

アントラセンのラット及びマウスを用いた
経口投与によるがん原性予備試験(混餌試験)報告書

試験番号

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Two-week studies	Thirteen-week Studies
<Method of Administration> Feed	Feed
<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> <i>Strain and Species</i> F344/DuCrj(Fischer)rat Crj:BDF1 mouse <i>Animal Source</i> Charles River Japan, Inc. <i>Duration of Time Held Before Study</i> 2 wk <i>Age When Placed on Study</i> 6 wk <i>Age When Killed</i> 8 wk	F344/DuCrj(Fischer)rat Crj:BDF1 mouse Charles River Japan, Inc. 2 wk 6 wk 19 wk
<Doses> Rat--0, 80, 400, 2000, 10000, or 500000ppm; Mouse--0, 80, 400, 2000, 10000, or 500000ppm;	Same as two-week studies Same as two-week studies
<Duration of Dosing> 7d/wk for 2wk	7d/wk for 13wk
<Animal Maintenance> <i>Feed</i> CRF-1 (Oriental Yeast Co.,Ltd.) Sterilized by γ -ray Available ad libitum <i>Water</i> Sterilized by ultraviolet rays Automatic watering system Available ad libitum <i>Animal per Cage</i> Single (stainless steel wire) <i>Animal Room Environment</i> Barrier system Temperature : $24 \pm 2^{\circ}\text{C}$ Humidity : $55 \pm 10\%$ Fluorescent light 12h/d 15-17 room air changes /h	Same as two-week studies Same as two-week studies Same as two-week studies Same as two-week studies
<Type and Frequency of Observation> <i>Clinical sign</i> Observed 1/d <i>Body weight</i> Weighed 0-0, 1-4, 1-7, 2-4, and 2-7 (wk-d) <i>Food Consumption</i> Weighed 1-7, 2-7 (wk-d)	Observed 1/d Weighed 1/wk for 13wk Weighed 1/wk for 13wk

TABLE 1
(Continued)

EXPERIMENTAL DESIGN AND MATERIALS AND METHODS IN THE FEEDING STUDIES OF ANTHRACENE

Two-week Studies	Thirteen-week Studies
<p><Hematology> Hematology measurement performed on at least five animals per sex per groups. Red blood cell (RBC), Hemoglobin, Hematocrit, Mean corpuscular volume (MCV), Mean corpuscular hemoglobin (MCH), Mean corpuscular hemoglobin concentration (MCHC), Reticulocyte<rat only>, Prothrombin time<rat only>, APTT<rat only>, Platelet, White blood cell (WBC), Differential WBC.</p>	<p>Hematology measurement performed on schedule sacrificed animals. 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, Reticulocyte<rat only>, Prothrombin time<rat only>, APTT<rat only>.</p>
<p><Biochemistry> Biochemistry measurement performed on at least five animals per sex per groups. 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), γ-Glutamyl transpeptidase (γ-GTP) <rat only>, Creatine phosphokinase (CPK), Urea nitrogen, Creatinine <rat only>, Calcium, Inorganic phosphorus.</p>	<p>Biochemistry measurement performed on schedule sacrificed animals. Total protein, Albumin, A/G ratio, T-bilirubin, Glucose, T-cholesterol, Triglyceride, Phospholipid <rat only>, Glutamic oxaloacetic transaminase (GOT), Glutamic pyruvic transaminase (GPT), Alkaline phosphatase(ALP), Lactate dehydrogenase(LDH), γ-Glutamyl transpeptidase (γ-GTP) <rat only>, Creatine phosphokinase (CPK), Urea nitrogen, Creatinine <rat only>, Sodium, Potassium, Chloride, Calcium, Inorganic phosphorus.</p>
<p><Urinalysis> None</p>	<p>Urinalysis measurement performed on schedule sacrificed animals. pH, Protein, Glucose, Ketone body, Bilirubin <rat only>, Occult blood, Urobilinogen.</p>
<p><Necropsy> Necropsy performed on all animals.</p>	<p>Same as two-week studies.</p>
<p><Organ weight> Organ weight measurement performed on at least five animals per sex per groups. The following organs were weighed: brain, lung, liver, spleen, heart, kidney, adrenal, testis, ovary, thymus.</p>	<p>Organ weight measurement performed on scheduled sacrificed animals. The following organs were weighed: brain, lung, liver, spleen, heart, kidney, adrenal, testis, ovary, thymus.</p>
<p><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, thyroid, adrenal, testis, ovary, brain.</p>	<p>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, mammary gland, brain, spinal cord, peripheral nerve, eye, Harderian gland, muscle, bone.</p>

