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SC-18862 AND 19192 (3:1 RATIO):
SEGMENT II TERATOLOGY STUDY IN THE RABBIT

P-T NO. 1002H72

FINAL REPORT

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HAZLETON LABORATORIES

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SPONSOR: Searle Laboratories; Division of
G. D. Searle and Company

DATE: October 12, 1972

MATERIAL: SC-18862 and SC-19192 (3:1 Ratio)

SUBJECT: FINAL REPORT
Segment II Teratology Study in the Rabbit
Project No. 700-263

SUMMARY

The purpose of this study was to evaluate the potential of SC-18862 and 19192 (3:1 ratio) for embryotoxic and/or teratogenic effects in albino rabbits. A 15% aqueous suspension of the test material in a 1% aqueous solution of Tween 80 was administered by oral intubation, twice daily (in divided doses) at intervals of four hours from Day 6 through Day 18 of gestation. Three test groups of 20 females each received total daily dosage levels of 1 g/kg/day (Group No. 2), 2 g/kg/day (Group No. 3), and 3 g/kg/day (Group No. 4). A control group of 20 females (Group No. 1) received a 1% aqueous solution of Tween 80 only, by oral intubation, twice daily from Day 6 through Day 18 in a volume equal to that received by the high level test females.

Four control, 12 Group No. 2, nine Group No. 3, and 12 Group No. 4 does died during the study. In most instances, gross necropsy findings did not reveal a definitive cause of death. There were no visible lesions in 0/4 control, 4/12 Group No. 2, 4/9 Group No. 3, and 9/12 Group No. 4 animals.



In some instances, lung findings were suggestive of aspiration or respiratory disease. Litters from these animals could not be evaluated for visceral or skeletal development. Pregnancy rates in the does found dead were as follows: 100% in the control group, 58% in Group No. 2, 89% in Group No. 3, and 75% in Group No. 4.

Pregnancy rates for animals delivered by Caesarean section at term were similar in all groups (94%, 88%, 82%, and 100% for the controls and Groups No. 2, No. 3, and No. 4, respectively). Findings for these animals and data for the litters obtained (14 control, seven Group No. 2, nine Group No. 3, and eight Group No. 4) are summarized below.

There were no apparent consistent compound-related effects with regard to the appearance and behavior of the maternal does or in gross visceral alterations observed at necropsy.

Following initiation of treatment, overall group mean body weight gain for Group No. 2 was similar to that for the control group. For Group No. 3, overall mean weight gain during the treatment period was also similar to that for the controls; however, due to posttreatment weight loss, these animals only maintained their mean weight from initiation of treatment to term. The Group No. 4 does exhibited overall group mean body weight losses both during the treatment period and from the initiation of treatment to term.

Decreases in food consumption were evident among both control and test animals during the treatment and posttreatment periods, as compared to respective pretreatment food consumption data. Percent decreases observed for Group No. 2 were similar to those for the control group. However, percent decreases observed for Groups No. 3 and No. 4 exceeded those for the controls, both during treatment and during the posttreatment period, in a dose-related manner.



Ratio of the number of implantation sites to the number of corpora lutea for Groups No. 2, No. 3, and No. 4, the mean number of implantation sites and of live and dead fetuses for each of the three test groups, and the mean number of resorption sites for Groups No. 2 and No. 3 were comparable to control values and were within normal limits. The mean number of resorption sites for Group No. 4 (2.5) was increased as compared to control (0.8) and to historical data, although the incidence of females with resorption sites was similar between Group No. 4 and the control group (50% each). Statistical analysis revealed no significant differences between the mean number of implantation sites, resorption sites, and live and dead fetuses for the control group and Groups No. 2, No. 3, and No. 4.

There were no statistical differences between live fetal mean weights and lengths for the controls and Group No. 2. There was a statistically significant decrease, apparently dose related, in the mean weight of the live fetuses in Groups No. 3 and No. 4 as compared to the control fetuses. Live fetal mean lengths were also slightly lower in these two test groups than in the control group, but not to a statistically significant degree.

Examination of external, visceral, and skeletal structures revealed no consistent unusual findings among the test fetuses, and the fetal skeletal evaluations revealed no trends toward lesser or greater development in the test groups as compared to the control group.



In conclusion, oral administration of SC-18862 and 19192 (3:1 ratio) to female albino rabbits from Day 6 through Day 18 of gestation at a dosage level of 1 g/kg/day (Group No. 2) resulted in no discernible embryotoxic and/or teratogenic effects in this study. Likewise, teratogenic effects were not discernible in Groups No. 3 and No. 4. At dosage levels of 2 g/kg/day (Group No. 3) and 3 g/kg/day (Group No. 4), live mean fetal weights were decreased to a statistically significant degree. An apparent increase in the mean number of resorptions at the high test level as compared to the controls was without statistical significance. These findings may be related to the greater decrease in food consumption which was evident among the maternal does in these two test groups.

INTRODUCTION

Artificial insemination of the does used for this study was performed on April 25, 1972. Sacrifice of females for Cesarean delivery on Day 28 or Day 29 of gestation was performed on May 23 and May 24, 1972.

MATERIALS

Compound

Identification: SC-18862 and 19192 (3:1 ratio).

Receipt Dates and Days Used:

Gestation Days 6 Thru 13 - SC-18862 and 19192 (3:1 ratio) prepared by sponsor and received April 11, 1972; Lot No. PD-16-180A.

Gestation Days 14 Thru 18 - SC-18862 (Lot No. 56057C received January 5, 1972) and SC-19192 (Lot No. 1R received November 23, 1971), 3:1 ratio, prepared by this laboratory.



Description: A fine, white powder with no noticeable odor.

Purity: Assumed 100% active ingredient.

Animals

Species: Rabbits; proven does and bucks.

Supplier: Camm Research Institute, Inc.

Strain: New Zealand White.

Basal Diet: Purina Rabbit Chow and water available ad libitum.

Housing: Individually in Hoeltge self-flushing, stainless steel,
hanging cages.

Stock Chemicals

Tissue Preservation: 10% neutral buffered formalin, Bouin's solution,
80% ethyl alcohol, and 100% glycerin.

Skeletal Staining: 1.0% potassium hydroxide; alizarin red S stain and
potassium hydroxide solution; one part benzyl alcohol, one part
glycerin, and two parts 70% alcohol; and 75% glycerin.

METHODS

Impregnation of Does

Each doe, following stimulation of ovulation by intravenous
administration of Pituitary Luteinizing Hormone, was impregnated by means
of artificial insemination using techniques similar to those described by



Gibson, J. P., Staples, R. E., and Newberne, J. W., Use of the Rabbit in Teratogenicity Studies, Toxicology and Applied Pharmacology, Vol. 9, No. 2, September 1966. The day of insemination was designated as Day 0 of gestation.

Animal Groups

From a total of 80 does impregnated by means of artificial insemination, the animals were arbitrarily selected and were placed into one control and three test groups to receive the indicated dosage levels.

<u>Group No.</u>	<u>No. of Does</u>	<u>Dosage Levels (Total Daily Dose*)</u> g/kg/day
1 (Control)	20	0
2	20	1
3	20	2
4	20	3

* The total daily dose administered was divided into two equal parts with each animal receiving one-half of the total dose at four hour intervals.

Administration of Compound

A 15% aqueous suspension (weight-per-volume) of the test material was prepared fresh daily in a 1% aqueous solution (weight-per-volume) of Tween 80. The test suspension was administered to the test animals by oral intubation, in equally divided doses at intervals of four hours, from Day 6 through Day 18 of gestation - a total of 13 consecutive days of treatment. The total daily volumes administered to the test females were 6.7, 13.3, and 20.0 ml/kg of body weight (Groups No. 2, No. 3, and No. 4, respectively).



One-half of the total daily volume was administered at each divided dose. Dosages were based on individual body weights on Days 6, 10, 15, and 18 of gestation. No compound was administered prior to Day 6 or subsequent to Day 18. Control animals were treated in a same manner and received a 1% aqueous solution (weight-per-volume) of Tween 80 only, from Day 6 through Day 18, by oral intubation in a volume equal to that received by the high test level females.

Observations and Records on Impregnated Females

Mortality, appearance, and behavior of all impregnated animals were observed daily throughout the expected gestation period. Body weight was recorded on the following days of gestation: Day 0 (initially); Days 6, 10, 15, 18, 22; and Day 28 or 29 (terminally). Food consumption was recorded daily throughout the study.

Termination

Does were sacrificed on Day 28 or Day 29 of gestation by means of intravenous air embolism. Caesarean sections were performed, and the following observations were recorded: number of corpora lutea, number and placement of uterine implantation and resorption sites, number and placement of live and dead fetuses at delivery, individual fetal weight and length (crown - rump), and external fetal anatomy. Gross necropsies (partial) with examination of uterine and visceral structures were performed on all sacrificed



does. The necropsy procedure employed included gross examination of the thoracic and abdominal cavities and of the following organs: heart, lung, stomach, small and large intestine, and urinary bladder. The same observations were recorded for all does which died during the study.

Visceral Examination of Fetuses by Wilson's Technique

Following delivery and after external observations were completed, approximately one-half of the fetuses from each litter* were fixed in Bouin's solution and evaluated for visceral changes using techniques similar to those described by Wilson, J. G. and Warkany, J., Teratology: Principles and Techniques. 1st ed. p. 267, The University of Chicago Press, Chicago, 1965. Whole body transverse sections of the head, thoracic, and abdominal regions were taken and examined for abnormalities under the dissecting microscope.

Skeletal Examination of Fetuses

Following completion of external observations, the remaining one-half of the fetuses were exsanguinated, placed in distilled water for approximately an hour, eviscerated, skinned, replaced in distilled water for approximately two hours, and then fixed in 80% ethyl alcohol for a minimum of three days. The fetuses were then placed in 1.0% potassium hydroxide for about 24 hours,

* With the exception of one Group No. 3 litter (Rabbit No. 673, 29-C) from which all fetuses, each dead, were processed for skeletal examination.



stained for approximately 24 hours in a solution of alizarin red S and potassium hydroxide, and then rinsed with distilled water. The stain was extracted from the soft tissue with one part benzyl alcohol, one part glycerin, and two parts 70% alcohol, for a minimum of 24 hours and the fetuses and pups were cleared with 75% glycerin for approximately 24 hours. Lamps were kept over the specimen containers throughout the staining and extraction procedures. Each skeleton was evaluated for relative differences in size, location, normal or abnormal bone structure, degree of ossification, and the presence or absence of bone structure. The fetuses were then stored in 100% glycerin.

Summary of Visceral and Skeletal Evaluations

A tabularized summary of the number of fetuses processed for visceral (Wilson's) or skeletal (alizarin) examinations is presented on the following page. All such fetuses (live plus dead) were obtained by Caesarean section. The only fetuses not evaluated from these litters were dead fetuses too small for processing procedures. As stated previously, fetuses from litters of females which died (not included in summary on following page) could not be evaluated due to small size and/or autolysis.



Group No.	Number of Fetuses			Number Evaluated		Number Not Evaluated
	Live	Dead	= Total	Wilson's	Alizarin	
1	91	8	99	52	45	2
2	57	4	61	30	27	4
3	59	8	67	32	35	0
4	47	3	50	24	24	2

Tissue Preservation

The ovaries and uterus of each female were preserved in 10% neutral buffered formalin. Dead fetuses not processed for visceral or skeletal evaluation were preserved in 10% neutral buffered formalin. Fetuses examined by Wilson's technique were preserved in Bouin's solution and fetuses stained with alizarin red S were preserved in 100% glycerin. All tissues are being held at Hazleton Laboratories, Inc., for possible future examination.

Statistical Analysis

Statistical analysis of the following parameters was performed by the t-test at the 5.0% probability level: mean number of implantation sites, resorption sites, live fetuses, and dead fetuses per group and mean weights and lengths of live fetuses by group. (Reference: Wilfred J. Dixon and Frank J. Massey, Jr., Introduction to Statistical Analysis, 123-124, McGraw Hill, 1957.)



RESULTS

Maternal Data Summary

A total of 19 of 20 control, 14 of 20 Group No. 2 (1 g/kg), 17 of 20 Group No. 3 (2 g/kg), and 17 of 20 Group No. 4 (3 g/kg) females placed on study were pregnant. Four control, seven Group No. 2, eight Group No. 3, and nine Group No. 4 pregnant females died during the study. The remaining pregnancies were among females which were sacrificed for Caesarean delivery. Conception rate data are summarized in Figure No. 1.

Figure No. 1 - Conception rate data.

	Group No. 1 <u>Control</u>	Group No. 2 <u>1 g/kg/day</u>	Group No. 3 <u>2 g/kg/day</u>	Group No. 4 <u>3 g/kg/day</u>
No. of Does Inseminated:	20	20	20	20
No. of Does Pregnant:	19	14	17	17
Caesarean Delivery	15	7	9	8
Death	4	7	8	9
No. of Does Not Pregnant:	1	6	3	3
Sacrificed for Caesarean Delivery	1	1	2	0
Death	0	5	1	3

Body weight and food consumption data for all inseminated rabbits are presented in appended Tables No. 1 and No. 1A, respectively. These and other pertinent data are discussed in detail in the following sections of this report.



