

Experiment Number: R92025B
Test Type: RACB
Route: Dosing in Feed
Species/Strain: Rat/Sprague-Dawley

PA06R: Organ Weights Summary
Test Compound: 4-Methylimidazole
CAS Number: 822-36-6

Date Report Requested: 05/17/2018
Time Report Requested: 13:06:56
Lab: RTI

C Number:	R92025B
Cage Range:	All
Date Range:	All
Reasons For Removal:	All
Removal Date Range:	All
Treatment Groups:	All
Study Gender:	Both

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F0 Males

Treatment Groups (ppm)

	0			750			2500			5000		
Terminal Body Wt.	504.5	± 4.4	(23) **	477.5	± 5.9	(23) **	459.2	± 7.5	(20) **	455.2	± 5.5	(21) **
Thyroid gland												
Absolute (g)	0.0257	± 0.0010	(23)	0.0240	± 0.0007	(22)	0.0231	± 0.0013	(20)	0.0236	± 0.0007	(21)
Relative (mg/g)	0.05	± 0.00	(23)	0.05	± 0.00	(22)	0.05	± 0.00	(20)	0.05	± 0.00	(21)
Thymus												
Absolute (g)	0.259	± 0.011	(23) **	0.221	± 0.006	(23) *	0.235	± 0.011	(20) *	0.205	± 0.009	(21) **
Relative (mg/g)	0.51	± 0.02	(23)	0.46	± 0.01	(23)	0.51	± 0.02	(20)	0.45	± 0.02	(21)
Liver												
Absolute (g)	19.18	± 0.27	(23) **	18.74	± 0.40	(23)	17.58	± 0.37	(20) *	18.18	± 0.35	(21) *
Relative (mg/g)	38.01	± 0.42	(23) *	39.21	± 0.59	(23)	38.27	± 0.45	(20)	39.92	± 0.48	(21) *
Kidneys												
Absolute (g)	3.65	± 0.04	(23) **	3.56	± 0.06	(23)	3.45	± 0.07	(20) *	3.43	± 0.05	(21) **
Relative (mg/g)	7.25	± 0.08	(23) *	7.46	± 0.11	(23)	7.50	± 0.11	(20)	7.54	± 0.08	(21)
Adrenal glands												
Absolute (g)	0.0556	± 0.0012	(23) **	0.0512	± 0.0012	(23)	0.0548	± 0.0014	(20)	0.0485	± 0.0012	(21) **
Relative (mg/g)	0.11	± 0.00	(23)	0.11	± 0.00	(23)	0.12	± 0.00	(20) *	0.11	± 0.00	(21)
R. Testis												
Absolute (g)	2.089	± 0.023	(23)	2.089	± 0.030	(23)	2.086	± 0.028	(20)	2.066	± 0.029	(21)
Relative (mg/g)	4.15	± 0.06	(23) **	4.38	± 0.06	(23) *	4.57	± 0.10	(20) **	4.55	± 0.08	(21) **
L. Testis												
Absolute (g)	2.075	± 0.026	(23)	2.074	± 0.026	(23)	2.096	± 0.026	(20)	2.062	± 0.029	(21)
Relative (mg/g)	4.12	± 0.06	(23) **	4.35	± 0.07	(23) *	4.59	± 0.10	(20) **	4.54	± 0.08	(21) **
R. Epididymis												
Absolute (g)	0.681	± 0.009	(23) **	0.686	± 0.010	(23)	0.628	± 0.009	(20) **	0.600	± 0.009	(21) **
Relative (mg/g)	1.35	± 0.02	(23)	1.44	± 0.03	(23) *	1.37	± 0.03	(20)	1.32	± 0.02	(21)
L. Epididymis												
Absolute (g)	0.704	± 0.009	(23) **	0.718	± 0.013	(23)	0.657	± 0.011	(20) **	0.635	± 0.010	(21) **
Relative (mg/g)	1.40	± 0.02	(23)	1.51	± 0.04	(23) *	1.44	± 0.03	(20)	1.40	± 0.02	(21)

