

Table 5 - continued
Urinary findings
Male, Female, 13w

Sex	Group and dose	Number of animals	Ketone body		Bilirubin		Occult blood		Urobilinogen (mg/dL)
			-	-	-	+++	-	+	<1
Male	Control	10	10	10	10	0	10	10	10
	0.1 mg/kg	10	10	10	10	0	10	10	10
	0.5 mg/kg	10	10	10	10	0	10	10	10
	2.5 mg/kg	9	9	9	8	1	9	9	9
Female	Control	10	10	10	10	0	10	10	10
	0.5 mg/kg	10	10	10	10	0	10	10	10
	2.5 mg/kg	10	10	10	10	0	10	10	10
	12.5 mg/kg	10	10	10	10	0	10	10	10

Not significantly different from control.
Grade sign: -, none; ±, trace; +, slight; ++, moderate; +++, severe.

Table 5 - continued

Urinary findings
Male, Female, 13w

Study No. P030097

Sex	Group and dose	Number of animals	Urinary sediment											
			Epithelial cells		Erythrocytes		Leukocytes		Casts	Crystals				
			-	+	-	+++	-	+		-	+	++	+++	
Male	Control	10	10	-	10	0	10	0	10	-	7	0	2	1
	0.1 mg/kg	10	10	-	10	0	10	0	10	-	9	1	0	0
	0.5 mg/kg	10	10	-	10	0	9	1	10	-	5	2	0	3
	2.5 mg/kg	9	9	-	8	1	9	0	9	-	8	0	1	0
Female	Control	10	10	-	10	0	10	0	10	-	10	0	0	0
	0.5 mg/kg	10	10	-	10	0	10	0	10	-	10	0	0	0
	2.5 mg/kg	10	10	-	10	0	10	0	10	-	10	0	0	0
	12.5 mg/kg	10	10	-	10	0	10	0	10	-	10	0	0	0

Not significantly different from control.

Grade signs are as follows.

Epithelial cells: -, < 3/field; +, 3/field \leq and < 10/field; ++, 10/field \leq and < 20/field; +++, \geq 20/field.
 Erythrocytes: -, < 10/field; +, 10/field \leq and < 30/field; ++, 30/field \leq and < 100/field; +++, countless.
 Leukocytes: -, < 3/field; +, 3/field \leq and < 20/field; ++, 20/field \leq and < 40/field; +++, \geq 40/field.
 Casts: -, none; +, \geq 1/all field.
 Crystals: -, < 10/field; +, 10/field \leq and < 20/field; ++, 20/field \leq and < 30/field; +++, countless.

Table 6 Urinary findings
Male, Female, 52w

Study No. P030097

Sex	Group and dose		Urine volume	Osmotic pressure	Specific gravity	Na	K	Cl
			(mL/24hr)	(Osm/kg)		(mEq/24hr)	(mEq/24hr)	(mEq/24hr)
Male	Control	N	10	10	10	10	10	10
		Mean	9.3	1.316	1.047	0.663	1.501	0.774
		S. D.	±3.3	±0.305	±0.013	±0.468	±0.480	±0.522
	0.1 mg/kg	N	8	8	8	8	8	8
		Mean	7.9	1.556	1.056	0.546	1.376	0.566
		S. D.	±3.0	±0.330	±0.013	±0.299	±0.270	±0.306
	0.5 mg/kg	N	10	10	10	10	10	10
		Mean	9.1	1.910**	1.065*	0.822	1.801	0.911
		S. D.	±2.5	±0.418	±0.016	±0.373	±0.286	±0.394
	2.5 mg/kg	N	10	10	10	10	10	10
		Mean	8.6	1.809*	1.061	0.870	1.517	0.831
		S. D.	±2.2	±0.417	±0.013	±0.461	±0.413	±0.353
Female	Control	N	10	10	10	10	10	10
		Mean	9.9	1.325	1.046	0.727	1.836	1.043
		S. D.	±5.0	±0.352	±0.013	±0.425	±0.648	±0.505
	0.5 mg/kg	N	10	10	10	10	10	10
		Mean	12.6	1.200	1.042	0.730	1.753	1.019
		S. D.	±7.3	±0.509	±0.017	±0.251	±0.226	±0.263
	2.5 mg/kg	N	10	10	10	10	10	10
		Mean	14.7	1.043	1.037	0.722	1.780	1.077
		S. D.	±8.7	±0.432	±0.018	±0.309	±0.393	±0.346
	12.5 mg/kg	N	9	9	9	9	9	9
		Mean	19.0*	0.840*	1.030	0.999	2.167	1.315
		S. D.	±9.0	±0.219	±0.008	±0.256	±0.418	±0.308

*: P<0.05, **: P<0.01 (significantly different from control).

Table 6 - continued

Urinary findings
Male, Female, 52w

Study No. P030097

Sex	Group and dose	Number of animals	Color		pH							Protein						Glucose	
			PY	Y	5.5	6.0	6.5	7.0	7.5	8.0	8.5	-	±	+	++	+++	++++	†	-
Male	Control	10	2	8	1	0	0	1	3	4	1	2	0	3	5	0	0	10	
	0.1 mg/kg	8	0	8	0	0	0	1	1	6	0	0	2	5	1	0	0	8	
	0.5 mg/kg	10	0	10	2	2	3	0	1	2	0	0	1	3	3	2	1	10	
	2.5 mg/kg	10	0	10	0	2	0	1	1	4	2	0	1	5	3	1	0	10	
Female	Control	10	0	10	2	0	0	0	1	3	4	6	2	2	0	0	0	10	
	0.5 mg/kg	10	1	9	1	1	1	1	1	4	1	6	1	3	0	0	0	10	
	2.5 mg/kg	10	3	7	2	0	3	1	1	2	1	5	1	3	1	0	0	10	
	12.5 mg/kg	9	2	7	1	1	3	1	1	2	0	3	4	2	0	0	0	9	

Not significantly different from control.

