

1 - ブロモブタンのラットを用いた
吸入による2週間毒性試験報告書

試験番号：0480

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(SACRIFICED ANIMALS)

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

Week-Day on Study	Control		500ppm			1000ppm			2000ppm			4000ppm			8000ppm		
	Av.Wt.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.
	< 5>		< 5>			< 5>			< 5>			< 5>			< 5>		
0-0	118 (5)	5/5	119 (5)	101	5/5	119 (5)	101	5/5	119 (5)	101	5/5	119 (5)	101	5/5	119 (5)	101	5/5
1-2	125 (5)	5/5	121 (5)	97	5/5	120 (5)	96	5/5	115 (5)	92	5/5	- (-)	-	0/5	- (-)	-	0/5
1-4	129 (5)	5/5	126 (5)	98	5/5	123 (5)	95	5/5	95 (5)	74	5/5	- (-)	-	0/5	- (-)	-	0/5
1-7	137 (5)	5/5	131 (5)	96	5/5	128 (5)	93	5/5	82 (4)	60	4/5	- (-)	-	0/5	- (-)	-	0/5
2-3	144 (5)	5/5	140 (5)	97	5/5	133 (5)	92	5/5	- (-)	-	0/5	- (-)	-	0/5	- (-)	-	0/5
2-7	156 (5)	5/5	152 (5)	97	5/5	142 (5)	91	5/5	- (-)	-	0/5	- (-)	-	0/5	- (-)	-	0/5

< > : No.of effective animals, () : No.of measured animals Av.Wt. : Average body weight (Unit : g).

TABLE 2
SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES OF FEMALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

Week-Day on Study	Control		500ppm			1000ppm			2000ppm			4000ppm			8000ppm		
	Av.Wt.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.	Av.Wt.	% of cont.	No.of Surviv.
	< 5>		< 5>			< 5>			< 5>			< 5>			< 5>		
0-0	95 (5)	5/5	95 (5)	100	5/5	95 (5)	100	5/5	95 (5)	100	5/5	95 (5)	100	5/5	95 (5)	100	5/5
1-2	99 (5)	5/5	97 (5)	98	5/5	96 (5)	97	5/5	95 (5)	96	5/5	- (-)	-	0/5	- (-)	-	0/5
1-4	102 (5)	5/5	98 (5)	96	5/5	97 (5)	95	5/5	92 (5)	90	5/5	- (-)	-	0/5	- (-)	-	0/5
1-7	106 (5)	5/5	102 (5)	96	5/5	101 (5)	95	5/5	94 (5)	89	5/5	- (-)	-	0/5	- (-)	-	0/5
2-3	109 (5)	5/5	105 (5)	96	5/5	105 (5)	96	5/5	- (-)	-	0/5	- (-)	-	0/5	- (-)	-	0/5
2-7	115 (5)	5/5	109 (5)	95	5/5	111 (5)	97	5/5	- (-)	-	0/5	- (-)	-	0/5	- (-)	-	0/5

< > : No.of effective animals, () : No.of measured animals Av.Wt. : Average body weight (Unit : g).

TABLE 3 FOOD CONSUMPTION CHANGES OF MALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

Week-Day on Study	Control		500ppm			1000ppm			2000ppm			4000ppm			8000ppm		
	Av.Fc.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.
	< 5>		< 5>			< 5>			< 5>			< 5>			< 5>		
1-7	13.8 (5)	5/ 5	11.7 (5)	85	5/ 5	12.0 (5)	87	5/ 5	3.7 (4)	27	4/ 5	- (-)	-	0/ 5	- (-)	-	0/ 5
2-7	13.3 (5)	5/ 5	13.7 (5)	103	5/ 5	14.0 (5)	105	5/ 5	- (-)	-	0/ 5	- (-)	-	0/ 5	- (-)	-	0/ 5

< > : No.of effective animals, () : No.of measured animals Av.Fc. : Average food consumption (Unit : g).

TABLE 4 FOOD CONSUMPTION CHANGES OF FEMALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

Week-Day on Study	Control		500ppm			1000ppm			2000ppm			4000ppm			8000ppm		
	Av.Fc.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.	Av.Fc.	% of cont.	No.of Surviv.
	< 5>		< 5>			< 5>			< 5>			< 5>			< 5>		
1-7	10.5 (5)	5/ 5	10.0 (5)	95	5/ 5	10.1 (5)	96	5/ 5	8.8 (5)	84	5/ 5	- (-)	-	0/ 5	- (-)	-	0/ 5
2-7	10.0 (5)	5/ 5	9.9 (5)	99	5/ 5	11.2 (5)	112	5/ 5	- (-)	-	0/ 5	- (-)	-	0/ 5	- (-)	-	0/ 5

< > : No.of effective animals, () : No.of measured animals Av.Fc. : Average food consumption (Unit : g).

