

Study Number: MOG002B
Test Type: MOG
Route: Dosing in Feed
Species/Strain: Rat/Sprague-Dawley

R16: Pubertal Markers Summary
Test Compound: 2-Hydroxy-4-methoxybenzophenone
CAS Number: 131-57-7

Date Report Requested: 03/26/2019
Time Report Requested: 06:25:26
Lab: RTI

C Number:

MOG002B

Study Gender:

Both

PWG Approval Date

See web page for date of PWG Approval

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			Male					
Generation	Litter	Cohort	Treatment Groups (ppm)					
			0	3000	10000	30000	0.05 ppm EE	
F1	A	All Males	No. Examined (litters)	64 (22)	59 (20)	62 (21)	60 (20)	45 (15)
			No. Removed (litters) ^a	0 (0)	1 (1)	0 (0)	0 (0)	0 (0)
			No. Not Attaining BPS (litters) ^b	0 (0)	0 (0)	0 (0)	3 (1)	0 (0)
			Day of BPS					
			Mean Analysis ^c					
			Litter Mean ± SE ^d	43.7 ± 0.3 **	44.0 ± 0.4	44.9 ± 0.3 *	47.1 ± 0.4 **	45.8 ± 0.3 **
			Litter Mean of Adjusted ± SE ^e	44.7 ± 0.3	44.7 ± 0.3	44.8 ± 0.3	45.4 ± 0.3	44.8 ± 0.3
			Proportional Hazards Analysis ^f					
			Litter-based Model ^g	p=0.112	p=0.956	p=0.956	p=0.852	p=0.138
BW at Attainment (g) ^h	204.4 ± 2.9 **	203.3 ± 2.9	196.4 ± 2.2	192.1 ± 2.8 **	184.7 ± 2.2 **			
BW at Weaning (g) ^h	90.1 ± 1.1 **	87.4 ± 1.6	81.4 ± 1.2 **	68.6 ± 1.9 **	80.3 ± 1.2 **			

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Generation	Litter	Cohort	Female					
			Treatment Groups (ppm)					
			0	3000	10000	30000	0.05 ppm EE	
F1	A	All Females	No. Examined (litters)	63 (22)	60 (20)	62 (22)	60 (20)	55 (15)
			No. Removed (litters) ^a	1 (1)	0 (0)	0 (0)	0 (0)	0 (0)
			No. Not Attaining VO (litters) ^b	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
			Day of VO					
			Mean Analysis ^c					
			Litter Mean ± SE ^d	35.3 ± 0.2 **	35.4 ± 0.4	35.9 ± 0.3	38.1 ± 0.4 **	24.3 ± 0.3 **
			Litter Mean of Adjusted ± SE ^e	35.9 ± 0.2 *	35.8 ± 0.3	35.9 ± 0.3	37.0 ± 0.3	24.3 ± 0.3 **
			Proportional Hazards Analysis ^f					
			Litter-based Model ^g	p=0.004	p=0.846	p=0.846	p=0.062	p<0.001
			BW at Attainment (g) ^h	115.7 ± 1.9 **	114.3 ± 1.6	111.5 ± 1.6	109.0 ± 1.9 *	59.0 ± 1.5 **
			BW at Weaning (g) ^h	80.6 ± 1.1 **	78.1 ± 1.8	73.6 ± 1.3 **	60.7 ± 1.6 **	74.5 ± 1.2 **

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LEGEND

BPS = Balanopreputal separation; BW = Body weight; VO = Vaginal opening

In multiple breeding/littering studies Litter A is the default designation for the first litter; subsequent litters would be B, C etc.

The All Males cohort includes the Fertility males and Prenatal males. The All Females cohort includes all females on study except those from the Bio&MGWM@VO, F1 Extra, PND 28 Bio and PND 56 Bio cohorts..

No. Examined (litters) = the number of animals or pups examined (number of litters represented)

^aAnimals that died or were removed prior to the end of the observation period and did not attain. These animals were excluded from all analyses.

^bAnimals that survived to the end of the observation period without attaining.

^cSummary statistics and mixed model results are presented for animals that attained during the observation period.

^dMeans of litter means presented. Trend and pairwise tests were based on mixed models for day of attainment with dose as a covariate and a random effect for litter. The Dunnett-Hsu adjustment was used for multiple comparisons.

^eMean adjusted day of attainment was calculated from the mean of the litter means of the weaning weight-adjusted attainment days for individual pups. Trend and pairwise tests were based on mixed models for day of attainment with dose and weaning weight as covariates and a random effect for litter. The Dunnett-Hsu adjustment was used for multiple comparisons.

^fAnimals that did not attain by the end of the observation period were included in the proportional hazards analysis.

^gP-values for trend and pairwise comparisons were calculated from a Cox proportional hazards model with dose and weaning weight as covariates and a random effect for litter, and a Hommel adjustment for multiple comparisons.

^hAnalysis of body weight at attainment and body weight at weaning were performed using mixed effects models with dose as covariate and a random effect for litter. The Dunnett-Hsu adjustment was used for multiple comparisons. Animals that attained during the observation period were used for analysis.

Statistical significance for the control group indicates a significant trend test

Statistical significance for a treatment group indicates a significant pairwise test compared to the vehicle control group

* Statistically significant at $P \leq 0.05$

** Statistically significant at $P \leq 0.01$

The EE group was not included in any trend analysis, it was included in the pairwise analysis to the control group.

EE = Ethinyl estradiol

**** END OF REPORT ****