

塩化メチルのラット及びマウスを用いた
吸入によるがん原性予備試験報告書

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TABLE 1 EXPERIMENTAL DESIGN AND MATERIALS AND METHODS IN THE INHALATION STUDIES OF METHYL CHLORIDE

Two-week studies	Thirteen-week Studies
<Method of Administration> Inhalation	Inhalation
<Number of Groups> Male 6, Female 6	Male 6, Female 6
<Size of Groups> 10 males and 10 females of each groups	10 males and 10 females of each groups
<Animals> Strain and Species F344/DuCrj(Fischer)rat Crj:BDF1 mouse Animal Source Charles River Japan, Inc. Quarantine and Acclimation 2 wk Age When Placed on Study 6 wk old Age When Killed 8 wk old	F344/DuCrj(Fischer)rat Crj:BDF1 mouse Charles River Japan, Inc. 2 wk 6 wk old 19 wk old
<Doses> Rat--0, 190, 380, 750, 1500 or 3000ppm Mouse--0, 190, 380, 750, 1500 or 3000ppm	Rat--0, 190, 380, 750, 1500 or 3000ppm Mouse--0, 300, 440, 670, 1000, or 1500ppm
<Duration of Dosing> 5d/wk for 2wk	5d/wk for 13wk
<Animal Maintenance> Feed CRF-1 (Oriental Yeast Co.,Ltd.) Sterilized by γ -ray Available <i>ad libitum</i> Water Sterilized by ultraviolet rays Automatic watering system Available <i>ad libitum</i> Animals per Cage Single (stainless steel wire) Animal Room Environment Barrier system Temperature: $21 \pm 2^{\circ}\text{C}$ Humidity : $60 \pm 10\%$ Fluorescent light 12h/d 15-17 room air changes /h Chamber Environment Temperature: $20 \sim 24^{\circ}\text{C}$ Humidity : $30 \sim 70\%$ Fluorescent light 12h/d 12 ± 1 room air changes /h	Same as two-week studies Same as two-week studies Same as two-week studies Single (stainless steel wire) Same as two-week studies Same as two-week studies
<Type and Frequency of Observation> Clinical sign Observed 1 \times d Body weight Weighed 0-0, 1-1, 1-7, and 2-7 (wk-d) Food Consumption Weighed 1-7, 2-7 (wk-d)	Observed 1 \times d Weighed 1 \times wk for 13wk Weighed 1 \times wk for 13wk

TABLE 1 EXPERIMENTAL DESIGN AND MATERIALS AND METHODS IN THE INHALATION STUDIES OF METHYL CHLORIDE
(Continued)

Two-week Studies	Thirteen-week Studies
<Hematology> None	Red blood cell (RBC), Hemoglobin, Hematocrit, Mean corpuscular volume (MCV), Mean corpuscular hemoglobin (MCH), Mean corpuscular hemoglobin concentration (MCHC), Platelet, White blood cell (WBC), Differential WBC.
<Blood Biochemistry> None	Total protein, Albumin, A/G ratio, T-bilirubin, Glucose, T-cholesterol, Triglyceride, Phospholipid <rat only>, Glutamic oxaloacetic transaminase (GOT), Glutamic pyruvic transaminase (GPT), Lactate dehydrogenase (LDH), Alkaline phosphatase (ALP), γ -Glutamyl transpeptidase (G-GTP) <rat only>, Creatine phosphokinase (CPK), Urea nitrogen, Creatinine <rat only>, Sodium, Potassium, Chloride, Calcium, Inorganic phosphorus.
<Urinalysis> None	pH, Protein, Glucose, Ketone body Bilirubin <rat only>, Occult blood Urobilinogen.
<Necropsy> Necropsy performed on all animals.	Same as two-week studies.
<Organ weight> None	Organ weight measurement performed on scheduled sacrificed animals. The following organs were weighed: brain, lung, liver, spleen, heart, kidney, adrenal, testis, ovary, thymus.
<Histopathologic Examination> Histopathologic examination performed on at least two animals per sex per groups. The following organs were examined: nasal cavity, trachea, lung, bone marrow, lymph node, thymus, spleen, heart, stomach, small intestine, large intestine, liver, pancreas, kidney, pituitary, adrenal, testis, ovary, brain, muscle.	Histopathologic examination performed on all animals. The following organs were examined: skin, nasal cavity trachea, lung, bone marrow, lymph node, thymus, spleen, heart, tongue, salivary gland, esophagus, stomach, small intestine, large intestine, liver, pancreas, kidney, urinary bladder, pituitary, thyroid, adrenal, testis, epididymis, seminal vesicle, prostate, ovary, uterus, vagina, mammary gland, brain, spinal cord, peripheral nerve, eye, Harderian gland, muscle, bone.