TABLE 5 HEMATOLOGY OF MALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

Group Name	Control	500 ppm	1000 ppm	2000 ppm	4000 ppm	8000 ppm
No. of examined animals	5	5	5	0	0	0
RED BLOOD CELL ($10^6/\mu\text{L}$)	8.89 \pm 0.17	8.97 \pm 0.20	9.21 \pm 0.15 *	—	—	—
HEMATOCRIT (%)	45.5 \pm 0.9	46.2 \pm 1.0	47.7 \pm 0.8 **	—	—	—
MCHC (g/dL)	34.8 \pm 0.3	34.2 \pm 0.2 **	34.2 \pm 0.2 **	—	—	—
PLATELET ($10^3/\mu\text{L}$)	805 \pm 69	973 \pm 55 **	936 \pm 35 **	—	—	—
APTT (sec)	22.2 \pm 0.4	21.3 \pm 1.5	19.5 \pm 0.5 **	—	—	—
WBC ($10^3/\mu\text{L}$)	3.43 \pm 1.02	5.69 \pm 1.13 *	5.66 \pm 1.12 *	—	—	—

Mean \pm S.D.
^{*}) Significant difference, $p < 0.05$ (Test of Dunnett)
^{**}) Significant difference, $p < 0.01$ (Test of Dunnett)

TABLE 6 HEMATOLOGY OF FEMALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

Group Name	Control	500 ppm	1000 ppm	2000 ppm	4000 ppm	8000 ppm
No. of examined animals	5	5	5	0	0	0
RETICULOCYTE (%)	1.4 \pm 0.2	1.6 \pm 0.2	2.0 \pm 0.1 **	—	—	—

Mean \pm S.D.
^{*}) Significant difference, $p < 0.05$ (Test of Dunnett)
^{**}) Significant difference, $p < 0.01$ (Test of Dunnett)

TABLE 7 BIOCHEMISTRY OF MALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

Group Name	Control	500 ppm	1000 ppm	2000 ppm	4000 ppm	8000 ppm
No. of examined animals	5	5	5	0	0	0
Total protein(g/dL)	5.7 ± 0.2	6.0 ± 0.1 **	6.2 ± 0.1 **	—	—	—
Albumin (g/dL)	3.4 ± 0.1	3.5 ± 0.1	3.7 ± 0.2 *	—	—	—
T-cholesterol(mg/dL)	49 ± 3	36 ± 4 **	32 ± 2 **	—	—	—
Triglyceride(mg/dL)	33 ± 11	14 ± 6 *	13 ± 2 *	—	—	—
Phospholipid (mg/dL)	99 ± 7	77 ± 6 **	78 ± 2 **	—	—	—
GOT(IU/L)	57 ± 9	66 ± 2 *	64 ± 4	—	—	—
ALP(IU/L)	677 ± 40	569 ± 25 **	517 ± 44 **	—	—	—
CPK(IU/L)	255 ± 47	188 ± 16 *	153 ± 27 **	—	—	—
Urea nitrogen(mg/dL)	16.5 ± 1.0	13.7 ± 1.4 *	11.5 ± 1.6 **	—	—	—
Sodium(mEq/L)	141 ± 1	141 ± 1	139 ± 1 *	—	—	—
Chloride (mEq/L)	103 ± 1	105 ± 1 **	107 ± 0 **	—	—	—
Calcium(mg/dL)	10.1 ± 0.2	10.3 ± 0.1 *	10.4 ± 0.1 **	—	—	—

Mean ± S.D.
^{*)} Significant difference, p<0.05 (Test of Dunnett)
^{**)} Significant difference, p<0.01 (Test of Dunnett)

TABLE 8 BIOCHEMISTRY OF FEMALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