Morbidity and Mortality Data

Incidence of mortality was as follows: control group - 4/20, Group No. 2 - 12/20, Group No. 3 - 9/20, and Group No. 4 - 12/20. General observations and necropsy findings for the animals which died are summarized in Figure No. 2; detailed findings are appended in Table No. 2. The necropsy procedure employed included gross examination of the thoracic and abdominal cavities and of the following organs: heart, lung, stomach, small and large intestine, and urinary bladder. In most instances, gross necropsy findings did not reveal a definitive cause of death. There were no visible lesions in 0/4 control, 4/12 Group No. 2, 4/9 Group No. 3, and 9/12 Group No. 4 animals. In some instances, lung findings which consisted of white purulent material or fluid present in the lungs or thoracic cavity (2/4 control, 4/12 Group No. 2, 1/9 Group No. 3, and 1/12 Group No. 4 animals) were suggestive of aspiration or respiratory disease. Other observations such as collapsed lungs (2/12 Group No. 2 and 1/9 Group No. 3 does) and red fluid present in the lungs (1/9 Group No. 3 does) or in the thoracic cavity (1/4 control and 2/12 Group No. 3 animals) were suggestive of mechanical problems encountered at dosing.



Figure No. 2 - General observations and necropsy findings for animals which died.

	Group No. 1 <u>Control</u>	Group No. 2 <u>1 g/kg/day</u>	Group No. 3 <u>2 g/kg/day</u>	Group No. 4 <u>3 g/kg/day</u>
No. of Does Inseminated:	20	20	20	20
No. of Deaths:	4	12	9	12
No. Pregnant*	4	7	8	9
No. Not Pregnant	0	5	1	3
Appearance and Behavior -				
Anorexia/Little or No Food Consumption	4	9	8	12
Body Weight Loss	3	6	4	12
Thin Appearance	1	3	2	9
Nasal Discharge	2	1	3	7
Wheezing	2	3	3	1
Labored or Rapid Respiration	3	5	6	5
Soft Feces	0	0	2	9
Depression**	1	1	2	6
Ataxia	0	0	1	3
Prostration	0	0	0	3
Weakened Condition	0	0	0	2
Bloated Appearance	0	2	0	1
Red Jelly-like Material Found in Cage Pan (Day 24)	0	0	0	1
Necropsy Findings -				
No Gross Lesions Observed	0	4	4	9
Lungs				
Appeared Collapsed	0	2	1	0
Small Portion White in Color	0	1	0	0
Red Fluid Present	0	0	1	0
White Purulent Material or White Fluid Present in Lungs or Thoracic Cavity	2	4	1	1

* Uterine and litter data for pregnant does are presented in appended Table No. 5. Further evaluation of litters obtained from animals which died for visceral and skeletal development was in all instances precluded by autolysis or the small size of the fetuses.

** Characterized by inactivity and a failure to respond to minimal agitation.



Figure No. 2 - Continued

	Group No. 1 <u>Control</u>	Group No. 2 <u>1 g/kg/day</u>	Group No. 3 <u>2 g/kg/day</u>	Group No. 4 <u>3 g/kg/day</u>
Thoracic Cavity				
Red Fluid Present	1	0	2	0
Abdominal Cavity				
Red Fluid Present	0	1	0	0
Large Intestine				
Filled with Air	0	0	0	1
Uterus				
Appeared Deformed	0	0	0	1
White Purulent Material				
Present in Both Horns	1	0	0	0
Ovaries				
Small	0	1	0	0
Absent	0	0	0	1

Data for Nonpregnant Females Sacrificed for Cesarean Delivery

There was a total of four does (one control, one Group No. 2, and two Group No. 3) which were found not pregnant at sacrifice for Cesarean delivery. General observations and necropsy findings for these nonpregnant females are summarized in Figure No. 3, with detailed findings following.



Figure No. 3 - General observations and necropsy findings for nonpregnant animals sacrificed at term.

	Group No. 1 <u>Control</u>	Group No. 2 <u>1 g/kg/day</u>	Group No. 3 <u>2 g/kg/day</u>	Group No. 4 <u>3 g/kg/day</u>
No. of Does Inseminated:	20	20	20	20
No. of Does Surviving to Time of Sacrifice:	16	8	11	8
No. of Does Found Not Pregnant:	1	1	2	0
Appearance and Behavior - Nasal Discharge	0	1	1	0
Necropsy Findings - No Gross Lesions Observed	1	1	2	0

Apart from nasal discharge exhibited by Group No. 2 Rabbit No. 648 for a 10 day period and by Group No. 3 Rabbit No. 660 for two days only, each of the nonpregnant does were normal in appearance and behavior throughout the study. No gross lesions were evident at necropsy of these females. Individual body weight and food consumption data for these animals are presented in appended Tables No. 1 and No. 1A, respectively; data for these animals were excluded from group mean calculations.

Data for Pregnant Females Sacrificed for Caesarean Delivery

Fifteen control, seven Group No. 2, nine Group No. 3, and eight Group No. 4 does were pregnant at term sacrifice. General observations and necropsy findings for these pregnant does are summarized in Figure No. 4. Detailed findings are appended in Table No. 3.



Figure No. 4 - General observations and necropsy findings for pregnant animals sacrificed at term.

	Group No. 1 Control	Group No. 2 1 g/kg/day	Group No. 3 2 g/kg/day	Group No. 4 3 g/kg/day
No. of Does Inseminated:	20	20	20	20
No. of Does Surviving to Time of Sacrifice:	16	8	11	8
No. of Does Pregnant:	15	7	9	8
Appearance and Behavior -				
Nasal Discharge	4	4	5	4
Bloody-appearing Fluid Around Nose	0	1	0	0
Wheezing	1	2	2	2
Labored or Rapid Respiration	0	1	0	1
Thin Appearance	3	0	3	3
Soft Feces	2	0	1	4
Depression*	1	1	1	2
Ataxia	0	0	0	1
Red Fluid in Cage Pan	0	1	0	0
Necropsy Findings -				
No Gross Visceral Lesions Observed	15	7	8	7
Lungs				
Right Lung Contained a White, Purulent Material	0	0	1	0
Liver				
Appeared Mottled	0	0	1	0
Gallbladder				
Distended	0	0	1	1
Filled with a Large Amount of Bile	0	0	0	1
Gastrointestinal Tract				
Stomach Contained a Large Amount of a Thick, White Substance** Intermingled with a Moderate Amount of Hair and Food-like Material; Intestine Contained Very Small Amount of Food	0	0	0	1

* Characterized by inactivity and a failure to respond to minimal agitation.

** Presumptively identified as the administered compound.



Individual and group mean body weight and food consumption data are presented in appended Tables No. 1 and No. 1A, respectively. Data for non-pregnant females and for all females which died were excluded from calculation of group mean values. A summary of the body weight and food consumption data for the pregnant females are presented in Figures No. 5 and No. 6, respectively. Numbers in parentheses indicate the sample size used in calculation of group mean data.

Figure No. 5 - Body weight data for pregnant animals sacrificed at term.

	Group No. 1 <u>Control</u> (14)	Group No. 2 <u>1 g/kg/day</u> (7)	Group No. 3 <u>2 g/kg/day</u> (9)	Group No. 4 <u>3 g/kg/day</u> (8)
<u>Group Mean Body Weights (g.)</u>				
Day 0: Wt.	3695	3907	4156	4306
±s.d.	444.2	462.7	282.1	406.5
Day 6: Wt.	3852	4280	4333	4456
±s.d.	420.8	560.8	311.1	445.1
Day 10: Wt.	3805	4287	4268	4367
±s.d.	425.1	530.6	392.5	586.8
Day 15: Wt.	3878	4336	4323	4276
±s.d.	436.5	471.5	283.1	690.3
Day 18: Wt.	3999	4430	4442	4358
±s.d.	416.5	530.8	284.9	681.2
Day 22: Wt.	3993	4571	4407	4237
±s.d.	425.9	526.5	310.8	844.8
Day 28/29: Wt.	4063	4441	4334	4197
±s.d.	402.6	668.5	487.4	752.5



Figure No. 5 - Continued

	Group No. 1 <u>Control</u> (14)	Group No. 2 <u>1 g/kg/day</u> (7)	Group No. 3 <u>2 g/kg/day</u> (9)	Group No. 4 <u>3 g/kg/day</u> (8)
<u>Group Mean Weight Changes at Each Weighing Interval (±%)</u>				
Days 0 thru 6:	+4%	+10%	+4%	+3%
Days 6 thru 10:	-1%	0	-2%	-2%
Days 10 thru 15:	+2%	+1%	+1%	-2%
Days 15 thru 18:	+3%	+2%	+3%	+2%
Days 18 thru 22:	0	+3%	-1%	-3%
Days 22 thru 28/29:	+2%	-3%	-2%	-1%
<u>Group Mean Weight Changes at Selected Intervals (±%)</u>				
During Treatment:				
Days 6 thru 18	+4%	+4%	+3%	-2%
During Treatment and Posttreatment:				
Days 6 thru 28/29	+5%	+4%	0	-6%
Entire Period of Gestation:				
Days 0 thru 28/29	+10%	+14%	+4%	-3%

Overall group mean body weight gains during treatment (Days 6 thru 18) and from initiation of treatment to term (Days 6 thru 28/29) were similar between the control group and Group No. 2 (+4% or +5% for each of these intervals). During treatment, the overall group mean weight gain for the



Group No. 3 animals (+3%) was similar to that for the controls; however, these animals exhibited mean weight losses at both posttreatment weighing intervals and thus during the interval from initiation of treatment to term, the Group No. 3 does maintained their mean weight only (0). The Group No. 4 rabbits exhibited overall mean body weight losses during treatment (-2%) and from initiation of treatment to term (-6%).

Figure No. 6 - Food consumption data for pregnant animals sacrificed at term.

	Group No. 1 Control (14)	Group No. 2 1 g/kg/day (7)	Group No. 3 2 g/kg/day (9)	Group No. 4 3 g/kg/day (8)
<u>Group Mean Values/Day at Selected Intervals (g.)</u>				
Day 0: Food	158	194	183	185
±s.d.	29.8	27.1	28.7	35.7
Day 6: Food	137	188	150	99
±s.d.	49.3	24.6	36.1	52.2
Day 10: Food	137	173	110	100
±s.d.	41.6	25.5	58.9	81.8
Day 15: Food	153	182	149	99
±s.d.	40.6	52.2	62.9	83.6
Day 18: Food	155	200	152	112
±s.d.	35.8	39.5	74.2	100.5
Day 22: Food	154	182	148	137
±s.d.	39.8	74.7	77.1	90.4
Day 27: Food	143	89	99	92
±s.d.	31.3	77.9	59.9	87.8



Figure No. 6 - Continued

	Group No. 1 <u>Control</u> (14)	Group No. 2 <u>1 g/kg/day</u> (7)	Group No. 3 <u>2 g/kg/day</u> (9)	Group No. 4 <u>3 g/kg/day</u> (8)
<u>Total Group Mean Values/Day (g.)</u>				
Pretreatment Period				
Days 0 thru 5: Food	179	205	185	184
±s.d.	26.2	23.0	23.7	30.1
Treatment Period				
Days 6 thru 18: Food	148	177	135	98
±s.d.	28.6	21.5	32.1	70.8
Posttreatment Period				
Days 19 thru 27/28: Food	141	151	124	108
±s.d.	28.6	63.5	52.0	73.4
<u>Percent Decreases in Food Consumption (Comparison with Pretreatment Total Group Mean Values)</u>				
Treatment Period	-17%	-14%	-27%	-47%
Posttreatment Period	-21%	-26%	-33%	-41%

Decreases in food consumption were evident among both control and test animals during the treatment and posttreatment periods, as compared to respective pretreatment total group mean values. The decreases observed for the Group No. 2 animals were similar to those for the controls. However, the decreases observed at the two higher test levels exceed those for the controls both during treatment and during the posttreatment period.

Uterine and Ovarian Data

Uterine and ovarian data for each animal are presented in appended Table No. 4. Ratio calculations (number of implantation sites per number of corpora lutea X 100) were based only on those females which were pregnant. The calculated ratio values were similar between the control and the test groups and were within normal limits for all groups. Ratio values were as follows: control group, 68.1%; Group No. 2, 73.3%; Group No. 3, 71.3%; and Group No. 4, 73.2%. The cumulative mean ratio value and standard deviation for this parameter obtained from 21 control groups from the most recent studies performed in this laboratory is 76.6%, \pm 9.29%.

Uterine and Litter Data

The results of uterine and litter data for all pregnant animals are presented in appended Table No. 5. All data for females which died were excluded from calculation of group mean values. Data for control Rabbit No. 630 was also excluded from group mean calculations for the control group, since uterine findings for this doe (total postimplantation loss) were not considered representative of this group. Uterine and litter data are summarized in Figure No. 7.

Figure No. 7 - Summary of uterine and litter data.

	Group No. 1 <u>Control</u>	Group No. 2 <u>1 g/kg/day</u>	Group No. 3 <u>2 g/kg/day</u>	Group No. 4 <u>3 g/kg/day</u>
Number of Does Included in Mean:	14	7	9	8
Implantation Sites:				
Mean Number	7.9	9.0	8.6	8.8
s.d.	2.68	1.91	3.36	3.11



Figure No. 7 - Continued

	Group No. 1 <u>Control</u> 14	Group No. 2 <u>1 g/kg/day</u> 7	Group No. 3 <u>2 g/kg/day</u> 9	Group No. 4 <u>3 g/kg/day</u> 8
Resorption Sites:				
Mean Number	0.8	0.3	1.1	2.5
±s.d.	1.12	0.49	1.62	3.63
Live Fetuses:				
Mean Number	6.5	8.1	6.6	5.9
±s.d.	2.95	1.77	4.61	4.09
Dead Fetuses:				
Mean Number	0.6	0.6	0.9	0.4
±s.d.	1.34	1.51	1.69	0.74
Incidence of Does with:				
Resorption Sites				
Number	7	2	5	4
Percent	50%	29%	56%	50%
Dead Fetuses				
Number	4	1	3	2
Percent	29%	14%	33%	25%

The mean number of implantation sites, live fetuses, and dead fetuses were considered comparable between the control and three test groups and were within normal limits for all groups; and statistical analysis of these parameters revealed no statistically significant differences between control and test data. Historical data for these parameters obtained from 41 control groups from similarly performed studies are as follows: implantation sites - cumulative mean (8.1) and range in values (5.0 to 10.0); live fetuses -



cumulative mean (7.3), range in values (4.6 to 9.7), and a value of 5.9 or less was experienced in four of these 41 control groups; and dead fetuses - cumulative mean (0.4) and range in values (0 to 1.2).

