

表(I)-5 血液学的検査(雄 続き)

Items of Examination	Dose ($\mu\text{g}/\text{kg}$ b.w.)				
	0(control)	10	30	100	300
No.of samples 2d	5	5	5	5	5
No.of samples 7d	5	5	5	5	5
No.of samples 36d	5	5	5	5	5
WBC ($10^3/\mu\text{L}$)2d	6.62 \pm 0.98	7.36 \pm 2.51	7.12 \pm 1.49	6.65 \pm 2.46	7.06 \pm 2.39
7d	8.06 \pm 2.59	6.94 \pm 0.96	5.98 \pm 1.90	7.98 \pm 2.41	9.60 \pm 3.21
36d	11.7 \pm 3.5	8.62 \pm 1.26	7.27 \pm 3.33	8.27 \pm 3.66	3.96 \pm 1.60 **
Differential					
NEUTRO	16.6 \pm 3.4	15.6 \pm 4.2	19.8 \pm 7.4	15.2 \pm 4.7	15.0 \pm 2.1
WBC (%)2d					
LYMPHO	78.5 \pm 3.8	77.8 \pm 6.0	74.4 \pm 7.4	79.1 \pm 6.5	79.7 \pm 4.2
MONO	2.6 \pm 0.8	3.8 \pm 1.5	3.7 \pm 0.8	3.3 \pm 1.5	3.6 \pm 1.6
EOSINO	1.2 \pm 0.3	1.2 \pm 0.4	1.0 \pm 0.4	0.8 \pm 0.2	0.5 \pm 0.3 *
BASO	0.2 \pm 0.1	0.2 \pm 0.1	0.2 \pm 0.1	0.1 \pm 0.1	0.2 \pm 0.1
LUC	0.9 \pm 0.5	1.5 \pm 0.7	1.0 \pm 0.8	1.4 \pm 0.7	1.0 \pm 0.4
Differential					
NEUTRO	13.3 \pm 4.4	16.7 \pm 3.4	15.5 \pm 2.5	15.6 \pm 2.1	17.4 \pm 8.8
WBC (%)7d					
LYMPHO	81.3 \pm 4.2	78.4 \pm 4.4	81.0 \pm 2.7	80.6 \pm 2.5	79.0 \pm 8.8
MONO	3.3 \pm 0.8	3.0 \pm 0.7	2.4 \pm 0.4	2.5 \pm 0.5	2.3 \pm 0.6
EOSINO	1.0 \pm 0.3	0.9 \pm 0.3	0.4 \pm 0.2 **	0.2 \pm 0.1 **	0.3 \pm 0.1
BASO	0.1 \pm 0.1	0.1 \pm 0.0	0.2 \pm 0.1	0.1 \pm 0.1	0.2 \pm 0.1
LUC	1.0 \pm 0.7	0.9 \pm 0.5	0.6 \pm 0.1	1.0 \pm 0.4	0.9 \pm 0.1
Differential					
NEUTRO	14.1 \pm 2.8	21.0 \pm 4.5	14.8 \pm 4.5	14.5 \pm 4.2	18.3 \pm 8.7
WBC (%)36d					
LYMPHO	81.5 \pm 2.7	73.8 \pm 4.0	81.5 \pm 5.1	81.5 \pm 4.6	78.6 \pm 8.9
MONO	2.0 \pm 0.7	2.8 \pm 1.2	1.9 \pm 0.7	2.3 \pm 0.8	1.8 \pm 0.7
EOSINO	1.3 \pm 0.2	1.4 \pm 0.5	0.6 \pm 0.3 *	0.6 \pm 0.6 *	0.5 \pm 0.2 *
BASO	0.3 \pm 0.1	0.2 \pm 0.1	0.2 \pm 0.1	0.2 \pm 0.1	0.1 \pm 0.1
LUC	0.9 \pm 0.2	0.8 \pm 0.3	1.0 \pm 0.5	0.8 \pm 0.4	0.7 \pm 0.4
Prothrombin Time(sec)36d	15.7 \pm 0.67	15.4 \pm 0.33	15.4 \pm 0.5	15.5 \pm 0.4	16.3 \pm 1.0 a)
APTT (sec)36d	27.3 \pm 16.2	21.3 \pm 0.8	21.5 \pm 4.8	21.2 \pm 1.2	19.0 \pm 3.1 a)

(Significant difference on the test of Dunnett, *: $p < 0.05$, **: $p < 0.01$)

a) : Number of summerized animals is 4. Clotting time of one animal is scale-out of the equipment range.

表(I)-6 血液学的検査(雌)

Items of Examination	Dose ($\mu\text{g}/\text{kg b.w.}$)				
	0(control)	10	30	100	300
No. of samples 2d	5	5	5	5	5
No. of samples 7d	5	5	5	5	5
No. of samples 36d	5	5	5	5	2
Red blood cell($10^6/\mu\text{L}$) 2d	6.45 \pm 0.38	6.42 \pm 0.35	6.54 \pm 0.49	6.59 \pm 0.41	6.70 \pm 0.30
7d	6.74 \pm 0.36	6.95 \pm 0.26	6.82 \pm 0.42	7.06 \pm 0.49	7.72 \pm 0.57 **
36d	8.00 \pm 0.51	7.66 \pm 0.30	7.46 \pm 0.29	6.95 \pm 0.67 **	5.97 \pm 0.31 a)
Hemoglobin (g/dL)2d	12.9 \pm 0.8	13.4 \pm 0.4	13.4 \pm 0.4	13.5 \pm 0.3	13.2 \pm 0.6
7d	13.6 \pm 0.5	13.9 \pm 0.5	13.5 \pm 0.9	13.7 \pm 1.0	15.2 \pm 1.2
36d	14.6 \pm 0.7	14.0 \pm 0.5	13.3 \pm 0.3 *	12.3 \pm 1.3 **	10.5 \pm 0.8 a)
Hematocrit (%)2d	37.0 \pm 1.7	38.4 \pm 1.1	38.3 \pm 3.2	38.6 \pm 2.1	38.5 \pm 1.3
7d	39.0 \pm 2.1	40.3 \pm 1.3	39.2 \pm 2.2	39.7 \pm 2.4	43.8 \pm 3.7
36d	41.2 \pm 1.9	39.9 \pm 1.5	37.5 \pm 0.6 **	35.3 \pm 3.8 **	30.9 \pm 1.8 a)
MCV (fL)2d	57.4 \pm 0.9	59.8 \pm 1.8	58.5 \pm 1.1	58.6 \pm 1.5	57.4 \pm 1.1
7d	57.9 \pm 1.6	58.0 \pm 0.6	57.5 \pm 1.3	56.3 \pm 2.2	56.6 \pm 1.4
36d	51.6 \pm 1.4	52.2 \pm 1.6	50.3 \pm 1.5	50.7 \pm 0.8	51.8 \pm 0.2 a)
MCH (pg)2d	20.0 \pm 0.5	20.8 \pm 0.8	20.5 \pm 1.1	20.5 \pm 0.8	19.8 \pm 0.4
7d	20.2 \pm 0.4	19.9 \pm 0.2	19.8 \pm 0.5	19.5 \pm 0.8	19.7 \pm 0.7
36d	18.3 \pm 0.5	18.4 \pm 0.6	17.8 \pm 0.5	17.7 \pm 0.4	17.6 \pm 0.4 a)
MCHC (g/dL)2d	34.8 \pm 0.7	34.9 \pm 0.7	35.1 \pm 2.2	35.1 \pm 1.3	34.4 \pm 0.7
7d	34.8 \pm 0.7	34.4 \pm 0.5	34.5 \pm 0.7	34.6 \pm 0.4	34.8 \pm 0.6
36d	35.4 \pm 0.2	35.3 \pm 1.3	35.5 \pm 0.3	34.8 \pm 0.4	34.0 \pm 0.6 a)
Platlet ($10^3/\mu\text{L}$)2d	1440 \pm 114	1349 \pm 146	1324 \pm 210	1182 \pm 169	1262 \pm 154
7d	1357 \pm 85	1311 \pm 86	1144 \pm 131 *	749 \pm 156 **	762 \pm 149 **
36d	1148 \pm 102	1112 \pm 158	998 \pm 149	821 \pm 152 **	354 \pm 192 a)
Reticulocyte(%)2d	2.44 \pm 0.43	2.44 \pm 0.27	2.56 \pm 0.15	2.70 \pm 0.55	3.78 \pm 0.55 **
7d	4.16 \pm 1.11	3.92 \pm 1.15	2.94 \pm 0.55	2.38 \pm 0.19 *	2.76 \pm 1.07
36d	2.52 \pm 0.58	2.76 \pm 0.51	2.80 \pm 0.29	4.54 \pm 1.61	5.30 \pm 0.71 a)

(Significant difference on the test of Dunnett, *: $p < 0.05$, **: $p < 0.01$)

a) : Number of animals is 2. Statistical analysis was not applied.

