

Appendix 2 for KER-3348

Table 4. Empirical evidence for Increased nipple/areola retention in males (AO-1786) by prenatal exposure to flutamide, procymidone, vinclozolin, finasteride, DEHP and DBP.

Species	Exposure window	Measurement timepoint	NOAEL (mg/kg bw/day)	LOAEL (mg/kg bw/day)	Reference
Flutamide					
Rat	GD14-18	PND13	--*	40	(You et al., 1998)
Rat	GD12-21	PND13	No	6.25	(McIntyre et al., 2001)
Rat	GD7-PND16	PND12(+/-2)	0.5	1.0	(Hass et al., 2007)
Rat	GD6-PND30	PND12	0.025	0.25	(Fussell et al., 2015)
Procymidone					
Rat	GD7-PND16	PND12	10	14.1	(Hass et al., 2007)
Rat	GD14-PND3	PND10-13	--*	100	(Wolf et al., 1999)
Vinclozolin					
Rat	GD14-49	PND13	No	200	(Wolf et al., 2000)
Rat	GD7-PND16	PND13	**	**	(Christiansen et al., 2009)
Rat	GD7-PND16	PND13	5	50	(Christiansen et al., 2009)
Rat	GD7-PND16	PND12	No	5	(Hass et al., 2007)
Rat	14DPC-PND3	PND12-13	3	12	(Hellwig et al., 2000)

Species	Exposure window	Measurement timepoint	NOAEL (mg/kg bw/day)	LOAEL (mg/kg bw/day)	Reference
Rat	14DPC-PND3	PND12-13	6	12	(Hellwig et al., 2000)
Rat	Premating, gestation, lactation	PND12	40 ppm	200 ppm	(Matsuura et al., 2005)
Rat	Weaning F1-through lactation	PND12	40 ppm	200 ppm	(Matsuura et al., 2005)
Rat	GD14-PND3	PND13	No	3.125	(Ostby et al., 1999)
Rat	Premating, gestation, lactation	PND12-13	4	20	(Schneider et al., 2011)
Rat	GD14-19	PND14	--*	200	(Wolf et al., 2004)
Finasteride					
Rat	GD7-21+PND1-16	PND13	No	0.01	(Christiansen et al., 2009)
DEHP					
Rat	GD6-PND21	PND13	135	405	(Andrade et al., 2006)
Rat	GD7-PND17	PND13	--*	750	(Borch et al., 2004)
Rat	GD7-PND16	PND12	No	10	(Christiansen et al., 2010)
Rat	GD7-PND16	PND12	100	No	(Christiansen et al., 2010)
Rat	GD7-PND16	PND13	30	No	(Christiansen et al., 2009)
Rat	GD14-PND3	PND13	--*	750	(Gray et al., 2000)

Species	Exposure window	Measurement timepoint	NOAEL (mg/kg bw/day)	LOAEL (mg/kg bw/day)	Reference
Rat	GD8-lactation day 17	PND13	100	300	(Gray et al., 2009)
Rat	GD14-18	PND14	500	No	(Howdeshell et al., 2007)
Rat	GD7-PND17	PND13	No	300	(Jarfelt et al., 2005)
Rat	GD13-21	PND13	150	No	(Martino-Andrade et al., 2009)
Rat	GD3-PND21	PND14	375	750	(Moore et al., 2001)
Rat	GD12-21	PND12-14	--*	500	(Saillenfait et al., 2009)
Rat	GD14-PND3	PND10-13	--*	750	(Wolf et al., 1999)
DBP					
Rat	GD12-PND14	PND14	--*	7600ppm	(Clewett et al., 2013)
Rat	GD14-PND3	PND10-13	--*	500	(Wolf et al., 1999)
Rat	GD13-21	PND13	100	500	(Martino-Andrade et al., 2009)
Rat	GD14-18	PND14	500	No	(Howdeshell et al., 2007)
Rat	GD12-21	PND12-14	--*	500	(Saillenfait et al., 2008)
Rat	GD12-21	PND14	50	100	(Mylchreest et al., 2000)
Rat	GD12-21	PND13	No	100	(Barlow et al., 2004)

--*One dose only

** No report of NR per dose, but instead a g.logit function of the dose response. Effect doses: 30.2

for 3 nipples and 42.5 for 6 nipples.

PND=Postnatal day, DPC=Day Post Coitum, GD=Gestational Day.