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F0 Males				
	Treatment Groups (ppm)			
	0	750	2500	5000
Seminal vesicles with coagulating gland				
Absolute (g)	1.946 ± 0.053 (21) **	1.647 ± 0.045 (22) **	1.520 ± 0.045 (19) **	1.253 ± 0.042 (19) **
Relative (mg/g)	3.87 ± 0.10 (21) **	3.44 ± 0.11 (22) **	3.29 ± 0.08 (19) **	2.75 ± 0.10 (19) **
Dorso-lateral prostate gland				
Absolute (g)	0.604 ± 0.026 (23) **	0.492 ± 0.023 (23) **	0.469 ± 0.015 (20) **	0.421 ± 0.022 (21) **
Relative (mg/g)	1.20 ± 0.05 (23) **	1.03 ± 0.05 (23) **	1.02 ± 0.03 (20) **	0.93 ± 0.05 (21) **
Ventral prostate gland				
Absolute (g)	0.935 ± 0.028 (23) **	0.785 ± 0.021 (23) **	0.748 ± 0.029 (20) **	0.515 ± 0.027 (21) **
Relative (mg/g)	1.85 ± 0.05 (23) **	1.65 ± 0.05 (23) **	1.63 ± 0.06 (20) **	1.13 ± 0.06 (21) **
Levator ani-bulbocavernosus muscles (LABC)				
Absolute (g)	1.438 ± 0.028 (23) **	1.369 ± 0.023 (23)	1.265 ± 0.034 (20) **	1.236 ± 0.022 (21) **
Relative (mg/g)	2.85 ± 0.06 (23)	2.87 ± 0.05 (23)	2.76 ± 0.06 (20)	2.72 ± 0.06 (21)

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F0 Females

Treatment Groups (ppm)

	0			750			2500			5000		
Terminal Body Wt.	339.3	± 6.3	(22) **	291.7	± 2.3	(21) **	300.7	± 5.1	(19) **	275.0	± 4.1	(12) **
Thyroid gland												
Absolute (g)	0.0187	± 0.0011	(21)	0.0177	± 0.0005	(22)	0.0167	± 0.0008	(18)	0.0182	± 0.0011	(12)
Relative (mg/g)	0.05	± 0.00	(21) *	0.06	± 0.00	(21)	0.06	± 0.00	(18)	0.07	± 0.00	(12) *
Thymus												
Absolute (g)	0.201	± 0.011	(22)	0.184	± 0.007	(22)	0.172	± 0.009	(19)	0.203	± 0.015	(12)
Relative (mg/g)	0.59	± 0.03	(22)	0.63	± 0.02	(21)	0.58	± 0.03	(19)	0.74	± 0.05	(12) **
Liver												
Absolute (g)	12.99	± 0.59	(22) **	10.60	± 0.21	(22) **	11.47	± 0.43	(19) **	9.89	± 0.23	(12) **
Relative (mg/g)	38.25	± 1.59	(22)	36.36	± 0.67	(21)	37.95	± 0.90	(19)	36.06	± 1.10	(12)
Kidneys												
Absolute (g)	2.34	± 0.09	(22) **	2.24	± 0.03	(22)	2.22	± 0.03	(19)	2.02	± 0.05	(12) **
Relative (mg/g)	6.93	± 0.28	(22)	7.68	± 0.08	(21) **	7.39	± 0.09	(19)	7.34	± 0.14	(12)
Adrenal glands												
Absolute (g)	0.0614	± 0.0018	(22)	0.0660	± 0.0019	(22)	0.0623	± 0.0027	(18)	0.0576	± 0.0022	(12)
Relative (mg/g)	0.18	± 0.01	(22)	0.23	± 0.01	(21) **	0.21	± 0.01	(18)	0.21	± 0.01	(12)
R. Ovary												
Absolute (g)	0.0656	± 0.0039	(22) **	0.0482	± 0.0037	(22) *	0.0613	± 0.0036	(19) *	0.0397	± 0.0034	(12) **
Relative (mg/g)	0.19	± 0.01	(22)	0.17	± 0.01	(21)	0.20	± 0.01	(19)	0.15	± 0.01	(12) *
L. Ovary												
Absolute (g)	0.0661	± 0.0044	(22) **	0.0532	± 0.0048	(22)	0.0660	± 0.0045	(19)	0.0389	± 0.0037	(12) **
Relative (mg/g)	0.20	± 0.01	(22)	0.18	± 0.02	(21)	0.22	± 0.01	(19)	0.14	± 0.01	(12) *

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F1 Males: F1 NonParental Male