表(I)-6 血液学的検査(雌 続き)

Items of Examination	Dose ($\mu\text{g}/\text{kg}$ b.w.)				
	0(control)	10	30	100	300
No.of samples 2d	5	5	5	5	5
No.of samples 7d	5	5	5	5	5
No.of samples 36d	5	5	5	5	2
WBC ($10^3/\mu\text{L}$)2d	3.83 ± 0.92	5.00 ± 1.17	6.10 ± 1.93	6.25 ± 1.11	4.99 ± 0.99
7d	4.52 ± 1.03	6.57 ± 1.36	5.82 ± 1.34	9.52 ± 3.38 *	9.63 ± 4.38 *
36d	7.87 ± 0.65	5.65 ± 1.78	4.64 ± 1.82 *	5.18 ± 1.53 *	4.14 ± 0.10 a)
Differential					
NEUTRO	20.3 ± 5.0	15.1 ± 4.0	16.9 ± 2.2	18.1 ± 7.1	17.1 ± 1.7
WBC (%)2d					
LYMPHO	72.9 ± 4.4	80.0 ± 6.1	74.8 ± 3.5	75.9 ± 7.1	76.2 ± 2.7
MONO	3.2 ± 0.8	2.8 ± 1.3	4.4 ± 0.7	3.4 ± 0.3	4.2 ± 1.0
EOSINO	2.5 ± 1.0	1.2 ± 0.5 *	1.8 ± 0.4	1.2 ± 0.5 **	1.2 ± 0.2 **
BASO	0.2 ± 0.1	0.1 ± 0.0	0.2 ± 0.1	0.2 ± 0.0	0.2 ± 0.2
LUC	1.0 ± 0.5	0.9 ± 0.6	2.0 ± 1.5	1.2 ± 1.0	1.1 ± 0.6
Differential					
NEUTRO	14.0 ± 3.7	14.9 ± 1.9	13.5 ± 2.7	12.2 ± 1.9	14.7 ± 4.5
WBC (%)7d					
LYMPHO	81.7 ± 4.5	80.1 ± 2.9	82.2 ± 2.8	83.6 ± 2.4	81.6 ± 4.4
MONO	1.9 ± 0.7	2.7 ± 0.9	2.3 ± 0.6	2.6 ± 0.5	2.4 ± 0.5
EOSINO	1.5 ± 0.4	1.2 ± 0.5	0.9 ± 0.4	0.5 ± 0.3 **	0.3 ± 0.1 **
BASO	0.2 ± 0.2	0.2 ± 0.1	0.2 ± 0.1	0.2 ± 0.1	0.2 ± 0.1
LUC	0.6 ± 0.1	0.9 ± 0.3	1.0 ± 0.4	1.0 ± 0.2	0.8 ± 0.1
Differential					
NEUTRO	19.9 ± 3.5	18.4 ± 5.1	19.0 ± 4.0	16.1 ± 4.1	19.1 ± 0.6 a)
WBC (%)36d					
LYMPHO	74.8 ± 4.4	76.7 ± 6.2	76.2 ± 4.2	80.3 ± 4.7	78.3 ± 1.3 a)
MONO	2.3 ± 0.2	2.6 ± 0.9	2.6 ± 0.4	2.4 ± 0.7	1.6 ± 0.5 a)
EOSINO	2.3 ± 1.0	1.6 ± 0.7	1.4 ± 0.2	0.6 ± 0.1 **	0.4 ± 0.0 a)
BASO	0.2 ± 0.0	0.2 ± 0.1	0.1 ± 0.1	0.1 ± 0.1	0.2 ± 0.1 a)
LUC	0.5 ± 0.1	0.6 ± 0.2	0.6 ± 0.3	0.7 ± 0.2	0.6 ± 0.4 a)
Prothrombin Time(sec)36d	18.1 ± 2.7	16.7 ± 0.3	16.4 ± 0.6	15.7 ± 0.4 **	15.6 ± 1.2 a)
APTT (sec)36d	20.4 ± 2.1	19.3 ± 1.4	20.3 ± 1.8	19.6 ± 2.1	18.8 ± 0.4 a)

(Significant difference on the test of Dunnett, *: $p < 0.05$, **: $p < 0.01$)

a) : Number of animals is 2. Statistical analysis was not applied.

表(I)- 7 血液生化学的检查(雄)

Items of Examination	Dose ($\mu\text{g}/\text{kg}$ b.w.)					
	0(control)	10	30	100	300	
No.of samples 2d	5	5	5	5	5	
No.of samples 7d	5	5	5	5	5	
No.of samples 36d	5	5	5	5	5	
Total protein (g/dL) 2d	5.38 \pm 0.15	5.4 \pm 0.1	5.5 \pm 0.1	5.7 \pm 0.2 **	5.5 \pm 0.1	
7d	5.48 \pm 0.23	5.7 \pm 0.2	5.9 \pm 0.3	6.1 \pm 0.2 *	6.4 \pm 0.4 **	
36d	5.76 \pm 0.21	6.2 \pm 0.3	6.0 \pm 0.2	6.4 \pm 0.3	6.2 \pm 0.5	
Albumin (g/dL) 2d	3.34 \pm 0.11	3.32 \pm 0.13	3.28 \pm 0.08	3.44 \pm 0.21	3.34 \pm 0.23	
7d	3.32 \pm 0.13	3.34 \pm 0.19	3.46 \pm 0.21	3.66 \pm 0.05 *	3.80 \pm 0.24 **	
36d	3.46 \pm 0.18	3.60 \pm 0.19	3.54 \pm 0.17	3.94 \pm 0.23 **	3.82 \pm 0.29	
um protein	Albumin	51.9 \pm 1.8	51.9 \pm 1.4	52.0 \pm 1.8	52.6 \pm 1.8	51.8 \pm 2.2
ctrophoresis	a1 globlin	20.6 \pm 1.2	20.1 \pm 0.7	19.1 \pm 0.7	18.6 \pm 1.6 *	18.7 \pm 0.6 *
2d(%)	a2 globlin	9.0 \pm 1.3	9.0 \pm 2.1	9.2 \pm 1.7	9.1 \pm 1.1	9.3 \pm 1.8
	b globlin	16.6 \pm 1.0	17.2 \pm 1.0	18.0 \pm 1.1	17.9 \pm 1.2	18.7 \pm 1.0
	g globlin	1.9 \pm 0.1	1.8 \pm 0.2	1.7 \pm 0.2	1.7 \pm 0.2	1.6 \pm 0.2
um protein	Albumin	50.4 \pm 1.1	49.2 \pm 1.7	48.6 \pm 1.3	48.5 \pm 1.3	47.7 \pm 1.2
ctrophoresis	a1 globlin	22.1 \pm 2.1	22.8 \pm 2.0	19.8 \pm 1.2	19.1 \pm 1.6	18.4 \pm 2.5 *
7d(%)	a2 globlin	9.0 \pm 1.0	8.9 \pm 1.3	10.8 \pm 0.6	11.5 \pm 1.5 *	12.3 \pm 1.4 **
	b globlin	16.9 \pm 1.0	17.5 \pm 1.2	19.3 \pm 0.9 *	19.5 \pm 1.2 *	20.0 \pm 1.7 **
	g globlin	1.6 \pm 0.3	1.6 \pm 0.5	1.6 \pm 0.5	1.4 \pm 0.4	1.5 \pm 0.3
um protein	Albumin	49.1 \pm 1.8	47.1 \pm 1.2	47.0 \pm 2.0	49.1 \pm 1.3	48.9 \pm 2.0
ctrophoresis	a1 globlin	23.1 \pm 2.3	24.1 \pm 1.4	23.7 \pm 1.6	20.8 \pm 0.6	17.3 \pm 2.1 **
36d(%)	a2 globlin	9.0 \pm 0.5	8.9 \pm 1.2	8.9 \pm 0.7	9.5 \pm 1.3	10.6 \pm 0.9
	b globlin	16.9 \pm 0.9	18.0 \pm 0.8	18.5 \pm 1.1	18.9 \pm 0.9 *	21.6 \pm 3.3 **
	g globlin	1.9 \pm 0.3	2.0 \pm 0.5	1.8 \pm 0.2	1.7 \pm 0.5	1.6 \pm 0.6
A/G ratio 2d		1.66 \pm 0.09	1.60 \pm 0.10	1.52 \pm 0.13	1.48 \pm 0.11	1.50 \pm 0.21
7d		1.54 \pm 0.15	1.42 \pm 0.16	1.44 \pm 0.11	1.52 \pm 0.13	1.48 \pm 0.08
36d		1.50 \pm 0.14	1.38 \pm 0.11	1.42 \pm 0.11	1.62 \pm 0.04	1.62 \pm 0.22
T-Bilirubin (mg/dL) 2d		0.12 \pm 0.02	0.13 \pm 0.03	0.12 \pm 0.02	0.12 \pm 0.02	0.14 \pm 0.03
7d		0.13 \pm 0.02	0.11 \pm 0.01	0.10 \pm 0.01	0.11 \pm 0.01	0.14 \pm 0.05
36d		0.13 \pm 0.02	0.16 \pm 0.04	0.12 \pm 0.02	0.15 \pm 0.03	0.46 \pm 0.57
Glucose (mg/dL) 2d		186 \pm 11	180 \pm 4	166 \pm 8	171 \pm 13	169 \pm 17
7d		199 \pm 11	191 \pm 19	177 \pm 6 *	163 \pm 13 **	147 \pm 12 **
36d		207 \pm 31	184 \pm 10	186 \pm 21	197 \pm 6	149 \pm 35
T-Cholesterol (mg/dL) 2d		78 \pm 10	87 \pm 6	98 \pm 9 *	107 \pm 14 **	96 \pm 12
7d		82 \pm 5	109 \pm 26	127 \pm 16 **	126 \pm 5 **	118 \pm 21 *
36d		62 \pm 9	96 \pm 12	117 \pm 15 **	139 \pm 38 **	131 \pm 26 **
Triglyceride (mg/dL) 2d		67 \pm 28	89 \pm 48	78 \pm 24	70 \pm 34	115 \pm 54
7d		80 \pm 45	57 \pm 25	26 \pm 7 **	30 \pm 13 *	43 \pm 24
36d		83 \pm 28	116 \pm 69	86 \pm 39	64 \pm 30	88 \pm 54
Phospholipid (mg/dL) 2d		171 \pm 19	184 \pm 24	200 \pm 17	206 \pm 11	201 \pm 26
7d		175 \pm 7	199 \pm 36	207 \pm 22	212 \pm 13	217 \pm 35
36d		135 \pm 15	186 \pm 8	213 \pm 23 **	231 \pm 53 **	235 \pm 47 **

(Significant difference on the test of Dunnett, *: $p < 0.05$, **: $p < 0.01$)

表(I)-7 血液生化学的検査(雄 続き)