Group Name	Control	500 ppm	1000 ppm	2000 ppm	4000 ppm	8000 ppm
No. of examined animals	5	5	5	0	0	0
Glucose(mg/dL)	138 ± 8	134 ± 8	161 ± 3 **	—	—	—
T-cholesterol(mg/dL)	67 ± 5	60 ± 4	56 ± 4 **	—	—	—
GOT(IU/L)	75 ± 2	70 ± 5	69 ± 4 *	—	—	—
GPT(IU/L)	32 ± 3	24 ± 1 **	27 ± 1 **	—	—	—
ALP(IU/L)	571 ± 31	479 ± 26 **	478 ± 31 **	—	—	—
CPK(IU/L)	258 ± 42	214 ± 45	158 ± 29 **	—	—	—
Urea nitrogen(mg/dL)	18.3 ± 1.6	14.0 ± 0.9 **	11.2 ± 1.0 **	—	—	—
Sodium(mEq/L)	139 ± 1	138 ± 1	137 ± 1 *	—	—	—
Chloride (mEq/L)	105 ± 1	106 ± 1	107 ± 1 **	—	—	—
Calcium(mg/dL)	9.6 ± 0.2	9.8 ± 0.2	10.1 ± 0.3 *	—	—	—

Mean ± S.D.
^{*)} Significant difference, p<0.05 (Test of Dunnett)
^{**)} Significant difference, p<0.01 (Test of Dunnett)

TABLE 9 ORGAN WEIGHTS OF MALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

Group Name	Control	500 ppm	1000 ppm	2000 ppm	4000 ppm	8000 ppm
No. of examined animals	5	5	5	0	0	0
Body weight (g)	140 ± 10	137 ± 5	127 ± 4 *	—	—	—
Thymus (g)	0.246 ± 0.035	0.238 ± 0.019	0.195 ± 0.034 *	—	—	—
Thymus (%)	0.175 ± 0.024	0.173 ± 0.016	0.154 ± 0.028	—	—	—
Adrenals (g)	0.039 ± 0.003	0.043 ± 0.005	0.045 ± 0.007	—	—	—
Adrenals (%)	0.028 ± 0.003	0.031 ± 0.003	0.035 ± 0.006 *	—	—	—
Heart (g)	0.578 ± 0.031	0.616 ± 0.020	0.618 ± 0.019 *	—	—	—
Heart (%)	0.413 ± 0.019	0.448 ± 0.006 **	0.488 ± 0.018 **	—	—	—
Lungs (g)	0.656 ± 0.037	0.697 ± 0.025	0.682 ± 0.023	—	—	—
Lungs (%)	0.469 ± 0.044	0.508 ± 0.022	0.538 ± 0.014 **	—	—	—
Kidneys (g)	1.138 ± 0.085	1.296 ± 0.066 **	1.314 ± 0.052 **	—	—	—
Kidneys (%)	0.811 ± 0.023	0.943 ± 0.024 **	1.038 ± 0.019 **	—	—	—
Spleen (g)	0.305 ± 0.032	0.301 ± 0.015	0.261 ± 0.015 *	—	—	—
Spleen (%)	0.217 ± 0.009	0.219 ± 0.005	0.206 ± 0.007	—	—	—
Liver (g)	4.089 ± 0.299	4.865 ± 0.235 **	5.177 ± 0.241 **	—	—	—
Liver (%)	2.913 ± 0.077	3.540 ± 0.085 **	4.088 ± 0.081 **	—	—	—
Brain (g)	1.699 ± 0.035	1.688 ± 0.027	1.606 ± 0.018 **	—	—	—
Brain (%)	1.217 ± 0.106	1.230 ± 0.057	1.270 ± 0.045	—	—	—

Mean ± S.D.

*) Significant difference, p<0.05 (Test of Dunnett)

**) Significant difference, p<0.01 (Test of Dunnett)

TABLE10 ORGAN WEIGHTS OF FEMALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE

Group Name	Control	500 ppm	1000 ppm	2000 ppm	4000 ppm	8000 ppm
No. of examined animals	5	5	5	0	0	0
Body weight (g)	103 ± 4	98 ± 3	99 ± 4	—	—	—
Thymus (g)	0.262 ± 0.014	0.208 ± 0.011 **	0.166 ± 0.009 **	—	—	—
Thymus (%)	0.253 ± 0.008	0.212 ± 0.016 **	0.168 ± 0.012 **	—	—	—
Adrenals (g)	0.043 ± 0.002	0.047 ± 0.004	0.054 ± 0.005 **	—	—	—
Adrenals (%)	0.042 ± 0.002	0.048 ± 0.003 *	0.055 ± 0.006 **	—	—	—
Heart (g)	0.453 ± 0.020	0.449 ± 0.006	0.491 ± 0.019 **	—	—	—
Heart (%)	0.439 ± 0.030	0.457 ± 0.014	0.496 ± 0.022 **	—	—	—
Kidneys (g)	0.869 ± 0.041	0.974 ± 0.013 **	1.028 ± 0.046 **	—	—	—
Kidneys (%)	0.841 ± 0.014	0.993 ± 0.032 **	1.036 ± 0.017 **	—	—	—
Spleen (g)	0.251 ± 0.015	0.239 ± 0.009	0.213 ± 0.013 **	—	—	—
Spleen (%)	0.243 ± 0.008	0.243 ± 0.011	0.215 ± 0.016 **	—	—	—
Liver (g)	3.089 ± 0.181	3.582 ± 0.133 **	4.242 ± 0.190 **	—	—	—
Liver (%)	2.987 ± 0.132	3.647 ± 0.044	4.276 ± 0.043 **	—	—	—

Mean ± S.D.