Abbreviation: PY, pale yellow; Y, yellow; YB, yellowish brown; B, brown.

Grade sign: -, none; ±, trace; +, slight; ++, moderate; +++, severe; +++++, very severe.

Table 6 - continued

Urinary findings
Male, Female, 52w

Study No. P030097

Sex	Group and dose	Number of animals	Ketone body		Bilirubin			Occult blood			Urobilinogen (mg/dL)
			-	-	-	±	+	++	+++	<1	
Male	Control	10	10	10	9	1	0	10	10	10	10
	0.1 mg/kg	8	8	8	7	0	1	8	8	8	8
	0.5 mg/kg	10	10	10	9	0	1	10	10	10	10
	2.5 mg/kg	10	10	10	10	0	0	10	10	10	10
Female	Control	10	10	10	10	0	0	10	10	10	10
	0.5 mg/kg	10	10	10	10	0	0	10	10	10	10
	2.5 mg/kg	10	10	10	10	0	0	10	10	10	10
	12.5 mg/kg	9	9	9	9	0	0	9	9	9	9

Not significantly different from control.

Grade sign: -, none; ±, trace; +, slight; ++, moderate; +++, severe; +++++, very severe.

Table 6 - continued

Urinary findings
Male, Female, 52w

Study No. P030097

Sex	Group and dose	Number of animals	Urinary sediment									
			Epithelial cells		Erythrocytes		Leukocytes		Casts	Crystals		
			-	+	-	+	-	+	-	+	+++	
Male	Control	10	10	0	10	0	9	1	10	10	0	0
	0.1 mg/kg	8	8	0	8	0	7	1	8	7	0	1
	0.5 mg/kg	10	10	1	9	1	9	1	10	9	1	0
	2.5 mg/kg	10	10	0	10	0	10	0	10	9	1	0
Female	Control	10	10	0	10	0	10	0	10	9	1	0
	0.5 mg/kg	10	10	0	10	0	10	0	10	8	2	0
	2.5 mg/kg	10	10	0	10	0	10	0	10	10	0	0
	12.5 mg/kg	9	9	0	9	0	9	0	9	9	0	0

Not significantly different from control.

Grade signs are as follows.

Epithelial cells: -, < 3/field; +, 3/field \leq and < 10/field; ++, 10/field \leq and < 20/field; +++, \geq 20/field.
 Erythrocytes: -, < 10/field; +, 10/field \leq and < 30/field; ++, 30/field \leq and < 100/field; +++, countless.
 Leukocytes: -, < 3/field; +, 3/field \leq and < 20/field; ++, 20/field \leq and < 40/field; +++, \geq 40/field.
 Casts: -, none; +, \leq 1/all field.
 Crystals: -, < 10/field; +, 10/field \leq and < 20/field; ++, 20/field \leq and < 30/field; +++, countless.

Table 7 Hematological findings
Male, Female, 13w

Study No. P030097

Sex	Group and dose		Leukocytes (10 ³ / μ L)	Erythrocytes (10 ⁴ / μ L)	Hemoglobin (g/dL)	Hematocrit (%)	MCV (fL)	MCH (pg)	MCHC (g/dL)	Reticulocyte (10 ⁴ / μ L)	Platelets (10 ⁴ / μ L)
Male	Control	N	10	10	10	10	10	10	10	10	10
		Mean	7.83	855	15.6	43.3	50.6	18.3	36.1	15.8	103.4
		S.D.	± 1.32	± 27	± 0.4	± 1.6	± 1.6	± 0.5	± 0.7	± 2.8	± 11.2
	0.1 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	8.63	870	15.5	43.2	49.6	17.8	35.9	16.3	108.7
		S.D.	± 2.72	± 29	± 0.5	± 1.2	± 1.1	± 0.5	± 0.4	± 1.8	± 8.2
	0.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	9.08	828	15.0*	41.6*	50.3	18.1	36.0	16.2	112.9
		S.D.	± 2.48	± 43	± 0.6	± 1.8	± 0.9	± 0.5	± 0.7	± 3.2	± 16.0
	2.5 mg/kg	N	9	9	9	9	9	9	9	9	9
		Mean	9.97	807**	14.3**	40.0**	49.6	17.7	35.7	14.8	130.5*
		S.D.	± 1.88	± 22	± 0.6	± 1.3	± 2.6	± 1.0	± 0.5	± 3.6	± 27.1
Female	Control	N	10	10	10	10	10	10	10	10	10
		Mean	4.94	768	13.9	40.1	52.2	18.1	34.6	16.5	106.1
		S.D.	± 1.16	± 38	± 0.5	± 1.7	± 1.1	± 0.4	± 0.5	± 3.4	± 12.1
	0.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	4.30	793	14.1	40.7	51.3	17.7	34.6	13.9	110.4
		S.D.	± 1.04	± 40	± 0.6	± 2.2	± 0.7	± 0.4	± 0.6	± 1.9	± 6.8
	2.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	4.95	762	13.8	39.5	51.9	18.1	35.0	14.8	117.4
		S.D.	± 0.99	± 23	± 0.4	± 0.9	± 1.3	± 0.5	± 0.3	± 3.4	± 11.6
	12.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	5.18	753	13.4	38.1*	50.6**	17.7	35.1*	13.7	106.2
		S.D.	± 1.70	± 25	± 0.5	± 1.2	± 1.0	± 0.5	± 0.5	± 1.6	± 9.9

*: P<0.05, **: P<0.01 (significantly different from control).