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

Week-Day on Study	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 ppm						
	Au.Wt. (10)	No. of Surviv. <10>	Au.Wt. (10)	% of cont. <10>	No. of Surviv.	Au.Wt. (10)	% of cont. <10>	No. of Surviv.	Au.Wt. (10)	% of cont. <10>	No. of Surviv.	Au.Wt. (10)	% of cont. <10>	No. of Surviv.	Au.Wt. (10)	% of cont. <10>	No. of Surviv.
0-0	120	10/10	120	100	10/10	120	100	10/10	120	100	10/10	120	100	10/10	120	100	10/10
1-1	123	10/10	122	99	10/10	122	99	10/10	121	98	10/10	118	96	10/10	117	95	10/10
1-3	130	10/10	130	100	10/10	129	99	10/10	126	97	10/10	123	95	10/10	123	95	10/10
1-7	147	10/10	146	99	10/10	145	99	10/10	145	99	10/10	142	97	10/10	140	95	10/10
2-3	159	10/10	158	99	10/10	157	99	10/10	157	99	10/10	154	97	10/10	152	96	10/10
2-7	171	10/10	169	99	10/10	168	98	10/10	167	98	10/10	165	96	10/10	161	94	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	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 ppm						
	Au.Wt. (10)	No. of Surviv. <10>	Au.Wt. (10)	% of cont. <10>	No. of Surviv.	Au.Wt. (10)	% of cont. <10>	No. of Surviv.	Au.Wt. (10)	% of cont. <10>	No. of Surviv.	Au.Wt. (10)	% of cont. <10>	No. of Surviv.	Au.Wt. (10)	% of cont. <10>	No. of Surviv.
0-0	99	10/10	99	100	10/10	99	100	10/10	99	100	10/10	99	100	10/10	99	100	10/10
1-1	101	10/10	101	100	10/10	100	99	10/10	99	98	10/10	96	95	10/10	94	93	10/10
1-3	105	10/10	106	101	10/10	105	100	10/10	103	98	10/10	101	96	10/10	97	92	10/10
1-7	113	10/10	114	101	10/10	114	101	10/10	112	99	10/10	110	97	10/10	107	95	10/10
2-3	120	10/10	120	100	10/10	121	101	10/10	117	98	10/10	116	97	10/10	113	94	10/10
2-7	124	10/10	125	101	10/10	127	102	10/10	121	98	10/10	121	98	10/10	117	94	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	Control		80 ppm		400 ppm			2000 ppm			10000 ppm			50000 ppm			
	Au.F.C. <10>	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.	Au.F.C.	% of cont. <10>	No. of Surviv.
1-3	12.3 (10)	10/10	12.5 (10)	102	10/10	12.0 (10)	98	10/10	11.4 (10)	93	10/10	10.3 (10)	84	10/10	9.4 (10)	76	10/10
1-7	13.4 (10)	10/10	13.4 (10)	100	10/10	13.1 (10)	98	10/10	12.8 (10)	96	10/10	12.8 (10)	96	10/10	12.8 (10)	96	10/10
2-3	12.8 (10)	10/10	12.9 (10)	101	10/10	12.8 (10)	100	10/10	12.8 (10)	100	10/10	12.7 (10)	99	10/10	13.1 (10)	102	10/10
2-7	13.4 (10)	10/10	13.1 (10)	98	10/10	13.2 (10)	99	10/10	13.2 (10)	99	10/10	13.0 (10)	97	10/10	13.4 (10)	100	10/10

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

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

Week-Day on Study	Control		80 ppm		400 ppm			2000 ppm			10000 ppm			50000 ppm			
	Au.F.C. <10>	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.	Au.F.C.	% of cont. <10>	No. of Surviv.
1-3	10.1 (10)	10/10	10.5 (10)	104	10/10	10.2 (10)	101	10/10	9.0 (10)	89	10/10	8.0 (10)	79	10/10	6.5 (10)	64	10/10
1-7	11.1 (10)	10/10	11.3 (10)	102	10/10	10.9 (10)	98	10/10	10.3 (10)	93	10/10	10.1 (10)	91	10/10	9.9 (10)	89	10/10
2-3	10.5 (10)	10/10	10.9 (9)	104	10/10	11.0 (10)	105	10/10	10.2 (10)	97	10/10	10.2 (10)	97	10/10	9.9 (10)	94	10/10
2-7	10.8 (10)	10/10	11.4 (10)	106	10/10	11.5 (10)	106	10/10	10.5 (10)	97	10/10	10.4 (10)	96	10/10	10.0 (10)	93	10/10

