

### 1020\_L\_MWNT Tissue Burden Analysis in Female HD Rats at 0 days

Parameter <sup>a</sup>	0 mg/m3	0.1 mg/m3	0.3 mg/m3	1.0 mg/m3	3.0 mg/m3	10.0 mg/m3	Trend <sup>b</sup>
Sample Weight (g) <sup>b</sup>	1.230 ± 0.033 [10]	1.222 ± 0.050 [10]	1.200 ± 0.025 [10]	1.212 ± 0.035 [10]	1.378 ± 0.041 [10]**	1.477 ± 0.040 [10]**	<.001
Sample Conc (ug Ni/g) <sup>c</sup>	0.002 ± 0.000 [10]	0.026 ± 0.001 [10]**	0.101 ± 0.009 [10]**	0.374 ± 0.013 [10]**	0.885 ± 0.028 [10]**	2.351 ± 0.079 [10]**	<.001
Total Lung Burden (ug Ni/lung) <sup>c</sup>	0.002 ± 0.000 [10]	0.032 ± 0.001 [10]**	0.120 ± 0.009 [10]**	0.452 ± 0.018 [10]**	1.212 ± 0.027 [10]**	3.468 ± 0.138 [10]**	<.001
Total Lung Burden (ug L-MWNT-1020/lung) <sup>c</sup>	0.382 ± 0.092 [10]	6.177 ± 0.219 [10]**	23.12 ± 1.778 [10]**	86.94 ± 3.465 [10]**	233.0 ± 5.252 [10]**	666.9 ± 26.61 [10]**	<.001
Norm. Lung Burden (ug LMWNT/lung per mg/m3) <sup>c</sup>		61.77 ± 2.19 [10]	77.07 ± 5.93 [10]	86.94 ± 3.46 [10]	77.67 ± 1.75 [10]	66.69 ± 2.66 [10]	

a: All values shown as mean ± standard error [number]

b: For sample weight, each dose was compared to the control with Williams' test when a trend was present (P <0.01 from Jonckheere's trend test).

Otherwise, Dunnett's test was applied. [\* = P<0.05, \*\* = P<0.01]

c: For endpoints other than sample weight, each dose was compared to the control with Shirley's test when a trend was present (P <0.01 from Jonckheere's trend test).

Otherwise, Dunn's test was applied. [\* = P<0.05, \*\* = P<0.01]

### 1020\_L\_MWNT Tissue Burden Analysis in Female HD Rats at 14 days

Parameter <sup>a</sup>	0 mg/m3	0.1 mg/m3	0.3 mg/m3	1.0 mg/m3	3.0 mg/m3	10.0 mg/m3	Trend <sup>b</sup>
Sample Weight (g) <sup>b</sup>	1.452 ± 0.070 [5]	1.298 ± 0.107 [5]	1.330 ± 0.080 [5]	1.421 ± 0.088 [5]	1.428 ± 0.066 [5]	1.571 ± 0.041 [5]	0.067
Sample Conc (ug Ni/g) <sup>c</sup>	0.004 ± 0.002 [5]	0.022 ± 0.002 [5]**	0.065 ± 0.007 [5]**	0.285 ± 0.030 [5]**	0.893 ± 0.048 [5]**	2.117 ± 0.069 [5]**	<.001
Total Lung Burden (ug Ni/lung) <sup>c</sup>	0.006 ± 0.002 [5]	0.028 ± 0.002 [5]**	0.085 ± 0.009 [5]**	0.395 ± 0.022 [5]**	1.276 ± 0.087 [5]**	3.335 ± 0.189 [5]**	<.001
Total Lung Burden (ug L-MWNT-1020/lung) <sup>c</sup>	1.11 ± 0.37 [5]	5.34 ± 0.44 [5]**	16.42 ± 1.70 [5]**	76.00 ± 4.27 [5]**	245.33 ± 16.71 [5]**	641.37 ± 36.38 [5]**	<.001
Norm. Lung Burden (ug LMWNT/lung per mg/m3) <sup>c</sup>		53.44 ± 4.41 [5]	54.74 ± 5.65 [5]	76.00 ± 4.27 [5]	81.78 ± 5.57 [5]	64.14 ± 3.64 [5]	

a: All values shown as mean ± standard error [number]

b: For sample weight, each dose was compared to the control with Williams' test when a trend was present (P <0.01 from Jonckheere's trend test).

Otherwise, Dunnett's test was applied. [\* = P<0.05, \*\* = P<0.01]

c: For endpoints other than sample weight, each dose was compared to the control with Shirley's test when a trend was present (P <0.01 from Jonckheere's trend test).