The mean number of resorption sites for Groups No. 2 and No. 3 were considered comparable to that for the control group and were within normal limits. The mean number of resorption sites for Group No. 4 (2.5) was increased as compared to the controls (0.8) and to historical data, although the incidence of females with resorption sites was similar between these two groups (50% each). Statistical analysis of the mean number of resorption sites revealed no significant differences between values for the control and each of the three test groups. Historical data for this parameter are as follows: resorption sites - cumulative mean (0.3) and range in values (0 to 1.1); data were obtained from 41 control groups.

Group mean fetal weights and lengths presented in Figure No. 8 were based on the number of litters. Some dead fetuses were too small for an accurate recording of measurements and the affected means are indicated in appended Table No. 3.

There were no statistically significant differences between the mean weights and lengths of the live fetuses in Group No. 2 and those in the control group. There was a statistically significant decrease, apparently dose related, in the mean weights of the live fetuses in Groups No. 3 and No. 4. Mean lengths were also slightly decreased in these two test groups, but not to a statistically significant degree.



Figure No. 8 - Group mean fetal weights and lengths.

	Group No. 1 <u>Control</u>	Group No. 2 <u>1 g/kg/day</u>	Group No. 3 <u>2 g/kg/day</u>	Group No. 4 <u>3 g/kg/day</u>
<u>Live Fetuses</u>				
Number of Litters Included				
in Mean:	13	7	7	6
Group Mean				
Weight (g.)	43.3	39.3	35.6 ^{S-}	33.2 ^{S-}
±s.d.	6.95	6.71	8.87	7.52
Length (cm.)	8.7	8.6	8.3	8.0
±s.d.	0.57	0.81	0.93	0.91
<u>Dead Fetuses</u>				
Number of Litters Included				
in Mean:	2	0	3	1
Group Mean				
Weight* (g.)	12.9	-	11.4	20.0
Length* (cm.)	6.1	-	5.3	6.5
S- = Significantly lower than control at $p < 0.05$.				
* = Statistical analysis not performed.				

External and Visceral Fetal Data

There was no indication of a compound-related effect with regard to the external anatomy of the test fetuses. Remarkable observations were confined to dead fetuses only and findings are presented in Figure No. 9. Litters are identified by maternal rabbit numbers. No gross external irregularities were evident in any live control and test fetuses delivered by Caesarean section.



Figure No. 9 - Fetal external examination data.

<u>Group No.</u>	<u>Maternal Rabbit No.</u>	<u>Observations</u>
3 2 g/kg	659 (28-C)	2/2 Dead Fetuses - one fetus: right forepaw and right hindpaw rotated inward, and other fetus: right hindpaw rotated inward.*
3 2 g/kg	673 (29-C)	5/5 Dead Fetuses - severely mutilated and misshapen.**
4 3 g/kg	683 (29-C)	1/2 Dead Fetuses - right forepaw rotated inward and left forepaw rotated outward.* ϕ

* Skeletal examination performed on each - gross external irregularity observed not evident at subsequent skeletal examinations.

** Skeletal evaluation of these (and all other control and test dead fetuses) generally revealed lesser development than that of live fetuses which were of larger size. Detailed skeletal findings are presented on Page No. 27.

ϕ Skeletal examination showed the absence of the 2nd, 3rd, and 4th digits from one hindpaw of this fetus.

Visceral examination of test fetuses revealed no evidence of a compound-related effect. Results of the visceral examinations by Wilson's technique are presented in Figure No. 10. Numbers in parentheses indicate the number of litters/number of fetuses examined.

Figure No. 10 - Fetal visceral examination data (Wilson's technique).

	<u>Group No. 1 Control (13/52)</u>	<u>Group No. 2 1 g/kg/day (7/30)</u>	<u>Group No. 3 2 g/kg/day (7/32)</u>	<u>Group No. 4 3 g/kg/day (6/24)</u>
All Viscera Appeared Normal	51	28	31	23
Dilation of One or Both Renal Pelves	1	1	1	1
Dilation of the Lateral Ventricles of the Brain	0	1	0	0



Skeletal Data for Fetuses

The results of the fetal skeletal examinations from control and test litters are presented in Figure No. 11. The skull, ribs, sternum, vertebrae, pelvic girdle, longbones, forepaws, and hindpaws of the animals were evaluated. Included in the figure are those bone structures which are consistently indicative of developmental variations. The skeletal evaluations are presented as the combined data, for fetuses delivered by Caesarean section on Days 28 or 29, which could be evaluated within each group. Variations in the number of fetuses examined (numbers in parentheses in Figure No. 11) reflect inability to accurately evaluate certain bone structures due to damage during processing.

EXPLANATION FOR FOOTNOTES

Figure No. 11

* Grading of development of skull:

0 = No skull surface stained.

1 = Approximately 25% of skull surface stained.

2 = Approximately 50% of skull surface stained.

3 = Approximately 75% of skull surface stained.

4 = Approximately 100% of skull surface stained.

** Single rib present on one side only of the vertebral column.

∅ Floating rib unattached to the vertebral column.

Figure No. 11 - Skeletal development of rabbit fetuses from litters of females serving as controls or receiving SC-18862 and 19192 (3:1 ratio). Numbers in parentheses indicate the number of fetuses evaluated. Incidence is expressed as number and percent of fetuses.

	INCIDENCE AS NUMBER					INCIDENCE AS PERCENT				
	GROUP NO.					GROUP NO.				
	1	2	3	4		1	2	3	4	
Number of Litters Examined	CONTROL	1 G/KG	2 G/KG	3 G/KG		CONTROL	1 G/KG	2 G/KG	3 G/KG	
(Number of Fetuses Examined)	13	7	8	6		13	7	8	6	
Skull	(45)	(27)	(35)	(24)		(45)	(27)	(35)	(24)	
Closure Grading*:	0	0	0	0	0					
	1	0	0	0	0					
	2	0	0	0	0					
	3	3	0	5	0	7		14		
	4	5	2	4	4	11	7	11	17	
		37	25	26	20	82	93	74	83	
Occipital Bone										
Underdeveloped	1	0	1	0	0	2		3		
Interparietal Bone										
Underdeveloped	0	0	2	0	0			6		
Hyoid Bone										
Nonossified	0	0	1	0	0			3		
Midportion Nonossified	4	0	4	0	0	9		11		
Maxilla, Nasal, and Orbit Bones										
Not Fully Ossified	3	0	0	0	0	7				
Premaxilla Bones										
Appeared Shortened	3	1	3	0	0	7	4	9		
Nasal Bones										
Appeared Shortened	3	1	4	0	0	7	4	11		
Triangular Bipartite Bone										
Present Between Frontal and										
Nasal Bones	1	0	0	0	0	2				
Frontal and Parietal Bones										
Overlapped at Sutures	1	0	2	0	0	2		6		
Sutures Wider Than Normal with										
Bottom Portions Overlapped	0	0	1	0	0			3		
Parietal Bones										
Sutures Wider Than Normal	0	0	1	0	0			3		
Most Head Bones Exhibited										
Moderate Ossification Only	0	0	1	0	0			3		

Figure No. 11 - Continued

	INCIDENCE AS NUMBER					INCIDENCE AS PERCENT				
	GROUP NO.					GROUP NO.				
	1	2	3	4		1	2	3	4	
Number of Litters Examined	CONTROL	1 G/KG	2 G/KG	3 G/KG		CONTROL	1 G/KG	2 G/KG	3 G/KG	
(Number of Fetuses Examined)	13	7	8	6		13	7	8	6	
Ribs	(45)	(27)	(35)	(24)		(45)	(27)	(35)	(24)	
Pairs 12	23	23	20	13		51	85	57	54	
Pairs 13	22	4	15	11		49	15	43	46	
Single**	8	4	5	1		18	15	14	4	
Small	6	2	3	1		13	7	9	4	
Floating	6	2	3	2		13	7	9	8	
Fused (7th and 8th ribs on one side at proximal end)	0	0	1	0						
1st Pair of Ribs Located at 7th Cervical Vertebra	1	0	0	0		2		3		
Vertebrae										
Total Number Nonossified										
Cervical:										
Centra 2	0	0	1	0				3		
Thoracic:										
Centra 2	0	0	1	0				3		
Sacral:										
Centra 2	0	0	1	0				3		
Dorsal Arches 3	0	0	1	0				3		
Caudal:										
Centra 1-2	(45)	(26)	(35)	(24)		(45)	(26)	(35)	(24)	
6-10	18	8	8	11		40	31	23	46	
16	2	0	3	0		4		9		
Dorsal Arches 8-10	0	0	1	0				3		
11	44	26	34	24		98	100	97	100	
16	1	0	0	0		2				
Total Number Incompletely Ossified										
Thoracic:	(45)	(27)	(35)	(24)		(45)	(27)	(35)	(24)	
Centra 1	0	0	0	1				3	4	

Figure No. 11 - Continued

	INCIDENCE AS NUMBER					INCIDENCE AS PERCENT				
	GROUP NO.					GROUP NO.				
	1	2	3	4		1	2	3	4	
Number of Litters Examined	CONTROL	1 G/KG	2 G/KG	3 G/KG		CONTROL	1 G/KG	2 G/KG	3 G/KG	
(Number of Fetuses Examined)	13	7	8	6		13	7	8	6	
Vertebrae (Continued)	(45)	(27)	(35)	(24)		(45)	(27)	(35)	(24)	
Total Number Small										
Cervical:										
Centra 5	0	0	1	0				3		
Thoracic:										
Centra 6	0	0	1	0				3		
Dorsal Arches 1	0	0	1	0				3		
Total Number Fused										
Thoracic:										
Dorsal Arches 2	0	0	1	0				3		
(7th and 8th on right)										
Total Number Out of Alignment										
Thoracic:										
Dorsal Arches 1	0	0	1	0				3		
(8th on right out of alignment vertically)										
Caudal:	0	0	1	0				3		
Centra 3	(45)	(26)	(35)	(24)		(45)	(26)	(35)	(24)	
(18th, 19th, and 20th aligned horizontally at tip of tail)										
Total Number Irregular in Shape	0	0	1	0				3		
Thoracic:	(45)	(27)	(35)	(24)		(45)	(27)	(35)	(24)	
Centra 1										
(small irregularly shaped ossification center between 7th and 9th thoracic centra on left side)										
Seven Lumbar Vertebrae Present	0	0	1	0				3		
17 Caudal Vertebrae Present	2	0	3	0		4		9		
	1	2	5	0		2	7	14		

Figure No. 11 - Continued

	INCIDENCE AS NUMBER					INCIDENCE AS PERCENT				
	GROUP NO.					GROUP NO.				
	1	2	1 G/KG	2 G/KG	3 G/KG	1	2	1 G/KG	2 G/KG	3 G/KG
Number of Litters Examined	CONTROL	13	7	8	6	CONTROL	13	7	8	6
(Number of Fetuses Examined)	(45)	(27)	(35)	(24)		(45)	(27)	(35)	(24)	
Sternum										
Ossification Centers:										
Absent - 1st	1	0	1	0	0	2		3		
2nd	0	0	1	0	0			3		
3rd	0	0	1	0	0			3		
4th	0	0	1	0	0			3		
5th	6	11	5	3	3	13	41	14	12	
6th	2	0	1	1	1	4		3	4	
Small - 1st	1	0	0	1	1	2			4	
2nd	0	0	0	1	1				4	
3rd and 4th	0	0	0	0	0				4	
5th	6	7	2	3	3	13	26	6	12	
6th	2	0	0	1	1	4			4	
Split - 1st thru 4th	0	0	0	0	0					
5th	0	1	1	0	0		4	3		
6th	0	0	0	0	0					
Fused - 4th and 5th	2	0	0	0	0	4				
Pelvis										
Ossification Centers:										
Absent - Right Pubis	4	0	7	0	0	9		20		
Left Pubis	4	0	7	0	0	9		20		
Forepaws										
Ossification Centers:										
Total Number Absent	45	27	35	24		100	100	100	100	
Carpus 16	5	1	8	7	24	11	4	23	29	
Metacarpus 1-2	2	0	0	0	0	4				
3-4	0	0	1	0	0			3		
5-10	5	7	5	7	7	11	26	14	29	
Phalanges 1-15	2	0	1	0	0	4		3		
16-20	0	0	0	0	0					
21-28	0	0	0	0	0					
Total Number Small	0	0	1	0	0					
Metacarpus 2								3		

Figure No. 11 - Continued

	INCIDENCE AS NUMBER					INCIDENCE AS PERCENT				
	GROUP NO.					GROUP NO.				
	1	2	3	4		1	2	3	4	
	CONTROL	1 G/KG	2 G/KG	3 G/KG		CONTROL	1 G/KG	2 G/KG	3 G/KG	
	13	7	8	6		13	7	8	6	
	(45)	(27)	(35)	(24)		(45)	(27)	(35)	(24)	
Number of Litters Examined (Number of Fetuses Examined)										
Forepaws (Continued)										
Flexure of One or Both Forepaws	3	0	0	0		7				
with Inward Rotation	1	0	0	0		2				
	(44)	(27)	(35)	(24)		(44)	(27)	(35)	(24)	
Hindpaws										
Ossification Centers:										
Total Number Absent										
Tarsus 8	39	27	28	19		89	100	80	79	
10	3	0	6	5		7		17	21	
12	2	0	1	0		5		3		
Metatarsus 1-2	0	0	1	0				3		
3-4	0	0	0	0						
5-8	0	0	0	0						
	(44)	(27)	(35)	(23)		(44)	(27)	(35)	(23)	
Phalanges 1-15	3	0	0	1		7			4	
16-20	2	0	0	0		4				
21-24	0	0	1	0				3		
Flexure of One or Both Hindpaws or Leg	2	0	1	0		4		3		
Inward Rotation of One Hindpaw	0	0	1	0				3		
2nd, 3rd, and 4th Digits										
Missing from One Hindpaw	0	0	0	1					4	



Skeletal evaluations revealed no consistent unusual findings among the test fetuses and no trends toward greater or lesser development were seen in the test groups, as compared to the control group. In general, lesser development and ossification was evident among dead control and test fetuses as compared to live fetuses which were of larger size. There were no meaningful differences between the developmental indices for the control and test groups.