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

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

Week on Study	Control			80 ppm			100 ppm			2000 ppm			10000 ppm			50000 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	124 (10)	10/10		124 (10)	100	10/10	124 (10)	100	10/10	124 (10)	100	10/10	124 (10)	100	10/10	124 (10)	100	10/10
1	149 (10)	10/10		149 (10)	100	10/10	149 (10)	100	10/10	146 (10)	98	10/10	142 (10)	95	10/10	141 (10)	95	10/10
2	182 (10)	10/10		185 (10)	102	10/10	185 (10)	102	10/10	180 (10)	99	10/10	175 (10)	96	10/10	172 (10)	95	10/10
3	209 (10)	10/10		212 (10)	101	10/10	210 (10)	100	10/10	204 (10)	98	10/10	198 (10)	95	10/10	196 (10)	94	10/10
4	229 (10)	10/10		229 (10)	100	10/10	227 (10)	99	10/10	221 (10)	97	10/10	217 (10)	95	10/10	214 (10)	93	10/10
5	245 (10)	10/10		247 (10)	101	10/10	244 (10)	100	10/10	237 (10)	97	10/10	234 (10)	96	10/10	232 (10)	95	10/10
6	259 (10)	10/10		261 (10)	101	10/10	256 (10)	99	10/10	252 (10)	97	10/10	248 (10)	96	10/10	245 (10)	95	10/10
7	275 (10)	10/10		276 (10)	100	10/10	270 (10)	98	10/10	265 (10)	96	10/10	260 (10)	95	10/10	260 (10)	95	10/10
8	290 (10)	10/10		294 (10)	101	10/10	286 (10)	99	10/10	280 (10)	97	10/10	275 (10)	95	10/10	274 (10)	94	10/10
9	298 (10)	10/10		304 (10)	102	10/10	296 (10)	99	10/10	291 (10)	98	10/10	286 (10)	96	10/10	285 (10)	96	10/10
10	310 (10)	10/10		316 (10)	102	10/10	307 (10)	99	10/10	301 (10)	97	10/10	295 (10)	95	10/10	295 (10)	95	10/10
11	321 (10)	10/10		326 (10)	102	10/10	317 (10)	99	10/10	313 (10)	98	10/10	307 (10)	96	10/10	307 (10)	96	10/10
12	328 (10)	10/10		337 (10)	103	10/10	328 (10)	100	10/10	322 (10)	98	10/10	314 (10)	96	10/10	315 (10)	96	10/10
13	337 (10)	10/10		345 (10)	102	10/10	335 (10)	99	10/10	329 (10)	98	10/10	322 (10)	96	10/10	323 (10)	96	10/10