TABLE 2 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES IN MALE RAT (TWO-WEEK STUDY)

Week-Day on Study	0 ppm			190 ppm			380 ppm			750 ppm			1500 ppm			3000 ppm		
	Au.Wt.	No. of Surviv. <10>		Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.
0-0	124 (10)	10/10		124 (10)	100	10/10	124 (10)	100	10/10	122 (10)	98	10/10	124 (10)	100	10/10	124 (10)	100	10/10
1-1	127 (10)	10/10		126 (10)	99	10/10	126 (10)	99	10/10	125 (10)	98	10/10	125 (10)	98	10/10	123 (10)	97	10/10
1-7	147 (10)	10/10		143 (10)	97	10/10	145 (10)	99	10/10	139 (10)	95	10/10	140 (10)	95	10/10	132 (10)	90	10/10
2-7	168 (10)	10/10		166 (10)	99	10/10	167 (10)	99	10/10	158 (10)	94	10/10	157 (10)	93	10/10	140 (10)	83	10/10

< >:No. of effective animals, ():No. of measured animals Au.Wt.: g

TABLE 3 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES IN FEMALE RAT (TWO-WEEK STUDY)

Week-Day on Study	0 ppm			190 ppm			380 ppm			750 ppm			1500 ppm			3000 ppm		
	Au.Wt.	No. of Surviv. <10>		Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.
0-0	102 (10)	10/10		102 (10)	100	10/10	103 (10)	101	10/10	102 (10)	100	10/10	102 (10)	100	10/10	102 (10)	100	10/10
1-1	103 (10)	10/10		103 (10)	100	10/10	104 (10)	101	10/10	102 (10)	99	10/10	102 (10)	99	10/10	101 (10)	98	10/10
1-7	111 (10)	10/10		110 (10)	99	10/10	111 (10)	100	10/10	108 (10)	97	10/10	107 (10)	96	10/10	105 (10)	95	10/10
2-7	121 (10)	10/10		121 (10)	100	10/10	123 (10)	102	10/10	117 (10)	97	10/10	115 (10)	95	10/10	110 (10)	91	10/10

< >:No. of effective animals, ():No. of measured animals Au.Wt.: g

TABLE 4 FOOD CONSUMPTION IN MALE RAT (TWO-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm		380 ppm		750 ppm		1500 ppm		3000 ppm						
	Au.FC. <10>	No.of Surviv. <10>	Au.FC. <10>	% of cont. <10>	No.of Surviv. <10>	Au.FC. <10>	% of cont. <10>	No.of Surviv. <10>	Au.FC. <10>	% of cont. <10>	No.of Surviv. <10>	Au.FC. <10>	% of cont. <10>	No.of Surviv. <10>			
1-7	14.1 (10)	10/10	14.3 (10)	101	10/10	14.2 (10)	101	10/10	14.0 (10)	99	10/10	13.5 (10)	96	10/10	12.9 (10)	91	10/10
2-7	14.9 (10)	10/10	15.4 (10)	103	10/10	14.8 (10)	99	10/10	14.5 (10)	97	10/10	14.2 (10)	95	10/10	13.8 (10)	93	10/10

< >:No.of effective animals,():No.of measured animals Au.FC.: g

TABLE 5 FOOD CONSUMPTION IN FEMALE RAT (TWO-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm		380 ppm		750 ppm		1500 ppm		3000 ppm						
	Au.FC. <10>	No.of Surviv. <10>	Au.FC. <10>	% of cont. <10>	No.of Surviv. <10>	Au.FC. <10>	% of cont. <10>	No.of Surviv. <10>	Au.FC. <10>	% of cont. <10>	No.of Surviv. <10>	Au.FC. <10>	% of cont. <10>	No.of Surviv. <10>			
1-7	11.3 (10)	10/10	11.1 (10)	98	10/10	11.1 (10)	98	10/10	11.0 (10)	97	10/10	10.5 (10)	93	10/10	10.7 (10)	95	10/10
2-7	11.5 (10)	10/10	11.7 (10)	102	10/10	11.4 (10)	99	10/10	10.9 (10)	95	10/10	11.0 (10)	96	10/10	10.5 (10)	91	10/10