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NOTE: The research described in this report involved animals maintained in animal care facilities fully accredited by the American Association for Accreditation of Laboratory Animal Care.

EXPLANATION FOR FOOTNOTES

Code placed after individual maternal rabbit numbers indicates the following.

P = Pregnant

NP = Not Pregnant (For purposes of this report, pregnancy is defined as "grossly visible implantation sites or products thereof in the endometrium.")

C = Caesarean delivery; sacrifice of doe on gestation day indicated.

D = Death of female on day indicated.

Tables No. 1 and No. 1A

* = Group mean data for females found pregnant at Caesarean delivery only; data for females which died or were nonpregnant were excluded from calculation of group mean values.

** = Body weight and food consumption data for control Rabbit No. 630 (P, 29-C) were excluded from calculation of group mean values since uterine findings for this female (total postimplantation loss) were not considered representative of this group.

() = Body weight in parentheses is terminal weight of animal which died; day recorded is indicated.

NR = Not Recorded.

Table No. 4

* = Only those females which were pregnant were included in ratio calculations.

** = Both ovaries absent.

EXPLANATION FOR FOOTNOTES (Continued)

Table No. 5

- * = Some dead fetuses were too small to record weight and length; statistical analysis was not performed on group mean weights and lengths of dead fetuses.
- ** = No weights or lengths recorded for any dead fetuses in litter.
- _/ = Indicates number of fetuses (sample size) for mean so designated when less than the total number of dead fetuses found.
- ϕ = Evidence of development of conceptus observed at each implantation site; however, further classification regarding stage of development could not be made.
- ϕϕ = Total postimplantation loss; no evidence of fetuses or resorption sites found; data for Rabbit No. 630 excluded from group mean calculations since findings were not considered representative of this group.
- # = One resorption site present, other implantation sites exhibited evidence of development.
- ## = One resorption site present, postimplantation loss at other sites.

Table No. 1 - Individual and mean body weight data in grams for rabbits serving as controls or receiving SC-18862 and 19192 (3:1 ratio)

MATERNAL RABBIT NO.		GROUP NO. 1 - CONTROL							
		DAY OF GESTATION							
		0	6	10	15	18	22	28/29	
616	P, 28-C	3879	3970	3910	3970	4137	4133	4160	
617	P, 28-C	3320	3468	3405	3441	3500	3492	3495	
618	P, 28-C	3412	3611	3482	3606	3744	3743	3800	
620	P, 29-C	3632	3760	3812	3868	3820	3775	3790	
621	P, 29-C	3715	3778	3792	3877	4016	3880	3975	
622	P, 29-C	3805	4050	4006	4158	4256	4262	4415	
623	P, 29-C	3716	3959	3964	4007	4250	4264	4245	
624	P, 29-C	3975	4063	4147	4175	4248	4253	4393	
625	P, 29-C	3438	3735	3494	3300	3538	3698	3870	
626	P, 29-C	3773	4019	3855	3974	4180	4220	4195	
627	P, 29-C	2975	3256	3351	3445	3543	3471	3580	
629	P, 29-C	4865	5056	5000	5070	5087	5115	5080	
631	P, 29-C	3964	3614	3526	3648	3772	3750	3885	
634	P, 29-C	3267	3586	3522	3755	3897	3843	4000	
Mean*		3695	3852	3805	3878	3999	3993	4063	
±s.d.		444.2	420.8	425.1	436.5	416.5	425.9	402.6	
628	P, 18-D	3707	3926	3810	3686	(3646 - Day 18)-	-	-	
630**	P, 29-C	4621	4787	4979	4953	4948	4456	3765	
632	P, 29-D	3934	4070	4126	4035	3869	3330	(2840 - Day 29)	
633	P, 17-D	4189	4407	4402	4135	(4095 - Day 17)-	-	-	
635	P, 9-D	3451	3603	(3240 - Day 9)	-	-	-	-	
619	NP, 28-C	4027	4281	4263	4372	4525	4673	4710	

Table No. 1 - Continued

GROUP NO. 2 - 1 G/KG/DAY

MATERNAL RABBIT NO.		DAY OF GESTATION						
		0	6	10	15	18	22	28/29
641	P, 28-C	3874	4263	4183	4265	4325	4368	4305
645	P, 28-C	3828	5025	4968	5090	5113	5195	5015
647	P, 28-C	3500	3780	3836	4035	4050	4296	4190
650	P, 29-C	4102	4293	4353	4230	4359	4558	4600
651	P, 29-C	3336	3564	3666	3730	3716	3828	3210
653	P, 29-C	3937	4040	3987	4157	4285	4416	4470
654	P, 29-C	4771	4995	5016	4845	5160	5333	5295
Mean*		3907	4280	4287	4336	4430	4571	4441
±s.d.		462.7	560.8	530.6	471.5	530.8	526.5	668.5
636	P, 10-D	3595	3810	(3796 - Day 10)-	-	-	-	-
637	P, 22-D	3469	3656	3574	3397	-	(2873 - Day 22)-	-
639	P, 17-D	3774	4025	4132	3750	3316	(2873 - Day 17)-	-
643	P, 25-D	4630	4749	4925	5128	(3460 - Day 17)-	4829	(4390 - Day 25)
644	P, 9-D	3220	3366	(3110 - Day 9) -	-	5090	-	-
646	P, 23-D	4146	4294	4460	4070	3846	3520	(3482 - Day 23)
652	P, 15-D	3937	4112	3835	(3626 - Day 15)-	-	-	-
638	NP, 6-D	3661	(3760 - Day 6) -	-	-	-	-	-
640	NP, 13-D	3137	3464	3255	(3041 - Day 13)-	-	-	-
642	NP, 4-D	3162	(3025 - Day 4) -	-	-	-	-	-
648	NP, 28-C	2874	3260	3326	3537	3590	3899	3920
649	NP, 6-D	4764	(4856 - Day 6) -	-	-	-	-	-
655	NP, 10-D	4261	4431	(4252 - Day 10)-	-	-	-	-

Table No. 1 - Continued

GROUP NO. 3 - 2 G/KG/DAY

MATERNAL RABBIT NO.		DAY OF GESTATION						
		0	6	10	15	18	22	28/29
659	P, 28-C	3975	4051	3700	4088	4193	4328	3710
661	P, 28-C	3975	4352	4285	4212	4518	4468	4495
663	P, 29-C	3985	4100	4078	4110	4251	4285	4215
664	P, 29-C	3920	4122	3795	4080	4247	4348	4405
667	P, 29-C	4583	4840	4881	4847	5069	5138	5210
668	P, 29-C	3957	4099	4215	4126	4185	4218	4200
670	P, 29-C	4141	4238	4271	4260	4396	4296	4350
672	P, 29-C	4199	4336	4420	4535	4655	4550	4780
673	P, 29-C	4668	4857	4766	4650	4460	4031	3640
Mean*		4156	4333	4268	4323	4442	4407	4334
±s.d.		282.1	311.1	392.5	283.1	284.9	310.8	487.4
656	P, 13-D	3838	3980	3690	(3342 - Day 10)-	(3342 - Day 13)-	-	-
658	P, 10-D	4357	4573	(3900 - Day 10)-	-	-	-	-
662	P, 18-D	3941	4069	4030	3637	(3400 - Day 18)-	-	-
665	P, 24-D	4663	4685	4410	4422	4596	4456	(4265 - Day 24)
666	P, 12-D	3895	4050	4153	(4060 - Day 12)-	-	-	-
671	P, 10-D	3741	3815	(3720 - Day 10)-	-	-	-	-
674	P, 15-D	4622	4794	4800	(4296 - Day 15)-	-	-	-
675	P, 16-D	5000	5206	4716	(4410 - Day 15)-	-	-	-
657	NP, 28-C	3913	4313	4400	4315	4455	4405	4200
660	NP, 28-C	5757	5937	5960	5950	6105	6110	6130
669	NP, 6-D	4273	(4233 - Day 6) -	-	-	-	-	-

Table No. 1 - Continued

GROUP NO. 4 - 3 G/KG/DAY

MATERNAL
RABBIT NO.

		DAY OF GESTATION						
		0	6	10	15	18	22	28/29
677	P, 28-C	3897	4032	3592	3395	3440	3050	3140
679	P, 28-C	3613	3724	3480	3243	3466	3287	3635
680	P, 28-C	4393	4510	4500	4422	4559	4534	4300
681	P, 28-C	4482	4542	4412	4162	4057	3544	3274
683	P, 29-C	4297	4482	4530	4405	4580	4574	4785
684	P, 29-C	4980	5257	5320	5380	5456	5444	5090
687	P, 29-C	4417	4578	4580	4680	4812	4902	4855
691	P, 29-C	4367	4520	4520	4520	4490	4558	4500
Mean*		4306	4456	4367	4276	4358	4237	4197
±s.d.		406.5	445.1	586.8	690.3	681.2	844.8	752.5
676	P, 27-D	4344	4542	4490	4100	4155	3792	(3383 - Day 27)
678	P, 18-D	3982	4170	3945	3625	(3310 - Day 18)-	-	-
682	P, 18-D	4289	4453	4415	3888	(3794 - Day 18)-	-	-
685	P, 29-D	4534	4774	4595	4705	4491	3915	(3530 - Day 29)
688	P, 13-D	4382	4535	4398	(3885 - Day 13)-	-	-	-
690	P, 18-D	3921	4005	3652	3208	(2992 - Day 18)-	-	-
692	P, 19-D	3959	3960	3800	3290	3201	(3135 - Day 19)-	-
694	P, 25-D	4150	4220	4106	3583	3590	3174	(3030 - Day 25)
695	P, 21-D	4510	4775	4540	4040	4040	3840	(3325 - Day 21)
686	NP, 10-D	3802	3217	(3194 - Day 10)-	-	-	-	-
689	NP, 11-D	3952	4142	3840	(3840 - Day 11)-	-	-	-
693	NP, 14-D	5237	5350	4972	(4483 - Day 14)-	-	-	-

Table No. 1A - Individual and mean food consumption data in grams for rabbits serving as controls or receiving SC-18862 and 19192 (3:1 ratio)

MATERNAL RABBIT NO.		PRETREATMENT					GROUP NO. 1 - CONTROL					TREATMENT PERIOD				
							DAY OF GESTATION									
		0	1	2	3	4	5	6	7	8	9	10	11			
616	P, 28-C	154	197	184	230	175	184	184	164	157	156	128	170			
617	P, 28-C	159	175	164	168	150	160	142	136	126	129	113	118			
618	P, 28-C	150	172	164	160	135	150	39	132	135	125	128	141			
620	P, 29-C	125	152	158	190	185	175	119	144	165	186	185	174			
621	P, 29-C	129	144	140	153	126	136	149	153	154	151	128	155			
622	P, 29-C	220	244	249	245	215	222	200	200	193	200	198	204			
623	P, 29-C	162	175	194	200	154	189	74	181	173	191	149	202			
624	P, 29-C	121	181	170	185	176	221	183	158	164	194	168	196			
625	P, 29-C	146	181	151	200	165	194	113	105	127	129	104	121			
626	P, 29-C	179	206	199	190	190	218	141	135	163	163	156	165			
627	P, 29-C	170	198	181	195	166	168	172	125	174	171	154	157			
629	P, 29-C	207	200	208	215	190	218	200	155	195	190	171	197			
631	P, 29-C	125	145	160	160	123	148	125	131	131	99	105	122			
634	P, 29-C	169	199	217	237	205	234	80	10	17	25	34	55			
Mean*		158	184	181	195	168	187	137	138	148	151	137	156			
±s.d.		29.8	26.9	29.6	29.1	28.2	32.0	49.3	43.9	43.8	47.5	41.6	41.7			
628	P, 18-D	189	193	200	205	194	187	156	146	102	120	104	108			
630**	P, 29-C	192	218	218	215	184	204	205	216	192	214	188	193			
632	P, 29-D	186	194	192	190	215	165	165	155	176	150	135	157			
633	P, 17-D	146	186	209	215	185	209	204	183	184	140	145	3			
635	P, 9-D	179	178	162	170	160	175	20	0	0	-	-	-			
619	NP, 28-C	257	253	251	245	190	202	224	212	201	177	194	200			