Table 7 - continued. Hematological findings
Male, Female, 13w

Study No. P030097

Sex	Group and dose		PT (sec)	APTT (sec)
Male	Control	N	10	10
		Mean	15.2	24.6
		S. D.	±2.7	±1.9
	0.1 mg/kg	N	10	10
		Mean	14.9	24.1
		S. D.	±1.1	±1.9
	0.5 mg/kg	N	10	10
		Mean	15.1	23.0
		S. D.	±1.5	±1.5
	2.5 mg/kg	N	9	9
		Mean	14.3	23.5
		S. D.	±1.4	±3.1
Female	Control	N	10	10
		Mean	11.7	19.2
		S. D.	±0.5	±1.5
	0.5 mg/kg	N	10	10
		Mean	11.7	19.7
		S. D.	±0.3	±0.9
	2.5 mg/kg	N	10	10
		Mean	11.7	19.0
		S. D.	±0.3	±1.6
	12.5 mg/kg	N	10	10
		Mean	11.8	19.2
		S. D.	±0.4	±1.5

Not significantly different from control.

Table 7 - continued

Hematological findings
Male, Female, 13w

Study No. P030097

Sex	Group and dose		Differential leukocyte count					
			Eosinophils	Neutrophils	Lymphocytes	Basophils	Monocytes	Large unstained cells
			(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)
Male	Control	N	10	10	10	10	10	10
		Mean	1.2	16.2	58.7	0.1	1.7	0.3
		S. D.	± 0.6	± 6.1	± 14.3	± 0.1	± 0.5	± 0.1
	0.1 mg/kg	N	10	10	10	10	10	10
		Mean	1.4	16.8	65.5	0.2	1.8	0.5
		S. D.	± 0.4	± 9.8	± 23.5	± 0.1	± 0.7	± 0.4
	0.5 mg/kg	N	10	10	10	10	10	10
		Mean	1.3	13.8	72.9	0.2	2.0	0.7*
		S. D.	± 0.7	± 5.1	± 21.7	± 0.1	± 1.0	± 0.3
	2.5 mg/kg	N	9	9	9	9	9	9
		Mean	1.1	14.4	81.1	0.2	2.2	0.7
		S. D.	± 0.4	± 5.5	± 19.9	± 0.1	± 0.7	± 0.5
Female	Control	N	10	10	10	10	10	10
		Mean	0.9	7.1	40.1	0.1	0.8	0.4
		S. D.	± 0.4	± 2.7	± 9.5	± 0.1	± 0.3	± 0.2
	0.5 mg/kg	N	10	10	10	10	10	10
		Mean	0.8	6.3	34.9	0.1	0.7	0.3
		S. D.	± 0.5	± 1.6	± 8.5	± 0.1	± 0.3	± 0.1
	2.5 mg/kg	N	10	10	10	10	10	10
		Mean	0.8	8.3	39.4	0.1	0.6	0.4
		S. D.	± 0.3	± 3.8	± 8.7	± 0.0	± 0.2	± 0.2
	12.5 mg/kg	N	10	10	10	10	10	10
		Mean	0.9	6.2	43.5	0.1	0.8	0.3
		S. D.	± 0.3	± 2.2	± 15.2	± 0.1	± 0.4	± 0.2

*: P<0.05 (significantly different from control).