< >: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 on Study	Control			80 ppm			100 ppm			2000 ppm			10000 ppm			50000 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	103 (10)	10/10		103 (10)	100	10/10	103 (10)	100	10/10	103 (10)	100	10/10	103 (10)	100	10/10	103 (10)	100	10/10
1	116 (10)	10/10		118 (10)	102	10/10	116 (10)	100	10/10	114 (10)	98	10/10	113 (10)	97	10/10	107 (10)	92	10/10
2	132 (10)	10/10		135 (10)	102	10/10	130 (10)	98	10/10	130 (10)	98	10/10	130 (10)	98	10/10	121 (10)	92	10/10
3	143 (10)	10/10		144 (10)	101	10/10	138 (10)	97	10/10	139 (10)	97	10/10	138 (10)	97	10/10	130 (10)	91	10/10
4	148 (10)	10/10		152 (10)	103	10/10	144 (10)	97	10/10	144 (10)	97	10/10	144 (10)	97	10/10	136 (10)	92	10/10
5	157 (10)	10/10		160 (10)	102	10/10	151 (10)	96	10/10	150 (10)	96	10/10	151 (10)	96	10/10	140 (10)	89	10/10
6	162 (10)	10/10		166 (10)	102	10/10	156 (10)	96	10/10	155 (10)	96	10/10	154 (10)	95	10/10	145 (10)	90	10/10
7	167 (10)	10/10		170 (10)	102	10/10	159 (10)	95	10/10	161 (10)	96	10/10	158 (10)	95	10/10	151 (10)	90	10/10
8	173 (10)	10/10		175 (10)	101	10/10	161 (10)	93	10/10	162 (10)	94	10/10	161 (10)	93	10/10	156 (10)	90	10/10
9	178 (10)	10/10		180 (10)	101	10/10	167 (10)	94	10/10	167 (10)	94	10/10	164 (10)	92	10/10	156 (10)	88	10/10
10	175 (10)	10/10		174 (10)	99	10/10	170 (10)	97	10/10	172 (10)	98	10/10	167 (10)	95	10/10	159 (10)	91	10/10
11	185 (10)	10/10		186 (10)	101	10/10	173 (10)	94	10/10	175 (10)	95	10/10	170 (10)	92	10/10	162 (10)	88	10/10
12	186 (10)	10/10		190 (10)	102	10/10	177 (10)	95	10/10	178 (10)	96	10/10	173 (10)	93	10/10	164 (10)	88	10/10
13	191 (10)	10/10		193 (10)	101	10/10	181 (10)	95	10/10	182 (10)	95	10/10	176 (10)	92	10/10	166 (10)	87	10/10

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

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

Week on Study	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 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	12.3 (10)	10/10	12.5 (10)	102	10/10	12.4 (10)	101	10/10	11.9 (10)	97	10/10	11.1 (10)	90	10/10	10.8 (10)	88	10/10
2	13.5 (10)	10/10	13.9 (10)	103	10/10	14.0 (10)	104	10/10	13.7 (10)	101	10/10	13.3 (10)	99	10/10	13.6 (10)	101	10/10
3	14.7 (10)	10/10	15.1 (10)	103	10/10	14.9 (10)	101	10/10	14.3 (10)	97	10/10	14.2 (10)	97	10/10	14.6 (10)	99	10/10
4	14.9 (10)	10/10	15.2 (10)	102	10/10	14.8 (10)	99	10/10	14.6 (10)	98	10/10	14.6 (10)	98	10/10	15.0 (10)	101	10/10
5	15.0 (10)	10/10	16.1 (10)	107	10/10	15.6 (10)	104	10/10	15.3 (10)	102	10/10	15.0 (10)	100	10/10	15.7 (10)	105	10/10
6	14.8 (10)	10/10	16.1 (10)	109	10/10	15.3 (10)	103	10/10	15.4 (10)	104	10/10	15.0 (10)	101	10/10	15.7 (10)	106	10/10
7	15.3 (10)	10/10	16.3 (10)	107	10/10	15.3 (10)	100	10/10	15.1 (10)	99	10/10	15.0 (10)	98	10/10	16.0 (10)	105	10/10
8	15.9 (10)	10/10	16.8 (10)	106	10/10	15.7 (10)	99	10/10	15.5 (10)	97	10/10	15.3 (10)	96	10/10	16.3 (10)	103	10/10
9	16.0 (10)	10/10	17.0 (10)	106	10/10	15.9 (10)	99	10/10	15.6 (10)	98	10/10	15.6 (10)	98	10/10	16.6 (10)	104	10/10
10	15.9 (10)	10/10	16.6 (10)	104	10/10	15.5 (10)	97	10/10	15.6 (10)	98	10/10	15.4 (10)	97	10/10	16.4 (10)	103	10/10
11	16.5 (10)	10/10	17.1 (10)	104	10/10	16.2 (10)	98	10/10	15.9 (10)	97	10/10	15.6 (10)	95	10/10	16.6 (10)	101	10/10
12	16.5 (10)	10/10	17.1 (10)	104	10/10	16.4 (10)	99	10/10	16.1 (10)	98	10/10	15.8 (10)	96	10/10	17.1 (10)	104	10/10
13	16.4 (10)	10/10	16.7 (10)	102	10/10	15.9 (10)	97	10/10	15.8 (10)	96	10/10	15.9 (10)	97	10/10	17.0 (10)	104	10/10