Table No. 1A - Continued

GROUP NO. 1 - CONTROL (Continued)

MATERNAL RABBIT NO.		TREATMENT PERIOD DAY OF GESTATION					POSTTREATMENT PERIOD			
		12	13	14	15	16	17	18	19	20
616	P, 28-C	172	135	191	171	151	170	175	125	171
617	P, 28-C	158	154	160	163	110	125	135	105	133
618	P, 28-C	172	142	161	155	40	150	168	150	162
620	P, 29-C	173	175	186	57	43	75	110	100	125
621	P, 29-C	151	153	172	176	150	173	153	50	59
622	P, 29-C	191	163	185	185	177	212	207	185	204
623	P, 29-C	195	172	172	187	170	200	212	183	193
624	P, 29-C	215	171	210	202	157	115	132	140	160
625	P, 29-C	108	112	121	124	122	155	175	173	202
626	P, 29-C	167	160	190	173	165	200	205	162	168
627	P, 29-C	134	140	158	150	130	160	140	125	97
629	P, 29-C	190	158	184	162	115	150	135	105	100
631	P, 29-C	113	95	95	80	80	105	110	102	134
634	P, 29-C	90	101	138	160	155	198	115	112	155
Mean*		159	145	166	153	126	156	155	130	147
±s.d.		36.4	26.1	30.7	40.6	44.6	40.3	35.8	38.1	42.2
628	P, 18-D	64	61	0	0	0	0	-	-	-
630**	P, 29-C	194	93	148	62	0	12	0	0	0
632	P, 29-D	160	125	35	11	20	15	0	0	0
633	P, 17-D	0	0	6	0	0	-	-	-	-
635	P, 9-D	-	-	-	-	-	-	-	-	-
619	NP, 28-C	206	187	212	210	170	200	200	170	199

Table No. 1A - Continued

GROUP NO. 1 - CONTROL (Continued)

MATERNAL RABBIT NO.	POSTTREATMENT PERIOD									
	DAY OF GESTATION									
	21	22	23	24	25	26	27	28		
616	P, 28-C	180	171	171	168	148	109	121	-	
617	P, 28-C	123	63	59	112	117	134	147	-	
618	P, 28-C	169	166	147	111	82	92	91	-	
620	P, 29-C	137	137	147	136	114	96	191	102	
621	P, 29-C	135	140	145	153	138	136	141	115	
622	P, 29-C	188	185	191	170	164	164	161	122	
623	P, 29-C	210	185	168	176	160	164	141	139	
624	P, 29-C	170	188	163	154	140	146	150	135	
625	P, 29-C	215	214	209	203	197	197	183	160	
626	P, 29-C	163	160	145	134	98	100	115	106	
627	P, 29-C	73	105	111	100	113	126	120	124	
629	P, 29-C	123	138	130	120	96	106	107	95	
631	P, 29-C	NR	123	110	105	108	130	144	125	
634	P, 29-C	162	186	174	172	151	180	194	174	
Mean*		158	154	148	144	130	134	143	127	
±s.d.		39.0	39.8	37.7	31.7	31.8	32.7	31.3	24.0	
628	P, 18-D	-	-	-	-	-	-	-	-	
630**	P, 29-C	0	0	0	0	0	0	0	0	
632	P, 29-D	0	0	0	NR	0	0	0	0	
633	P, 17-D	-	-	-	-	-	-	-	-	
635	P, 9-D	-	-	-	-	-	-	-	-	
619	NP, 28-C	200	230	212	216	227	218	222	-	

Table No. 1A - Continued

GROUP NO. 1 - CONTROL (Continued)

MATERNAL RABBIT NO.	TOTAL MEAN VALUES/DAY					
	DAYS 0 THRU 5		DAYS 6 THRU 18		DAYS 19 THRU 27/28	
	MEAN	±S.D.	MEAN	±S.D.	MEAN	±S.D.
616 P, 28-C	187	25.3	163	17.9	152	26.6
617 P, 28-C	163	8.5	136	18.1	110	30.7
618 P, 28-C	155	13.0	130	42.7	130	35.6
620 P, 29-C	164	24.2	138	51.8	128	28.4
621 P, 29-C	138	9.9	155	12.5	121	36.5
622 P, 29-C	232	15.1	193	13.3	173	22.9
623 P, 29-C	179	18.4	175	34.7	172	22.3
624 P, 29-C	176	32.2	174	29.8	155	16.2
625 P, 29-C	173	22.4	124	20.2	195	18.0
626 P, 29-C	197	13.8	168	20.4	135	28.2
627 P, 29-C	180	14.1	151	16.3	109	16.5
629 P, 29-C	206	10.2	169	26.3	112	14.8
631 P, 29-C	144	16.3	107	17.2	120	14.6
634 P, 29-C	210	25.2	91	60.7	166	23.1
Mean*	179	-	148	-	141	-
±s.d.	26.2	-	28.6	-	28.6	-
628 P, 18-D	195	6.8	72	59.5	-	-
630**P, 29-C	205	14.5	132	86.3	0	-
632 P, 29-D	190	16.0	100	70.7	0	-
633 P, 17-D	192	25.7	79	90.3	-	-
635 P, 9-D	171	8.1	7	11.5	-	-
619 NP, 28-C	233	29.2	199	14.8	210	18.6

Table No. 1A - Continued

GROUP NO. 2 - 1 G/KG/DAY

MATERNAL RABBIT NO.		PRETREATMENT					TREATMENT PERIOD									
		DAY OF GESTATION														
		0	1	2	3	4	5	6	7	8	9	10	11			
641	P, 28-C	220	226	209	220	174	225	205	183	100	180	151	173			
645	P, 28-C	226	228	245	245	204	232	170	175	179	173	158	174			
647	P, 28-C	217	263	248	225	183	251	217	220	208	220	205	208			
650	P, 29-C	181	206	205	185	190	246	175	187	201	203	200	164			
651	P, 29-C	171	193	188	190	183	192	164	203	195	170	156	162			
653	P, 29-C	156	155	175	165	170	180	165	121	136	150	148	152			
654	P, 29-C	189	209	224	208	217	205	218	192	215	193	196	202			
Mean*		194	211	213	205	189	219	188	183	176	184	173	176			
±s.d.		27.1	33.4	27.4	27.3	16.7	27.1	24.6	31.0	42.7	23.2	25.5	20.9			
636	P, 10-D	193	199	200	185	163	175	163	142	161	160	-	-			
637	P, 22-D	187	206	200	195	185	207	175	170	153	130	100	102			
639	P, 17-D	202	209	222	212	198	234	213	218	217	180	86	11			
643	P, 25-D	237	267	252	240	230	225	210	221	216	214	213	217			
644	P, 9-D	190	205	197	190	181	189	160	162	0	0	-	-			
646	P, 23-D	202	203	205	200	190	208	190	215	225	210	188	0			
652	P, 15-D	180	204	177	190	190	191	165	165	174	0	15	30			
638	NP, 6-D	208	198	179	157	145	125	-	-	-	-	-	-			
640	NP, 13-D	142	156	190	195	170	217	132	101	108	80	115	0			
642	NP, 4-D	183	4	0	0	-	-	-	-	-	-	-	-			
648	NP, 28-C	164	192	178	238	250	258	230	230	209	230	222	232			
649	NP, 6-D	217	251	250	255	201	201	-	-	-	-	-	-			
655	NP, 10-D	206	207	225	223	185	193	124	76	22	0	-	-			

Table No. 1A - Continued

GROUP NO. 2 - 1 G/KG/DAY (Continued)

MATERNAL RABBIT NO.		TREATMENT PERIOD						POSTTREATMENT PERIOD		
		DAY OF GESTATION								
		12	13	14	15	16	17	18	19	20
641	P, 28-C	190	190	185	128	95	155	165	160	163
645	P, 28-C	195	149	209	196	127	187	206	196	153
647	P, 28-C	162	165	175	201	146	195	212	220	222
650	P, 29-C	188	128	255	284	245	260	270	278	280
651	P, 29-C	145	121	182	160	114	175	162	116	135
653	P, 29-C	158	111	170	164	107	190	165	172	174
654	P, 29-C	218	102	66	141	157	212	220	205	204
Mean*		179	138	177	182	142	196	200	192	190
±s.d.		25.4	31.5	57.1	52.2	50.5	33.1	39.5	50.9	49.5
636	P, 10-D	-	-	-	-	-	-	-	-	-
637	P, 22-D	134	91	21	0	0	0	0	0	0
639	P, 17-D	80	33	56	7	0	-	-	-	-
643	P, 25-D	207	211	234	240	207	215	217	211	170
644	P, 9-D	-	-	-	-	-	-	-	-	-
646	P, 23-D	0	16	28	0	0	15	4	10	0
652	P, 15-D	12	0	0	-	-	-	-	-	-
638	NP, 6-D	-	-	-	-	-	-	-	-	-
640	NP, 13-D	0	-	-	-	-	-	-	-	-
642	NP, 4-D	-	-	-	-	-	-	-	-	-
648	NP, 28-C	220	229	252	210	165	186	205	212	225
649	NP, 6-D	-	-	-	-	-	-	-	-	-
655	NP, 10-D	-	-	-	-	-	-	-	-	-

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Table No. 1A - Continued

GROUP NO. 2 - 1 G/KG/DAY (Continued)

MATERNAL RABBIT NO.		POSTTREATMENT PERIOD							
		DAY OF GESTATION							
		21	22	23	24	25	26	27	28
641	P, 28-C	136	130	61	3	107	140	53	-
645	P, 28-C	162	167	178	157	145	34	0	-
647	P, 28-C	208	205	161	156	136	144	95	-
650	P, 29-C	258	291	245	260	225	132	200	206
651	P, 29-C	96	56	15	0	0	0	0	0
653	P, 29-C	210	202	198	192	153	193	170	191
654	P, 29-C	231	225	185	192	158	160	107	115
Mean*		186	182	149	137	132	115	89	128
±s.d.		56.9	74.7	81.2	98.9	68.3	70.3	77.9	94.2
636	P, 10-D	-	-	-	-	-	-	-	-
637	P, 22-D	0	-	-	-	-	-	-	-
639	P, 17-D	-	-	-	-	-	-	-	-
643	P, 25-D	30	21	0	0	-	-	-	-
644	P, 9-D	-	-	-	-	-	-	-	-
646	P, 23-D	0	0	-	-	-	-	-	-
652	P, 15-D	-	-	-	-	-	-	-	-
638	NP, 6-D	-	-	-	-	-	-	-	-
640	NP, 13-D	-	-	-	-	-	-	-	-
642	NP, 4-D	-	-	-	-	-	-	-	-
648	NP, 28-C	225	217	263	250	216	252	220	-
649	NP, 6-D	-	-	-	-	-	-	-	-
655	NP, 10-D	-	-	-	-	-	-	-	-

Table No. 1A - Continued

GROUP NO. 2 - 1 G/KG/DAY (Continued)

MATERNAL RABBIT NO.	TOTAL MEAN VALUES/DAY					
	DAYS 0 THRU 5		DAYS 6 THRU 18		DAYS 19 THRU 27/28	
	MEAN	±S.D.	MEAN	±S.D.	MEAN	±S.D.
641 P, 28-C	212	19.7	162	34.8	106	55.0
645 P, 28-C	230	15.2	177	23.1	132	67.6
647 P, 28-C	231	29.2	195	24.6	172	44.1
650 P, 29-C	202	23.8	212	46.6	238	48.3
651 P, 29-C	186	8.2	162	25.4	42	54.6
653 P, 29-C	167	10.1	149	24.3	186	17.5
654 P, 29-C	209	11.9	179	49.1	178	42.7
Mean*	205	-	177	-	151	-
±s.d.	23.0	-	21.5	-	63.5	-
636 P, 10-D	186	14.6	156	9.7	-	-
637 P, 22-D	197	9.4	83	69.4	0	-
639 P, 17-D	213	13.3	100	89.7	-	-
643 P, 25-D	242	15.4	217	9.8	72	93.4
644 P, 9-D	192	8.1	80	93.0	-	-
646 P, 23-D	201	6.2	84	100.9	2	5.0
652 P, 15-D	189	9.5	62	79.9	-	-
638 NP, 6-D	169	32.0	-	-	-	-
640 NP, 13-D	178	27.6	77	54.6	-	-
642 NP, 4-D	47	90.9	-	-	-	-
648 NP, 28-C	213	40.2	217	22.5	231	18.7
649 NP, 6-D	229	25.7	-	-	-	-
655 NP, 10-D	206	15.9	56	55.7	-	-