Items of Examination	Dose($\mu\text{g}/\text{kg b.w.}$)				
	0(control)	10	30	100	300
No.of samples 2d	5	5	5	5	5
No.of samples 7d	5	5	5	5	5
No.of samples 36d	5	5	5	5	5
GOT (IU/L) 2d	56.4 \pm 2.4	58 \pm 5	58 \pm 8	63 \pm 4	62 \pm 4
7d	63.0 \pm 4.4	59 \pm 5	57 \pm 5	76 \pm 26	81 \pm 33
36d	60.4 \pm 7.3	62 \pm 7	73 \pm 17	81 \pm 21	181 \pm 195
GPT (IU/L) 2d	35.0 \pm 5.7	34 \pm 6	35 \pm 4	40 \pm 4	38 \pm 8
7d	32.6 \pm 1.8	29 \pm 4	24 \pm 3 **	21 \pm 4 **	19 \pm 3 **
36d	30.4 \pm 4.2	29 \pm 4	30 \pm 10	28 \pm 3	43 \pm 32
LDH (IU/L) 2d	240 \pm 76	245 \pm 74	367 \pm 237	279 \pm 112	278 \pm 134
7d	313 \pm 49	286 \pm 86	303 \pm 71	324 \pm 38	438 \pm 91 *
36d	197 \pm 46	249 \pm 121	224 \pm 56	326 \pm 194	374 \pm 221
ALP (IU/L) 2d	704 \pm 66	777 \pm 125	698 \pm 149	825 \pm 136	743 \pm 151
7d	711 \pm 64	541 \pm 85	560 \pm 92	460 \pm 139	560 \pm 139
36d	360 \pm 82	325 \pm 65	357 \pm 83	339 \pm 73	550 \pm 181 *
g-GTP(IU/L) 2d	1.2 \pm 0.8	1.0 \pm 0.0	1.0 \pm 0.7	1.2 \pm 0.8	1.4 \pm 0.5
7d	1.2 \pm 0.4	1.2 \pm 0.4	1.2 \pm 0.4	0.8 \pm 0.4	1.2 \pm 0.8
36d	1.8 \pm 0.4	1.8 \pm 0.4	1.6 \pm 0.9	2.0 \pm 0.7	5.8 \pm 8.5
CPK (IU/L) 2d	201 \pm 26	213 \pm 52	264 \pm 90	245 \pm 81	214 \pm 56
7d	260 \pm 70	232 \pm 37	180 \pm 10	192 \pm 21	191 \pm 41
36d	143 \pm 20	138 \pm 46	128 \pm 24	129 \pm 41	108 \pm 15
Urea Nitrogen(mg/L) 2d	18.1 \pm 2.3	17.9 \pm 2.7	15.2 \pm 3.5	15.1 \pm 7.0	13.1 \pm 3.3
7d	15.2 \pm 2.7	14.4 \pm 1.9	14.3 \pm 2.4	16.2 \pm 4.3	18.8 \pm 4.3
36d	21.3 \pm 3.6	20.8 \pm 2.6	17.5 \pm 1.7	20.1 \pm 4.5	32.0 \pm 15.0
Creatinine(mg/dL) 2d	0.4 \pm 0.0	0.4 \pm 0.0	0.4 \pm 0.0	0.4 \pm 0.0	0.4 \pm 0.1
7d	0.4 \pm 0.1	0.4 \pm 0.1	0.5 \pm 0.1	0.5 \pm 0.1	0.5 \pm 0.0
36d	0.5 \pm 0.1	0.5 \pm 0.1	0.5 \pm 0.0	0.5 \pm 0.1	0.5 \pm 0.1
Sodium (mEq/L) 2d	139 \pm 1	139 \pm 1	139 \pm 1	140 \pm 3	139 \pm 3
7d	141 \pm 2	141 \pm 1	141 \pm 1	141 \pm 2	141 \pm 2
36d	138 \pm 2	139 \pm 1	138 \pm 1	138 \pm 1	139 \pm 1
Potassium (mEq/L) 2d	4.94 \pm 0.48	5.02 \pm 0.78	4.86 \pm 0.63	4.76 \pm 0.17	4.80 \pm 0.38
7d	4.22 \pm 0.48	4.16 \pm 0.40	4.68 \pm 0.40	4.72 \pm 0.40	4.68 \pm 0.33
36d	3.86 \pm 0.31	4.02 \pm 0.75	4.18 \pm 0.31	4.26 \pm 0.30	4.80 \pm 0.52
Chloride (mEq/L) 2d	103 \pm 1	103 \pm 1	104 \pm 1	104 \pm 1	104 \pm 2
7d	104 \pm 2	105 \pm 2	105 \pm 2	105 \pm 2	106 \pm 2
36d	103 \pm 3	103 \pm 3	103 \pm 1	104 \pm 3	104 \pm 3
Calcium(mg/dL) 2d	11.0 \pm 0.2	11.0 \pm 0.2	10.8 \pm 0.3	10.9 \pm 0.6	10.9 \pm 0.3
7d	10.8 \pm 0.3	10.7 \pm 0.2	10.6 \pm 0.3	10.8 \pm 0.3	11.1 \pm 0.4
36d	10.6 \pm 0.2	11.1 \pm 0.3	10.8 \pm 0.1	10.8 \pm 0.3	11.1 \pm 0.4
Inorganic phosphorus 2d	9.06 \pm 0.62	9.12 \pm 0.40	9.18 \pm 0.50	8.44 \pm 0.57	7.36 \pm 0.55 **
(mg/dL) 7d	8.94 \pm 0.61	9.32 \pm 0.61	9.44 \pm 0.52	8.52 \pm 0.66	8.48 \pm 0.43
36d	7.28 \pm 1.05	7.20 \pm 0.45	7.24 \pm 0.61	8.02 \pm 0.49	7.46 \pm 0.50

(Significant difference on the test of Dunnett, *: $p < 0.05$, **: $p < 0.01$)

表(I)-8 血液生化学的检查(雌)

Items of Examination	Dose($\mu\text{g}/\text{kg b.w.}$)				
	0(control)	10	30	100	300
No.of samples 2d	5	5	5	5	5
No.of samples 7d	5	5	5	5	5
No.of samples 36d	5	5	5	5	2
Total protein (g/dL) 2d	5.6 \pm 0.2	5.6 \pm 0.2	5.7 \pm 0.2	5.9 \pm 0.2	5.7 \pm 0.1
7d	5.6 \pm 0.3	5.9 \pm 0.2	6.0 \pm 0.3	6.2 \pm 0.3 *	6.4 \pm 0.4 **
36d	6.0 \pm 0.3	6.2 \pm 0.3	6.4 \pm 0.1	6.2 \pm 0.6	5.9 \pm 0.2 a)
Albumin (g/dL) 2d	3.58 \pm 0.19	3.48 \pm 0.23	3.56 \pm 0.21	3.58 \pm 0.15	3.68 \pm 0.04
7d	3.58 \pm 0.27	3.72 \pm 0.15	3.76 \pm 0.21	3.94 \pm 0.09	3.80 \pm 0.16
36d	3.86 \pm 0.27	4.06 \pm 0.26	4.12 \pm 0.22	3.80 \pm 0.54	3.50 \pm 0.28 a)
Serum protein electrophoresis 2d(%)					
Albumin	57.1 \pm 1.2	55.2 \pm 2.3	54.1 \pm 2.2	55.2 \pm 1.8	54.3 \pm 1.7
α 1 globlin	17.8 \pm 0.7	18.0 \pm 1.1	17.0 \pm 0.8	17.3 \pm 1.2	16.0 \pm 1.6
α 2 globlin	7.7 \pm 1.2	8.3 \pm 2.3	8.6 \pm 1.4	8.4 \pm 1.4	9.6 \pm 1.9
β globlin	15.6 \pm 1.5	16.5 \pm 1.5	18.7 \pm 1.3 **	17.5 \pm 1.1	18.2 \pm 1.5 *
γ globlin	1.8 \pm 0.2	2.0 \pm 0.3	1.6 \pm 0.3	1.7 \pm 0.3	1.8 \pm 0.5
Serum protein electrophoresis 7d(%)					
Albumin	54.4 \pm 0.9	52.7 \pm 1.4	52.6 \pm 1.8	51.3 \pm 1.7 *	48.4 \pm 1.5 **
α 1 globlin	18.9 \pm 1.6	17.5 \pm 1.2	16.6 \pm 1.7	17.5 \pm 1.1	20.3 \pm 1.5
α 2 globlin	8.6 \pm 1.3	10.3 \pm 1.9	11.0 \pm 1.6	11.5 \pm 1.4	11.4 \pm 2.1
β globlin	16.0 \pm 0.8	17.6 \pm 0.4	18.3 \pm 0.5 **	18.2 \pm 2.3 *	18.5 \pm 1.1 **
γ globlin	2.0 \pm 0.3	1.9 \pm 0.3	1.5 \pm 0.2 **	1.5 \pm 0.2 **	1.5 \pm 0.1 **
Serum protein electrophoresis 36d(%)					
Albumin	52.7 \pm 0.8	52.7 \pm 2.9	54.7 \pm 3.1	50.0 \pm 2.4	46.8 \pm 4.0 a)
α 1 globlin	19.3 \pm 1.5	19.2 \pm 0.7	17.7 \pm 1.5	17.6 \pm 1.6	20.3 \pm 2.1 a)
α 2 globlin	8.3 \pm 1.4	8.7 \pm 1.3	9.1 \pm 1.2	10.7 \pm 0.6 *	10.4 \pm 3.4 a)
β globlin	16.9 \pm 1.3	17.5 \pm 2.1	16.5 \pm 2.0	20.1 \pm 2.5	20.8 \pm 1.1 a)
γ globlin	2.7 \pm 0.2	2.0 \pm 0.3 **	2.1 \pm 0.3 **	1.6 \pm 0.3 **	1.8 \pm 0.4 a)
A/G ratio 2d	1.78 \pm 0.22	1.66 \pm 0.21	1.66 \pm 0.15	1.58 \pm 0.16	1.84 \pm 0.09
7d	1.78 \pm 0.16	1.68 \pm 0.13	1.68 \pm 0.15	1.74 \pm 0.15	1.46 \pm 0.11 **
36d	1.82 \pm 0.13	1.86 \pm 0.15	1.80 \pm 0.25	1.62 \pm 0.22	1.50 \pm 0.14 a)
T-Bilirubin(mg/dL) 2d	0.10 \pm 0.01	0.11 \pm 0.01	0.12 \pm 0.01	0.11 \pm 0.01	0.11 \pm 0.01
7d	0.11 \pm 0.01	0.11 \pm 0.02	0.12 \pm 0.01	0.13 \pm 0.01	0.18 \pm 0.06
36d	0.16 \pm 0.01	0.15 \pm 0.02	0.16 \pm 0.03	0.18 \pm 0.02	0.21 \pm 0.02 a)
Glucose (mg/dL) 2d	168 \pm 10	159 \pm 11	158 \pm 11	148 \pm 11 *	141 \pm 8 **
7d	189 \pm 7	183 \pm 7	169 \pm 5 *	152 \pm 12 **	129 \pm 13 **
36d	172 \pm 7	173 \pm 13	167 \pm 19	156 \pm 26	149 \pm 7 a)
T-Cholesterol (mg/dL) 2d	68 \pm 6	82 \pm 3	81 \pm 12	79 \pm 8	82 \pm 9
7d	77 \pm 5	129 \pm 20 **	115 \pm 7 **	128 \pm 18 **	163 \pm 18 **
36d	69 \pm 11	93 \pm 7 *	110 \pm 14 **	117 \pm 21 **	120 \pm 70 a)
Triglyceride (mg/dL) 2d	34 \pm 15	41 \pm 21	49 \pm 16	48 \pm 4	25 \pm 12
7d	52 \pm 12	28 \pm 12	38 \pm 30	20 \pm 5 *	16 \pm 7 **
36d	99 \pm 56	91 \pm 49	43 \pm 14	18 \pm 3 **	35 \pm 26 a)
Phospholipid (mg/dL) 2d	143 \pm 11	170 \pm 10	169 \pm 23	164 \pm 15	160 \pm 10
7d	157 \pm 10	231 \pm 30 **	211 \pm 17 *	224 \pm 30 **	292 \pm 36 **
36d	163 \pm 25	201 \pm 14 *	218 \pm 7 **	212 \pm 29 **	226 \pm 110 a)