< >:No.of effective animals,():No.of measured animals Au.FC.: g

TABLE 6 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES IN MALE RAT (THIRTEEN-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm		380 ppm		750 ppm		1500 ppm		3000 ppm						
	Au.Wt.	No. of Surviv. <10>	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.			
0-0	127 (10)	10/10	127 (10)	100	10/10	128 (10)	101	10/10	127 (10)	100	10/10	127 (10)	100	10/10	126 (10)	99	10/10
1-7	158 (10)	10/10	157 (10)	99	10/10	154 (10)	97	10/10	153 (10)	97	10/10	146 (10)	92	10/10	133 (10)	84	10/10
2-7	192 (10)	10/10	185 (10)	96	10/10	183 (10)	95	10/10	178 (10)	93	10/10	165 (10)	86	10/10	130 (10)	68	10/10
3-7	213 (10)	10/10	205 (10)	96	10/10	202 (10)	95	10/10	195 (10)	92	10/10	179 (10)	84	10/10	145 (10)	68	10/10
4-7	231 (10)	10/10	225 (10)	97	10/10	219 (10)	95	10/10	206 (10)	89	10/10	186 (10)	81	10/10	139 (10)	60	10/10
5-7	249 (10)	10/10	249 (10)	100	10/10	237 (10)	95	10/10	217 (10)	87	10/10	191 (10)	77	10/10	139 (10)	56	10/10
6-7	261 (10)	10/10	262 (10)	100	10/10	247 (10)	95	10/10	222 (10)	85	10/10	191 (10)	73	10/10	139 (10)	53	10/10
7-7	272 (10)	10/10	272 (10)	100	10/10	257 (10)	94	10/10	227 (10)	83	10/10	190 (10)	70	10/10	138 (10)	51	10/10
8-7	285 (10)	10/10	284 (10)	100	10/10	267 (10)	94	10/10	232 (10)	81	10/10	190 (10)	67	10/10	138 (10)	48	10/10
9-7	299 (10)	10/10	302 (10)	101	10/10	280 (10)	94	10/10	241 (10)	81	10/10	195 (10)	65	10/10	144 (10)	48	10/10
10-7	306 (10)	10/10	310 (10)	101	10/10	287 (10)	94	10/10	246 (10)	80	10/10	195 (10)	64	10/10	143 (10)	47	10/10
11-7	317 (10)	10/10	318 (10)	100	10/10	297 (10)	94	10/10	254 (10)	80	10/10	199 (10)	63	10/10	144 (10)	45	10/10
12-7	325 (10)	10/10	323 (10)	99	10/10	303 (10)	93	10/10	260 (10)	80	10/10	203 (10)	62	10/10	139 (10)	43	10/10
13-7	335 (10)	10/10	330 (10)	99	10/10	310 (10)	93	10/10	266 (10)	79	10/10	210 (10)	63	10/10	145 (9)	43	9/10

< >:No. of effective animals, ():No. of measured animals Au.Wt.: g

TABLE 7 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES IN FEMALE RAT (THIRTEEN-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm		380 ppm		750 ppm		1500 ppm		3000 ppm						
	Au.Wt.	No. of Surviv. <10>	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.			
0-0	102 (10)	10/10	103 (10)	101	10/10	103 (10)	101	10/10	102 (10)	100	10/10	102 (10)	100	10/10	102 (10)	100	10/10
1-7	117 (10)	10/10	118 (10)	101	10/10	118 (10)	101	10/10	116 (10)	99	10/10	112 (10)	96	10/10	102 (10)	87	10/10
2-7	134 (10)	10/10	130 (10)	97	10/10	130 (10)	97	10/10	126 (10)	94	10/10	120 (10)	90	10/10	97 (10)	72	10/10
3-7	140 (10)	10/10	139 (10)	99	10/10	137 (10)	98	10/10	132 (10)	94	10/10	127 (10)	91	10/10	105 (10)	75	10/10
4-7	151 (10)	10/10	147 (10)	97	10/10	146 (10)	97	10/10	138 (10)	91	10/10	130 (10)	86	10/10	99 (10)	66	10/10
5-7	160 (10)	10/10	158 (10)	99	10/10	156 (10)	98	10/10	146 (10)	91	10/10	134 (10)	84	10/10	99 (10)	62	10/10
6-7	166 (10)	10/10	164 (10)	99	10/10	159 (10)	96	10/10	148 (10)	89	10/10	134 (10)	81	10/10	99 (10)	60	10/10
7-7	169 (10)	10/10	168 (10)	99	10/10	163 (10)	96	10/10	150 (10)	89	10/10	134 (10)	79	10/10	100 (10)	59	10/10
8-7	172 (10)	10/10	172 (10)	100	10/10	167 (10)	97	10/10	154 (10)	90	10/10	132 (10)	77	10/10	98 (10)	57	10/10
9-7	177 (10)	10/10	178 (10)	101	10/10	171 (10)	97	10/10	156 (10)	88	10/10	136 (10)	77	10/10	100 (10)	56	10/10
10-7	180 (10)	10/10	180 (10)	100	10/10	172 (10)	96	10/10	157 (10)	87	10/10	135 (10)	75	10/10	98 (10)	54	10/10
11-7	182 (10)	10/10	184 (10)	101	10/10	177 (10)	97	10/10	162 (10)	89	10/10	136 (10)	75	10/10	99 (10)	54	10/10
12-7	186 (10)	10/10	188 (10)	101	10/10	180 (10)	97	10/10	163 (10)	88	10/10	137 (10)	74	10/10	98 (10)	53	10/10
13-7	189 (10)	10/10	192 (10)	102	10/10	181 (10)	96	10/10	166 (10)	88	10/10	138 (10)	73	10/10	99 (10)	52	10/10