^{*)} Significant difference, p<0.05 (Test of Dunnett)

^{**)} Significant difference, p<0.01 (Test of Dunnett)

TABLE 11 INCIDENCES OF SELECTED LESIONS OF MALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE
(DEAD AND MORIBUND ANIMALS)

Group Name No. of Animals on Study Grade	Control 0				500ppm 0				1000ppm 0				2000ppm 5				4000ppm 5				8000ppm 5			
	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
nasal cavity	< 5>																							
disarrangement:olfactory epithelium	—				—				—				4	0	0	0	0	0	0	0	0	0	0	0
atrophy:olfactory epithelium	—				—				—				1	1	3	0	0	0	0	0	0	0	0	0
necrosis:olfactory epithelium	—				—				—				0	4	0	0	0	0	5	0	0	0	0	0
lung	< 5>																							
congestion	—				—				—				0	1	0	0	0	0	0	0	2	3	0	0
hemorrhage	—				—				—				0	0	0	0	0	0	0	0	2	0	0	0
edema: alveolus	—				—				—				0	1	0	0	0	4	1	0	5	0	0	0
edema:perivascular	—				—				—				4	1	0	0	0	5	0	0	0	5	0	0
bone marrow	< 5>																							
congestion	—				—				—				0	4	0	0	0	0	0	0	0	0	0	0
thymus	< 5>																							
atrophy	—				—				—				1	0	4	0	0	0	0	0	0	0	0	0
congestion	—				—				—				3	0	0	0	0	0	0	0	0	0	0	0
spleen	< 5>																							
atrophy	—				—				—				1	4	0	0	0	0	0	0	0	0	0	0
heart	< 5>																							
hemorrhage	—				—				—				0	0	0	0	3	2	0	0	0	0	0	0
stomach	< 5>																							
erosion:forestomach	—				—				—				1	0	0	0	0	0	0	0	0	0	0	0
liver	< 5>																							
necrosis:centeral	—				—				—				0	0	0	0	1	0	4	0	0	0	0	0

TABLE 11 INCIDENCES OF SELECTED LESIONS OF MALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE
(DEAD AND MORIBUND ANIMALS) (CONTINUE)

Group Name No. of Animals on Study Grade	Control				500ppm				1000ppm				2000ppm				4000ppm				8000ppm			
	0				0				0				5				5				5			
	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
adrenal gland																								
hemorrhage	—				—				—				< 5> 0 0 0 0				< 5> 0 1 0 0				< 5> 0 0 0 0			
testis																								
germ cell necrosis	—				—				—				< 5> 1 4 0 0				< 5> 0 0 0 0				< 5> 0 0 0 0			
epididymis																								
decreased sperma	—				—				—				< 5> 0 1 4 0				< 5> 0 0 0 0				< 5> 0 0 0 0			
debris of spermatic elements	—				—				—				2 3 0 0				0 0 0 0				0 0 0 0			
brain																								
degeneration granular cell	—				—				—				< 5> 0 2 3 0				< 5> 0 0 0 0				< 5> 0 0 0 0			
Grade	1: Slight 2: Moderate 3: Marked 4: Severe																							
< >	: Number of animals examined at the site																							
—	: All animals survived at the terminal necropsy																							

TABLE 12 INCIDENCES OF SELECTED LESIONS OF MALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE (SACRIFICED ANIMALS)

Group Name	Control				500ppm				1000ppm				2000ppm				4000ppm				8000ppm			
No. of Animals on Study	5				5				5				0				0				0			
Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
nasal cavity	< 5>				< 5>				< 5>															
disarrangement:olfactory epithelium	0	0	0	0	4	0	0	0	5	0	0	0	—				—				—			
atrophy:olfactory epithelium	0	0	0	0	0	0	0	0	1	0	0	0	—				—				—			

Grade 1: Slight 2: Moderate 3: Marked 4: Severe
 < > : Number of animals examined at the site
 — : All animals died before the terminal necropsy

TABLE 13 INCIDENCES OF SELECTED LESIONS OF FEMALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE
(DEAD AND MORIBUND ANIMALS)