(Significant difference on the test of Dunnett, *: $p < 0.05$, **: $p < 0.01$)

a) : Number of animals is 2. Statistical analysis was not applied.

表(I)-8 血液生化学的检查(雌 続き)

Items of Examination	Dose ($\mu\text{g}/\text{kg}$ b.w.)				
	0(control)	10	30	100	300
No. of samples 2d	5	5	5	5	5
No. of samples 7d	5	5	5	5	5
No. of samples 36d	5	5	5	5	2
GOT (IU/L) 2d	54 \pm 5	62 \pm 7	62 \pm 6	64 \pm 5	61 \pm 4
7d	53 \pm 5	57 \pm 5	61 \pm 8 *	78 \pm 15 **	311 \pm 281 **
36d	48 \pm 8	60 \pm 18	62 \pm 10	97 \pm 16 **	186 \pm 81 a)
GPT (IU/L) 2d	27 \pm 3	32 \pm 4	38 \pm 5 **	36 \pm 4 *	31 \pm 5
7d	24 \pm 4	25 \pm 3	26 \pm 3	23 \pm 6	24 \pm 9
36d	26 \pm 3	27 \pm 5	25 \pm 4	33 \pm 6	49 \pm 19 a)
LDH (IU/L) 2d	296 \pm 124	405 \pm 217	293 \pm 98	317 \pm 133	310 \pm 68
7d	383 \pm 90	296 \pm 97	328 \pm 68	313 \pm 123	505 \pm 345
36d	235 \pm 105	262 \pm 73	339 \pm 92	388 \pm 82	462 \pm 15 a)
ALP (IU/L) 2d	590 \pm 155	543 \pm 70	682 \pm 132	706 \pm 70	689 \pm 171
7d	439 \pm 49	412 \pm 79	429 \pm 115	454 \pm 98	436 \pm 68
36d	308 \pm 47	296 \pm 67	236 \pm 25	386 \pm 178	502 \pm 23 a)
g-GTP(IU/L) 2d	1.4 \pm 0.5	1.2 \pm 0.4	1.2 \pm 0.4	1.6 \pm 0.5	1.2 \pm 0.4
7d	1.0 \pm 0.7	1.0 \pm 0.0	1.0 \pm 0.7	0.6 \pm 0.5	1.0 \pm 1.0
36d	1.2 \pm 0.4	1.4 \pm 0.5	1.6 \pm 0.5	2.2 \pm 0.4	3.5 \pm 0.7 a)
CPK (IU/L) 2d	210 \pm 67	235 \pm 61	213 \pm 31	268 \pm 106	212 \pm 39
7d	238 \pm 22	194 \pm 62	163 \pm 43	146 \pm 34	169 \pm 48
36d	115 \pm 27	122 \pm 34	132 \pm 38	144 \pm 25	127 \pm 17 a)
Urea Nitrogen(mg/L) 2d	18.1 \pm 2.9	17.5 \pm 1.5	18.3 \pm 3.1	18.9 \pm 2.3	16.4 \pm 6.0
7d	19.2 \pm 3.7	21.5 \pm 3.1	14.6 \pm 5.2	18.7 \pm 2.4	28.1 \pm 12.4
36d	22.6 \pm 2.2	22.3 \pm 2.5	20.4 \pm 2.2	30.9 \pm 10.7	22.6 \pm 2.1 a)
Creatinine(mg/dL) 2d	0.4 \pm 0.0	0.4 \pm 0.0	0.4 \pm 0.0	0.4 \pm 0.1	0.4 \pm 0.0
7d	0.5 \pm 0.1	0.5 \pm 0.0	0.4 \pm 0.1	0.5 \pm 0.0	0.6 \pm 0.1 *
36d	0.5 \pm 0.1	0.5 \pm 0.0	0.5 \pm 0.1	0.6 \pm 0.1	0.4 \pm 0.0 a)
Sodium (mEq/L) 2d	140 \pm 2	139 \pm 1	139 \pm 1	139 \pm 1	139 \pm 2
7d	140 \pm 1	140 \pm 2	139 \pm 2	138 \pm 2	139 \pm 2
36d	138 \pm 2	138 \pm 1	138 \pm 1	137 \pm 1	138 \pm 1 a)
Potassium (mEq/L) 2d	4.22 \pm 0.59	4.70 \pm 0.39	4.52 \pm 0.36	4.72 \pm 0.25	4.66 \pm 0.27
7d	3.70 \pm 0.46	3.76 \pm 0.19	4.46 \pm 0.52 *	4.32 \pm 0.27	4.34 \pm 0.54
36d	3.66 \pm 0.42	3.68 \pm 0.22	4.16 \pm 0.56	4.46 \pm 0.72	4.85 \pm 0.21 a)
Chloride (mEq/L) 2d	104 \pm 2	104 \pm 2	103 \pm 1	104 \pm 2	104 \pm 2
7d	107 \pm 2	106 \pm 2	106 \pm 1	104 \pm 2	104 \pm 3
36d	105 \pm 2	104 \pm 1	105 \pm 2	107 \pm 2	108 \pm 0 a)
Calcium(mg/dL) 2d	10.9 \pm 0.3	11.0 \pm 0.3	11.1 \pm 0.3	11.1 \pm 0.2	10.8 \pm 0.2
7d	10.7 \pm 0.2	10.7 \pm 0.3	10.8 \pm 0.5	11.1 \pm 0.4	11.2 \pm 0.3
36d	10.7 \pm 0.3	11.1 \pm 0.4	11.1 \pm 0.1	11.1 \pm 0.4	10.5 \pm 0.4 a)
Inorganic phosphorus 2d	8.24 \pm 0.15	8.54 \pm 0.66	8.84 \pm 0.89	7.86 \pm 0.40	8.88 \pm 0.47
(mg/dL) 7d	7.52 \pm 1.24	8.12 \pm 1.16	8.58 \pm 0.65	7.74 \pm 0.32	8.28 \pm 0.55
36d	5.06 \pm 1.47	6.06 \pm 0.73	6.38 \pm 0.32	6.88 \pm 0.80	6.30 \pm 0.00 a)

(Significant difference on the test of Dunnett, *: $p < 0.05$, **: $p < 0.01$)

a) : Number of animals is 2. Statistical analysis was not applied.