< >:No. of effective animals, ():No. of measured animals Au.Wt.: g

TABLE 8 FOOD CONSUMPTION IN MALE RAT (THIRTEEN-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm		380 ppm		750 ppm		1500 ppm		3000 ppm						
	Au.FC.	No. of Surviv. <10>	Au.FC.	% of cont. <10>	No. of Surviv.	Au.FC.	% of cont. <10>	No. of Surviv.	Au.FC.	% of cont. <10>	No. of Surviv.	Au.FC.	% of cont. <10>	No. of Surviv.			
1-7	15.8 (10)	10/10	15.7 (10)	99	10/10	15.3 (10)	97	10/10	15.2 (10)	96	10/10	14.1 (10)	89	10/10	12.8 (10)	81	10/10
2-7	17.4 (10)	10/10	16.5 (10)	95	10/10	16.5 (10)	95	10/10	16.0 (10)	92	10/10	14.6 (10)	84	10/10	11.4 (10)	66	10/10
3-7	17.2 (10)	10/10	16.3 (10)	95	10/10	16.5 (10)	96	10/10	15.5 (10)	90	10/10	14.9 (10)	87	10/10	12.2 (10)	71	10/10
4-7	17.5 (10)	10/10	17.4 (10)	99	10/10	17.1 (10)	98	10/10	16.3 (10)	93	10/10	14.9 (10)	85	10/10	11.6 (10)	66	10/10
5-7	17.7 (10)	10/10	18.0 (10)	102	10/10	17.4 (10)	98	10/10	15.2 (10)	86	10/10	14.3 (10)	81	10/10	11.2 (10)	63	10/10
6-7	17.2 (10)	10/10	18.4 (10)	107	10/10	17.1 (10)	99	10/10	15.0 (10)	87	10/10	13.7 (10)	80	10/10	11.7 (10)	68	10/10
7-7	16.7 (10)	10/10	17.6 (10)	105	10/10	16.1 (10)	96	10/10	14.3 (10)	86	10/10	13.3 (10)	80	10/10	10.8 (10)	65	10/10
8-7	16.8 (10)	10/10	16.8 (10)	100	10/10	15.9 (10)	95	10/10	13.8 (10)	82	10/10	12.4 (10)	74	10/10	10.3 (10)	61	10/10
9-7	17.7 (10)	10/10	17.0 (10)	96	10/10	16.1 (10)	91	10/10	13.3 (10)	75	10/10	11.4 (10)	64	10/10	10.7 (10)	60	10/10
10-7	16.8 (10)	10/10	16.6 (10)	99	10/10	15.7 (10)	93	10/10	13.2 (10)	79	10/10	10.4 (10)	62	10/10	9.7 (10)	58	10/10
11-7	16.9 (10)	10/10	17.4 (10)	103	10/10	16.9 (10)	100	10/10	14.6 (10)	86	10/10	12.4 (10)	73	10/10	10.2 (10)	60	10/10
12-7	16.5 (10)	10/10	16.6 (10)	101	10/10	16.1 (10)	98	10/10	14.3 (10)	87	10/10	12.6 (10)	76	10/10	9.7 (10)	59	10/10
13-7	16.7 (10)	10/10	16.4 (10)	98	10/10	16.0 (10)	96	10/10	14.6 (10)	87	10/10	13.1 (10)	78	10/10	10.7 (10)	64	9/10