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

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

Week on Study	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 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	10.8 (10)	10/10	10.9 (10)	101	10/10	10.4 (10)	96	10/10	9.8 (10)	91	10/10	9.3 (10)	86	10/10	8.1 (10)	75	10/10
2	11.0 (10)	10/10	11.3 (10)	103	10/10	10.7 (10)	97	10/10	10.6 (10)	96	10/10	10.6 (10)	96	10/10	9.9 (10)	90	10/10
3	11.4 (10)	10/10	11.6 (10)	102	10/10	11.0 (9)	96	10/10	11.1 (10)	97	10/10	10.8 (10)	95	10/10	10.0 (10)	88	10/10
4	11.3 (10)	10/10	11.5 (10)	102	10/10	10.6 (10)	94	10/10	10.8 (10)	96	10/10	10.8 (10)	96	10/10	10.0 (10)	88	10/10
5	12.0 (10)	10/10	12.2 (10)	102	10/10	11.4 (10)	95	10/10	11.3 (10)	94	10/10	11.0 (10)	92	10/10	10.1 (10)	84	10/10
6	11.4 (10)	10/10	11.6 (10)	102	10/10	10.6 (10)	93	10/10	10.6 (10)	93	10/10	10.6 (10)	93	10/10	9.7 (10)	85	10/10
7	11.6 (10)	10/10	11.7 (10)	101	10/10	10.5 (10)	91	10/10	11.4 (10)	98	10/10	10.5 (10)	91	10/10	9.8 (10)	84	10/10
8	11.8 (10)	10/10	11.4 (10)	97	10/10	9.7 (10)	82	10/10	10.1 (10)	86	10/10	10.4 (10)	88	10/10	9.9 (10)	84	10/10
9	11.9 (10)	10/10	11.9 (10)	100	10/10	11.1 (10)	93	10/10	11.1 (10)	93	10/10	10.7 (10)	90	10/10	9.7 (10)	82	10/10
10	11.5 (10)	10/10	11.0 (10)	95	10/10	10.8 (10)	94	10/10	11.0 (10)	96	10/10	10.3 (10)	90	10/10	9.6 (10)	83	10/10
11	12.5 (10)	10/10	12.3 (10)	98	10/10	11.0 (10)	88	10/10	11.5 (10)	92	10/10	10.5 (10)	84	10/10	9.9 (10)	79	10/10
12	11.6 (10)	10/10	12.3 (10)	106	10/10	11.2 (10)	97	10/10	11.0 (10)	95	10/10	10.5 (10)	91	10/10	9.8 (10)	84	10/10
13	12.3 (10)	10/10	12.0 (10)	98	10/10	11.1 (10)	90	10/10	11.4 (10)	93	10/10	10.7 (10)	87	10/10	9.9 (10)	80	10/10

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

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

Week-Day on Study	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 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	23.5 (10)	10/10	23.6 (10)	100	10/10	23.5 (10)	100	10/10	23.5 (10)	100	10/10	23.5 (10)	100	10/10
1-1	23.2 (10)	10/10	23.1 (10)	100	10/10	23.3 (10)	100	10/10	23.3 (10)	100	10/10	23.3 (10)	100	10/10
1-3	22.7 (10)	10/10	22.9 (10)	101	10/10	22.9 (10)	101	10/10	23.1 (10)	102	10/10	22.9 (10)	101	10/10
1-7	23.7 (10)	10/10	23.6 (10)	100	10/10	23.7 (10)	100	10/10	23.6 (10)	100	10/10	23.3 (10)	98	10/10
2-3	23.7 (10)	10/10	23.9 (10)	101	10/10	23.9 (10)	101	10/10	23.5 (10)	99	10/10	23.3 (10)	98	10/10
2-7	24.7 (10)	10/10	24.4 (10)	99	10/10	24.5 (10)	99	10/10	24.2 (10)	98	10/10	24.0 (10)	97	10/10

< >: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	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 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	18.9 (10)	10/10	18.9 (10)	100	10/10	18.9 (10)	100	10/10	18.9 (10)	100	10/10	18.9 (10)	100	10/10
1-1	18.7 (10)	10/10	18.8 (10)	101	10/10	18.5 (10)	99	10/10	18.5 (10)	99	10/10	18.2 (10)	97	10/10
1-3	18.4 (10)	10/10	18.6 (10)	101	10/10	18.3 (10)	99	10/10	18.3 (10)	99	10/10	18.1 (10)	98	10/10
1-7	18.9 (10)	10/10	18.8 (10)	99	10/10	18.6 (10)	98	10/10	18.5 (10)	98	10/10	18.5 (10)	98	10/10
2-3	18.9 (10)	10/10	19.4 (10)	103	10/10	19.8 (10)	105	10/10	19.0 (10)	101	10/10	18.8 (10)	99	10/10
2-7	19.3 (10)	10/10	19.8 (10)	103	10/10	19.3 (10)	100	10/10	19.3 (10)	100	10/10	18.5 (10)	96	10/10