	Treatment Groups (ppm)								
	0			750			2500		
Terminal Body Wt.	395.0	± 5.7	(49) **	383.4	± 4.9	(56)	338.9	± 4.4	(20) **
Thyroid gland									
Absolute (g)	0.0164	± 0.0007	(49)	0.0153	± 0.0004	(56)	0.0163	± 0.0005	(20)
Relative (mg/g)	0.04	± 0.00	(49) **	0.04	± 0.00	(56)	0.05	± 0.00	(20) **
Thymus									
Absolute (g)	0.512	± 0.012	(49)	0.485	± 0.017	(56)	0.479	± 0.025	(20)
Relative (mg/g)	1.30	± 0.03	(49)	1.27	± 0.04	(56)	1.41	± 0.06	(20)
Liver									
Absolute (g)	17.81	± 0.29	(49) **	17.76	± 0.33	(56)	15.39	± 0.38	(20) **
Relative (mg/g)	45.06	± 0.34	(49)	46.26	± 0.40	(56) *	45.33	± 0.59	(20)
Kidneys									
Absolute (g)	3.25	± 0.06	(49) **	3.24	± 0.06	(56)	2.88	± 0.07	(20) **
Relative (mg/g)	8.23	± 0.08	(49)	8.44	± 0.10	(56)	8.48	± 0.14	(20)
Adrenal glands									
Absolute (g)	0.0573	± 0.0016	(49) **	0.0548	± 0.0012	(56)	0.0466	± 0.0019	(20) **
Relative (mg/g)	0.14	± 0.00	(49)	0.14	± 0.00	(56)	0.14	± 0.00	(20)
R. Testis									
Absolute (g)	1.928	± 0.029	(49)	1.928	± 0.023	(56)	1.897	± 0.047	(20)
Relative (mg/g)	4.89	± 0.07	(49) **	5.04	± 0.07	(56)	5.62	± 0.17	(20) **
L. Testis									
Absolute (g)	1.931	± 0.029	(49)	1.911	± 0.023	(56)	1.884	± 0.038	(20)
Relative (mg/g)	4.90	± 0.06	(49) **	4.99	± 0.06	(56)	5.57	± 0.13	(20) **
R. Epididymis									
Absolute (g)	0.571	± 0.009	(49) **	0.564	± 0.008	(56)	0.513	± 0.012	(20) **
Relative (mg/g)	1.45	± 0.02	(49)	1.47	± 0.02	(56)	1.52	± 0.05	(20)
L. Epididymis									
Absolute (g)	0.579	± 0.010	(49) **	0.561	± 0.007	(56)	0.521	± 0.012	(20) **
Relative (mg/g)	1.47	± 0.02	(49) *	1.47	± 0.01	(56)	1.54	± 0.04	(20) *

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F1 Males: F1 NonParental Male

	Treatment Groups (ppm)		
	0	750	2500
Seminal vesicles with coagulating gland			
Absolute (g)	1.304 ± 0.032 (48) **	1.186 ± 0.028 (55) *	1.016 ± 0.035 (20) **
Relative (mg/g)	3.31 ± 0.06 (48) *	3.10 ± 0.07 (55)	3.00 ± 0.09 (20) *
Dorso-lateral prostate gland			
Absolute (g)	0.402 ± 0.011 (49) **	0.382 ± 0.012 (56)	0.330 ± 0.013 (20) **
Relative (mg/g)	1.02 ± 0.03 (49)	1.00 ± 0.03 (56)	0.97 ± 0.03 (20)
Ventral prostate gland			
Absolute (g)	0.561 ± 0.020 (49) **	0.446 ± 0.014 (56) **	0.355 ± 0.021 (20) **
Relative (mg/g)	1.42 ± 0.05 (49) **	1.16 ± 0.03 (56) **	1.05 ± 0.06 (20) **
Levator ani-bulbocavernosus muscles (LABC)			
Absolute (g)	1.161 ± 0.027 (49) *	1.095 ± 0.024 (56)	1.052 ± 0.035 (20)
Relative (mg/g)	2.94 ± 0.05 (49)	2.86 ± 0.06 (56)	3.11 ± 0.11 (20)