表(I)-9 血清中のホルモン濃度(雄)

Items of Examination	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 2d	3	3	3	3	3
No.of samples 7d	3	3	3	3	3
No.of samples 36d	3	3	3	3	3
Thyroid stimulating hormone (ng/mL) 2d	25.5 ± 2.7	21.7 ± 9.7	24.2 ± 8.3	23.4 ± 3.5	21.3 ± 5.5
7d	21.7 ± 4.9	21.8 ± 2.3	16.5 ± 2.9	19.9 ± 1.4	18.7 ± 0.9
36d	26.7 ± 3.7	21.9 ± 2.5	18.3 ± 1.4	20.4 ± 4.9	17.7 ± 9.1
Triiodothyronine(T3) (ng/mL) 2d	2.1 ± 0.3	2.3 ± 0.4	2.0 ± 0.1	2.0 ± 0.1	2.0 ± 0.3
7d	2.7 ± 0.2	2.5 ± 0.3	2.8 ± 0.4	2.6 ± 0.3	3.1 ± 0.3
36d	2.4 ± 0.4	2.5 ± 0.4	2.6 ± 0.4	2.7 ± 0.3	2.5 ± 0.1
Thyroxin(T4) (ng/mL) 2d	144.7 ± 21.9	117.3 ± 0.4	96.7 ± 3.1 **	87.9 ± 5.8 **	81.9 ± 6.7 **
7d	178.6 ± 16.5	176.8 ± 16.2	131.2 ± 24.5 *	110.0 ± 5.2 **	108.8 ± 19.0 **
36d	143.5 ± 31.9	165.2 ± 44.9	153.3 ± 18.0	119.8 ± 27.0	81.9 ± 10.0

(Significant difference on the test of Dunnett, *: p<0.05, **: p<0.01)

表(I)-10 血清中のホルモン濃度(雌)

Items of Examination	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 2d	3	3	3	3	3
No.of samples 7d	3	3	3	3	3
No.of samples 36d	3	3	3	3	2
Thyroid stimulating hormone (ng/mL) 2d	29.2 ± 5.9	42.5 ± 16.5	26.9 ± 2.9	27.8 ± 5.7	29.2 ± 4.0
7d	22.0 ± 5.4	65.3 ± 75.9	44.3 ± 50.5	22.5 ± 4.5	15.2 ± 4.8
36d	18.2 ± 3.0	29.0 ± 4.1 **	22.8 ± 2.3	19.4 ± 3.3	21.1 ± 3.3 a)
Triiodothyronine(T3) (ng/mL) 2d	2.2 ± 0.3	2.5 ± 0.5	2.4 ± 0.2	2.3 ± 0.4	2.4 ± 0.4
7d	2.9 ± 0.6	2.9 ± 0.7	3.0 ± 0.4	3.5 ± 0.5	3.7 ± 0.6
36d	2.6 ± 0.2	2.8 ± 0.2	3.0 ± 0.4	2.5 ± 0.3	2.6 ± 0.2 a)
Thyroxin(T4) (ng/mL) 2d	122.2 ± 15.8	110.9 ± 27.2	101.6 ± 17.6	85.9 ± 17.9	88.4 ± 23.7
7d	137.2 ± 37.4	153.9 ± 63.1	115.6 ± 22.0	139.7 ± 33.5	118.6 ± 5.1
36d	101.4 ± 12.6	137.8 ± 22.9	116.9 ± 16.4	88.0 ± 15.5	87.7 ± 2.9 a)

(Significant difference on the test of Dunnett, *: p<0.05, **: p<0.01)

a) : Number of animals is 2. Statistical analysis was not applied.

表(I)-11 肝臓中の誘導酵素量 (雄)

Items of Examination		Dose ($\mu\text{g}/\text{kg}$ b.w.)				
		0(control)	10	30	100	300
No. of samples	2d	2	2	2	2	2
No. of samples	7d	2	2	2	2	2
No. of samples	36d	2	2	2	2	2
AHH	2d	14.9,13.6(14.3)	28.1,26.0(27.1)	26.4,19.3(22.9)	25.5,14.4(20.0)	16.1,22.4(19.3)
(nmol/g liver/min)	7d	6.4,9.3(7.9)	12.0,26.3(19.2)	21.8,12.5(17.2)	17.0,27.1(22.1)	14.1,9.1(11.6)
	36d	15.4,14.1(14.8)	22.5,25.2(23.9)	32.6,20.4(26.5)	21.7,26.5(24.1)	21.7,16.9(19.3)
AHH	2d	0.56,0.58(0.57)	1.19,1.08(1.14)	1.22,1.10(1.16)	1.16,1.17(1.17)	0.96,1.31(1.14)
(nmol/mg protein/min)	7d	0.36,0.51(0.44)	0.63,1.01(0.82)	1.25,0.69(0.97)	1.09,1.19(1.14)	1.06,1.05(1.06)
	36d	0.76,0.76(0.76)	1.06,1.10(1.08)	1.18,0.93(1.06)	1.27,1.08(1.18)	1.27,1.08(1.12)
ECOD	2d	29.8,38.8(34.3)	234,246(240)	232,210(221)	221,138(179)	160,198(179)
(nmol/g liver/min)	7d	23.3,21.8(22.6)	94.6,215(155)	208,204(206)	182,282(232)	145,86.2(116)
	36d	20.3,24.6(22.5)	76.1,94.6(85.4)	231,124(178)	206,293(250)	194,167(181)
ECOD	2d	1.12,1.66(1.39)	9.92,10.2(10.1)	10.7,12.0(11.4)	10.1,11.2(10.6)	9.55,11.6(10.6)
(nmol/mg protein/min)	7d	1.30,1.19(1.25)	4.98,8.26(6.62)	11.9,11.3(11.6)	11.6,12.4(12.0)	10.9,9.91(10.4)
	36d	1.00,1.32(1.16)	3.59,4.13(3.86)	8.41,5.65(7.03)	12.1,12.0(12.1)	11.3,9.62(10.5)
EROD	2d	4.3,4.9(4.6)	97.9,108(103)	103,103(103)	95.5,80.0(87.8)	81.8,104(92.8)
(nmol/g liver/min)	7d	2.9,3.1(3.0)	48.1,92.0(70.1)	83.3,94.1(88.7)	78.3,93.7(86.0)	45.0,39.4(42.2)
	36d	2.2,2.8(2.5)	29.5,41.0(35.3)	101,51.9(76.5)	67.7,105(86.6)	56.8,42.5(49.7)
EROD	2d	0.16,0.21(0.19)	4.15,4.49(4.32)	4.78,5.89(5.34)	4.34,6.50(5.42)	4.87,6.07(5.47)
(nmol/mg protein/min)	7d	0.16,0.17(0.17)	2.53,3.54(3.04)	4.79,5.20(5.00)	5.02,4.11(4.57)	3.38,4.53(3.96)
	36d	0.11,0.15(0.13)	1.39,1.79(1.59)	3.66,2.37(3.02)	3.96,4.30(4.13)	3.32,2.44(2.88)

表(I)-12 肝臓中の誘導酵素量 (雌)

Items of Examination		Dose (µg/kg b.w.)				
		0(control)	10	30	100	300
No. of samples	2d	2	2	2	2	2
No. of samples	7d	2	2	2	2	2
No. of samples	36d	2	2	2	2	2
AHH	2d	2.5,1.6(2.1)	14.3,17.5(15.9)	13.1,19.4(16.3)	7.3,10.8(9.1)	18.1,12.2(15.2)
(nmol/g liver/min)	7d	2.1,2.1(2.1)	25.9,25.9(25.9)	18.1,17.1(17.6)	20.6,23.4(22.0)	19.7,14.4(17.1)
	36d	1.8,2.4(2.1)	14.6,10.1(12.4)	18.4,22.4(20.4)	11.1,24.8(18.0)	14.3,11.6(13.0)
AHH	2d	0.12,0.10(0.11)	0.93,0.80(0.87)	1.01,1.09(1.05)	0.53,0.68(0.61)	1.04,0.95(1.00)
(nmol/mg protein/min)	7d	0.12,0.12(0.12)	1.16,1.10(1.13)	1.12,1.18(1.15)	1.16,1.17(1.17)	1.01,1.24(1.13)
	36d	0.11,0.13(0.12)	0.73,0.54(0.64)	0.89,0.95(0.92)	0.94,1.04(0.99)	1.06,1.01(1.04)
ECOD	2d	11.9,8.4(10.2)	133,208(171)	127,171(149)	146,168(157)	186,149(168)
(nmol/g liver/min)	7d	11.5,10.8(11.2)	231,259(245)	196,165(181)	249,280(265)	221,152(187)
	36d	10.4,10.0(10.2)	127,78.0(103)	175,218(196)	99.6,246(173)	146,131(138)
ECOD	2d	0.57,0.54(0.56)	8.65,9.51(9.08)	9.78,9.60(9.69)	10.6,10.6(10.6)	10.7,11.7(11.2)
(nmol/mg protein/min)	7d	0.64,0.62(0.63)	10.4,11.0(10.7)	12.1,11.4(11.8)	14.0,14.0(14.0)	11.4,13.1(12.2)
	36d	0.64,0.55(0.60)	6.35,4.17(5.26)	8.43,9.22(8.83)	8.44,10.3(9.38)	10.8,11.4(11.1)
EROD	2d	4.8,3.0(3.9)	77.5,103(90.3)	58.6,88.8(73.7)	59.2,71.4(65.3)	78.5,64.4(71.5)
(nmol/g liver/min)	7d	3.4,3.3(3.4)	67.6,79.0(73.3)	56.4,53.4(54.9)	65.0,66.0(65.5)	46.0,44.2(45.1)
	36d	2.4,2.9(2.7)	47.6,30.9(39.3)	61.3,81.9(71.6)	28.8,60.7(44.8)	46.8,51.1(49.0)
EROD	2d	0.23,0.19(0.21)	5.03,4.71(4.87)	4.51,4.99(4.75)	4.29,4.49(4.39)	4.51,5.03(4.77)
(nmol/mg protein/min)	7d	0.19,0.19(0.19)	3.03,3.36(3.20)	3.48,3.68(3.58)	3.65,3.30(3.48)	2.36,3.81(3.09)
	36d	0.15,0.16(0.16)	2.38,1.65(2.02)	2.96,3.47(3.22)	2.44,2.55(2.50)	3.47,4.44(3.96)

表(I)-13 体内負荷量(雄)

Items of Examination		Dose (µg/kg b.w.)				
		0(control)	10	30	100	300
No. of samples	2d	2	0	2	2	2
No. of samples	7d	0	0	0	2	2
No. of samples	36d	0	0	0	2	2
Liver TBDD	2d	0.0095,0.012(0.0108)	-	61,74(68)	250,250(250)	540,630(585)
(ng/g liver)	7d	-	-	-	230,200(215)	650,280(465)
	36d	-	-	-	27,56(42)	110,130(120)
Fat TBDD	2d	0.0045,0.025(0.0148)	-	41,52(47)	130,120(125)	470,410(440)
(ng/g Fat)	7d	-	-	-	210,190(200)	730,270(500)
	36d	-	-	-	27,51(39)	110,130(120)

