	18.7 (5)	101	5/5	- ( -)	_	0/5
1-7	18.7 (5)	5/5	19.4 (5)	104	5/5	18.7 (5)	100	5/5	19.6 (5)	105	5/5	18.8 (5)	101	5/5	- ( -)	_	0/5
2-3	19.5 (5)	5/5	19.2 (5)	98	5/5	19.4 (5)	99	5/5	19.9 (5)	102	5/5	20.1 (5)	103	5/5	- ( -)	_	0/5
2-7	19.8 (5)	5/5	20.0 (5)	101	5/5	19.9 (5)	101	5/5	20.7 (5)	105	5/5	20.2 (5)	102	5/5	- ( -)	_	0/5

TABLE 3 FOOD CONSUMPTION CHANGES OF MALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE

	0ppm	6.3ppm	12.5ppm	25ppm	50ppm	100ppm
Week-Day on Study	Av.Fc. No.of Surviv.	Av.Fc. % of No.of cont. Surviv. < 5>	Av.Fc. % of No.of cont. Surviv. < 5>	Av.Fc. % of No.of cont. Surviv. < 5>	Av.Fc. % of No.of cont. Surviv. < 5>	Av.Fc. % of No.of cont. Surviv.
1-7 2-7	4.4 ( 5) 5/ 5 4.4 ( 5) 5/ 5	4.6 ( 5) 105 5/5 4.5 ( 5) 102 5/5	4.3 (5) 98 5/5 4.3 (5) 98 5/5	4.5 ( 5) 102 5/5 4.6 ( 5) 105 5/5	4.1 (5) 93 5/5 4.4 (5) 100 5/5	3.7 ( 2) 84 2/5 4.7 ( 2) 107 2/5
	< > : No.of 6	effective animals, ( ): N	o.of measured animals	Av.Fc. : Average food	consumption (Unit : g).	

TABLE 4 FOOD CONSUMPTION CHANGES OF FEMALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE

	0ppm	6.3ppm	12.5ppm	25ppm	50ppm	100ppm
Week-Day on Study	Av.Fc. No.of Surviv. < 5>	Av.Fc. % of No.of cont. Surviv. < 5>	Av.Fc. % of No.of cont. Surviv. < 5>	Av.Fc. % of No.of cont. Surviv. < 5>	Av.Fc. % of No.of cont. Surviv. < 5>	Av.Fc. % of No.of cont. Surviv. < 5>
1-7 2-7	3.4 ( 5) 5/5 3.6 ( 5) 5/5	3.8 ( 5) 112 5/5 3.8 ( 5) 106 5/5	3.4 ( 5) 100 5/5 3.8 ( 5) 106 5/5	3.6 ( 5) 106 5/5 3.8 ( 5) 106 5/5	3.5 ( 5) 103 5/ 5 4.0 ( 5) 111 5/ 5	- ( -) - 0/5 - ( -) - 0/5
	< > : No.of 6	effective animals, ( ): N	o.of measured animals	Av.Fc. : Average food	consumption (Unit : g).	

TABLE 5 HEMATOLOGY OF MALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE

Group Name	Control	6.3 ppm	12.5 ppm	25 ppm	50 ppm	100 ppm
No. of examined animals	4	4	5	5	5	2
Platelet $(10^3/\mu\mathrm{L})$	$1412~\pm~41$	$1356~\pm~48$	$1336~\pm~69$	1215 ± 160 *	1220 ± 90 *	1460 ?

Mean  $\pm$  S.D.

TABLE 6 HEMATOLOGY OF FEMALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE

Group Name	Control	6.3 ppm	12.5 ppm	25 ppm	50 ppm	100 ppm
No. of examined animals	4	5	5	5	5	0
Red blood cell $(10^6/\mu$ L)	$10.98 \pm 0.40$	$10.75 \pm 0.40$	$10.82 \pm 0.39$	$11.01 \pm 0.30$	$10.36 \pm 0.38$	_
Hemoglobin (g/dL)	$17.0~\pm~0.6$	$16.4~\pm~0.5$	$16.8~\pm~0.4$	$16.9~\pm~0.4$	$16.0~\pm~0.6$	_
Hematocrit (%)	$50.3~\pm~1.8$	$49.1 \pm 1.4$	$49.2 ~\pm~ 0.8$	$50.2 \pm 1.0$	47.6 ± 1.4 *	_

Mean  $\pm$  S.D.

 $<sup>^{*)}</sup>$  Significant difference, p<0.05 (Test of Dunnett)

 $<sup>^{\</sup>star\star)}$  Significant difference, p<0.01 (Test of Dunnett)

<sup>?)</sup> The test of Dunnett could not be applied because number of data was less than three.