Table No. 1A - Continued

GROUP NO. 3 - 2 G/KG/DAY

MATERNAL RABBIT NO.	PRETREATMENT										TREATMENT PERIOD									
	DAY OF GESTATION																			
	0	1	2	3	4	5	6	7	8	9	10	11								
659	P, 28-C	187	131	156	185	152	113	86	35	4	0	2								
661	P, 28-C	191	206	198	206	183	205	153	142	137	160	82								
663	P, 29-C	164	180	183	157	145	154	128	120	153	112	52								
664	P, 29-C	196	193	217	215	212	208	190	65	3	8	100								
667	P, 29-C	209	211	229	234	196	215	200	204	212	188	186								
668	P, 29-C	205	215	225	212	177	195	170	191	179	173	177								
670	P, 29-C	149	157	160	170	165	172	114	102	121	121	131								
672	P, 29-C	133	157	184	180	167	167	155	158	139	161	153								
673	P, 29-C	217	195	182	202	189	196	154	185	165	162	155								
Mean*		183	183	193	196	176	181	150	134	124	121	115								
±s.d.		28.7	28.8	26.6	24.4	21.4	32.6	36.1	58.3	73.1	70.3	58.9	61.5							
656	P, 13-D	163	174	184	173	148	170	153	141	170	0	0								
658	P, 10-D	258	232	224	196	170	167	112	0	0	0	-								
662	P, 18-D	147	166	164	156	140	148	110	110	132	121	11								
665	P, 24-D	206	207	204	200	195	206	190	200	35	2	26								
666	P, 12-D	148	156	167	172	157	168	150	175	180	160	20								
671	P, 10-D	157	125	144	118	100	123	133	135	27	0	-								
674	P, 15-D	228	221	238	220	198	207	150	177	191	190	37								
675	P, 16-D	168	188	200	200	192	192	99	156	40	0	0								
657	NP, 28-C	200	206	220	205	175	194	166	161	175	182	168								
660	NP, 28-C	220	234	240	237	215	212	196	192	182	194	213								
669	NP, 6-D	170	178	115	50	60	142	-	-	-	-	-								

Table No. 1A - Continued

GROUP NO. 3 - 2 G/KG/DAY (Continued)

MATERNAL RABBIT NO.		TREATMENT PERIOD						POSTTREATMENT PERIOD		
		DAY OF GESTATION								
		12	13	14	15	16	17	18	19	20
659	P, 28-C	56	142	162	161	147	195	210	185	187
661	P, 28-C	91	151	162	181	158	200	200	186	198
663	P, 29-C	64	71	118	136	107	154	165	133	148
664	P, 29-C	190	175	213	211	166	188	185	190	180
667	P, 29-C	174	174	196	202	156	200	235	173	205
668	P, 29-C	166	154	181	181	123	150	75	136	101
670	P, 29-C	123	124	150	129	100	132	125	125	103
672	P, 29-C	150	155	172	142	125	143	173	111	95
673	P, 29-C	160	156	0	0	0	0	0	6	0
Mean*		130	145	150	149	120	151	152	138	135
±s.d.		49.4	31.6	62.6	62.9	50.7	62.6	74.2	57.9	66.9
656	P, 13-D	-	-	-	-	-	-	-	-	-
658	P, 10-D	-	-	-	-	-	-	-	-	-
662	P, 18-D	0	0	0	0	0	0	-	-	-
665	P, 24-D	30	61	36	140	80	95	85	98	85
666	P, 12-D	-	-	-	-	-	-	-	-	-
671	P, 10-D	-	-	-	-	-	-	-	-	-
674	P, 15-D	0	0	0	-	-	-	-	-	-
675	P, 16-D	0	0	0	0	-	-	-	-	-
657	NP, 28-C	197	190	180	184	152	205	107	91	150
660	NP, 28-C	218	208	136	200	160	177	144	130	177
669	NP, 6-D	-	-	-	-	-	-	-	-	-

Table No. 1A - Continued

GROUP NO. 3 - 2 G/KG/DAY (Continued)

MATERNAL RABBIT NO.	POSTTREATMENT PERIOD DAY OF GESTATION								
	21	22	23	24	25	26	27	28	
659	P, 28-C	200	241	166	4	0	17	15	-
661	P, 28-C	183	75	158	155	109	140	120	-
663	P, 29-C	138	100	157	149	96	80	60	80
664	P, 29-C	191	210	173	175	151	167	161	156
667	P, 29-C	211	205	163	155	149	129	140	155
668	P, 29-C	185	178	145	170	147	169	152	154
670	P, 29-C	110	136	112	110	97	110	109	98
672	P, 29-C	93	183	156	152	147	141	137	140
673	P, 29-C	0	0	0	0	0	0	0	0
Mean*		146	148	137	119	100	106	99	112
±s.d.		68.5	77.1	54.2	68.7	60.7	61.7	59.9	57.9
656	P, 13-D	-	-	-	-	-	-	-	-
658	P, 10-D	-	-	-	-	-	-	-	-
662	P, 18-D	-	-	-	-	-	-	-	-
665	P, 24-D	32	0	0	-	-	-	-	-
666	P, 12-D	-	-	-	-	-	-	-	-
671	P, 10-D	-	-	-	-	-	-	-	-
674	P, 15-D	-	-	-	-	-	-	-	-
675	P, 16-D	-	-	-	-	-	-	-	-
657	NP, 28-C	202	191	163	187	173	95	59	-
660	NP, 28-C	185	216	215	224	222	240	207	-
669	NP, 6-D	-	-	-	-	-	-	-	-

Table No. 1A - Continued

GROUP NO. 3 - 2 G/KG/DAY (Continued)

MATERNAL RABBIT NO.	TOTAL MEAN VALUES/DAY					
	DAYS 0 THRU 5 MEAN	±S.D.	DAYS 6 THRU 18 MEAN	±S.D.	DAYS 19 THRU 27/28 MEAN	±S.D.
659 P, 28-C	154	29.2	92	80.1	113	100.6
661 P, 28-C	198	9.5	154	35.9	147	40.3
663 P, 29-C	164	15.0	111	38.5	114	34.8
664 P, 29-C	207	10.1	135	77.0	175	18.0
667 P, 29-C	216	13.9	189	25.5	168	29.1
668 P, 29-C	205	16.9	161	31.1	154	24.2
670 P, 29-C	162	8.6	123	12.8	111	11.8
672 P, 29-C	165	18.3	151	14.0	136	28.2
673 P, 29-C	197	12.0	97	80.9	1	1.9
Mean*	185	-	135	-	124	-
±s.d.	23.7	-	32.1	-	52.0	-
656 P, 13-D	169	12.2	77	85.2	-	-
658 P, 10-D	208	36.3	28	56.0	-	-
662 P, 18-D	154	10.3	49	59.4	-	-
665 P, 24-D	203	4.6	78	63.7	43	46.4
666 P, 12-D	161	9.1	142	60.8	-	-
671 P, 10-D	128	20.1	74	70.4	-	-
674 P, 15-D	219	14.4	104	91.1	-	-
675 P, 16-D	190	11.8	30	54.4	-	-
657 NP, 28-C	200	15.0	171	24.8	146	51.3
660 NP, 28-C	226	12.1	186	25.8	202	33.2
669 NP, 6-D	119	54.5	-	-	-	-

Table No. 1A - Continued

GROUP NO. 4 - 3 G/KG/DAY

MATERNAL RABBIT NO.		PRETREATMENT					DAY OF GESTATION					TREATMENT PERIOD				
		0	1	2	3	4	5	6	7	8	9	10	11			
677	P, 28-C	150	151	160	155	135	158	31	0	0	0	0	10			
679	P, 28-C	187	190	162	185	160	172	62	40	50	26	12	0			
680	P, 28-C	177	131	232	195	185	192	120	106	110	144	123	123			
681	P, 28-C	131	136	148	145	132	137	42	71	101	81	0	0			
683	P, 29-C	217	174	239	221	195	211	90	168	200	180	157	71			
684	P, 29-C	237	252	235	225	220	226	185	182	180	177	193	171			
687	P, 29-C	168	180	199	193	185	195	125	153	172	173	159	98			
691	P, 29-C	211	204	196	201	165	201	134	162	160	181	156	136			
Mean*		185	177	196	190	172	186	99	110	122	120	100	76			
±s.d.		35.7	39.7	36.7	28.3	30.1	29.1	52.2	67.2	69.7	74.4	81.8	66.8			
676	P, 27-D	216	232	248	234	180	243	165	185	191	186	206	22			
678	P, 18-D	169	170	170	175	160	171	93	50	60	0	0	0			
682	P, 18-D	176	187	200	190	175	180	66	121	140	65	3	0			
685	P, 29-D	158	181	170	177	190	189	125	110	150	112	100	92			
688	P, 13-D	174	191	198	182	171	179	75	125	144	91	25	0			
690	P, 18-D	186	180	191	185	165	184	46	25	10	22	13	0			
692	P, 19-D	163	180	197	180	170	186	113	131	130	91	100	7			
694	P, 25-D	156	155	200	189	177	189	89	110	157	78	61	8			
695	P, 21-D	231	237	239	150	180	225	67	70	120	29	12	0			
686	NP, 10-D	88	29	15	0	0	0	0	0	0	0	-	-			
689	NP, 11-D	196	139	143	135	135	147	29	29	31	10	0	-			
693	NP, 14-D	164	181	172	188	140	189	28	7	8	3	0	0			

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Table No. 1A - Continued
GROUP NO. 4 - 3 G/KG/DAY (Continued)

MATERNAL RABBIT NO.		TREATMENT PERIOD							POSTTREATMENT PERIOD		
		DAY OF GESTATION									
		12	13	14	15	16	17	18	19	20	
677	P, 28-C	0	0	0	0	0	0	0	0	0	0
679	P, 28-C	0	0	0	0	0	0	0	24	60	
680	P, 28-C	90	101	100	130	120	165	205	93	174	
681	P, 28-C	37	0	6	4	0	0	0	0	0	
683	P, 29-C	107	59	152	133	124	193	207	201	234	
684	P, 29-C	166	171	190	175	150	184	190	135	100	
687	P, 29-C	165	100	182	157	152	183	205	180	173	
691	P, 29-C	152	129	170	194	82	35	87	90	90	
Mean*		90	70	100	99	78	95	112	90	104	
±s.d.		70.2	65.9	85.6	83.6	68.5	93.2	100.5	78.4	84.8	
676	P, 27-D	33	0	0	20	0	5	0	8	0	
678	P, 18-D	0	0	0	0	0	0	-	-	-	
682	P, 18-D	0	0	0	0	0	0	-	-	-	
685	P, 29-D	155	129	169	186	35	80	5	0	0	
688	P, 13-D	0	-	-	-	-	-	-	-	-	
690	P, 18-D	0	0	0	0	0	0	-	-	-	
692	P, 19-D	0	0	0	0	0	0	0	-	-	
694	P, 25-D	11	0	0	0	0	0	0	0	0	
695	P, 21-D	0	0	0	0	0	0	0	0	0	
686	NP, 10-D	-	-	-	-	-	-	-	-	-	
689	NP, 11-D	-	-	-	-	-	-	-	-	-	
693	NP, 14-D	0	0	-	-	-	-	-	-	-	

Table No. 1A - Continued

GROUP NO. 4 - 3 G/KG/DAY (Continued)

MATERNAL RABBIT NO.	POSTTREATMENT PERIOD									
	DAY OF GESTATION									
	21	22	23	24	25	26	27	28		
P, 28-C	0	0	0	34	17	55	69	-		
P, 28-C	115	145	145	191	175	205	175	-		
P, 28-C	185	157	115	98	109	160	12	-		
P, 28-C	0	0	0	0	0	0	0	-		
P, 29-C	240	246	232	234	220	214	234	175		
P, 29-C	130	190	127	134	82	0	0	0		
P, 29-C	207	202	152	172	143	158	147	136		
P, 29-C	123	155	124	107	66	94	96	81		
Mean*	125	137	112	121	102	111	92	98		
±s.d.	88.6	90.4	78.0	78.7	75.9	86.3	87.8	75.9		
P, 27-D	0	0	0	0	0	0	-	-		
P, 18-D	-	-	-	-	-	-	-	-		
P, 18-D	-	-	-	-	-	-	-	-		
P, 29-D	37	28	0	25	8	30	11	0		
P, 13-D	-	-	-	-	-	-	-	-		
P, 18-D	-	-	-	-	-	-	-	-		
P, 19-D	-	-	-	-	-	-	-	-		
P, 25-D	0	0	0	0	-	-	-	-		
P, 21-D	-	-	-	-	-	-	-	-		
NP, 10-D	-	-	-	-	-	-	-	-		
NP, 11-D	-	-	-	-	-	-	-	-		
NP, 14-D	-	-	-	-	-	-	-	-		

Table No. 1A - Continued
GROUP NO. 4 - 3 G/KG/DAY (Continued)

MATERNAL RABBIT NO.	TOTAL MEAN VALUES/DAY					
	DAYS 0 THRU 5		DAYS 6 THRU 18		DAYS 19 THRU 27/28	
	MEAN	±S.D.	MEAN	±S.D.	MEAN	±S.D.
677 P, 28-C	152	9.0	3	8.8	19	27.0
679 P, 28-C	176	13.2	15	22.3	137	61.0
680 P, 28-C	185	32.7	126	30.9	123	53.7
681 P, 28-C	138	6.9	26	36.5	0	-
683 P, 29-C	210	22.5	142	49.1	223	21.4
684 P, 29-C	232	11.5	178	11.7	90	67.8
687 P, 29-C	187	11.5	156	31.5	167	24.3
691 P, 29-C	196	16.1	137	44.7	103	25.6
Mean*	184	-	98	-	108	-
±s.d	30.1	-	70.8	-	73.4	-
676 P, 27-D	226	24.8	78	90.4	1	2.8
678 P, 18-D	169	5.0	17	32.1	-	-
682 P, 18-D	185	9.6	33	52.1	-	-
685 P, 29-D	178	12.1	111	51.2	14	14.6
688 P, 13-D	182	10.3	66	58.5	-	-
690 P, 18-D	182	9.0	10	14.7	-	-
692 P, 19-D	179	11.9	44	57.7	-	-
694 P, 25-D	178	18.7	40	53.5	0	-
695 P, 21-D	210	36.7	23	38.7	0	-
686 NP, 10-D	22	34.4	0	-	-	-
689 NP, 11-D	149	23.4	20	14.0	-	-
693 NP, 14-D	172	18.5	6	9.6	-	-

Table No. 2 - Detailed observations and necropsy findings
for animals which died.