表(I)-14 体内負荷量(雌)

Items of Examination		Dose (µg/kg b.w.)				
		0(control)	10	30	100	300
No. of samples	2d	2	0	2	2	2
No. of samples	7d	0	0	0	2	2
No. of samples	36d	0	0	0	2	2
Liver TBDD	2d	0.024,0.0077(0.0159)	-	44,48(46)	160,240(200)	720,700(710)
(ng/g liver)	7d	-	-	-	110,210(160)	430,470(450)
	36d	-	-	-	42,47(45)	110,63(87)
Fat TBDD	2d	0.011,0.0086(0.0098)	-	34,33(34)	110,170(140)	530,500(515)
(ng/g Fat)	7d	-	-	-	140,280(210)	500,590(545)
	36d	-	-	-	55,68(62)	130,110(120)

表(I)-15 病理学的検査(剖検)

途中死亡動物		
性	Dose(µg/kg b.w.)	動物番号(死亡日) 剖検所見
雌	300	2413(21d) 肺:赤色斑, 胸腺:萎縮, 頭蓋腔と脊椎腔:出血(胃;内容物なし)
雌	300	2415(23d) 胸腺:萎縮, 皮下(頸部):出血, 頭蓋腔と脊椎腔:出血, 脳:貧血様, 鼻腔:出血
雌	300	2412(27d) 肺:赤色斑, 胸腺:萎縮
定期解剖動物		
性	Dose (µg/kg b.w.)	動物番号(解剖日) 剖検所見
雄	10	1109(7d) 肺:赤色斑
雄	300	1412(36d) 皮下~筋肉(右上肢):出血, 肝:黒色斑
雄	300	1414(36d) 胸水(中等度,透明), 心:肥大(右心室壁, 心室中隔及び右心房壁の肥厚)
雌	100	2311(36d) 肺:赤色斑

表(I)-16 臟器重量 (雄)

Items of Examination	Dose ($\mu\text{g}/\text{kg}$ b. w.)				
	0(control)	10	30	100	300
No. of samples 2d	5	5	5	5	5
No. of samples 7d	5	5	5	5	5
No. of samples 36d	5	5	5	5	5
Body weight (g) 2d	220 \pm 9	219 \pm 13	220 \pm 7	212 \pm 9	209 \pm 11
7d	259 \pm 9	264 \pm 9	242 \pm 13	242 \pm 12	224 \pm 16
36d	387 \pm 20	401 \pm 18	414 \pm 26	364 \pm 42	306 \pm 54 **
Thymus (mg) 2d	615 \pm 78	591 \pm 151	428 \pm 47 *	577 \pm 86	423 \pm 111 *
7d	652 \pm 130	599 \pm 71	441 \pm 87 **	290 \pm 83 **	263 \pm 65 **
36d	666 \pm 128	639 \pm 139	602 \pm 60	360 \pm 136 **	187 \pm 79 **
Thymus (%) 2d	0.279 \pm 0.024	0.268 \pm 0.052	0.195 \pm 0.022 *	0.273 \pm 0.043	0.201 \pm 0.047 *
7d	0.252 \pm 0.047	0.227 \pm 0.026	0.181 \pm 0.031 *	0.120 \pm 0.033 **	0.116 \pm 0.024 **
36d	0.172 \pm 0.028	0.159 \pm 0.033	0.146 \pm 0.019	0.098 \pm 0.037 **	0.060 \pm 0.020 **
Adrenals (mg) 2d	57.2 \pm 4.764	62.2 \pm 8.643	63.4 \pm 5.459	61.2 \pm 6.535	55.2 \pm 5.675
7d	66.6 \pm 9.317	70 \pm 3.536	66.6 \pm 6.542	62.8 \pm 8.526	62.4 \pm 7.925
36d	73.4 \pm 4.037	84.6 \pm 7.369	80.4 \pm 9.839	69.8 \pm 11.61	61.2 \pm 7.259
Adrenals (%) 2d	0.026 \pm 0.002	0.028 \pm 0.003	0.029 \pm 0.002	0.029 \pm 0.004	0.026 \pm 0.003
7d	0.026 \pm 0.003	0.027 \pm 0.002	0.027 \pm 0.002	0.026 \pm 0.003	0.028 \pm 0.003
36d	0.019 \pm 0.002	0.021 \pm 0.002	0.019 \pm 0.002	0.019 \pm 0.003	0.020 \pm 0.002
Testes (mg) 2d	2239 \pm 166	2098 \pm 99	2190 \pm 110	2104 \pm 203	2205 \pm 122
7d	2464 \pm 97	2586 \pm 81	2562 \pm 94	2764 \pm 188 **	2584 \pm 169
36d	3562 \pm 150	3566 \pm 313	3665 \pm 325	3452 \pm 236	3458 \pm 252
Testes (%) 2d	1.022 \pm 0.104	0.963 \pm 0.075	0.999 \pm 0.067	0.993 \pm 0.072	1.054 \pm 0.052
7d	0.952 \pm 0.053	0.978 \pm 0.018	1.062 \pm 0.095 *	1.146 \pm 0.091 **	1.157 \pm 0.118 **
36d	0.922 \pm 0.035	0.891 \pm 0.100	0.885 \pm 0.063	0.956 \pm 0.096	1.157 \pm 0.215
Heart (mg) 2d	831 \pm 66	855 \pm 74	780 \pm 168	815 \pm 45	772 \pm 44
7d	939 \pm 50	1055 \pm 159	877 \pm 66	897 \pm 98	806 \pm 73 *
36d	1179 \pm 96	1197 \pm 112	1276 \pm 65	1315 \pm 251	1203 \pm 401
Heart (%) 2d	0.378 \pm 0.017	0.391 \pm 0.017	0.354 \pm 0.069	0.386 \pm 0.022	0.369 \pm 0.008
7d	0.363 \pm 0.018	0.398 \pm 0.051	0.362 \pm 0.013	0.371 \pm 0.027	0.359 \pm 0.016
36d	0.305 \pm 0.026	0.298 \pm 0.022	0.309 \pm 0.031	0.362 \pm 0.063	0.392 \pm 0.118
Lungs (mg) 2d	1129 \pm 115	1124 \pm 112	1089 \pm 41	1038 \pm 61	1030 \pm 52
7d	1203 \pm 29	1463 \pm 263 *	1227 \pm 143	1272 \pm 85	1208 \pm 74
36d	1396 \pm 76	1229 \pm 388	1473 \pm 108	1505 \pm 212	1468 \pm 272
Lungs (%) 2d	0.513 \pm 0.031	0.514 \pm 0.041	0.496 \pm 0.020	0.491 \pm 0.041	0.493 \pm 0.024
7d	0.465 \pm 0.005	0.552 \pm 0.088 *	0.506 \pm 0.050	0.527 \pm 0.027 *	0.539 \pm 0.023 *
36d	0.362 \pm 0.023	0.307 \pm 0.096	0.356 \pm 0.023	0.415 \pm 0.049	0.485 \pm 0.084 **

(Significant difference on the test of Dunnett, *: $p < 0.05$, **: $p < 0.01$)

表(I)-16 臓器重量(雄 続き)