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F1 Males: F1 Parental Males

	Treatment Groups (ppm)								
	0			750			2500		
Terminal Body Wt.	497.3	± 7.7	(40) **	486.8	± 6.7	(44)	443.8	± 10.3	(39) **
Thyroid gland									
Absolute (g)	0.0209 ± 0.0006 (39)			0.0206 ± 0.0006 (43)			0.0218 ± 0.0010 (39)		
Relative (mg/g)	0.04 ± 0.00 (39) **			0.04 ± 0.00 (43)			0.05 ± 0.00 (39) **		
Thymus									
Absolute (g)	0.295 ± 0.011 (40) *			0.269 ± 0.009 (44)			0.252 ± 0.012 (39) *		
Relative (mg/g)	0.59 ± 0.02 (40)			0.55 ± 0.02 (44)			0.57 ± 0.03 (39)		
Liver									
Absolute (g)	20.84 ± 0.46 (40) **			20.89 ± 0.47 (44)			18.30 ± 0.62 (39) **		
Relative (mg/g)	41.85 ± 0.59 (40)			42.94 ± 0.86 (44)			41.15 ± 0.72 (39)		
Kidneys									
Absolute (g)	4.02 ± 0.09 (40) **			4.08 ± 0.08 (44)			3.72 ± 0.08 (39) *		
Relative (mg/g)	8.08 ± 0.13 (40)			8.40 ± 0.14 (44)			8.40 ± 0.11 (39)		
Adrenal glands									
Absolute (g)	0.0635 ± 0.0013 (40) **			0.0613 ± 0.0013 (44)			0.0576 ± 0.0019 (39) *		
Relative (mg/g)	0.13 ± 0.00 (40)			0.13 ± 0.00 (44)			0.13 ± 0.00 (39)		
R. Testis									
Absolute (g)	2.095 ± 0.036 (40)			2.121 ± 0.026 (44)			2.182 ± 0.049 (39)		
Relative (mg/g)	4.22 ± 0.07 (40) **			4.37 ± 0.07 (44)			4.94 ± 0.10 (39) **		
L. Testis									
Absolute (g)	2.090 ± 0.038 (40)			2.103 ± 0.030 (44)			2.150 ± 0.049 (39)		
Relative (mg/g)	4.21 ± 0.07 (40) **			4.34 ± 0.08 (44)			4.87 ± 0.12 (39) **		
R. Epididymis									
Absolute (g)	0.697 ± 0.009 (40) **			0.697 ± 0.009 (44)			0.651 ± 0.011 (39) **		
Relative (mg/g)	1.41 ± 0.02 (40) *			1.44 ± 0.02 (44)			1.47 ± 0.02 (39)		
L. Epididymis									
Absolute (g)	0.697 ± 0.013 (40) **			0.701 ± 0.009 (44)			0.648 ± 0.012 (39) **		
Relative (mg/g)	1.41 ± 0.02 (40)			1.45 ± 0.02 (44)			1.47 ± 0.02 (39)		

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F1 Males: F1 Parental Males

	Treatment Groups (ppm)		
	0	750	2500
Seminal vesicles with coagulating gland			
Absolute (g)	1.761 ± 0.037 (40) **	1.688 ± 0.041 (44)	1.457 ± 0.051 (39) **
Relative (mg/g)	3.56 ± 0.08 (40)	3.48 ± 0.09 (44)	3.29 ± 0.10 (39)
Dorso-lateral prostate gland			
Absolute (g)	0.539 ± 0.016 (40) **	0.475 ± 0.019 (44) *	0.449 ± 0.019 (39) **
Relative (mg/g)	1.08 ± 0.03 (40)	0.98 ± 0.04 (44)	1.01 ± 0.03 (39)
Ventral prostate gland			
Absolute (g)	0.825 ± 0.022 (40) **	0.796 ± 0.027 (44)	0.591 ± 0.018 (39) **
Relative (mg/g)	1.67 ± 0.05 (40) **	1.64 ± 0.06 (44)	1.34 ± 0.05 (39) **
Levator ani-bulbocavernosus muscles (LABC)			
Absolute (g)	1.341 ± 0.029 (40) *	1.301 ± 0.034 (44)	1.222 ± 0.041 (39)
Relative (mg/g)	2.70 ± 0.05 (40)	2.68 ± 0.06 (44)	2.75 ± 0.07 (39)

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F1 Females: F1 NonParent Female