< >:No. of effective animals, ():No. of measured animals Au.FC.: g

TABLE 9 FOOD CONSUMPTION IN FEMALE RAT (THIRTEEN-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm		380 ppm		750 ppm		1500 ppm		3000 ppm						
	Au.FC.	No. of Surviv. <10>	Au.FC.	% of cont. <10>	No. of Surviv.	Au.FC.	% of cont. <10>	No. of Surviv.	Au.FC.	% of cont. <10>	No. of Surviv.	Au.FC.	% of cont. <10>	No. of Surviv.			
1-7	12.4 (10)	10/10	12.4 (10)	100	10/10	12.3 (10)	99	10/10	12.0 (10)	97	10/10	11.0 (10)	89	10/10	9.9 (10)	80	10/10
2-7	13.1 (10)	10/10	12.0 (10)	92	10/10	12.4 (10)	95	10/10	12.4 (10)	95	10/10	11.0 (10)	84	10/10	7.9 (10)	60	10/10
3-7	12.7 (10)	10/10	12.2 (10)	96	10/10	12.5 (10)	98	10/10	11.9 (10)	94	10/10	11.7 (10)	92	10/10	8.6 (10)	68	10/10
4-7	13.0 (10)	10/10	12.6 (10)	97	10/10	12.9 (10)	99	10/10	12.4 (10)	95	10/10	11.4 (10)	88	10/10	7.3 (10)	56	10/10
5-7	14.4 (10)	10/10	13.1 (10)	91	10/10	12.8 (10)	89	10/10	12.4 (10)	86	10/10	10.7 (10)	74	10/10	7.4 (10)	51	10/10
6-7	12.8 (10)	10/10	12.9 (10)	101	10/10	12.3 (10)	96	10/10	11.9 (10)	93	10/10	10.3 (10)	80	10/10	7.9 (10)	62	10/10
7-7	11.7 (10)	10/10	11.6 (10)	99	10/10	11.5 (10)	98	10/10	10.7 (10)	91	10/10	9.7 (10)	83	10/10	8.2 (10)	70	10/10
8-7	11.0 (10)	10/10	11.3 (10)	103	10/10	11.3 (10)	103	10/10	10.6 (10)	96	10/10	9.3 (10)	85	10/10	7.2 (10)	65	10/10
9-7	12.1 (10)	10/10	12.2 (10)	101	10/10	11.3 (10)	93	10/10	10.6 (10)	88	10/10	8.9 (10)	74	10/10	7.2 (10)	60	10/10
10-7	11.3 (10)	10/10	11.0 (10)	97	10/10	11.1 (10)	98	10/10	10.3 (10)	91	10/10	8.3 (10)	73	10/10	6.5 (10)	58	10/10
11-7	11.3 (10)	10/10	11.3 (10)	100	10/10	11.6 (10)	103	10/10	10.7 (10)	95	10/10	9.6 (10)	85	10/10	7.1 (10)	63	10/10
12-7	11.6 (10)	10/10	11.7 (10)	101	10/10	11.4 (10)	98	10/10	10.4 (10)	90	10/10	9.0 (10)	78	10/10	7.2 (10)	62	10/10
13-7	11.7 (10)	10/10	12.1 (10)	103	10/10	11.2 (10)	96	10/10	11.0 (10)	94	10/10	8.8 (10)	75	10/10	7.6 (10)	65	10/10

< >:No. of effective animals, ():No. of measured animals Au.FC.: g

TABLE 10 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES IN MALE MOUSE (TWO-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm			380 ppm			750 ppm			1500 ppm			3000 ppm		
	Au.Wt.	No. of Surviv. < 9 >	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.
0-0	22.9 (9)	10/10	22.7 (10)	99	10/10	22.7 (10)	99	10/10	22.8 (10)	100	10/10	22.9 (10)	100	10/10	22.7 (10)	99	10/10
1-1	23.6 (9)	10/10	22.5 (10)	95	10/10	23.1 (10)	98	10/10	22.7 (10)	96	10/10	22.2 (10)	94	10/10	20.9 (10)	89	8/10
1-7	24.6 (9)	10/10	23.7 (10)	96	10/10	24.1 (10)	98	10/10	24.3 (10)	99	10/10	23.2 (9)	94	9/10	- (-)	-	0/10
2-7	25.2 (9)	9/9	24.9 (10)	99	10/10	24.9 (10)	99	10/10	24.8 (10)	98	10/10	23.6 (9)	94	9/10	- (-)	-	0/10

< >:No. of effective animals, ():No. of measured animals Au.Wt.: g

TABLE 11 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES IN FEMALE MOUSE (TWO-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm			380 ppm			750 ppm			1500 ppm			3000 ppm		
	Au.Wt.	No. of Surviv. <10>	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.
0-0	18.9 (10)	10/10	19.0 (10)	101	10/10	19.0 (10)	101	10/10	18.7 (10)	99	10/10	18.9 (10)	100	10/10	18.6 (10)	98	10/10
1-1	18.8 (10)	10/10	19.0 (10)	101	10/10	18.9 (10)	101	10/10	18.4 (10)	98	10/10	18.4 (10)	98	10/10	18.0 (10)	96	10/10
1-7	19.6 (10)	10/10	19.7 (10)	101	10/10	19.2 (10)	98	10/10	19.4 (10)	99	10/10	18.4 (10)	94	10/10	- (-)	-	0/10
2-7	20.4 (10)	10/10	20.9 (10)	102	10/10	20.6 (10)	101	10/10	20.5 (10)	100	10/10	19.6 (10)	96	10/10	- (-)	-	0/10

< >:No. of effective animals, ():No. of measured animals Au.Wt.: g

TABLE 12 FOOD CONSUMPTION IN MALE MOUSE (TWO-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm		380 ppm		750 ppm		1500 ppm		3000 ppm						
	Au.F.C.	No. of Surviv. < 9 >	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.			
1-7	4.1 (9)	10/10	4.0 (10)	98	10/10	4.1 (10)	100	10/10	4.1 (10)	100	10/10	3.5 (10)	85	9/10	0.8 (7)	20	0/10
2-7	4.2 (6)	9/ 9	4.0 (10)	95	10/10	4.2 (10)	100	10/10	4.4 (10)	105	10/10	4.3 (9)	102	9/10	- (-)	-	0/10