Table 8 Hematological findings
Male, Female, 52w

Study No. P030097

Sex	Group and dose		Leukocytes ($10^3 / \mu\text{L}$)	Erythrocytes ($10^4 / \mu\text{L}$)	Hemoglobin (g/dL)	Hematocrit (%)	MCV (fL)	MCH (pg)	MCHC (g/dL)	Reticulocyte ($10^4 / \mu\text{L}$)	Platelets ($10^4 / \mu\text{L}$)	
Male	Control	N	10	10	10	10	10	10	10	10	10	
		Mean	7.98	840	14.0	44.2	52.7	16.7	31.7	18.2	106.5	
		S. D.	± 1.24	± 68	± 1.1	± 2.9	± 2.1	± 0.8	± 0.8	± 8.4	± 12.6	
	0.1 mg/kg	N	8	8	8	8	8	8	8	8	8	8
		Mean	8.22	780	13.1	41.3	53.1	16.7	31.5	20.1	110.2	
		S. D.	± 3.46	± 145	± 2.7	± 7.4	± 1.2	± 0.7	± 1.5	± 9.8	± 28.5	
	0.5 mg/kg	N	10	10	10	10	10	10	10	10	10	10
		Mean	6.69	754*	12.7	40.3	53.9	16.9	31.3	27.1	123.7	
		S. D.	± 0.98	± 133	± 2.1	± 5.7	± 4.5	± 1.0	± 1.1	± 20.4	± 28.5	
	2.5 mg/kg	N	10	10	10	10	10	10	10	10	10	10
		Mean	7.90	778*	12.9	40.7*	52.3	16.6	31.8	15.7	140.1**	
		S. D.	± 2.19	± 66	± 1.1	± 3.6	± 2.3	± 0.7	± 0.3	± 3.3	± 13.6	
Female	Control	N	10	10	10	10	10	10	10	10	10	
		Mean	3.83	707	13.2	40.3	57.5	18.8	32.7	14.9	90.2	
		S. D.	± 1.30	± 100	± 1.4	± 3.8	± 4.3	± 1.0	± 0.9	± 8.9	± 10.0	
	0.5 mg/kg	N	10	10	10	10	10	10	10	10	10	10
		Mean	3.73	708	13.5	41.0	58.1	19.1	33.0	16.4	94.2	
		S. D.	± 0.96	± 62	± 0.8	± 2.5	± 2.3	± 0.7	± 0.5	± 9.6	± 14.7	
	2.5 mg/kg	N	10	10	10	10	10	10	10	10	10	10
		Mean	3.98	730	13.5	41.3	56.6	18.5	32.7	13.9	101.5	
		S. D.	± 1.44	± 55	± 1.0	± 3.0	± 2.4	± 0.8	± 0.6	± 5.8	± 13.9	
	12.5 mg/kg	N	9	9	9	9	9	9	9	9	9	9
		Mean	3.69	673	12.3	37.3	56.1	18.4	32.9	17.1	105.6*	
		S. D.	± 0.58	± 115	± 1.5	± 4.4	± 4.8	± 1.4	± 0.4	± 15.1	± 11.9	

*: $P < 0.05$, **: $P < 0.01$ (significantly different from control).

Table 8 - continued

Hematological findings
Male, Female, 52w

Study No. P030097

Sex	Group and dose		PT	APTT
			(sec)	(sec)
Male	Control	N	10	10
		Mean	13.5	21.5
		S. D.	±1.0	±1.5
	0.1 mg/kg	N	8	8
		Mean	13.8	20.9
		S. D.	±1.0	±2.7
	0.5 mg/kg	N	10	10
		Mean	14.5	21.2
		S. D.	±1.9	±2.6
	2.5 mg/kg	N	10	10
		Mean	21.8**	29.5
		S. D.	±9.0	±9.3
Female	Control	N	10	10
		Mean	12.3	18.4
		S. D.	±0.8	±0.9
	0.5 mg/kg	N	10	10
		Mean	12.9	18.5
		S. D.	±0.7	±0.9
	2.5 mg/kg	N	10	10
		Mean	12.5	17.7
		S. D.	±0.5	±1.4
	12.5 mg/kg	N	9	9
		Mean	12.1	17.7
		S. D.	±0.5	±1.2

** : P<0.01 (significantly different from control).

Table 8 - continued

Hematological findings
Male, Female, 52w

Study No. P030097

Sex	Group and dose	Differential leukocyte count						
		Bosinophils	Neutrophils	Lymphocytes	Basophils	Monocytes	Large unstained cells	
		(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	(10 ² / μ L)	
Male	Control	N	10	10	10	10	10	10
		Mean	1.2	26.2	47.8	0.1	3.0	1.5
		S. D.	± 0.3	± 9.9	± 11.9	± 0.0	± 0.9	± 2.0
	0.1 mg/kg	N	8	8	8	8	8	8
		Mean	1.0	30.8	46.4	0.1	2.5	1.5
		S. D.	± 0.6	± 29.1	± 6.5	± 0.1	± 0.5	± 2.4
	0.5 mg/kg	N	10	10	10	10	10	10
		Mean	0.9	18.6	44.3	0.1	2.3	0.7
		S. D.	± 0.3	± 7.3	± 6.3	± 0.0	± 0.7	± 0.2
	2.5 mg/kg	N	10	10	10	10	10	10
		Mean	1.1	17.6	56.2	0.1	2.9	1.1
		S. D.	± 0.5	± 7.5	± 17.5	± 0.1	± 0.8	± 0.7
Female	Control	N	10	10	10	10	10	10
		Mean	0.6	10.1	25.4	0.0	1.4	0.7
		S. D.	± 0.2	± 8.0	± 8.2	± 0.0	± 0.8	± 0.3
	0.5 mg/kg	N	10	10	10	10	10	10
		Mean	0.7	11.0	23.3	0.0	1.6	0.6
		S. D.	± 0.2	± 4.0	± 7.2	± 0.0	± 0.5	± 0.3
	2.5 mg/kg	N	10	10	10	10	10	10
		Mean	0.9	12.1	24.6	0.0	1.6	0.6
		S. D.	± 0.4	± 6.7	± 8.2	± 0.0	± 0.6	± 0.3
	12.5 mg/kg	N	9	9	9	9	9	9
		Mean	0.8	9.2	24.8	0.0	1.6	0.5
		S. D.	± 0.2	± 1.9	± 6.3	± 0.0	± 0.5	± 0.2

Not significantly different from control.