< >: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	Control		80 ppm			400 ppm			2000 ppm			10000 ppm			50000 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.	Au.F.C.	% of cont. <10>	No. of Surviv.
1-3	4.9 (10)	10/10	4.4 (10)	90	10/10	4.9 (10)	100	10/10	5.1 (10)	104	10/10	4.6 (10)	94	10/10	4.5 (10)	92	10/10
1-7	5.4 (10)	10/10	4.9 (10)	91	10/10	5.3 (10)	98	10/10	5.3 (10)	98	10/10	5.1 (10)	94	10/10	5.1 (10)	94	10/10
2-3	4.7 (10)	10/10	4.3 (10)	91	10/10	4.6 (10)	98	10/10	4.5 (10)	96	10/10	4.4 (10)	94	10/10	4.3 (10)	91	10/10
2-7	5.1 (10)	10/10	5.0 (10)	98	10/10	4.9 (10)	96	10/10	4.8 (10)	94	10/10	4.8 (10)	94	10/10	4.6 (10)	90	10/10

< >: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	Control		80 ppm			400 ppm			2000 ppm			10000 ppm			50000 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.	Au.F.C.	% of cont. <10>	No. of Surviv.
1-3	4.3 (10)	10/10	4.8 (10)	112	10/10	4.5 (10)	105	10/10	4.2 (10)	98	10/10	4.0 (10)	93	10/10	3.9 (10)	91	10/10
1-7	4.6 (10)	10/10	4.7 (10)	102	10/10	4.7 (10)	102	10/10	4.5 (10)	98	10/10	4.2 (10)	91	10/10	4.5 (10)	98	10/10
2-3	4.1 (10)	10/10	4.4 (10)	107	10/10	4.4 (10)	107	10/10	4.1 (10)	100	10/10	3.9 (10)	95	10/10	4.0 (10)	98	10/10
2-7	4.5 (10)	10/10	4.4 (10)	98	10/10	4.5 (10)	100	10/10	4.4 (10)	98	10/10	4.0 (10)	89	10/10	4.1 (10)	91	10/10

< >: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 on Study	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 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	23.0 (10)	10/10	23.0 (10)	100	10/10	23.0 (10)	100	10/10	23.0 (10)	100	10/10	23.0 (10)	100	10/10
1	24.1 (10)	10/10	23.5 (10)	98	10/10	23.4 (10)	97	10/10	23.3 (10)	97	10/10	23.1 (10)	96	10/10
2	25.2 (10)	10/10	24.5 (10)	97	10/10	24.4 (10)	97	10/10	23.8 (10)	94	10/10	23.3 (10)	92	10/10
3	26.0 (10)	10/10	25.5 (10)	98	10/10	24.9 (10)	96	10/10	24.4 (10)	94	10/10	23.9 (10)	92	10/10
4	26.8 (10)	10/10	26.3 (10)	98	10/10	25.8 (10)	96	10/10	25.5 (10)	95	10/10	24.5 (10)	91	10/10
5	27.6 (10)	10/10	27.0 (10)	98	10/10	26.6 (10)	96	10/10	25.9 (10)	94	10/10	24.9 (10)	90	10/10
6	28.5 (10)	10/10	28.1 (10)	99	10/10	27.5 (10)	96	10/10	26.7 (10)	94	10/10	25.5 (10)	89	10/10
7	28.8 (10)	10/10	28.0 (10)	97	10/10	27.6 (10)	96	10/10	26.9 (10)	93	10/10	25.9 (10)	90	10/10
8	30.4 (10)	10/10	29.7 (10)	98	10/10	28.9 (10)	95	10/10	27.9 (10)	92	10/10	26.4 (10)	87	10/10
9	31.2 (10)	10/10	30.3 (10)	97	10/10	29.5 (10)	95	10/10	28.7 (10)	92	10/10	26.8 (10)	86	10/10
10	32.2 (10)	10/10	31.3 (10)	97	10/10	30.6 (10)	95	10/10	29.6 (10)	92	10/10	27.3 (10)	85	10/10
11	32.7 (10)	10/10	32.1 (10)	98	10/10	31.2 (10)	95	10/10	30.1 (10)	92	10/10	27.8 (10)	85	10/10
12	32.6 (10)	10/10	31.6 (10)	97	10/10	31.4 (10)	96	10/10	29.9 (10)	92	10/10	27.8 (10)	85	10/10
13	33.6 (10)	10/10	32.7 (10)	97	10/10	32.5 (10)	97	10/10	31.0 (10)	92	10/10	28.4 (10)	85	10/10