< >:No.of effective animals,():No.of measured animals Au.F.C.: g

TABLE 13 FOOD CONSUMPTION IN FEMALE MOUSE (TWO-WEEK STUDY)

Week-Day on Study	0 ppm		190 ppm		380 ppm		750 ppm		1500 ppm		3000 ppm						
	Au.F.C.	No. of Surviv. <10>	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.			
1-7	3.2 (10)	10/10	3.4 (10)	106	10/10	3.2 (10)	100	10/10	3.2 (10)	100	10/10	3.1 (10)	97	10/10	1.0 (10)	31	0/10
2-7	3.9 (10)	10/10	4.0 (10)	103	10/10	3.9 (10)	100	10/10	3.8 (10)	97	10/10	3.7 (10)	95	10/10	- (-)	-	0/10

< >:No.of effective animals,():No.of measured animals Au.F.C.: g

TABLE 14 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES IN MALE MOUSE (THIRTEEN-WEEK STUDY)

Week-Day on Study	0 ppm		300 ppm		440 ppm		670 ppm		1000 ppm		1500 ppm						
	Au.Wt.	No. of Surviv. <10>	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.			
0-0	23.3 (10)	10/10	23.1 (10)	99	10/10	23.2 (10)	100	10/10	23.5 (10)	101	10/10	23.1 (10)	99	10/10	23.3 (10)	100	10/10
1-7	25.2 (10)	10/10	25.0 (10)	99	10/10	24.9 (10)	99	10/10	24.4 (10)	97	10/10	23.4 (10)	93	10/10	22.9 (6)	91	6/10
2-7	26.1 (10)	10/10	25.5 (10)	98	10/10	25.2 (10)	97	10/10	24.8 (10)	95	10/10	23.9 (10)	92	10/10	22.5 (6)	86	6/10
3-7	27.0 (10)	10/10	26.2 (10)	97	10/10	25.8 (10)	96	10/10	25.1 (10)	93	10/10	24.2 (10)	90	10/10	23.4 (6)	87	6/10
4-7	27.8 (10)	10/10	26.5 (10)	95	10/10	26.2 (10)	94	10/10	25.2 (10)	91	10/10	24.3 (10)	87	10/10	23.2 (6)	83	6/10
5-7	29.3 (10)	10/10	27.3 (10)	93	10/10	26.7 (10)	91	10/10	25.6 (10)	87	10/10	24.6 (10)	84	10/10	22.7 (6)	77	6/10
6-7	30.3 (10)	10/10	27.5 (10)	91	10/10	27.2 (10)	90	10/10	25.8 (10)	85	10/10	24.7 (10)	82	10/10	24.4 (5)	81	5/10
7-7	30.9 (10)	10/10	27.8 (10)	90	10/10	26.8 (10)	87	10/10	25.7 (10)	83	10/10	24.7 (10)	80	10/10	24.0 (5)	78	5/10
8-7	31.8 (10)	10/10	27.9 (10)	88	10/10	26.9 (10)	85	10/10	25.7 (10)	81	10/10	24.2 (10)	76	10/10	23.4 (5)	74	5/10
9-7	33.2 (10)	10/10	28.7 (10)	86	10/10	27.5 (10)	83	10/10	26.2 (10)	79	10/10	24.9 (10)	75	10/10	23.6 (4)	71	4/10
10-7	33.6 (10)	10/10	28.8 (10)	86	10/10	27.0 (10)	80	10/10	26.1 (10)	78	10/10	24.8 (10)	74	10/10	- (-)	-	0/10
11-7	34.7 (10)	10/10	29.3 (10)	84	10/10	27.7 (10)	80	10/10	26.6 (10)	77	10/10	25.5 (8)	73	8/10	- (-)	-	0/10
12-7	35.3 (10)	10/10	30.1 (10)	85	10/10	28.2 (10)	80	10/10	26.8 (10)	76	10/10	26.2 (8)	74	8/10	- (-)	-	0/10
13-7	36.5 (10)	10/10	30.7 (10)	84	10/10	29.1 (10)	80	10/10	27.5 (10)	75	10/10	26.8 (8)	73	8/10	- (-)	-	0/10

< >:No. of effective animals, ():No. of measured animals Au.Wt.: g

TABLE 15 SURVIVAL ANIMAL NUMBERS AND BODY WEIGHT CHANGES IN FEMALE MOUSE (THIRTEEN-WEEK STUDY)