<sup>\*)</sup> Significant difference, p<0.05 (Test of Dunnett)

 $<sup>^{**)}</sup>$  Significant difference, p<0.01 (Test of Dunnett)

TABLE 7 BIOCHEMISTRY OF MALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE

Group Name	Control	6.3 ppm	12.5 ppm	25 ppm	50 ppm	100 ppm
No. of examined animals	4	5	5	5	5	2
Total protein (g/dL)	$5.3~\pm~0.1$	$5.2~\pm~0.1$	$5.2~\pm~0.2$	$5.1~\pm~0.2$	$5.0~\pm~0.2$	4.6 ?
Albumin (g/dL)	$3.4~\pm~0.2$	$3.2~\pm~0.1$	$3.3 \pm 0.1$	$3.1~\pm~0.2$	$3.1~\pm~0.2$	2.8 ?
Triglyceride (mg/dL)	$20 \pm 8$	$14 \pm 5$	10 ± 3 *	$13 \pm 5$	11 ± 3 *	14 ?

Mean  $\pm$  S.D.

<sup>\*)</sup> Significant difference, p<0.05 (Test of Dunnett)

<sup>\*\*)</sup> Significant difference, p<0.01 (Test of Dunnett)

<sup>?)</sup> The test of Dunnett could not be applied because number of data was less than three.

TABLE 8 ORGAN WEIGHTS OF MALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE

Group Name	Control	6.3 ppm	12.5 ppm	25 ppm	50 ppm	100 ppm	
No. of examined animals	5	5	5	5	5	2	
Body weight (g)	$21.2~\pm~0.6$	$21.7~\pm~0.9$	$20.8~\pm~0.7$	$21.1~\pm~0.4$	$21.1 \pm 1.1$	21.7	?
Thymus (g)	$0.044 \pm 0.004$	$0.046 \pm 0.003$	$0.041 \pm 0.007$	$0.039 \pm 0.009$	$0.041 \pm 0.004$	0.022	?
Thymus (%)	$0.207 \pm 0.016$	$0.212 \pm 0.019$	$0.196 \pm 0.030$	$0.184 \pm 0.041$	$0.193 \pm 0.022$	0.099	?
Liver (g)	$0.987 \pm 0.037$	$1.017 \pm 0.051$	$0.994 \pm 0.057$	$1.024 \pm 0.043$	$0.977 \pm 0.063$	1.148	?
Liver (%)	$4.650 \pm 0.143$	$4.687 \pm 0.065$	$4.789 \pm 0.238$	$4.848 \pm 0.162$	$4.636 \pm 0.171$	5.302	?

Mean  $\pm$  S.D.

TABLE 9 ORGAN WEIGHTS OF FEMALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE

Group Name	Control	6.3 ppm	12.5 ppm	25 ppm	50 ppm	100 ppm
No. of examined animals	5	5	5	5	5	0
ody weight (g)	$16.2~\pm~0.4$	$16.4~\pm~0.8$	$16.1~\pm~0.5$	$16.7~\pm~0.9$	$16.5~\pm~0.6$	_
iver (g)	$0.739 \pm 0.022$	$0.747 \pm 0.014$	$0.764 \pm 0.009$	0.801 ± 0.033 *	0.800 ± 0.042 *	_
iver (%)	$4.577 \pm 0.121$	$4.552 \pm 0.144$	$4.753 \pm 0.169$	$4.810 \pm 0.176$	$4.838 \pm 0.173$	_

Mean  $\pm$  S.D.

<sup>\*)</sup> Significant difference, p<0.05 (Test of Dunnett)

<sup>\*\*)</sup> Significant difference, p<0.01 (Test of Dunnett)

 $<sup>^{?)}</sup>$  The test of Dunnett could not be applied because number of data was less than three.

<sup>\*)</sup> Significant difference, p<0.05 (Test of Dunnett)

 $<sup>^{**)}</sup>$  Significant difference, p<0.01 (Test of Dunnett)

TABLE 10 INCIDENCES OF SELECTED LESIONS OF MALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE (DEAD AND MORIBUND ANIMALS)

Group			100 ppm
Number of examine	d animals		3
Organ Findings		Grade	
Nasal cavity Congestion		1+	3
Lung Congestion		1+	3
Grade	1+: Slight		

TABLE 11 INCIDENCES OF SELECTED LESIONS OF FEMALE MICE IN THE 2-WEEK INHALATION STUDY OF PROPIONONITRILE (DEAD AND MORIBUND ANIMALS)

Group		100 ppm
Number of examined animals		5
Organ Findings	Grade	
Nasal cavity		
Congestion	1+	2
Lung		
Congestion	1+	2
Hemorrhage	1+	2
Grade 1+: Slight		