Table 9 Biochemical findings
Male, Female, 13w

Study No. P030097

Sex	Group and dose		T. Protein (g/dL)	A/G ratio	α_1 -Globulin (%)	α_2 -Globulin (%)	β -Globulin (%)	γ -Globulin (%)	Albumin (%)
Male	Control	N	10	10	10	10	10	10	10
		Mean	5.8	1.22	18.7	7.1	15.2	4.2	54.8
		S.D.	±0.3	±0.12	±1.6	±0.7	±0.8	±0.5	±2.3
	0.1 mg/kg	N	10	10	10	10	10	10	10
		Mean	5.8	1.30	17.9	6.8	14.4	4.3	56.6
		S.D.	±0.2	±0.09	±1.6	±0.6	±0.6	±0.6	±1.6
	0.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	5.7	1.67**	15.6**	5.9**	11.5**	4.6	62.4**
		S.D.	±0.5	±0.23	±1.3	±0.6	±1.0	±0.8	±2.9
	2.5 mg/kg	N	9	9	9	9	9	9	9
		Mean	5.8	2.09**	12.1**	5.6**	9.9**	5.0	67.4**
		S.D.	±0.5	±0.27	±2.4	±0.6	±0.7	±1.4	±3.0
Female	Control	N	10	10	10	10	10	10	10
		Mean	6.2	1.78	13.8	5.6	12.6	3.9	64.0
		S.D.	±0.4	±0.16	±1.0	±0.8	±0.9	±0.8	±2.0
	0.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	6.3	1.87	12.9	5.6	12.4	4.3	64.9
		S.D.	±0.2	±0.22	±1.7	±0.2	±1.2	±1.0	±2.8
	2.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	6.4	1.93	12.6	5.5	12.1	4.2	65.7
		S.D.	±0.4	±0.19	±1.6	±0.6	±1.4	±1.0	±2.2
	12.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	6.7*	2.24**	12.9	4.7*	9.9**	3.6	68.9**
		S.D.	±0.5	±0.31	±1.8	±0.5	±0.8	±1.1	±2.9

*: P<0.05, **: P<0.01 (significantly different from control).

Table 9 - continued

Biochemical findings
Male, Female, 13w

Study No. P030097

Sex	Group and dose		T. Bilirubin (mg/dL)	AST (IU/L)	ALT (IU/L)	ALP (IU/L)	T. Cholesterol (mg/dL)	Triglycerides (mg/dL)	Phospholipids (mg/dL)	Glucose (mg/dL)
Male	Control	N	10	10	10	10	10	10	10	10
		Mean	0.0	90	35	164	68	48	119	121
		S. D.	±0.0	±22	±29	±23	±12	±21	±21	±9
	0.1 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	0.0	118	47	216	64	43	116	120
		S. D.	±0.0	±66	±58	±57	±12	±12	±23	±7
	0.5 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	0.0	83	23	373**	59	58	125	154**
		S. D.	±0.0	±14	±4	±60	±13	±29	±27	±13
	2.5 mg/kg	N	9	9	9	9	9	9	9	9
		Mean	0.0	94	32	619**	58	65	131	151**
		S. D.	±0.0	±16	±9	±115	±9	±19	±16	±9
Female	Control	N	10	10	10	10	10	10	10	10
		Mean	0.1	75	20	92	75	29	150	119
		S. D.	±0.1	±20	±7	±30	±12	±20	±21	±13
	0.5 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	0.0*	69	18	107	78	24	155	117
		S. D.	±0.0	±8	±6	±25	±10	±9	±15	±10
	2.5 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	0.0	77	17	101	72	23	147	118
		S. D.	±0.0	±14	±2	±31	±12	±10	±23	±15
	12.5 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	0.0*	68	18	136	83	35	169	130
		S. D.	±0.0	±10	±2	±81	±7	±13	±16	±10

*: P<0.05, **: P<0.01 (significantly different from control).

Table 9 - continued

Biochemical findings
Male, Female, 13w

Study No. P030097

Sex	Group and dose		BUN (mg/dL)	Creatinine (mg/dL)	IP (mg/dL)	Ca (mg/dL)	Na (mEq/L)	K (mEq/L)	Cl (mEq/L)
Male	Control	N	10	10	10	10	10	10	10
		Mean	12.3	0.4	6.1	10.2	147.1	4.14	106.5
		S.D.	±1.1	±0.1	±0.6	±0.2	±0.6	±0.13	±1.0
	0.1 mg/kg	N	10	10	10	10	10	10	10
		Mean	11.8	0.4	6.4	10.3	147.3	4.36	105.7
		S.D.	±1.7	±0.0	±0.3	±0.3	±0.7	±0.21	±1.1
	0.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	14.2*	0.4	6.6	10.1	145.9*	4.53**	105.3
		S.D.	±1.7	±0.0	±0.5	±0.4	±0.9	±0.28	±1.8
	2.5 mg/kg	N	9	9	9	9	9	9	9
		Mean	14.8**	0.4	6.8*	10.0	144.9**	4.66**	105.7
		S.D.	±1.8	±0.0	±0.6	±0.3	±1.1	±0.32	±1.7
Female	Control	N	10	10	10	10	10	10	10
		Mean	14.5	0.5	4.9	10.4	145.7	4.02	108.6
		S.D.	±1.7	±0.1	±0.6	±0.2	±0.7	±0.17	±0.9
	0.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	14.3	0.5	5.1	10.4	145.8	4.14	108.6
		S.D.	±1.7	±0.1	±0.6	±0.2	±1.1	±0.20	±1.2
	2.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	13.6	0.5	5.2	10.5	145.6	3.98	108.5
		S.D.	±1.1	±0.1	±0.7	±0.4	±0.8	±0.13	±1.7
	12.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	14.1	0.4	5.4	10.7	145.1	3.98	108.9
		S.D.	±1.8	±0.1	±0.7	±0.3	±1.1	±0.19	±1.2

*: P<0.05, **: P<0.01 (significantly different from control).