Week-Day on Study	0 ppm		300 ppm		440 ppm		670 ppm		1000 ppm		1500 ppm						
	Au.Wt.	No. of Surviv. <10>	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.	Au.Wt.	% of cont. <10>	No. of Surviv.			
0-0	18.7 (10)	10/10	18.5 (10)	99	10/10	18.6 (10)	99	10/10	18.2 (10)	97	10/10	18.4 (10)	98	10/10	18.7 (10)	100	10/10
1-7	20.4 (10)	10/10	19.9 (10)	98	10/10	20.0 (10)	98	10/10	19.0 (10)	93	10/10	18.8 (10)	92	10/10	18.8 (10)	92	10/10
2-7	21.2 (10)	10/10	20.1 (10)	95	10/10	20.4 (10)	96	10/10	20.1 (10)	95	10/10	19.2 (10)	91	10/10	18.3 (10)	86	10/10
3-7	22.2 (10)	10/10	21.0 (10)	95	10/10	20.7 (10)	93	10/10	20.5 (10)	92	10/10	19.9 (10)	90	10/10	19.3 (10)	87	10/10
4-7	22.8 (10)	10/10	21.7 (10)	95	10/10	21.6 (10)	95	10/10	21.2 (10)	93	10/10	20.5 (10)	90	10/10	19.9 (10)	87	10/10
5-7	24.2 (10)	10/10	22.7 (10)	94	10/10	22.0 (10)	91	10/10	21.4 (10)	88	10/10	20.6 (10)	85	10/10	19.8 (10)	82	10/10
6-7	23.9 (10)	10/10	22.9 (10)	96	10/10	22.9 (10)	96	10/10	22.1 (10)	92	10/10	21.4 (10)	90	10/10	20.5 (10)	86	10/10
7-7	24.5 (10)	10/10	23.3 (10)	95	10/10	22.7 (10)	93	10/10	21.9 (10)	89	10/10	21.2 (10)	87	10/10	20.8 (10)	85	10/10
8-7	24.6 (10)	10/10	23.8 (10)	97	10/10	22.7 (10)	92	10/10	22.2 (10)	90	10/10	21.2 (10)	86	10/10	20.6 (9)	84	9/10
9-7	25.3 (10)	10/10	24.1 (10)	95	10/10	23.1 (10)	91	10/10	22.6 (10)	89	10/10	21.4 (10)	85	10/10	20.9 (8)	83	8/10
10-7	25.4 (10)	10/10	23.9 (10)	94	10/10	23.0 (10)	91	10/10	22.5 (10)	89	10/10	21.1 (10)	83	10/10	20.3 (2)	80	2/10
11-7	25.4 (10)	10/10	24.4 (10)	96	10/10	23.8 (10)	94	10/10	23.1 (10)	91	10/10	21.2 (9)	83	9/10	- (-)	-	0/10
12-7	26.2 (10)	10/10	24.2 (10)	92	10/10	24.1 (10)	92	10/10	23.2 (10)	89	10/10	22.3 (9)	85	9/10	- (-)	-	0/10
13-7	26.4 (10)	10/10	24.7 (10)	94	10/10	24.3 (10)	92	10/10	23.5 (10)	89	10/10	22.9 (9)	87	9/10	- (-)	-	0/10

< >:No. of effective animals, ():No. of measured animals Au.Wt.: g

TABLE 16 FOOD CONSUMPTION IN MALE MOUSE (THIRTEEN-WEEK STUDY)