< >: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 on Study	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 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	19.1 (10)	10/10	19.1 (10)	100	10/10	19.1 (10)	100	10/10	19.1 (10)	100	10/10	19.1 (10)	100	10/10
1	19.7 (10)	10/10	19.3 (10)	98	10/10	19.5 (10)	99	10/10	19.4 (10)	98	10/10	18.8 (10)	95	10/10
2	20.3 (10)	10/10	20.0 (10)	99	10/10	19.9 (10)	98	10/10	19.8 (10)	98	10/10	19.2 (10)	95	10/10
3	20.7 (10)	10/10	20.5 (10)	99	10/10	20.9 (9)	101	9/10	20.2 (10)	98	10/10	19.5 (10)	94	10/10
4	21.2 (10)	10/10	21.5 (10)	101	10/10	21.4 (9)	101	9/10	20.8 (10)	98	10/10	19.9 (10)	94	10/10
5	21.7 (10)	10/10	21.2 (10)	98	10/10	21.5 (9)	99	9/10	21.3 (10)	98	10/10	20.1 (10)	93	10/10
6	22.4 (10)	10/10	22.4 (10)	100	10/10	22.6 (9)	101	9/10	22.4 (10)	100	10/10	21.1 (10)	94	10/10
7	22.2 (10)	10/10	22.2 (10)	100	10/10	22.5 (9)	101	9/10	21.9 (10)	99	10/10	20.9 (10)	94	10/10
8	22.6 (10)	10/10	22.6 (10)	100	10/10	22.8 (9)	101	9/10	22.4 (10)	99	10/10	21.4 (10)	95	10/10
9	22.6 (10)	10/10	22.6 (10)	100	10/10	22.6 (9)	100	9/10	22.5 (10)	100	10/10	21.5 (10)	95	10/10
10	23.7 (10)	10/10	23.3 (10)	98	10/10	23.4 (9)	99	9/10	22.9 (10)	97	10/10	22.2 (10)	94	10/10
11	23.9 (10)	10/10	23.7 (10)	99	10/10	23.7 (9)	99	9/10	23.6 (10)	99	10/10	22.4 (10)	94	10/10
12	23.8 (10)	10/10	23.9 (10)	100	10/10	24.0 (9)	101	9/10	23.5 (10)	99	10/10	22.7 (10)	95	10/10
13	24.0 (10)	10/10	23.4 (10)	98	10/10	23.7 (9)	99	9/10	23.3 (10)	97	10/10	22.3 (10)	93	10/10