Table 10 Biochemical findings
Male, Female, 52w

Sex	Group and dose		T. Protein (g/dL)	A/G ratio	α_1 -Globulin (%)	α_2 -Globulin (%)	β -Globulin (%)	γ -Globulin (%)	Albumin (%)
Male	Control	N	10	10	10	10	10	10	10
		Mean	5.8	1.01	19.2	7.5	17.9	5.7	49.7
		S.D.	± 0.2	± 0.21	± 2.2	± 0.5	± 2.3	± 2.3	± 5.4
	0.1 mg/kg	N	8	8	8	8	8	8	8
		Mean	5.8	1.01	18.2	7.1	18.5	6.9	49.3
		S.D.	± 0.3	± 0.29	± 1.8	± 1.4	± 4.5	± 3.1	± 8.4
	0.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	5.8	1.42**	15.2**	6.1*	15.3	5.2	58.1**
		S.D.	± 0.5	± 0.31	± 2.4	± 1.3	± 3.0	± 1.7	± 5.4
	2.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	5.8	1.75**	13.4**	5.0**	12.7**	5.8	63.2**
		S.D.	± 0.2	± 0.30	± 2.0	± 1.1	± 2.2	± 1.2	± 4.7
Female	Control	N	10	10	10	10	10	10	10
		Mean	6.4	1.79	13.5	4.8	13.2	4.6	63.9
		S.D.	± 0.3	± 0.25	± 1.6	± 0.6	± 1.5	± 0.9	± 3.1
	0.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	6.7	1.69	14.2	4.8	13.5	4.9	62.6
		S.D.	± 0.2	± 0.17	± 1.6	± 0.5	± 0.7	± 1.2	± 2.5
	2.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	6.7	1.73	12.8	5.0	13.6	5.4	63.3
		S.D.	± 0.3	± 0.17	± 1.4	± 0.9	± 1.6	± 1.2	± 2.3
	12.5 mg/kg	N	9	9	9	9	9	9	9
		Mean	6.5	2.00	12.1	4.1	12.2	5.0	66.5
		S.D.	± 0.5	± 0.19	± 1.0	± 0.4	± 1.2	± 1.2	± 2.1

*: P<0.05, **: P<0.01 (significantly different from control).

Table 10 - continued

Biochemical findings
Male, Female, 52w

Study No. P030097

Sex	Group and dose		T. Bilirubin (mg/dL)	AST (IU/L)	ALT (IU/L)	ALP (IU/L)	T. Cholesterol (mg/dL)	Triglycerides (mg/dL)	Phospholipids (mg/dL)	Glucose (mg/dL)	
Male	Control	N	10	10	10	10	10	10	10	10	
		Mean	0.0	85	32	141	80	92	134	125	
		S. D.	±0.0	±17	±15	±42	±11	±41	±12	±27	
	0.1 mg/kg	N	8	8	8	8	8	8	8	8	8
		Mean	0.0	91	31	165	78	69	123	115	
		S. D.	±0.0	±18	±8	±56	±22	±28	±26	±11	
	0.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	0.0	89	42	364**	82	98	152	139	
		S. D.	±0.0	±23	±31	±87	±21	±32	±33	±17	
	2.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	0.0	77	86	565**	75	77	143	125	
		S. D.	±0.0	±15	±11	±137	±13	±32	±15	±16	
Female	Control	N	10	10	10	10	10	10	10	10	
		Mean	0.1	112	37	57	100	95	196	103	
		S. D.	±0.1	±98	±36	±26	±23	±71	±41	±9	
	0.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	0.0	92	27	59	104	92	207	110	
		S. D.	±0.0	±48	±12	±16	±13	±51	±17	±9	
	2.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	0.0	132	76	57	108	91	208	106	
		S. D.	±0.0	±103	±103	±14	±22	±61	±36	±16	
	12.5 mg/kg	N	9	9	9	9	9	9	9	9	9
		Mean	0.0*	69	25	86**	99	77	196	119*	
		S. D.	±0.0	±13	±8	±20	±21	±26	±31	±16	

*: P<0.05, **: P<0.01 (significantly different from control).