Week-Day on Study	0 ppm		300 ppm		440 ppm		670 ppm		1000 ppm		1500 ppm						
	Au.F.C.	No. of Surviv. <10>	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.			
1-7	4.3 (10)	10/10	4.1 (10)	95	10/10	4.1 (10)	95	10/10	3.8 (10)	88	10/10	3.6 (10)	84	10/10	2.3 (9)	53	6/10
2-7	4.2 (10)	10/10	4.0 (10)	95	10/10	3.9 (10)	93	10/10	4.2 (10)	100	10/10	4.3 (10)	102	10/10	5.1 (6)	121	6/10
3-7	4.2 (10)	10/10	4.0 (10)	95	10/10	4.0 (10)	95	10/10	4.6 (10)	110	10/10	4.6 (10)	110	10/10	5.1 (6)	121	6/10
4-7	4.6 (10)	10/10	4.3 (10)	93	10/10	4.3 (10)	93	10/10	4.5 (10)	98	10/10	4.6 (10)	100	10/10	4.9 (6)	107	6/10
5-7	4.6 (10)	10/10	4.3 (10)	93	10/10	4.3 (10)	93	10/10	4.4 (10)	96	10/10	4.6 (10)	100	10/10	4.6 (6)	100	6/10
6-7	4.6 (10)	10/10	4.1 (10)	89	10/10	4.2 (10)	91	10/10	4.4 (10)	96	10/10	4.4 (10)	96	10/10	5.3 (5)	115	5/10
7-7	4.4 (10)	10/10	4.2 (10)	95	10/10	4.2 (10)	95	10/10	4.6 (10)	105	10/10	4.3 (10)	98	10/10	4.9 (5)	111	5/10
8-7	4.6 (10)	10/10	4.3 (10)	93	10/10	4.4 (10)	96	10/10	4.6 (10)	100	10/10	4.5 (10)	98	10/10	5.1 (5)	111	5/10
9-7	4.9 (10)	10/10	4.3 (10)	88	10/10	4.3 (10)	88	10/10	4.6 (10)	94	10/10	4.7 (10)	96	10/10	4.9 (5)	100	4/10
10-7	4.7 (10)	10/10	4.4 (10)	94	10/10	4.4 (10)	94	10/10	4.7 (10)	100	10/10	4.6 (10)	98	10/10	1.7 (4)	36	0/10
11-7	4.8 (10)	10/10	4.3 (10)	90	10/10	4.5 (10)	94	10/10	4.7 (10)	98	10/10	4.5 (10)	94	8/10	- (-)	-	0/10
12-7	4.8 (10)	10/10	4.5 (10)	94	10/10	4.6 (10)	96	10/10	4.6 (10)	96	10/10	4.7 (8)	98	8/10	- (-)	-	0/10
13-7	4.8 (10)	10/10	4.4 (10)	92	10/10	4.7 (10)	98	10/10	4.5 (10)	94	10/10	4.6 (8)	96	8/10	- (-)	-	0/10

< >:No. of effective animals, ():No. of measured animals Au.F.C.: g

TABLE 17 FOOD CONSUMPTION IN FEMALE MOUSE (THIRTEEN-WEEK STUDY)

Week-Day on Study	0 ppm		300 ppm		440 ppm		670 ppm		1000 ppm		1500 ppm						
	Au.F.C.	No. of Surviv. <10>	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.	Au.F.C.	% of cont. <10>	No. of Surviv.			
1-7	3.6 (10)	10/10	3.5 (10)	97	10/10	3.4 (10)	94	10/10	3.1 (10)	86	10/10	3.1 (10)	86	10/10	3.0 (10)	83	10/10
2-7	3.6 (10)	10/10	3.4 (10)	94	10/10	3.5 (10)	97	10/10	3.7 (10)	103	10/10	3.7 (10)	103	10/10	3.6 (10)	100	10/10
3-7	3.9 (10)	10/10	3.8 (10)	97	10/10	3.7 (10)	95	10/10	4.2 (10)	108	10/10	4.2 (10)	108	10/10	4.5 (10)	115	10/10
4-7	4.1 (10)	10/10	4.1 (10)	100	10/10	4.0 (10)	98	10/10	4.1 (10)	100	10/10	4.3 (10)	105	10/10	4.4 (10)	107	10/10
5-7	4.4 (10)	10/10	4.2 (10)	95	10/10	4.0 (10)	91	10/10	4.0 (10)	91	10/10	4.3 (10)	98	10/10	4.4 (10)	100	10/10
6-7	4.2 (10)	10/10	4.1 (10)	98	10/10	4.0 (10)	95	10/10	4.2 (10)	100	10/10	4.3 (10)	102	10/10	4.4 (10)	105	10/10
7-7	4.4 (10)	10/10	4.3 (10)	98	10/10	4.2 (10)	95	10/10	4.3 (10)	98	10/10	4.2 (10)	95	10/10	4.3 (10)	98	10/10
8-7	4.5 (10)	10/10	4.6 (10)	102	10/10	4.3 (10)	96	10/10	4.6 (10)	102	10/10	4.4 (5)	98	10/10	4.2 (10)	93	9/10
9-7	4.6 (10)	10/10	4.5 (10)	98	10/10	4.3 (10)	93	10/10	4.6 (10)	100	10/10	4.3 (10)	93	10/10	4.3 (9)	93	8/10
10-7	4.7 (10)	10/10	4.7 (10)	100	10/10	4.5 (10)	96	10/10	4.8 (10)	102	10/10	4.4 (10)	94	10/10	3.1 (8)	66	2/10
11-7	4.5 (10)	10/10	4.7 (10)	104	10/10	4.6 (10)	102	10/10	4.8 (10)	107	10/10	3.9 (10)	87	9/10	1.1 (2)	24	0/10
12-7	4.7 (10)	10/10	4.7 (10)	100	10/10	4.4 (10)	94	10/10	4.7 (10)	100	10/10	4.4 (9)	94	9/10	- (-)	-	0/10
13-7	4.7 (10)	10/10	4.7 (10)	100	10/10	4.3 (10)	91	10/10	4.5 (10)	96	10/10	4.3 (9)	91	9/10	- (-)	-	0/10

< >:No. of effective animals, ():No. of measured animals Au.F.C.: g