Items of Examination	Dose($\mu\text{g}/\text{kg}$ b.w.)				
	0(control)	10	30	100	300
No.of samples 2d	5	5	5	5	5
No.of samples 7d	5	5	5	5	5
No.of samples 36d	5	5	5	5	5
Kidneys (mg) 2d	2039 \pm 160	2063 \pm 160	1944 \pm 142	1906 \pm 105	1888 \pm 70
7d	2308 \pm 174	2344 \pm 154	2213 \pm 147	2244 \pm 229	2040 \pm 248
36d	2641 \pm 239	2820 \pm 371	2989 \pm 129	2716 \pm 146	2453 \pm 370
Kidneys (%) 2d	0.927 \pm 0.043	0.944 \pm 0.059	0.885 \pm 0.043	0.901 \pm 0.046	0.903 \pm 0.037
7d	0.891 \pm 0.065	0.887 \pm 0.058	0.914 \pm 0.058	0.928 \pm 0.067	0.906 \pm 0.052
36d	0.683 \pm 0.051	0.701 \pm 0.075	0.722 \pm 0.023	0.752 \pm 0.061	0.809 \pm 0.093
Spleen (mg) 2d	677 \pm 70	758 \pm 62	770 \pm 103	671 \pm 61	583 \pm 198
7d	950 \pm 165	948 \pm 105	818 \pm 102	827 \pm 200	825 \pm 181
36d	856 \pm 115	900 \pm 118	1043 \pm 179	1039 \pm 128	1198 \pm 513
Spleen (%) 2d	0.308 \pm 0.020	0.346 \pm 0.011 *	0.351 \pm 0.044	0.317 \pm 0.021	0.277 \pm 0.091
7d	0.366 \pm 0.057	0.360 \pm 0.051	0.337 \pm 0.030	0.345 \pm 0.092	0.367 \pm 0.072
36d	0.222 \pm 0.031	0.224 \pm 0.024	0.251 \pm 0.038	0.286 \pm 0.027 *	0.401 \pm 0.181 **
Liver (mg) 2d	11090 \pm 437	13496 \pm 1601	14366 \pm 1265 *	14405 \pm 1201 **	15678 \pm 2422 **
7d	12024 \pm 1567	14939 \pm 534 **	15543 \pm 890 **	17411 \pm 1773 **	15853 \pm 1290 **
36d	14819 \pm 1571	17371 \pm 1806	19353 \pm 1607	21171 \pm 4295 **	18731 \pm 3276
Liver (%) 2d	5.050 \pm 0.176	6.164 \pm 0.479	6.541 \pm 0.501 **	6.814 \pm 0.616 **	7.494 \pm 1.155 **
7d	4.634 \pm 0.513	5.656 \pm 0.287 **	6.432 \pm 0.513 **	7.201 \pm 0.518 **	7.062 \pm 0.192 **
36d	3.828 \pm 0.269	4.322 \pm 0.311	4.671 \pm 0.282 *	5.792 \pm 0.694 **	6.120 \pm 0.258 **
Brain (mg) 2d	1724 \pm 96	1781 \pm 43	1808 \pm 125	1774 \pm 62	1853 \pm 300
7d	1844 \pm 37	1681 \pm 281	1733 \pm 365	1795 \pm 51	1756 \pm 46
36d	1889 \pm 48	1956 \pm 47	1970 \pm 77	1843 \pm 73	1798 \pm 55
Brain (%) 2d	0.784 \pm 0.026	0.817 \pm 0.039	0.825 \pm 0.067	0.839 \pm 0.028	0.837 \pm 0.154
7d	0.712 \pm 0.020	0.638 \pm 0.119	0.717 \pm 0.153	0.744 \pm 0.037	0.785 \pm 0.055
36d	0.489 \pm 0.022	0.488 \pm 0.032	0.476 \pm 0.019	0.512 \pm 0.059	0.601 \pm 0.102
Thyroids (mg) 2d	26 \pm 5	33 \pm 4	29 \pm 7	28 \pm 8	31 \pm 6
7d	27 \pm 4	38 \pm 4 **	34 \pm 3 **	37 \pm 4 **	32 \pm 3
36d	28 \pm 1	28 \pm 5	29 \pm 5	34 \pm 8	24 \pm 7
Thyroids (%) 2d	0.012 \pm 0.002	0.015 \pm 0.002	0.013 \pm 0.003	0.013 \pm 0.004	0.015 \pm 0.003
7d	0.011 \pm 0.001	0.014 \pm 0.002 **	0.014 \pm 0.002 **	0.015 \pm 0.002 **	0.014 \pm 0.001 **
36d	0.007 \pm 0.000	0.007 \pm 0.001	0.007 \pm 0.001	0.009 \pm 0.002	0.008 \pm 0.003
Pituitary (mg) 2d	11.2 \pm 1.8	12.6 \pm 1.5	14.6 \pm 2.6 *	14.0 \pm 1.2	12.8 \pm 1.5
7d	12.8 \pm 2.4	15.2 \pm 1.3	12.2 \pm 0.4	13.0 \pm 1.2	12.8 \pm 1.5
36d	10.0 \pm 2.4	11.2 \pm 2.3	11.6 \pm 2.1	11.0 \pm 1.2	10.4 \pm 2.1
Pituitary (%) 2d	0.005 \pm 0.001	0.006 \pm 0.001	0.007 \pm 0.001 *	0.007 \pm 0.001 *	0.006 \pm 0.001
7d	0.005 \pm 0.001	0.006 \pm 0.001	0.005 \pm 0.000	0.005 \pm 0.000	0.006 \pm 0.000
36d	0.003 \pm 0.001	0.003 \pm 0.001	0.003 \pm 0.000	0.003 \pm 0.000	0.004 \pm 0.001

(Significant difference on the test of Dunnett, *: $p < 0.05$, **: $p < 0.01$)

表(I)-17 臟器重量 (雌)

Items of Examination	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 2d	5	5	5	5	5
No.of samples 7d	5	5	5	5	5
No.of samples 36d	5	5	5	5	2
Body weight (g) 2d	167 ± 11	166 ± 7	171 ± 8	166 ± 7	164 ± 7
7d	189 ± 17	190 ± 11	189 ± 9	175 ± 16	163 ± 9 *
36d	262 ± 9	279 ± 16	231 ± 11	230 ± 21	226 ± 25 a)
Thymus (mg) 2d	431 ± 100	459 ± 118	388 ± 75	425 ± 53	446 ± 107
7d	501 ± 57	485 ± 98	355 ± 98 *	240 ± 48 **	207 ± 59 **
36d	526 ± 86	504 ± 152	341 ± 37 *	177 ± 94 **	138 ± 42 a)
Thymus (%) 2d	0.256 ± 0.049	0.275 ± 0.062	0.226 ± 0.035	0.256 ± 0.029	0.271 ± 0.063
7d	0.266 ± 0.020	0.254 ± 0.038	0.187 ± 0.050 **	0.136 ± 0.018 **	0.126 ± 0.031 **
36d	0.201 ± 0.037	0.180 ± 0.051	0.148 ± 0.024	0.078 ± 0.041 **	0.061 ± 0.012 a)
Adrenals (mg) 2d	64.8 ± 4.919	66.8 ± 7.53	66.4 ± 6.066	62.6 ± 7.403	67.6 ± 10.85
7d	78.4 ± 7.765	75 ± 10.32	74.2 ± 6.834	66.6 ± 5.273	71.4 ± 11.46
36d	91 ± 13.78	99 ± 9.247	97.2 ± 18.98	80.8 ± 15.96	81.5 ± 7.778 a)
Adrenals (%) 2d	0.039 ± 0.001	0.040 ± 0.006	0.039 ± 0.004	0.038 ± 0.006	0.041 ± 0.007
7d	0.042 ± 0.004	0.040 ± 0.005	0.039 ± 0.003	0.038 ± 0.003	0.044 ± 0.007
36d	0.035 ± 0.006	0.036 ± 0.002	0.042 ± 0.008	0.035 ± 0.005	0.037 ± 0.007 a)
Ovaries (mg) 2d	127 ± 19	128 ± 11	120 ± 20	118 ± 19	108 ± 10
7d	150 ± 20	132 ± 22	126 ± 21	135 ± 23	123 ± 13
36d	181 ± 22	196 ± 12	170 ± 23	146 ± 37	161 ± 25 a)
Ovaries (%) 2d	0.076 ± 0.010	0.077 ± 0.007	0.070 ± 0.012	0.071 ± 0.010	0.066 ± 0.006
7d	0.080 ± 0.014	0.070 ± 0.010	0.067 ± 0.012	0.077 ± 0.009	0.076 ± 0.008
36d	0.069 ± 0.008	0.070 ± 0.004	0.073 ± 0.007	0.063 ± 0.015	0.071 ± 0.003 a)
Heart (mg) 2d	710 ± 65	682 ± 65	716 ± 53	693 ± 33	697 ± 53
7d	765 ± 107	738 ± 67	760 ± 82	679 ± 63	628 ± 57
36d	958 ± 82	989 ± 119	853 ± 71	856 ± 88	950 ± 100 a)
Heart (%) 2d	0.424 ± 0.017	0.410 ± 0.032	0.418 ± 0.016	0.418 ± 0.029	0.424 ± 0.024
7d	0.404 ± 0.026	0.388 ± 0.014	0.401 ± 0.027	0.389 ± 0.032	0.386 ± 0.019
36d	0.366 ± 0.030	0.354 ± 0.030	0.369 ± 0.023	0.372 ± 0.025	0.421 ± 0.002 a)
Lungs (mg) 2d	952 ± 94	931 ± 26	1007 ± 61	953 ± 59	909 ± 118
7d	1049 ± 98	1085 ± 67	1062 ± 86	1034 ± 110	990 ± 82
36d	1198 ± 31	1212 ± 89	1109 ± 29	1274 ± 113	1312 ± 74 a)
Lungs (%) 2d	0.568 ± 0.029	0.560 ± 0.019	0.588 ± 0.028	0.574 ± 0.028	0.554 ± 0.069
7d	0.556 ± 0.022	0.573 ± 0.036	0.561 ± 0.025	0.592 ± 0.036	0.609 ± 0.033
36d	0.458 ± 0.009	0.436 ± 0.035	0.481 ± 0.036	0.555 ± 0.051 **	0.587 ± 0.097 a)

(Significant difference on the test of Dunnett, *: p<0.05, **: p<0.01)

a) : Number of animals is 2. Statistical analysis was not applied.

表(I)-17 臓器重量 (雌 続き)

