### Male Hematology Data Summary

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| **Endpoint** | 0.0 mg/kgN = 6 | 0.15 mg/kgN = 2 | 0.5 mg/kgN = 4 | 1.4 mg/kgN = 4 | 4.0 mg/kgN = 5 | 12.0 mg/kgN = 3 | 37.0 mg/kgN = 2 | 111.0 mg/kgN = 4 | BMD1Std (mg/kg) | BMDL1Std (mg/kg) |
| Erythrocyte Count (M/uL) | 8.53 ± 0.28\* | 8.40 ± 0.29 | 8.79 ± 0.33 | 8.61 ± 0.21 | 8.23 ± 0.10 | 8.44 ± 0.26 | 7.99 ± 0.00 | 8.03 ± 0.14 | NVM | NVM |
| Mean Cell Hemoglobin (pg) | 18.0 ± 0.2 | 18.1 ± 0.4 | 17.9 ± 0.4 | 19.3 ± 0.1 | 19.5 ± 0.4\* | 18.6 ± 0.5 | 18.3 ± 0.1 | 18.5 ± 0.2 | 1.32 | 0.761 |
| Mean Cell HGB Concentration (g/dL) | 29.3 ± 0.3 | 29.8 ± 0.1 | 29.4 ± 0.3 | 31.4 ± 0.4 | 31.8 ± 0.5\* | 30.4 ± 0.9 | 29.3 ± 0.6 | 29.5 ± 0.2 | NVM | NVM |
| Neutrophil Count (K/uL) | 0.84 ± 0.12\*\* | 1.08 ± 0.22 | 0.85 ± 0.06 | 1.34 ± 0.16 | 1.54 ± 0.21 | 1.25 ± 0.19 | 1.23 ± 0.01 | 1.24 ± 0.16\* | 0.811 | 0.305 |

### Female Hematology Data Summary

|  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| **Endpoint** | 0.0 mg/kgN = 8 | 0.15 mg/kgN = 5 | 0.5 mg/kgN = 5 | 1.4 mg/kgN = 3 | 4.0 mg/kgN = 3-4 | 12.0 mg/kgN = 4 | 37.0 mg/kgN = 4 | 111.0 mg/kgN = 5 | BMD1Std (mg/kg) | BMDL1Std (mg/kg) |
| Manual hematocrit (%) | 44.6 ± 0.5\*\* | 47.4 ± 2.7 | 45.0 ± 0.7 | 44.3 ± 0.9 | 44.3 ± 0.5 | 43.5 ± 0.6 | 43.0 ± 0.0\* | 42.6 ± 0.7\* | NVM | NVM |
| Erythrocyte Count (M/uL) | 7.68 ± 0.09\* | 8.27 ± 0.48 | 7.85 ± 0.19 | 7.80 ± 0.07 | 7.46 ± 0.19 | 7.62 ± 0.22 | 7.63 ± 0.14 | 7.38 ± 0.13 | NVM | NVM |
| Mean Cell Hemoglobin (pg) | 19.4 ± 0.2 | 18.2 ± 0.1\* | 18.5 ± 0.2 | 18.2 ± 0.4 | 19.7 ± 0.1 | 18.5 ± 0.2 | 18.6 ± 0.6 | 19.7 ± 0.2 | <0.05 | NR |
| Mean Cell HGB Concentration (g/dL) | 32.2 ± 0.3 | 30.1 ± 0.3\* | 30.4 ± 0.2 | 30.3 ± 0.8 | 32.5 ± 0.1 | 30.3 ± 0.2 | 30.5 ± 0.7 | 32.3 ± 0.3 | <0.05 | NR |
| Platelet count (K/uL) | 802 ± 42 | 967 ± 76 | 933 ± 83 | 1038 ± 61 | 869 ± 77 | 958 ± 77 | 1073 ± 46\* | 921 ± 26 | 30.346 | 17.239 |
| Eosinophil count (K/uL) | 0.10 ± 0.01\* | 0.07 ± 0.01 | 0.09 ± 0.00 | 0.08 ± 0.00 | 0.10 ± 0.02 | 0.10 ± 0.04 | 0.07 ± 0.01 | 0.06 ± 0.01 | NVM | NVM |

Values given as mean ± SEM (N) of animals that survived to study termination.

Statistical analysis were performed by Jonckheere (trend) and Shirley or Dunn (pairwise) tests.

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 ≤ 0.05

\*\* Statistically significant at p ≤ 0.01

BMD1Std and BMDL1Std: Benchmark response (BMR) set at 1 standard deviation from the mean.

NVM = no viable model

NR = BMDL1Std is not reportable because the BMD is below the lower limit of extrapolation (less than 1/3 of the lowest tested dose in this study.).

<0.05 = A best-fit model was identified and a BMD was estimated that was less than 1/3 of the lowest tested dose in this study.