	Treatment Groups (ppm)								
	0			750			2500		
Terminal Body Wt.	242.6	± 3.9	(47) **	233.8	± 2.7	(58)	217.8	± 4.4	(27) **
Thyroid gland									
Absolute (g)	0.0145	± 0.0006	(47)	0.0148	± 0.0007	(58)	0.0134	± 0.0007	(25)
Relative (mg/g)	0.06	± 0.00	(47)	0.06	± 0.00	(58)	0.06	± 0.00	(25)
Thymus									
Absolute (g)	0.344	± 0.013	(47)	0.327	± 0.008	(58)	0.341	± 0.017	(27)
Relative (mg/g)	1.42	± 0.04	(47) *	1.40	± 0.04	(58)	1.56	± 0.06	(27)
Liver									
Absolute (g)	9.86	± 0.21	(47)	9.95	± 0.18	(58)	9.44	± 0.26	(27)
Relative (mg/g)	40.63	± 0.47	(47) **	42.55	± 0.54	(58) **	43.30	± 0.53	(27) **
Kidneys									
Absolute (g)	1.86	± 0.03	(47)	1.93	± 0.03	(58)	1.81	± 0.04	(27)
Relative (mg/g)	7.67	± 0.08	(47) **	8.26	± 0.10	(58) **	8.33	± 0.12	(27) **
Adrenal glands									
Absolute (g)	0.0691	± 0.0014	(47)	0.0700	± 0.0014	(58)	0.0683	± 0.0020	(27)
Relative (mg/g)	0.29	± 0.01	(47) *	0.30	± 0.01	(58)	0.31	± 0.01	(27) *
R. Ovary									
Absolute (g)	0.0552	± 0.0014	(46) *	0.0528	± 0.0022	(58)	0.0478	± 0.0016	(27) *
Relative (mg/g)	0.23	± 0.01	(46)	0.22	± 0.01	(58)	0.22	± 0.01	(27)
L. Ovary									
Absolute (g)	0.0585	± 0.0015	(46) *	0.0548	± 0.0024	(58)	0.0511	± 0.0027	(27)
Relative (mg/g)	0.24	± 0.01	(46)	0.23	± 0.01	(58)	0.23	± 0.01	(27)

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F1 Females: F1 Parental Females

	Treatment Groups (ppm)								
	0			750			2500		
Terminal Body Wt.	349.9	± 6.5	(28) **	325.5	± 4.1	(34) **	311.9	± 3.8	(23) **
Thyroid gland									
Absolute (g)	0.0193	± 0.0007	(27) **	0.0175	± 0.0003	(34) **	0.0165	± 0.0007	(23) **
Relative (mg/g)	0.06	± 0.00	(27)	0.05	± 0.00	(34)	0.05	± 0.00	(23)
Thymus									
Absolute (g)	0.213	± 0.010	(28)	0.189	± 0.009	(34)	0.193	± 0.007	(23)
Relative (mg/g)	0.61	± 0.02	(28)	0.58	± 0.03	(34)	0.62	± 0.02	(23)
Liver									
Absolute (g)	16.16	± 0.33	(28)	17.38	± 0.49	(34)	16.64	± 0.67	(23)
Relative (mg/g)	46.34	± 0.85	(28) *	53.41	± 1.35	(34) **	53.36	± 2.01	(23) **
Kidneys									
Absolute (g)	2.81	± 0.05	(28)	2.92	± 0.05	(34)	2.72	± 0.06	(23)
Relative (mg/g)	8.07	± 0.13	(28)	8.97	± 0.16	(34) **	8.73	± 0.17	(23) *
Adrenal glands									
Absolute (g)	0.0709	± 0.0017	(28)	0.0682	± 0.0015	(34)	0.0718	± 0.0029	(23)
Relative (mg/g)	0.20	± 0.01	(28) **	0.21	± 0.00	(34)	0.23	± 0.01	(23) **
R. Ovary									
Absolute (g)	0.0843	± 0.0043	(28)	0.0736	± 0.0032	(34)	0.0708	± 0.0033	(23)
Relative (mg/g)	0.24	± 0.01	(28)	0.23	± 0.01	(34)	0.23	± 0.01	(23)
L. Ovary									
Absolute (g)	0.0834	± 0.0041	(28) *	0.0754	± 0.0047	(34)	0.0711	± 0.0024	(23)
Relative (mg/g)	0.24	± 0.01	(28)	0.23	± 0.01	(34)	0.23	± 0.01	(23)

Experiment Number: R92025B
Test Type: RACB
Route: Dosing in Feed
Species/Strain: Rat/Sprague-Dawley

PA06R: Organ Weights Summary
Test Compound: 4-Methylimidazole
CAS Number: 822-36-6

Date Report Requested: 05/17/2018
Time Report Requested: 13:06:56
Lab: RTI

LEGEND

Data are displayed as mean \pm SEM (N) unless otherwise noted.

Reported means and standard errors for the F1 animals are the means and standard errors of the litter means, and the reported N values are the number of litters used in the calculation.

Relative organ weights (organ-weight-to-body-weight ratios) are given as mg organ weight/g body weight

Statistical analysis performed by Jonckheere (trend) and Williams or Dunnett (pairwise) tests unless otherwise noted.

Statistical analysis for F1 animals was performed by Generalized Linear Mixed Models, with dam ID as random effect, and using Dunnett-Hsu adjustment, for both trend and pairwise analyses.

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 $P \leq 0.01$

Non-pregnant females in the parental female cohort were excluded from the analysis.

**** END OF REPORT ****