Otherwise, Dunn's test was applied. [\* = P<0.05, \*\* = P<0.01]

### 1020\_L\_MWNT Tissue Burden Analysis in Female HD Rats at 42 days

Parameter <sup>a</sup>	0 mg/m3	0.1 mg/m3	0.3 mg/m3	1.0 mg/m3	3.0 mg/m3	10.0 mg/m3	Trend <sup>b</sup>
Sample Weight (g) <sup>b</sup>	1.359 ± 0.062 [5]	1.324 ± 0.056 [5]	1.295 ± 0.034 [5]	1.371 ± 0.037 [5]	1.574 ± 0.087 [5]*	1.509 ± 0.037 [5]*	0.004
Sample Conc (ug Ni/g) <sup>c</sup>	0.002 ± 0.001 [5]	0.015 ± 0.003 [5]**	0.048 ± 0.005 [5]**	0.268 ± 0.022 [5]**	0.827 ± 0.101 [5]**	2.147 ± 0.065 [5]**	<.001
Total Lung Burden (ug Ni/lung) <sup>c</sup>	0.003 ± 0.001 [5]	0.020 ± 0.004 [5]**	0.062 ± 0.006 [5]**	0.367 ± 0.034 [5]**	1.301 ± 0.178 [5]**	3.240 ± 0.139 [5]**	<.001
Total Lung Burden (ug L-MWNT-1020/lung) <sup>c</sup>	0.592 ± 0.243 [5]	3.756 ± 0.772 [5]**	11.92 ± 1.146 [5]**	70.65 ± 6.536 [5]**	250.2 ± 34.24 [5]**	623.1 ± 26.82 [5]**	<.001
Norm. Lung Burden (ug LMWNT/lung per mg/m3) <sup>c</sup>		37.56 ± 7.72 [5]	39.72 ± 3.82 [5]	70.65 ± 6.54 [5]	83.41 ± 11.41 [5]	62.31 ± 2.68 [5]	

a: All values shown as mean ± standard error [number]

b: For sample weight, each dose was compared to the control with Williams' test when a trend was present (P <0.01 from Jonckheere's trend test).

Otherwise, Dunnett's test was applied. [\* = P<0.05, \*\* = P<0.01]

c: For endpoints other than sample weight, each dose was compared to the control with Shirley's test when a trend was present (P <0.01 from Jonckheere's trend test).

Otherwise, Dunn's test was applied. [\* = P<0.05, \*\* = P<0.01]

### 1020\_L\_MWNT Tissue Burden Analysis in Female HD Rats at 126 days

Parameter <sup>a</sup>	0 mg/m3	0.1 mg/m3	0.3 mg/m3	1.0 mg/m3	3.0 mg/m3	10.0 mg/m3	Trend <sup>b</sup>
Sample Weight (g) <sup>b</sup>	1.404 ± 0.058 [5]	1.457 ± 0.071 [5]	1.468 ± 0.082 [5]	1.397 ± 0.088 [5]	1.665 ± 0.152 [5]	1.715 ± 0.053 [5]	0.011
Sample Conc (ug Ni/g) <sup>c</sup>	0.001 ± 0.000 [5]	0.005 ± 0.000 [5]**	0.014 ± 0.002 [5]**	0.105 ± 0.015 [5]**	0.489 ± 0.083 [5]**	1.776 ± 0.137 [5]**	<.001
Total Lung Burden (ug Ni/lung) <sup>c</sup>	0.002 ± 0.001 [5]	0.007 ± 0.000 [5]**	0.020 ± 0.002 [5]**	0.149 ± 0.029 [5]**	0.774 ± 0.094 [5]**	3.062 ± 0.291 [5]**	<.001
Total Lung Burden (ug L-MWNT-1020/lung) <sup>c</sup>	0.320 ± 0.133 [5]	1.320 ± 0.084 [5]**	3.886 ± 0.437 [5]**	28.59 ± 5.605 [5]**	148.8 ± 18.10 [5]**	588.8 ± 55.90 [5]**	<.001
Norm. Lung Burden (ug LMWNT/lung per mg/m3) <sup>c</sup>		13.20 ± 0.84 [5]	12.95 ± 1.46 [5]	28.59 ± 5.61 [5]	49.60 ± 6.03 [5]	58.88 ± 5.59 [5]	

a: All values shown as mean ± standard error [number]

b: For sample weight, each dose was compared to the control with Williams' test when a trend was present (P <0.01 from Jonckheere's trend test).

Otherwise, Dunnett's test was applied. [\* = P<0.05, \*\* = P<0.01]

c: For endpoints other than sample weight, each dose was compared to the control with Shirley's test when a trend was present (P <0.01 from Jonckheere's trend test).

Otherwise, Dunn's test was applied. [\* = P<0.05, \*\* = P<0.01]