Group No. 1 - Control (Four Deaths):

Rabbit No. 628 (18-D) - Nasal discharge and/or wheezing beginning Day 9; 6% cumulative body weight loss by Day 15, no food consumed for four days prior to death. Necropsy - thoracic cavity filled with white, purulent material.

Rabbit No. 632 (29-D) - Little or no food consumption from Day 14; 19% cumulative body weight loss by Day 22; depression, nasal discharge, wheezing, labored or rapid respiration, and/or a thin appearance exhibited from Day 19. Necropsy - both uterine horns filled with white, purulent material.

Rabbit No. 633 (17-D) - Little or no food consumed from Day 11; 6% body weight loss on Day 15; labored respiration Day 16. Necropsy - lungs filled with thick, white, purulent material.

Rabbit No. 635 (9-D) - Little or no food consumed from Day 6; labored respiration Day 8. Necropsy - dark red fluid present in thoracic cavity.

Group No. 2 - 1 g/kg (12 Deaths):

Rabbit No. 636 (10-D) - Labored respiration Day 9. Necropsy - lungs appeared collapsed and ovaries small.

Rabbit No. 637 (22-D) - Little or no food consumed from Day 14; 9% cumulative body weight loss by Day 18; and thin appearance. Necropsy - white, purulent, material present in thoracic cavity.

Rabbit No. 638 (6-D) - Found dead after dosing; no unusual signs noted. Necropsy - no gross lesions observed.

Rabbit No. 639 (17-D) - Reduced or no food consumption from Day 10; 9% body weight loss on Day 15; and wheezing from Day 14. Necropsy - white, purulent, material present in thoracic cavity.

Rabbit No. 640 (13-D) - 6% body weight loss on Day 10; no food consumed from Day 11; thin appearance; labored respiration and/or wheezing from Day 10; and a bloated appearance Day 12. Necropsy - white fluid present in thoracic cavity.

Rabbit No. 642 (4-D) - Died prior to initiation of treatment; appeared bloated and depressed; consumed little or no food. Necropsy - red fluid present in abdominal cavity.

Table No. 2 - Continued)

Rabbit No. 643 (25-D) - Nasal discharge observed Days 11 through 18; 6% cumulative body weight loss by Day 22; little or no food consumed from Day 21; wheezing and/or labored respiration Days 23 and 24. Necropsy - no gross lesions observed.

Rabbit No. 644 (9-D) - Labored respiration and no food consumed Day 8. Necropsy - lungs appeared collapsed.

Rabbit No. 646 (23-D) - Little or no food consumed from Day 11; 21% cumulative body weight loss by Day 22; thin appearance and labored respiration Day 22. Necropsy - no gross lesions observed.

Rabbit No. 649 (6-D) - Found dead after dosing; no unusual signs noted. Necropsy - no gross lesions observed.

Rabbit No. 652 (15-D) - Little or no food consumption from Day 9; 7% body weight loss Day 10; and thin appearance. Necropsy - white fluid present in thoracic cavity.

Rabbit No. 655 (10-D) - Reduced or no food consumption from Day 7. Necropsy - Small portion of lungs white in color.

Group No. 3 - 2 g/kg (Nine Deaths):

Rabbit No. 656 (13-D) - 7% body weight loss Day 10; no food consumed from Day 9; thin appearance; wheezing and/or nasal discharge Day 6 and Days 10 through 12; and depression Day 12. Necropsy - fluid present in thoracic cavity.

Rabbit No. 658 (10-D) - No food consumed from Day 7; nasal discharge Day 7; and labored respiration Days 8 and 9. Necropsy - lungs appeared collapsed.

Rabbit No. 662 (18-D) - Little or no food consumed from Day 11; 11% body weight loss on Day 15; thin appearance; soft feces Days 11 through 17; depression and labored respiration Day 17. Necropsy - no gross lesions observed.

Rabbit No. 665 (24-D) - Generally reduced food consumption from Day 8; 5% cumulative body weight loss by Day 22; ataxia and labored respiration Day 23. Necropsy - no gross lesions observed.

Rabbit No. 666 (12-D) - Little food consumed Day 11. Necropsy - dark red fluid present in thoracic cavity.

Table No. 2 - Continued

Rabbit No. 669 (6-D) - Found dead after dosing; no unusual signs noted. Necropsy - no gross lesions observed.

Rabbit No. 671 (10-D) - Little or no food consumed from Day 8; wheezing and labored respiration Day 9. Necropsy - red fluid present in lungs.

Rabbit No. 674 (15-D) - Little or no food consumed from Day 11; nasal discharge and labored respiration Days 13 and 14. Necropsy - no gross lesions observed.

Rabbit No. 675 (16-D) - Little or no food consumed from Day 8; 15% cumulative body weight loss by Day 15; soft feces Days 10 through 15; and wheezing and labored respiration Days 12 through 15. Necropsy - dark red fluid present in thoracic cavity.

Group No. 4 - 3 g/kg (12 Deaths):

Rabbit No. 676 (27-D) - Little or no food consumed from Day 11; 17% cumulative body weight loss by Day 22; thin appearance; red jelly-like material found in cage Day 24; and nasal discharge Day 26. Necropsy - no gross lesions observed.

Rabbit No. 678 (18-D) - No food consumed from Day 9; 13% cumulative weight loss by Day 15; thin appearance; soft feces Days 9 through 17; labored respiration and prostration Days 16 and 17. Necropsy - no gross lesions observed.

Rabbit No. 682 (18-D) - Little or no food consumed from Day 10; 13% cumulative weight loss by Day 15; thin appearance; soft feces Days 12 through 16; depression Day 16; nasal discharge Days 16 and 17; prostration and labored respiration Days 17 and 18. Necropsy - no gross lesions observed.

Rabbit No. 685 (29-D) - Reduced or no food consumption from Day 16; 18% cumulative body weight loss by Day 22; thin appearance; nasal discharge generally from Day 7; soft feces Days 9 through 14; and depression generally from Day 17. Necropsy - no gross lesions observed.

Rabbit No. 686 (10-D) - Reduced or no food consumption and body weight loss observed prior to treatment and a generally weakened condition. Necropsy - uterus appeared deformed; both ovaries absent.

Table No. 2 - Continued

Rabbit No. 688 (13-D) - 3% body weight loss Day 10; no food consumed Days 11 or 12; and wheezing, labored respiration, and/or soft feces Days 10 through 12. Necropsy - fluid present in thoracic cavity.

Rabbit No. 689 (11-D) - Little food consumed from Day 6; 7% body weight loss Day 10; thin appearance; soft feces Day 9; and weakened condition, slight ataxia, bloated appearance, and labored respiration Day 10. Necropsy - large intestine filled with air.

Rabbit No. 690 (18-D) - Little or no food consumed from Day 6; 20% cumulative body weight loss by Day 15; soft feces and thin appearance from Day 9; and depression Days 16 and 17. Necropsy - no gross lesions observed.

Rabbit No. 692 (19-D) - No food consumed from Day 12; 19% cumulative body weight loss by Day 18; thin appearance from Day 10; nasal discharge Day 12; depression and soft feces Days 16 and 17; and prostration and labored respiration Day 18. Necropsy - no gross lesions observed.

Rabbit No. 693 (14-D) - Little or no food consumed from Day 6; 7% body weight loss Day 10; soft feces and nasal discharge from Day 9. Necropsy - no gross lesions observed.

Rabbit No. 694 (25-D) - Little or no food consumed from Day 9; 25% cumulative body weight loss by Day 22; intermittent nasal discharge; thin appearance and depression from Day 15; ataxia Days 18 through 21; and labored respiration Days 23 and 24. Necropsy - no gross lesions observed.

Rabbit No. 695 (21-D) - No food consumed from Day 11; 20% cumulative body weight loss by Day 18; soft feces Days 8 through 13; nasal discharge Day 12; and thin appearance, depression, and/or ataxia from Day 15. Necropsy - no gross lesions observed.

Table No. 3 - Detailed observations and necropsy findings
for pregnant animals sacrificed at term.

Group No. 1 - Control (15 Animals):

Rabbit No. 618 (28-C) - Soft feces Day 7.

Rabbit No. 622 (29-C) - Nasal discharge Day 12.

Rabbit No. 625 (29-C) - Thin appearance Days 11 to 18.

Rabbit No. 627 (29-C) - Nasal discharge generally Days 11 to 28.

Rabbit No. 630 (29-C) - Wheezing and/or nasal discharge generally
Days 22 to 29, thin appearance from Day 24, and depression
Day 29.

Rabbit No. 631 (29-C) - Nasal discharge Days 9 to 29 and thin
appearance Days 12 to 14.

Rabbit No. 634 (29-C) - Soft feces Days 9 to 18.

Rabbits No. 616 (28-C), No. 617 (28-C), No. 620 (29-C), No. 621
(29-C), No. 623 (29-C), No. 624 (29-C), No. 626 (29-C), and
No. 629 (29-C) - Appeared normal throughout the study.

No gross visceral lesions were observed among any pregnant control animals
sacrificed at term.

Group No. 2 - 1 g/kg (Seven Animals):

Rabbit No. 645 (28-C) - Nasal discharge Days 12 to 15.

Rabbit No. 647 (28-C) - Nasal discharge Days 12 to 28 and wheezing
and a bloody-appearing fluid around the nose Day 21.

Rabbit No. 650 (29-C) - Red fluid in cage pan Day 24.

Rabbit No. 651 (29-C) - Nasal discharge generally from Day 14 and
depression Day 29.

Rabbit No. 653 (29-C) - Wheezing Days 11 to 16 and Days 27 to 29,
nasal discharge Days 14 to 16, and labored or rapid respiration
from Day 23.

Rabbits No. 641 (28-C) and No. 654 (29-C) - Appeared normal through-
out the study.

No gross visceral lesions were observed among any pregnant Group No. 2
animals sacrificed at term.

Table No. 3 - Continued

Group No. 3 - 2 g/kg (Nine Animals):

Rabbit No. 659 (28-C) - Thin appearance Days 12 to 14 and nasal discharge Days 26 to 28.

Rabbit No. 663 (29-C) - Nasal discharge Days 15 to 18 and Days 25 to 28 and wheezing Day 17.

Rabbit No. 664 (29-C) - Thin appearance Days 10 to 15.

Rabbit No. 667 (29-C) - Nasal discharge Days 10 to 18 and wheezing Day 17.

Rabbit No. 668 (29-C) - Nasal discharge generally Days 6 to 18 and Day 26 and thin appearance Day 16.

Rabbit No. 670 (29-C) - Soft feces Day 11.

Rabbit No. 673 (29-C) - Nasal discharge Days 26 to 29 and depression Day 29.

Rabbits No. 661 (28-C) and No. 672 (29-C) - Appeared normal throughout the study.

At necropsy, gross visceral lesions were evident in Group No. 3 Rabbit No. 673 - Purulent material present in right lung, liver mottled, and gallbladder distended. No other gross visceral lesions were observed among the pregnant Group No. 3 animals sacrificed at term.

Group No. 4 - 3 g/kg (Eight Animals):

Rabbit No. 677 (28-C) - Soft feces Days 9 to 17 and thin appearance Days 9 to 17 and Days 22 to 28.

Rabbit No. 679 (28-C) - Soft feces Days 9 to 12 and Day 17, thin appearance Days 9 to 17, and depression Days 11 to 17.

Rabbit No. 681 (28-C) - Soft feces Days 15 to 17 and Day 21, nasal discharge Days 15 and 17 and Days 24 to 28, ataxia Days 18 to 22 and Days 27 and 28, depression Days 18 to 26, labored respiration Days 19 to 21 and Days 27 and 28, and thin appearance Days 22 to 28.

Rabbit No. 684 (29-C) - Nasal discharge Days 16 and 17.

Rabbit No. 687 (29-C) - Soft feces generally Days 9 to 17, nasal discharge Days 10 to 17 and Days 21 and 22, and wheezing Day 21.

Table No. 3 - Continued

Rabbit No. 691 (29-C) - Nasal discharge Days 10 to 29 and wheezing Days 13 to 17, Day 24, and Days 27 and 28.

Rabbits No. 680 (28-C) and No. 683 (29-C) - Appeared normal throughout the study.

At necropsy, gross visceral lesions were evident in Group No. 4 Rabbit No. 681 - Large amount of a thick, white substance, presumptively identified as the administered compound, intermingled with a moderate amount of hair and food-like material present in the stomach, very small amount of food present in the intestinal tract, and gallbladder distended and filled with a large amount of bile. No other gross visceral lesions were observed among the pregnant Group No. 4 animals sacrificed at term.