Table 10 - continued

Biochemical findings
Male, Female, 52w

Study No. P030097

Sex	Group and dose		BUN	Creatinine	IP	Ca	Na	K	Cl
			(mg/dL)	(mg/dL)	(mg/dL)	(mg/dL)	(mEq/L)	(mEq/L)	(mEq/L)
Male	Control	N	10	10	10	10	10	10	10
		Mean	9.1	0.5	5.2	10.0	147.4	4.40	107.7
		S.D.	±1.5	±0.1	±0.6	±0.2	±1.3	±0.20	±1.1
	0.1 mg/kg	N	8	8	8	8	8	8	8
		Mean	8.8	0.4	5.2	10.0	147.3	4.54	108.0
		S.D.	±0.9	±0.1	±0.5	±0.3	±0.9	±0.16	±2.1
	0.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	10.4	0.4	5.8*	10.2	146.5	4.64	107.6
		S.D.	±1.9	±0.0	±0.5	±0.5	±0.8	±0.32	±1.8
	2.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	12.8**	0.4	5.6	9.9	146.2	4.63	107.1
		S.D.	±1.5	±0.1	±0.6	±0.3	±1.3	±0.26	±1.7
Female	Control	N	10	10	10	10	10	10	10
		Mean	13.4	0.5	4.7	10.6	145.8	4.04	107.2
		S.D.	±2.7	±0.1	±1.0	±0.4	±1.2	±0.27	±1.6
	0.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	12.6	0.5	4.6	10.9	146.4	3.80	107.0
		S.D.	±2.8	±0.0	±0.7	±0.3	±1.1	±0.33	±2.5
	2.5 mg/kg	N	10	10	10	10	10	10	10
		Mean	12.7	0.5	4.6	10.6	146.0	4.19	107.4
		S.D.	±3.1	±0.1	±0.6	±0.3	±1.5	±0.32	±1.8
	12.5 mg/kg	N	9	9	9	9	9	9	9
		Mean	12.1	0.4	5.0	10.5	145.7	3.96	107.9
		S.D.	±2.0	±0.1	±0.8	±0.4	±1.1	±0.21	±1.9

*: P<0.05, **: P<0.01 (significantly different from control).

Table 11 Necropsy findings
Male, Female, 13w

Study No. P030097

Sex	Group and dose	Male				Female			
		Control	0.1 mg/kg	0.5 mg/kg	2.5 mg/kg	Control	0.5 mg/kg	2.5 mg/kg	12.5 mg/kg
Organs and findings	Number of animals	10	10	10	9	10	10	10	10
Digestive system									
Liver	Enlargement	0	0	0	5*	0	0	0	1
Hematopoietic system									
Thymus	Coloration, dark red	0	0	0	0	1	0	0	0
Urinary system									
Kidney	Pit.	1	0	0	0	0	0	0	0
	Dilatation, pelvic cavity	0	0	0	0	0	0	0	0
	Mass, light gray	0	0	0	0	1	0	0	0
Integumentary system									
Integument	Loss, hair	0	0	0	0	0	0	0	1

*: $P < 0.05$ (significantly different from control).
No appreciable changes in all other organs and tissues.
One male in the 2.5 mg/kg group died.

Table 12 Necropsy findings
Male, Female, 52w

Study No. P030097

Organs and findings	Sex Group and dose Number of animals	Male				Female			
		Control	0.1 mg/kg	0.5 mg/kg	2.5 mg/kg	Control	0.5 mg/kg	2.5 mg/kg	12.5 mg/kg
		10	8	10	10	10	10	10	9
Digestive system									
Liver									
Macule, light gray		0	0	0	2	0	0	0	1
Macule, dark red		0	0	0	1	0	0	1	0
Discoloration		2	0	0	0	0	0	0	0
Enlargement		0	0	0	0	0	0	0	0
Hernia, diaphragmatic		0	0	7**	9**	0	0	0	5*
				0	0	1	0	0	1
Respiratory system									
Lung									
Spot, white		1	0	0	4	1	0	0	0
Coloration, dark red		0	0	0	1	0	0	0	0
Hematopoietic system									
Thymus									
Small		10	8	10	9	8	8	10	9
Popliteal lymph node									
Enlargement		1	0	1	1	0	0	0	0
Spleen									
Macule, light gray		1	1	0	0	0	0	0	0
Cardiovascular system									
Heart									
Coloration, light gray		1	0	1	1	0	0	0	0
Urinary system									
Kidney									
Pit		0	0	1	0	0	0	0	0
Genital system									
Testis									
Small		0	0	0	1	NA	NA	NA	NA
Epididymis									
Small		0	0	0	1	NA	NA	NA	NA
Prostate									
Spot, dark red		0	0	0	1	NA	NA	NA	NA
Ovary									
Enlargement		NA	NA	NA	NA				
Dilatation, ovarian bursa						0	0	2	0
Cyst						0	0	0	1
						2	3	1	2

*: P<0.05, **: P<0.01 (significantly different from control).

NA: not applicable.

No appreciable changes in all other organs and tissues.

Two males in the 0.1 mg/kg group and one female in the 12.5 mg/kg group died.

Table 12 - continued

Necropsy findings
Male, Female, 52w

Study No. P030097

Organs and findings	Sex Group and dose Number of animals	Male				Female			
		Control	0.1 mg/kg	0.5 mg/kg	2.5 mg/kg	Control	0.5 mg/kg	2.5 mg/kg	12.5 mg/kg
		10	8	10	10	10	10	10	9
Genital system									
Uterus		NA	NA	NA	NA				
Polyp, endometrium									
Mammary gland						0	0	0	1
Retention, milk		0	0	0	0	2	2	3	3
Endocrine system									
Pituitary									
Spot, dark red		0	0	0	0	2	1	3	1
Enlargement		0	0	0	0	1	1	2	1
Nodule, light gray		0	0	0	0	0	0	1	1
Thyroid									
Defect, left		0	1	0	0	0	0	0	0
Enlargement, right		0	1	0	0	0	0	0	0
Adrenal									
Spot, brown		0	0	0	0	1	2	1	1
Enlargement		0	0	0	0	0	0	1	0
Integumentary system									
Integument									
Nodule, cutis, white		0	0	1	0	0	0	0	0
Nodule, subcutis, white		0	0	0	0	0	0	0	1
Mass, subcutis, light gray		0	0	0	0	3	0	1	3
Others									
Extremity									
Swelling, hindlimb		0	0	0	1	0	0	0	0
Corn, hindlimb		5	5	7	1	1	0	2	2

Not significantly different from control.

