**Proportional Response Addition**

**Supplemental Materials**

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**Table S1. Prop-RA Results for Binary Mixture of CHCl3 and BDCM1,2**

**0.5:0.5 CHCl3:BDCM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **10** | **4.66** | **0.48** | **10** | **4.72** | **0.40** | **11** | **4.98** | **0.49** | **-0.28717** | **-0.74** | **0.16** |
| **Log(ALT)** | **10** | **1.41** | **0.10** | **10** | **1.40** | **0.11** | **11** | **1.40** | **0.11** | **0.007** | **-0.10** | **0.11** |
| **Log(AST)** | **10** | **1.61** | **0.09** | **10** | **1.65** | **0.11** | **11** | **1.63** | **0.11** | **0.0025** | **-0.10** | **0.10** |
| **Log(SDH)** | **10** | **1.16** | **0.04** | **10** | **1.08** | **0.16** | **11** | **1.16** | **0.06** | **-0.0395** | **-0.14** | **0.06** |

**0.5:0.5 CHCl3:BDCM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **9** | **5.40** | **0.31** | **7** | **6.15** | **0.83** | **10** | **5.76** | **0.37** | **0.018606** | **-0.53** | **0.56** |
| **Log(ALT)** | **9** | **1.62** | **0.13** | **7** | **1.91** | **0.26** | **11** | **1.63** | **0.14** | **0.13** | **-0.05** | **0.31** |
| **Log(AST)** | **9** | **1.64** | **0.08** | **7** | **1.78** | **0.17** | **11** | **1.63** | **0.07** | **0.083** | **-0.02** | **0.19** |
| **Log(SDH)** | **9** | **1.33** | **0.07** | **7** | **1.48** | **0.23** | **11** | **1.37** | **0.11** | **0.0325** | **-0.11** | **0.18** |

**0.5:0.5 CHCl3:BDCM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **7** | **7.72** | **0.95** | **7** | **8.16** | **1.59** | **9** | **8.42** | **0.93** | **-0.47626** | **-1.80** | **0.85** |
| **Log(ALT)** | **7** | **2.66** | **0.42** | **7** | **2.91** | **0.27** | **9** | **2.80** | **0.48** | **-0.014** | **-0.48** | **0.45** |
| **Log(AST)** | **7** | **2.42** | **0.46** | **7** | **2.85** | **0.35** | **9** | **2.67** | **0.54** | **-0.0365** | **-0.56** | **0.49** |
| **Log(SDH)** | **7** | **1.94** | **0.28** | **7** | **1.95** | **0.41** | **9** | **2.05** | **0.37** | **-0.1045** | **-0.51** | **0.30** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S1 (cont). Prop-RA Results for Binary Mixture of CHCl3 and BDCM1,2**

**0.730:0.270 CHCl3:BDCM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **9** | **5.40** | **0.31** | **7** | **6.15** | **0.83** | **11** | **5.68** | **0.44** | **-0.07441** | **-0.63** | **0.48** |
| **Log(ALT)** | **9** | **1.62** | **0.13** | **7** | **1.91** | **0.26** | **11** | **1.78** | **0.17** | **-0.0847** | **-0.28** | **0.11** |
| **Log(AST)** | **9** | **1.64** | **0.08** | **7** | **1.78** | **0.17** | **11** | **1.81** | **0.07** | **-0.13112** | **-0.24** | **-0.02** |
| **Log(SDH)** | **9** | **1.33** | **0.07** | **7** | **1.48** | **0.23** | **11** | **1.40** | **0.15** | **-0.03123** | **-0.20** | **0.13** |

**0.730:0.270 CHCl3:BDCM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **7** | **7.72** | **0.95** | **7** | **8.16** | **1.59** | **11** | **8.11** | **0.85** | **-0.26488** | **-1.51** | **0.98** |
| **Log(ALT)** | **7** | **2.66** | **0.42** | **7** | **2.91** | **0.27** | **11** | **2.91** | **0.40** | **-0.18388** | **-0.60** | **0.23** |
| **Log(AST)** | **7** | **2.42** | **0.46** | **7** | **2.85** | **0.35** | **11** | **2.72** | **0.39** | **-0.18571** | **-0.63** | **0.26** |
| **Log(SDH)** | **7** | **1.94** | **0.28** | **7** | **1.95** | **0.41** | **11** | **2.07** | **0.27** | **-0.12465** | **-0.48** | **0.23** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S2. Prop-RA Results for Binary Mixture of CHCl3 and BDCM-rep1,2**