Table No. 4 - Comparison of uterine implantation sites and ovarian corpora lutea in rabbits serving as controls or receiving SC-18862 and 19192 (3:1 ratio)

MATERNAL RABBIT NO.	DAY OF GESTATION	GROUP NO. 1 - CONTROL				OVARIAN			
		UTERINE		CORPORA LUTEA		CORPORA LUTEA		CORPORA LUTEA	
		IMPLANTATION SITES		RIGHT HORN		LEFT OVARY		RIGHT OVARY	
		LEFT HORN	RIGHT HORN	LEFT HORN	RIGHT HORN	LEFT OVARY	RIGHT OVARY	LEFT OVARY	RIGHT OVARY
616	P, 28-C	5		1		9	4		
617	P, 28-C	3		1		6	7		
618	P, 28-C	4		4		6	5		
619	NP, 28-C	0		0		0	0		
620	P, 29-C	0		6		2	8		
621	P, 29-C	0		3		2	6		
622	P, 29-C	3		7		4	8		
623	P, 29-C	3		2		6	5		
624	P, 29-C	5		6		5	7		
625	P, 29-C	3		4		5	6		
626	P, 29-C	7		3		9	5		
627	P, 29-C	2		7		5	7		
628	P, 18-D	5		3		6	5		
629	P, 29-C	6		4		9	8		
630	P, 29-C	6		4		8	4		
631	P, 29-C	2		9		4	9		
632	P, 29-D	6		3		6	4		
633	P, 17-D	3		1		5	3		
634	P, 29-C	6		4		7	4		
635	P, 9-D	4		2		5	2		
Subtotal		73		74		109	107		
Total			147				216		
Ratio*							147/216 = 68.1%		

Table No. 4 - Continued

GROUP NO. 2 - 1 G/KG/DAY

MATERNAL RABBIT NO.	DAY OF GESTATION	UTERINE		OVARIAN	
		IMPLANTATION SITES		CORPORA LUTEA	
		LEFT HORN	RIGHT HORN	LEFT OVARY	RIGHT OVARY
636	P, 10-D	2	4	6	5
637	P, 22-D	6	5	7	5
638	NP, 6-D	0	0	5	6
639	P, 17-D	1	1	1	1
640	NP, 13-D	0	0	0	0
641	P, 28-C	3	7	5	7
642	NP, 4-D	0	0	2	0
643	P, 25-D	6	5	7	6
644	P, 9-D	2	6	2	6
645	P, 28-C	4	7	5	7
646	P, 23-D	4	6	6	8
647	P, 28-C	4	5	4	5
648	NP, 28-C	0	0	5	4
649	NP, 6-D	0	0	3	9
650	P, 29-C	7	4	9	6
651	P, 29-C	3	6	4	7
652	P, 15-D	4	6	5	6
653	P, 29-C	3	3	9	8
654	P, 29-C	3	4	6	12
655	NP, 10-D	0	0	7	4
Subtotal		52	69	76	89
Total			121		165
Ratio*					121/165 = 73.3%

Table No. 4 - Continued
GROUP NO. 3 - 2 G/KG/DAY

MATERNAL RABBIT NO.	DAY OF GESTATION	UTERINE		OVARIAN	
		IMPLANTATION SITES		CORPORA LUTEA	
		LEFT HORN	RIGHT HORN	LEFT OVARY	RIGHT OVARY
656	P, 13-D	4	4	4	6
657	NP, 28-C	0	0	2	1
658	P, 10-D	7	8	7	7
659	P, 28-C	4	5	4	5
660	NP, 28-C	0	0	3	5
661	P, 28-C	3	7	4	12
662	P, 18-D	7	4	7	5
663	P, 29-C	5	4	8	5
664	P, 29-C	6	3	8	6
665	P, 24-D	2	5	6	10
666	P, 12-D	1	1	4	5
667	P, 29-C	4	6	4	8
668	P, 29-C	0	1	9	4
669	NP, 6-D	0	0	3	7
670	P, 29-C	2	8	2	10
671	P, 10-D	1	8	1	8
672	P, 29-C	5	8	7	10
673	P, 29-C	1	5	2	8
674	P, 15-D	5	6	5	7
675	P, 16-D	5	4	5	6
Subtotal		62	87	87	122
Total			149		209
Ratio*					149/209 = 71.3%

Table No. 4 - Continued

GROUP NO. 4 - 3 G/KG/DAY

MATERNAL RABBIT NO.	DAY OF GESTATION	UTERINE		OVARIAN	
		IMPLANTATION SITES		CORPORA LUTEA	
		LEFT HORN	RIGHT HORN	LEFT OVARY	RIGHT OVARY
676	P, 27-D	2	4	3	4
677	P, 28-C	3	7	3	8
678	P, 18-D	5	8	7	10
679	P, 28-C	5	1	6	2
680	P, 28-C	5	4	5	5
681	P, 28-C	4	0	8	10
682	P, 18-D	7	4	8	6
683	P, 29-C	6	7	8	8
684	P, 29-C	7	4	9	6
685	P, 29-D	8	2	4	2
686	NP, 10-D	0	0	-**	-**
687	P, 29-C	2	4	6	5
688	P, 13-D	3	0	3	3
689	NP, 11-D	0	0	6	3
690	P, 18-D	4	5	4	6
691	P, 29-C	5	6	6	7
692	P, 19-D	2	2	4	8
693	NP, 14-D	0	0	5	3
694	P, 25-D	5	2	5	2
695	P, 21-D	4	5	5	8
Subtotal		77	65	94	100
Total			142		194
Ratio*					142/194 = 73.2%

Table No. 5 - Uterine and litter data for pregnant rabbits serving as controls or receiving SC-18862 and 19192 (3:1 ratio)

GROUP NO. 1 - CONTROL

MATERNAL RABBIT NO.	DAY OF GESTATION	IMPLANTATION SITES	RESORPTION SITES		NUMBER OF LIVE FETUSES		NUMBER OF DEAD FETUSES		LIVE FETUSES		DEAD FETUSES*	
			LEFT	RIGHT	LEFT	RIGHT	LEFT	RIGHT	MEAN WEIGHT g.	MEAN LENGTH cm.	MEAN WEIGHT g.	MEAN LENGTH cm.
616	28-C	6	1	0	4	1	0	0	37.0	8.4	-	-
617	28-C	4	3	1	0	0	0	0	-	-	-	-
618	28-C	8	0	1	4	3	0	0	38.8	8.6	-	-
620	29-C	6	0	1	0	5	0	0	34.3	8.1	-	-
621	29-C	3	0	0	0	3	0	0	59.2	10.0	-	-
622	29-C	10	0	0	3	7	0	0	45.0	9.4	-	-
623	29-C	5	0	0	3	2	0	0	40.7	9.2	-	-
624	29-C	11	1	0	4	5	0	1	44.6	8.7	9.5	6.0
625	29-C	7	1	1	2	3	0	0	51.7	9.0	-	-
626	29-C	10	0	0	6	3	1	0	43.9	8.0	-**	-
627	29-C	9	0	1	2	5	0	1	38.7	8.4	-**	-
629	29-C	10	0	0	6	4	0	0	50.0	8.7	-	-
631	29-C	10	0	0	6	4	0	0	42.1	8.5	-	-
634	29-C	11	0	0	2	4	0	5	37.3	8.1	16.3	6.1
Subtotal			6	5	42	49	1	7				
Total		110	11		91			8				
Mean		7.9	0.8		6.5			0.6	43.3	8.7	12.9	6.1
±s.d.		2.68	1.12		2.95			1.34	6.95	0.57	-	-
628	18-D	8φ	-	-	-	-	-	-	-	-	-	-
630	29-C	10φφ	0	0	0	0	0	0	-	-	-	-
632	29-D	9	6	3	0	0	0	0	-	-	-	-
633	17-D	4φ	-	-	-	-	-	-	-	-	-	-
635	9-D	6φ	-	-	-	-	-	-	-	-	-	-

Table No. 5 - Continued
GROUP NO. 2 - 1 G/KG/DAY

MATERNAL RABBIT NO.	DAY OF GESTATION	IMPLANTATION SITES	RESORPTION SITES		NUMBER OF LIVE FETUSES		NUMBER OF DEAD FETUSES		LIVE FETUSES MEAN WEIGHT g. MEAN LENGTH cm.		DEAD FETUSES* MEAN WEIGHT g. MEAN LENGTH cm.	
			LEFT	RIGHT	LEFT	RIGHT	LEFT	RIGHT	WEIGHT	LENGTH	WEIGHT	LENGTH
641	28-C	10	0	0	3	7	0	0	35.4	8.0	-	-
645	28-C	11	0	0	4	3	0	4	35.3	8.1	-**	-
647	28-C	9	0	0	4	5	0	0	36.3	8.2	-	-
650	29-C	11	0	1	7	3	0	0	42.5	8.8	-	-
651	29-C	9	0	0	3	6	0	0	30.6	7.9	-	-
653	29-C	6	0	0	3	3	0	0	44.7	9.2	-	-
654	29-C	7	1	0	2	4	0	0	50.0	10.1	-	-
Subtotal			1	1	26	31	0	4				
Total		63	2	57			4					
Mean		9.0	0.3	8.1			0.6		39.3	8.6	-	-
±s.d.		1.91	0.49	1.77			1.51		6.71	0.81	-	-
636	10-D	6φ	-	-	-	-	-	-	-	-	-	-
637	22-D	11φ	-	-	-	-	-	-	-	-	-	-
639	17-D	2φ	-	-	-	-	-	-	-	-	-	-
643	25-D	11	0	0	0	0	6	5	-	-	13.6	7.2
644	9-D	8φ	-	-	-	-	-	-	-	-	-	-
646	23-D	10	0	0	0	0	4	6	-	-	-**	-
652	15-D	10φ	-	-	-	-	-	-	-	-	-	-

Table No. 5 - Continued
GROUP NO. 3 - 2 G/KG/DAY

MATERNAL RABBIT NO.	DAY OF GESTATION	IMPLANTATION SITES	RESORPTION SITES		NUMBER OF LIVE FETUSES		NUMBER OF DEAD FETUSES		LIVE FETUSES		DEAD FETUSES*	
			LEFT	RIGHT	LEFT	RIGHT	LEFT	RIGHT	MEAN WEIGHT g.	MEAN LENGTH cm.	MEAN WEIGHT g.	MEAN LENGTH cm.
659	28-C	9	3	2	0	2	1	1	17.2	6.5	19.8	6.5
661	28-C	10	0	0	3	7	0	0	36.8	8.0	-	-
663	29-C	9	1	0	4	4	0	0	33.4	8.2	-	-
664	29-C	9	0	0	6	3	0	0	45.0	9.3	-	-
667	29-C	10	0	0	4	6	0	0	40.9	9.2	-	-
668	29-C	1	0	1	0	0	0	0	-	-	-	-
670	29-C	10	0	2	2	6	0	0	38.7	8.5	-	-
672	29-C	13	0	0	4	8	1	0	36.9	8.1	5.6	4.0
673	29-C	6	0	1	0	0	1	4	-	-	8.7	5.3
Subtotal			4	6	23	36	3	5				
Total		77	10		59		8					
Mean		8.6	1.1		6.6		0.9		35.6 ^{S-}	8.3	11.4	5.3
±s.d.		3.36	1.62		4.61		1.69		8.87	0.93	-	-
656	13-D	8φ	-	-	-	-	-	-	-	-	-	-
658	10-D	15φ	-	-	-	-	-	-	-	-	-	-
662	18-D	11φ	-	-	-	-	-	-	-	-	-	-
665	24-D	7#	0	1	-	-	-	-	-	-	-	-
666	12-D	2φ	-	-	-	-	-	-	-	-	-	-
671	10-D	9φ	-	-	-	-	-	-	-	-	-	-
674	15-D	11φ	-	-	-	-	-	-	-	-	-	-
675	16-D	9φ	-	-	-	-	-	-	-	-	-	-

S- = Significantly lower than control at $p < 0.05$.

Table No. 5 - Continued

GROUP NO. 4 - 3 G/KG/DAY

MATERNAL RABBIT NO.	DAY OF GESTATION	IMPLANTATION SITES	RESORPTION SITES		NUMBER OF LIVE FETUSES		NUMBER OF DEAD FETUSES		LIVE FETUSES		DEAD FETUSES*	
			LEFT	RIGHT	LEFT	RIGHT	LEFT	RIGHT	MEAN WEIGHT g.	MEAN LENGTH cm.	MEAN WEIGHT g.	MEAN LENGTH cm.
677	28-C	10	3	7	0	0	0	0	-	-	-	-
679	28-C	6	0	0	5	1	0	0	23.7	7.3	-	-
680	28-C	9	0	0	5	3	0	1	26.9	7.4	-**	-
681	28-C	4	4	0	0	0	0	0	-	-	-1/	6.51/
683	29-C	13	2	3	3	3	1	1	33.1	7.8	20.01/	-
684	29-C	11	0	0	7	4	0	0	33.0	8.0	-	-
687	29-C	6	0	0	2	4	0	0	44.6	9.8	-	-
691	29-C	11	0	1	5	5	0	0	38.0	7.8	-	-
Subtotal			9	11	27	20	1	2				
Total		70	20		47		3					
Mean		8.8	2.5		5.9		0.4		33.2 S-	8.0	20.0	6.5
±s.d.		3.11	3.63		4.09		0.74		7.52	0.91	-	-
676	27-D	6	0	0	0	0	2	4	-	-	-**	-
678	18-D	13	-	-	-	-	-	-	-	-	-	-
682	18-D	11	-	-	-	-	-	-	-	-	-	-
685	29-D	10	8	2	0	0	0	0	-	-	-	-
688	13-D	3	-	-	-	-	-	-	-	-	-	-
690	18-D	9	-	-	-	-	-	-	-	-	-	-
692	19-D	4	-	-	-	-	-	-	-	-	-	-
694	25-D	7##	0	1	0	0	0	0	-	-	-	-
695	21-D	9	-	-	-	-	-	-	-	-	-	-

S- = Significantly lower than control at $p < 0.05$.