NA: not applicable.

No appreciable changes in all other organs and tissues.

Two males in the 0.1 mg/kg group and one female in the 12.5 mg/kg group died.

Table 13 Organ weights
Male, Female, 13w

Study No. P030097

Sex	Group and dose		Final body weight	Brain		Pituitary		Thyroids		Heart	
			(g)	(g)	(g/100 gB.W.)	(mg)	(mg/100 gB.W.)	(mg)	(mg/100 gB.W.)	(g)	(g/100 gB.W.)
Male	Control	N	10	10	10	10	10	10	10	10	10
		Mean	530.1	2.22	0.42	14.0	2.7	20.4	3.8	1.54	0.29
		S.D.	±32.1	±0.08	±0.02	±1.6	±0.3	±5.8	±1.0	±0.19	±0.03
	0.1 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	566.3	2.28	0.40	13.9	2.5	26.6*	4.7	1.61	0.29
		S.D.	±42.2	±0.05	±0.03	±1.3	±0.2	±5.2	±0.8	±0.13	±0.02
	0.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	546.5	2.29	0.42	14.2	2.6	24.5	4.5	1.63	0.30
		S.D.	±40.3	±0.07	±0.03	±1.2	±0.2	±5.7	±1.1	±0.10	±0.02
	2.5 mg/kg	N	9	9	9	9	9	9	9	9	9
		Mean	450.1**	2.19	0.49**	12.8	2.8	18.7	4.1	1.49	0.33**
		S.D.	±27.8	±0.07	±0.03	±0.5	±0.2	±4.1	±0.7	±0.09	±0.02
Female	Control	N	10	10	10	10	10	10	10	10	10
		Mean	304.1	2.05	0.68	16.9	5.6	16.7	5.5	0.96	0.32
		S.D.	±26.9	±0.06	±0.06	±1.7	±0.5	±4.1	±1.1	±0.07	±0.02
	0.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	303.0	2.06	0.69	18.4	6.1	17.7	5.9	0.91	0.30
		S.D.	±31.0	±0.09	±0.05	±1.9	±0.7	±2.2	±0.8	±0.08	±0.02
	2.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	297.0	2.09	0.70	19.1	6.4*	19.2	6.5*	0.95	0.32
		S.D.	±17.5	±0.08	±0.03	±2.5	±1.0	±2.7	±1.1	±0.06	±0.02
	12.5 mg/kg	N	10	10	10	10	10	10	10	10	10
		Mean	299.8	2.09	0.70	18.6	6.2	18.7	6.2	0.94	0.32
		S.D.	±23.1	±0.07	±0.05	±2.9	±0.8	±2.4	±0.7	±0.07	±0.03

*: P<0.05, **: P<0.01 (significantly different from control).
One male in the 2.5 mg/kg group died.

Table 13 - continued Organ weights
Male, Female, 13w

Study No. P030097

Sex	Group and dose		Lungs		Thymus		Liver		Spleen	
			(g)	(g/100 gB.W.)	(g)	(g/100 gB.W.)	(g)	(g/100 gB.W.)	(g)	(g/100 gB.W.)
Male	Control	N	10	10	10	10	10	10	10	10
		Mean	1.54	0.29	0.28	0.06	14.56	2.75	0.77	0.14
		S.D.	±0.11	±0.02	±0.05	±0.01	±0.81	±0.10	±0.12	±0.02
	0.1 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	1.59	0.28	0.31	0.05	15.96	2.82	0.84	0.15
		S.D.	±0.13	±0.03	±0.08	±0.02	±2.02	±0.23	±0.10	±0.01
	0.5 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	1.62	0.30	0.26	0.05	20.25**	3.71**	0.76	0.14
		S.D.	±0.11	±0.02	±0.07	±0.02	±2.00	±0.21	±0.06	±0.01
	2.5 mg/kg	N	9	9	9	9	9	9	9	9
		Mean	1.41	0.31	0.25	0.05	23.16**	5.12**	0.68	0.15
		S.D.	±0.09	±0.03	±0.05	±0.01	±4.45	±0.72	±0.07	±0.01
Female	Control	N	10	10	10	10	10	10	10	10
		Mean	1.11	0.37	0.25	0.08	7.97	2.63	0.54	0.18
		S.D.	±0.09	±0.03	±0.04	±0.01	±0.70	±0.14	±0.06	±0.02
	0.5 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	1.12	0.37	0.28	0.09	7.97	2.63	0.45**	0.15**
		S.D.	±0.09	±0.02	±0.07	±0.02	±0.94	±0.18	±0.06	±0.01
	2.5 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	1.08	0.36	0.27	0.09	8.32	2.80	0.50	0.17
		S.D.	±0.06	±0.02	±0.04	±0.01	±0.52	±0.18	±0.07	±0.02
	12.5 mg/kg	N	10	10	10	10	10	10	10	10
		Mean	1.14	0.38	0.27	0.09	11.63**	3.88**	0.49	0.17
		S.D.	±0.11	±0.03	±0.06	±0.02	±1.72	±0.50	±0.05	±0.01

** : P<0.01 (significantly different from control).
One male in the 2.5 mg/kg group died.