Items of Examination	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No. of samples 2d	5	5	5	5	5
No. of samples 7d	5	5	5	5	5
No. of samples 36d	5	5	5	5	2
Kidneys (mg) 2d	1609 ± 142	1524 ± 96	1668 ± 112	1624 ± 124	1636 ± 197
7d	1698 ± 144	1694 ± 173	1827 ± 164	1636 ± 207	1536 ± 134
36d	2076 ± 115	2077 ± 88	1867 ± 138 *	1815 ± 144 *	2213 ± 264 a)
Kidneys (%) 2d	0.961 ± 0.057	0.916 ± 0.025	0.974 ± 0.033	0.978 ± 0.050	0.994 ± 0.095
7d	0.900 ± 0.028	0.891 ± 0.045	0.964 ± 0.057	0.936 ± 0.076	0.944 ± 0.057
36d	0.794 ± 0.052	0.746 ± 0.027	0.807 ± 0.035	0.790 ± 0.055	0.981 ± 0.010 a)
Spleen (mg) 2d	562 ± 76	487 ± 59	593 ± 49	502 ± 90	546 ± 92
7d	638 ± 140	609 ± 43	601 ± 49	587 ± 114	551 ± 51
36d	700 ± 85	729 ± 115	565 ± 43	685 ± 78	688 ± 33 a)
Spleen (%) 2d	0.335 ± 0.034	0.292 ± 0.029	0.347 ± 0.043	0.302 ± 0.049	0.332 ± 0.055
7d	0.336 ± 0.049	0.321 ± 0.005	0.317 ± 0.020	0.334 ± 0.044	0.339 ± 0.028
36d	0.268 ± 0.037	0.262 ± 0.036	0.245 ± 0.017	0.298 ± 0.025	0.306 ± 0.019 a)
Liver (mg) 2d	8474 ± 984	9558 ± 1298	11208 ± 713 **	11545 ± 1074 **	10680 ± 1607 **
7d	8658 ± 820	10754 ± 1146	12560 ± 1274 **	12427 ± 2370 **	11342 ± 1189 **
36d	10663 ± 723	12417 ± 1651	10463 ± 928	11991 ± 1214	14396 ± 1366 a)
Liver (%) 2d	5.051 ± 0.274	5.730 ± 0.564	6.552 ± 0.462 **	6.957 ± 0.546 **	6.482 ± 0.739 **
7d	4.589 ± 0.225	5.656 ± 0.349 **	6.629 ± 0.480 **	7.067 ± 0.799 **	6.966 ± 0.412 **
36d	4.072 ± 0.215	4.451 ± 0.453	4.521 ± 0.239	5.203 ± 0.240 **	6.389 ± 0.095 a)
Brain (mg) 2d	1707 ± 103	1801 ± 290	1690 ± 54	1746 ± 55	1653 ± 42
7d	1732 ± 54	1758 ± 29	1720 ± 89	1699 ± 48	1662 ± 30
36d	1823 ± 26	1871 ± 111	1785 ± 45	1780 ± 73	1787 ± 93 a)
Brain (%) 2d	1.021 ± 0.043	1.081 ± 0.145	0.989 ± 0.058	1.054 ± 0.056	1.008 ± 0.038
7d	0.922 ± 0.060	0.929 ± 0.068	0.910 ± 0.055	0.977 ± 0.065	1.025 ± 0.057
36d	0.697 ± 0.019	0.673 ± 0.046	0.774 ± 0.048	0.777 ± 0.065 *	0.795 ± 0.046 a)
Thyroids (mg) 2d	28 ± 6	25 ± 8	26 ± 8	23 ± 6	30 ± 7
7d	27 ± 3	29 ± 4	29 ± 4	29 ± 4	25 ± 6
36d	24 ± 5	27 ± 6	26 ± 6	25 ± 8	23 ± 6 a)
Thyroids (%) 2d	0.017 ± 0.004	0.015 ± 0.005	0.015 ± 0.004	0.014 ± 0.003	0.018 ± 0.004
7d	0.014 ± 0.002	0.015 ± 0.001	0.015 ± 0.001	0.016 ± 0.002	0.016 ± 0.003
36d	0.009 ± 0.002	0.010 ± 0.002	0.011 ± 0.003	0.011 ± 0.004	0.010 ± 0.002 a)
Pituitary (mg) 2d	15.6 ± 2.6	13.4 ± 1.7	14.4 ± 1.8	12.6 ± 2.5	12.6 ± 1.8
7d	16.6 ± 2.1	15.8 ± 1.3	14.2 ± 3.1	14.2 ± 3.1	12.8 ± 0.8
36d	13.8 ± 1.3	15.0 ± 2.3	14.0 ± 3.8	12.4 ± 1.8	10.5 ± 0.7 a)
Pituitary (%) 2d	0.009 ± 0.002	0.008 ± 0.001	0.008 ± 0.001	0.008 ± 0.001	0.008 ± 0.001
7d	0.009 ± 0.002	0.008 ± 0.000	0.008 ± 0.002	0.008 ± 0.002	0.008 ± 0.001
36d	0.005 ± 0.000	0.005 ± 0.001	0.006 ± 0.002	0.005 ± 0.001	0.005 ± 0.000 a)

(Significant difference on the test of Dunnett, *: p<0.05, **: p<0.01)

a) : Number of animals is 2. Statistical analysis was not applied.

表(I)-18 病理組織学的検査 (途中死亡動物:雌)

Pathological findings	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 2d	0	0	0	0	0
No.of samples 7d	0	0	0	0	0
No.of samples 36d	0	0	0	0	3
liver					
irregular shape: liver cell(+)					2
irregular shape: liver cell(2+)					1
disarrangement: liver cell(2+)					2
disarrangement: liver cell(3+)					1
duct-like arrangement: liver cell(+)					1
duct-like arrangement: liver cell(2+)					2
peliosis(+)					1
peliosis(2+)					1
fibrosis:central(+)					3
inflammatory infiltration(+)					2
inflammatory infiltration(2+)					1
bone marrow					
decreased hematopoiesis(+)					2
decreased hematopoiesis(2+)					1
fibrosis(+)					2
fibrosis(2+)					1
thymus atrophy(4+)					
					3
spleen atrophy: white pulp(+)					
					2
bone thickening of bone(+)					
					2
brain hemorrhage(2+)					
					2
spinal cord hemorrhage(+)					
					1
hemorrhage(2+)					
					1
subcutis hemorrhage(+)					
					(1) 1
nasal cavity hemorrhage(2+)					
					(2) 2
lung congestion(2+)					
					1
congestion(3+)					
					1
edema(+)					
					2

(註)+:軽度、2+:中等度、3+:重度、4+:極重度

表(I)-19 病理組織学的検査 (定期解剖動物:雄 投与後2日)

Pathological findings	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 2d	5	5	5	5	5
liver					
cytoplasmic basophilia:tigroid(+)	0	5	5	5	5

(註) + : 軽度、2+ : 中等度、3+ : 重度

表(I)-20 病理組織学的検査 (定期解剖動物:雄 投与後7日)

Pathological findings	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 7d	5	5	5	5	5
liver					
cytoplasmic basophilia:tigroid(+)	0	5	5	5	5
hepatocellular hypertrophy(+)	0	0	5	5	5
cytoplasmic inclusion:liver cell(+)	0	0	0	0	3
bone marrow					
decreased hematopoiesis(+)	0	0	0	0	2
thymus					
atrophy(+)	0	0	0	2	3
atrophy(2+)	0	0	0	1	2

(註) + : 軽度、2+ : 中等度、3+ : 重度

表(I)-21 病理組織学的検査 (定期解剖動物:雄 投与後36日)

Pathological findings	Dose(μg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 36d	5	5	5	5	5
liver					
cytoplasmic basophilia:tigroid(+)	0	5	5	5	5
hepatocellular hypertrophy(+)	0	5	5	5	4
cytoplasmic inclusion:liver cell(+)	0	0	0	1	0
disarrangement:liver cell(2+)	0	0	0	0	1
duct-like arrangement:liver cell(2+)	0	0	0	0	1
cyst-like arrangement:liver cell(2+)	0	0	0	0	1
fibrosis: central(+)	0	0	0	0	1
bone marrow					
decreased hematopoiesis(+)	0	0	0	2	1
decreased hematopoiesis(2+)	0	0	0	1	2
decreased hematopoiesis(3+)	0	0	0	0	2
fibrosis(+)	0	0	0	1	1
fibrosis(2+)	0	0	0	0	2
thymus					
atrophy(+)	0	0	0	3	0
atrophy(2+)	0	0	0	0	2
atrophy(3+)	0	0	0	0	3
spleen					
extramedullary hematopoiesis(+)	0	0	0	1	4
bone					
thickening of bone(+)	0	0	0	0	1
thickening of bone(2+)	0	0	0	2	3
thyroid					
vacuolar change(2+)	0	0	0	1	0
subcutis					
hemorrhage(2+)	0	0	0	0	(1)1
muscle					
hemorrhage(+)	0	0	0	0	(1)1

(註) + : 軽度、2+ : 中等度、3+ : 重度

表(I)-22 病理組織学的検査 (定期解剖動物：雌 投与後2日)

Pathological findings	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 2d	5	5	5	5	5
liver					
cytoplasmic basophilia: tigroid(+)	0	5	5	5	2

(註) + : 軽度、2+ : 中等度、3+ : 重度

表(I)-23 病理組織学的検査 (定期解剖動物：雌 投与後7日)

Pathological findings	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 7d	5	5	5	5	5
liver					
cytoplasmic basophilia: tigroid(+)	0	5	5	5	5
hepatocellular hypertrophy(+)	0	5	5	5	5
cytoplasmic inclusion: liver cell(+)	0	0	0	0	2
necrosis: single cell(+)	0	0	0	4	4
fibrosis: central(+)	0	0	0	4	4
bone marrow					
decreased hematopoiesis(+)	0	0	0	0	2
thymus					
atrophy(+)	0	0	1	4	1
atrophy(2+)	0	0	0	1	4
karyorrhexis(+)	0	0	0	2	0

(註) + : 軽度、2+ : 中等度、3+ : 重度

表(I)-24 病理組織学的検査 (定期解剖動物:雌 投与後36日)

Pathological findings	Dose (µg/kg b.w.)				
	0(control)	10	30	100	300
No.of samples 36d	5	5	5	5	2
liver					
cytoplasmic basophilia:tigroid(+)	0	5	5	5	2
hepatocellular hypertrophy(+)	0	5	5	5	2
vacuolic change:liver cell(+)	0	0	0	0	1
multinuclear:liver cell(+)	0	0	0	4	2
disarrangement:liver cell(+)	0	0	0	0	1
disarrangement:liver cell(2+)	0	0	0	1	1
duct-like arrangement:liver cell(+)	0	0	0	0	1
duct-like arrangement:liver cell(2+)	0	0	0	1	0
cyst-like arrangement:liver cell(2+)	0	0	0	0	0
fibrosis: central(+)	0	1	4	5	2
bone marrow					
decreased hematopoiesis(+)	0	0	0	2	0
decreased hematopoiesis(2+)	0	0	0	3	0
decreased hematopoiesis(3+)	0	0	0	0	2
fibrosis(+)	0	0	0	1	0
fibrosis(2+)	0	0	0	2	2
thymus					
atrophy(+)	0	0	0	3	0
atrophy(2+)	0	0	0	0	0
atrophy(3+)	0	0	0	2	2
spleen					
atrophy: white pulp(+)	0	0	0	0	1
atrophy: white pulp(2+)	0	0	0	0	1
extramedullary hematopoiesis(+)	0	0	0	0	1
bone					
thickening of bone(+)	0	0	0	2	0
thickening of bone(2+)	0	0	0	3	2
thyroid					
vacuolar change(3+)	0	0	1	0	0

(註) + : 軽度、2+ : 中等度、3+ : 重度