**0.5:0.5 CHCl3:BDCM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **8** | **5.18** | **0.39** | **10** | **4.91** | **1.59** | **10** | **5.23** | **0.37** | **-0.1796** | **-1.21** | **0.85** |
| **Log(ALT)** | **8** | **1.38** | **0.17** | **10** | **1.35** | **0.14** | **10** | **1.39** | **0.11** | **-0.027** | **-0.17** | **0.12** |
| **Log(AST)** | **8** | **1.60** | **0.08** | **10** | **1.61** | **0.11** | **10** | **1.58** | **0.08** | **0.021** | **-0.07** | **0.12** |
| **Log(SDH)** | **8** | **1.13** | **0.10** | **10** | **1.09** | **0.07** | **10** | **1.12** | **0.07** | **-0.0095** | **-0.09** | **0.07** |

**0.5:0.5 CHCl3:BDCM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **9** | **6.34** | **0.54** | **8** | **6.22** | **0.50** | **10** | **6.30** | **0.63** | **-0.01859** | **-0.61** | **0.57** |
| **Log(ALT)** | **9** | **1.70** | **0.17** | **7** | **1.84** | **0.22** | **9** | **1.72** | **0.26** | **0.047** | **-0.19** | **0.29** |
| **Log(AST)** | **9** | **1.66** | **0.05** | **7** | **1.76** | **0.13** | **9** | **1.73** | **0.14** | **-0.0285** | **-0.15** | **0.09** |
| **Log(SDH)** | **9** | **1.42** | **0.16** | **7** | **1.61** | **0.18** | **9** | **1.48** | **0.19** | **0.041** | **-0.15** | **0.24** |

**0.5:0.5 CHCl3:BDCM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **5** | **8.29** | **1.78** | **7** | **8.41** | **0.55** | **9** | **9.22** | **1.77** | **-0.86813** | **-2.62** | **0.88** |
| **Log(ALT)** | **3** | **2.96** | **0.42** | **7** | **2.78** | **0.18** | **10** | **3.02** | **0.42** | **-0.154** | **-0.59** | **0.29** |
| **Log(AST)** | **3** | **2.65** | **0.46** | **7** | **2.65** | **0.16** | **10** | **2.81** | **0.50** | **-0.16** | **-0.67** | **0.35** |
| **Log(SDH)** | **3** | **2.35** | **0.44** | **7** | **2.23** | **0.24** | **10** | **2.34** | **0.42** | **-0.0505** | **-0.52** | **0.41** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S2 (cont). Prop-RA Results for Binary Mixture of CHCl3 and BDCM-rep1,2**

**0.730:0.270 CHCl3:BDCM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **9** | **6.34** | **0.54** | **8** | **6.22** | **0.50** | **10** | **6.40** | **0.72** | **-0.0947** | **-0.74** | **0.55** |
| **Log(ALT)** | **9** | **1.70** | **0.17** | **7** | **1.84** | **0.22** | **10** | **1.63** | **0.13** | **0.10842** | **-0.07** | **0.29** |
| **Log(AST)** | **9** | **1.66** | **0.05** | **7** | **1.76** | **0.13** | **10** | **1.65** | **0.09** | **0.02827** | **-0.07** | **0.13** |
| **Log(SDH)** | **9** | **1.42** | **0.16** | **7** | **1.61** | **0.18** | **10** | **1.36** | **0.13** | **0.1133** | **-0.05** | **0.28** |

**0.730:0.270 CHCl3:BDCM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **5** | **8.29** | **1.78** | **7** | **8.41** | **0.55** | **7** | **8.43** | **0.98** | **-0.11181** | **-1.66** | **1.43** |
| **Log(ALT)** | **3** | **2.96** | **0.42** | **7** | **2.78** | **0.18** | **7** | **2.74** | **0.46** | **0.1634** | **-0.40** | **0.73** |
| **Log(AST)** | **3** | **2.65** | **0.46** | **7** | **2.65** | **0.16** | **7** | **2.58** | **0.52** | **0.067** | **-0.56** | **0.69** |
| **Log(SDH)** | **3** | **2.35** | **0.44** | **7** | **2.23** | **0.24** | **7** | **2.21** | **0.42** | **0.10433** | **-0.46** | **0.67** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S3. Prop-RA Results for Binary Mixture of CHCl3 and CHBr31,2**

