

**Black Cohosh Chronic Perinatal Study-Rats (C00058-03)**

**Table 1: PND 1 Litter Data**

Parameter <sup>d</sup>	Control	75 mg/kg	250 mg/kg	750 mg/kg	Trend <sup>a</sup>
<b>Litter Size<sup>b</sup></b>					
<b>Total (Live + Dead)<sup>j</sup></b>	13.10 ± 0.45 [31]	12.52 ± 0.61 [31]	13.09 ± 0.41 [32]	11.69 ± 0.47 [35]*	0.015-
<b>Live Males<sup>j,k</sup></b>	6.61 ± 0.30 [28]	6.30 ± 0.54 [23]	6.46 ± 0.47 [24]	6.00 ± 0.40 [34]	0.224-
<b>Live Females<sup>j,k</sup></b>	6.57 ± 0.38 [28]	5.87 ± 0.49 [23]	6.33 ± 0.42 [24]	5.44 ± 0.37 [34]	0.116-
<b>Live (Male + Female)<sup>j</sup></b>	13.23 ± 0.40 [30]	12.45 ± 0.60 [31]	12.97 ± 0.42 [32]	11.51 ± 0.46 [35]**	0.007-
<b>Total Dead pups<sup>f,i</sup></b>	9 [3]	2 [2]	4 [4]	6 [5]	
<b>Dead per litter</b>	0.29 ± 0.18 [31]	0.06 ± 0.04 [32]	0.13 ± 0.06 [32]	0.17 ± 0.08 [35]	0.455+
<b>% Males</b>					
<b>% Male in Dose<sup>e,f,j,k</sup></b>	50.1 [369]	51.8 [280]	50.5 [307]	52.4 [389]	
<b>% Male per Litter<sup>b,j,k</sup></b>	50.5 ± 2.1 [28]	49.4 ± 3.5 [23]	50.0 ± 2.8 [24]	52.8 ± 3.0 [34]	0.881+
<b>Pup Body Weights(g)<sup>c,g,j</sup></b>					
<b>Male<sup>k</sup></b>	7.08 ± 0.10 [28]	6.85 ± 0.11 [22]	6.94 ± 0.17 [24]	6.64 ± 0.12 [34]*	0.007-
<b>Female<sup>k</sup></b>	6.79 ± 0.11 [28]	6.51 ± 0.08 [23]	6.59 ± 0.16 [24]	6.38 ± 0.12 [33]*	0.012-
<b>Male + Female</b>	6.95 ± 0.10 [30]	6.65 ± 0.07 [31]	6.68 ± 0.12 [32]	6.48 ± 0.11 [35]**	0.001-
<b>Adjusted Pup Body Weights(g)<sup>c,h,j</sup></b>					
<b>Male<sup>k</sup></b>	7.15 ± 0.09 [28]	6.88 ± 0.11 [22]	6.97 ± 0.14 [24]	6.54 ± 0.12 [34]**	<0.001-
<b>Female<sup>k</sup></b>	6.86 ± 0.09 [28]	6.49 ± 0.09 [23]	6.63 ± 0.14 [24]	6.31 ± 0.11 [33]**	0.001-
<b>Male + Female</b>	7.00 ± 0.09 [30]	6.64 ± 0.08 [31]*	6.72 ± 0.10 [32]*	6.40 ± 0.11 [35]**	<0.001-

a: P-value and direction of trend

b: Each dose was compared to the control with Shirley's test when a trend was present (P<.01 from Jonckheere's trend test) or with Dunn's test when no trend was present. [\* = P<0.05, \*\* = P<0.01]

c: Each dose was compared to the control with Williams' test when a trend was present (P<.01 from Jonckheere's trend test) or with Dunnett's test when no trend was present [\* = P<0.05, \*\* = P<0.01]

d: Mean ± standard error [Number of litters] unless otherwise specified

e: 100 x (Number of live males in dose group)/(number of live males and females in dose group) [Number of pups]

f: No statistics were performed on this endpoint.

g: (Litter weight of live pups at PND 1) / (Live litter size on PND 1)

h: (Litter weight of live pups at PND 1) / (Live litter size on PND 1) to obtain a mean pup weight per litter. Mean values are then adjusted for total number of live pups at PND 1.

i: Number of dead pups in dose [Number of litters with dead pups]

j: Dams that had differing total counts on PND 1 and PND 4 not included

k: Litters where the male/female pup counts were inconsistent between PND 1 and PND 4 were excluded from the male/female-specific endpoints.