Group Name No. of Animals on Study Grade	Control				500ppm				1000ppm				2000ppm				4000ppm				8000ppm			
	0				0				0				5				5				5			
	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
nasal cavity																								
thrombus	—	—	—	—	—	—	—	—	—	—	—	—	0	0	1	0	0	0	0	0	0	0	0	0
disarrangement:olfactory epithelium	—	—	—	—	—	—	—	—	—	—	—	—	3	0	0	0	0	0	0	0	0	0	0	0
atrophy:olfactory epithelium	—	—	—	—	—	—	—	—	—	—	—	—	2	3	0	0	0	0	0	0	0	0	0	0
necrosis:olfactory epithelium	—	—	—	—	—	—	—	—	—	—	—	—	5	0	0	0	0	1	4	0	0	0	0	0
lung																								
congestion	—	—	—	—	—	—	—	—	—	—	—	—	1	2	0	0	0	0	0	0	3	2	0	0
hemorrhage	—	—	—	—	—	—	—	—	—	—	—	—	0	0	0	0	0	0	0	0	0	1	0	0
edema: alveolus	—	—	—	—	—	—	—	—	—	—	—	—	3	2	0	0	0	5	0	0	2	1	0	0
thrombus	—	—	—	—	—	—	—	—	—	—	—	—	0	1	0	0	0	0	0	0	0	0	0	0
edema:perivascular	—	—	—	—	—	—	—	—	—	—	—	—	3	1	0	0	0	5	0	0	0	5	0	0
bone marrow																								
congestion	—	—	—	—	—	—	—	—	—	—	—	—	1	0	0	0	0	0	0	0	0	0	0	0
thymus																								
atrophy	—	—	—	—	—	—	—	—	—	—	—	—	3	1	0	0	0	0	0	0	0	0	0	0
congestion	—	—	—	—	—	—	—	—	—	—	—	—	0	0	0	0	1	0	0	0	0	0	0	0
spleen																								
atrophy	—	—	—	—	—	—	—	—	—	—	—	—	4	1	0	0	0	0	0	0	0	0	0	0
heart																								
hemorrhage	—	—	—	—	—	—	—	—	—	—	—	—	0	1	0	0	2	3	0	0	0	0	0	0
stomach																								
ulcer:forestomach	—	—	—	—	—	—	—	—	—	—	—	—	1	0	0	0	0	0	0	0	0	0	0	0
erosion:glandular stomach	—	—	—	—	—	—	—	—	—	—	—	—	1	0	0	0	0	0	0	0	0	0	0	0

TABLE 13 INCIDENCES OF SELECTED LESIONS OF FEMALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE
(DEAD AND MORIBUND ANIMALS) (CONTINUE)

Group Name No. of Animals on Study Grade	Control				500ppm				1000ppm				2000ppm				4000ppm				8000ppm			
	0				0				0				5				5				5			
	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
liver																								
necrosis:central	—				—				—				< 5>				< 5>				< 5>			
	—				—				—				1 0 0 0				1 0 0 0				0 0 0 0			
vacuolic change	—				—				—				0 0 0 0				5 0 0 0				0 0 0 0			
urinary bladder																								
hemorrhage	—				—				—				< 5>				< 5>				< 5>			
	—				—				—				0 1 0 0				0 0 0 0				0 0 0 0			
adrenal gland																								
hemorrhage	—				—				—				< 5>				< 5>				< 5>			
	—				—				—				0 0 0 0				0 1 2 0				0 0 0 0			
brain																								
degeneration:granular cell	—				—				—				< 5>				< 5>				< 5>			
	—				—				—				5 0 0 0				0 0 0 0				0 0 0 0			
Grade	1: Slight 2: Moderate 3: Marked 4: Severe																							
< >	: Number of animals examined at the site																							
—	: All animals survived at the terminal necropsy																							

TABLE 14 INCIDENCES OF SELECTED LESIONS OF FEMALE RATS IN THE 2-WEEK INHALATION STUDY OF 1-BROMOBUTANE (SACRIFICED ANIMALS)

Group Name	Control				500ppm				1000ppm				2000ppm				4000ppm				8000ppm			
No. of Animals on Study	5				5				5				0				0				0			
Grade	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4	1	2	3	4
nasal cavity	< 5>				< 5>				< 5>				—				—				—			
disarrangement:olfactory epithelium	0	0	0	0	5	0	0	0	5	0	0	0	—				—				—			

Grade 1: Slight 2: Moderate 3: Marked 4: Severe
 < > : Number of animals examined at the site
 — : All animals died before the terminal necropsy