**0.5:0.5 CHCl3:CHBr3**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **10** | **4.97** | **0.62** | **9** | **5.06** | **0.33** | **11** | **4.75** | **0.40** | **0.268194** | **-0.19** | **0.73** |
| **Log(ALT)** | **10** | **1.41** | **0.14** | **9** | **1.46** | **0.08** | **10** | **1.44** | **0.14** | **-0.0045** | **-0.13** | **0.12** |
| **Log(AST)** | **10** | **1.63** | **0.10** | **9** | **1.64** | **0.11** | **10** | **1.62** | **0.08** | **0.012** | **-0.09** | **0.11** |
| **Log(SDH)** | **9** | **1.28** | **0.08** | **9** | **1.30** | **0.07** | **10** | **1.31** | **0.08** | **-0.0265** | **-0.11** | **0.05** |

**0.5:0.5 CHCl3:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **9** | **5.70** | **0.56** | **9** | **6.42** | **0.63** | **9** | **6.08** | **0.39** | **-0.01351** | **-0.58** | **0.56** |
| **Log(ALT)** | **9** | **1.78** | **0.17** | **9** | **1.68** | **0.21** | **9** | **1.68** | **0.22** | **0.055** | **-0.16** | **0.27** |
| **Log(AST)** | **9** | **1.67** | **0.11** | **9** | **1.73** | **0.10** | **9** | **1.70** | **0.14** | **0.001** | **-0.12** | **0.12** |
| **Log(SDH)** | **9** | **1.63** | **0.13** | **9** | **1.57** | **0.15** | **9** | **1.53** | **0.08** | **0.075** | **-0.06** | **0.21** |

**0.5:0.5 CHCl3:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **8** | **7.69** | **1.14** | **8** | **7.47** | **0.70** | **4** | **7.12** | **0.29** | **0.461538** | **-0.84** | **1.76** |
| **Log(ALT)** | **8** | **2.71** | **0.39** | **8** | **2.40** | **0.20** | **4** | **2.10** | **0.24** | **0.458** | **0.02** | **0.90** |
| **Log(AST)** | **8** | **2.47** | **0.47** | **8** | **2.41** | **0.15** | **4** | **2.04** | **0.18** | **0.398** | **-0.09** | **0.89** |
| **Log(SDH)** | **8** | **2.19** | **0.35** | **8** | **1.84** | **0.06** | **4** | **1.80** | **0.06** | **0.216** | **-0.12** | **0.55** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S3 (cont). Prop-RA Results for Binary Mixture of CHCl3 and CHBr31,2**

**0.985:0.015 CHCl3:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **9** | **5.70** | **0.56** | **9** | **6.42** | **0.63** | **10** | **6.09** | **0.49** | **-0.38271** | **-1.05** | **0.28** |
| **Log(ALT)** | **9** | **1.78** | **0.17** | **9** | **1.68** | **0.21** | **10** | **1.70** | **0.22** | **0.07462** | **-0.16** | **0.31** |
| **Log(AST)** | **9** | **1.67** | **0.11** | **9** | **1.73** | **0.10** | **10** | **1.60** | **0.15** | **0.06196** | **-0.08** | **0.21** |
| **Log(SDH)** | **9** | **1.63** | **0.13** | **9** | **1.57** | **0.15** | **10** | **1.54** | **0.15** | **0.0951** | **-0.08** | **0.27** |

**0.985:0.015 CHCl3:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **8** | **7.69** | **1.14** | **8** | **7.47** | **0.70** | **6** | **7.23** | **1.77** | **0.456933** | **-1.28** | **2.19** |
| **Log(ALT)** | **8** | **2.71** | **0.39** | **8** | **2.40** | **0.20** | **5** | **2.84** | **0.30** | **-0.12974** | **-0.59** | **0.33** |
| **Log(AST)** | **8** | **2.47** | **0.47** | **8** | **2.41** | **0.15** | **5** | **2.65** | **0.41** | **-0.1869** | **-0.74** | **0.36** |
| **Log(SDH)** | **8** | **2.19** | **0.35** | **8** | **1.84** | **0.06** | **5** | **2.17** | **0.42** | **0.01781** | **-0.42** | **0.46** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S4. Prop-RA Results for Binary Mixture of BDCM and CHBr31,2**