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

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

Week on Study	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 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	5.2 (10)	10/10	5.1 (10)	98	10/10	4.9 (10)	94	10/10	4.8 (10)	92	10/10	4.7 (10)	90	10/10	4.6 (10)	88	10/10
2	4.9 (10)	10/10	5.0 (10)	102	10/10	4.8 (10)	98	10/10	4.7 (10)	96	10/10	4.3 (10)	88	10/10	4.3 (10)	88	10/10
3	4.8 (10)	10/10	4.9 (10)	102	10/10	4.7 (10)	98	10/10	4.4 (10)	92	10/10	4.1 (10)	85	10/10	4.0 (10)	83	10/10
4	4.9 (10)	10/10	4.9 (10)	100	10/10	4.9 (10)	100	10/10	4.6 (10)	94	10/10	4.4 (10)	90	10/10	4.2 (10)	86	10/10
5	5.3 (10)	10/10	5.2 (10)	98	10/10	5.3 (10)	100	10/10	4.8 (10)	91	10/10	4.7 (10)	89	10/10	4.4 (10)	83	10/10
6	5.2 (10)	10/10	5.3 (10)	102	10/10	5.3 (10)	102	10/10	4.9 (10)	94	10/10	4.6 (10)	88	10/10	4.1 (10)	79	10/10
7	5.0 (10)	10/10	5.2 (10)	104	10/10	5.0 (10)	100	10/10	4.8 (10)	96	10/10	4.5 (10)	90	10/10	4.2 (10)	84	10/10
8	5.7 (10)	10/10	5.9 (10)	104	10/10	5.7 (10)	100	10/10	5.2 (10)	91	10/10	5.2 (10)	91	10/10	4.4 (10)	77	10/10
9	5.7 (10)	10/10	5.8 (10)	102	10/10	5.9 (10)	104	10/10	5.4 (10)	95	10/10	5.0 (10)	88	10/10	4.5 (10)	79	10/10
10	5.0 (10)	10/10	5.3 (10)	106	10/10	5.2 (10)	104	10/10	4.9 (10)	98	10/10	4.6 (10)	92	10/10	4.2 (10)	84	10/10
11	4.7 (10)	10/10	4.8 (10)	102	10/10	5.0 (10)	106	10/10	4.6 (10)	98	10/10	4.6 (10)	98	10/10	4.2 (10)	89	10/10
12	5.2 (10)	10/10	5.1 (10)	98	10/10	5.1 (10)	98	10/10	4.8 (10)	92	10/10	4.6 (10)	88	10/10	4.2 (10)	81	10/10
13	5.1 (10)	10/10	5.5 (10)	108	10/10	5.5 (10)	108	10/10	5.2 (10)	102	10/10	4.7 (10)	92	10/10	4.3 (10)	84	10/10

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

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

Week on Study	Control		80 ppm		400 ppm		2000 ppm		10000 ppm		50000 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	4.9 (10)	10/10	5.2 (10)	106	10/10	5.3 (10)	108	10/10	4.9 (10)	100	10/10	4.5 (10)	92	10/10	4.4 (10)	90	10/10
2	4.5 (10)	10/10	4.8 (10)	107	10/10	4.8 (10)	107	10/10	4.6 (10)	102	10/10	3.9 (10)	87	10/10	4.1 (10)	91	10/10
3	4.8 (10)	10/10	5.1 (10)	106	10/10	5.0 (9)	104	9/10	4.7 (10)	98	10/10	4.0 (10)	83	10/10	3.8 (10)	79	10/10
4	4.8 (10)	10/10	4.7 (10)	98	10/10	5.1 (9)	106	9/10	4.9 (10)	102	10/10	4.5 (10)	94	10/10	4.0 (10)	83	10/10
5	5.3 (10)	10/10	5.6 (10)	106	10/10	5.6 (9)	106	9/10	5.5 (10)	104	10/10	4.7 (10)	89	10/10	4.5 (10)	85	10/10
6	5.1 (10)	10/10	5.4 (10)	106	10/10	5.3 (9)	104	9/10	5.5 (10)	108	10/10	4.8 (10)	94	10/10	4.5 (10)	88	10/10
7	5.0 (10)	10/10	5.6 (10)	112	10/10	5.4 (9)	108	9/10	5.3 (10)	106	10/10	4.6 (10)	92	10/10	4.5 (10)	90	10/10
8	5.6 (10)	10/10	6.1 (10)	109	10/10	6.0 (9)	107	9/10	5.9 (10)	105	10/10	5.4 (10)	96	10/10	4.8 (10)	86	10/10
9	5.8 (10)	10/10	6.3 (10)	109	10/10	6.2 (9)	107	9/10	6.2 (10)	107	10/10	5.3 (10)	91	10/10	5.2 (10)	90	10/10
10	5.0 (10)	10/10	5.7 (10)	114	10/10	5.3 (9)	106	9/10	5.5 (10)	110	10/10	4.8 (10)	96	10/10	4.6 (10)	92	10/10
11	5.3 (10)	10/10	5.5 (10)	104	10/10	5.3 (9)	100	9/10	5.6 (10)	106	10/10	4.9 (10)	92	10/10	4.7 (10)	89	10/10
12	5.2 (10)	10/10	5.7 (10)	110	10/10	5.6 (9)	108	9/10	5.8 (10)	112	10/10	4.9 (10)	94	10/10	4.9 (10)	94	10/10
13	5.5 (10)	10/10	6.0 (10)	109	10/10	5.6 (9)	102	9/10	5.9 (10)	107	10/10	5.1 (10)	93	10/10	5.0 (10)	91	10/10

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