**0.5:0.5 BDCM:CHBr3**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **9** | **4.98** | **0.32** | **11** | **5.23** | **1.29** | **10** | **5.00** | **0.65** | **0.10586** | **-0.79** | **1.00** |
| **Log(ALT)** | **9** | **1.32** | **0.15** | **11** | **1.48** | **0.27** | **10** | **1.37** | **0.13** | **0.032** | **-0.17** | **0.23** |
| **Log(AST)** | **9** | **1.53** | **0.07** | **11** | **1.65** | **0.21** | **10** | **1.67** | **0.25** | **-0.0855** | **-0.28** | **0.11** |
| **Log(SDH)** | **9** | **1.20** | **0.06** | **11** | **1.25** | **0.19** | **10** | **1.20** | **0.07** | **0.025** | **-0.10** | **0.15** |

**0.5:0.5 BDCM:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **10** | **5.99** | **0.45** | **9** | **6.39** | **0.37** | **7** | **6.87** | **0.92** | **-0.68016** | **-1.36** | **0.002** |
| **Log(ALT)** | **10** | **1.96** | **0.29** | **8** | **1.77** | **0.29** | **6** | **1.87** | **0.45** | **-0.001** | **-0.41** | **0.41** |
| **Log(AST)** | **10** | **1.79** | **0.22** | **8** | **1.75** | **0.19** | **6** | **1.86** | **0.31** | **-0.088** | **-0.38** | **0.20** |
| **Log(SDH)** | **10** | **1.73** | **0.15** | **8** | **1.54** | **0.17** | **6** | **1.56** | **0.29** | **0.0725** | **-0.17** | **0.32** |

**0.5:0.5 BDCM:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **5** | **8.34** | **1.16** | **4** | **7.69** | **1.13** | **6** | **7.69** | **0.49** | **0.323436** | **-1.05** | **1.69** |
| **Log(ALT)** | **5** | **2.84** | **0.19** | **4** | **2.44** | **0.35** | **6** | **2.37** | **0.22** | **0.2725** | **-0.10** | **0.64** |
| **Log(AST)** | **5** | **2.75** | **0.23** | **4** | **2.40** | **0.34** | **6** | **2.27** | **0.18** | **0.3** | **-0.06** | **0.66** |
| **Log(SDH)** | **5** | **2.53** | **0.33** | **4** | **2.05** | **0.46** | **6** | **1.79** | **0.09** | **0.5005** | **0.05** | **0.95** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S4 (cont). Prop-RA Results for Binary Mixture of BDCM and CHBr31,2**

**0.96:0.040 BDCM:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **10** | **5.99** | **0.45** | **9** | **6.39** | **0.37** | **8** | **5.98** | **0.33** | **0.026164** | **-0.45** | **0.50** |
| **Log(ALT)** | **10** | **1.96** | **0.29** | **8** | **1.77** | **0.29** | **8** | **2.03** | **0.35** | **-0.0766** | **-0.45** | **0.30** |
| **Log(AST)** | **10** | **1.79** | **0.22** | **8** | **1.75** | **0.19** | **8** | **1.83** | **0.23** | **-0.04944** | **-0.31** | **0.21** |
| **Log(SDH)** | **10** | **1.73** | **0.15** | **8** | **1.54** | **0.17** | **8** | **1.73** | **0.19** | **-0.0124** | **-0.22** | **0.20** |

**0.96:0.040 BDCM:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **5** | **8.34** | **1.16** | **4** | **7.69** | **1.13** | **6** | **8.75** | **0.97** | **-0.43673** | **-2.21** | **1.34** |
| **Log(ALT)** | **5** | **2.84** | **0.19** | **4** | **2.44** | **0.35** | **6** | **2.70** | **0.31** | **0.1248** | **-0.35** | **0.60** |
| **Log(AST)** | **5** | **2.75** | **0.23** | **4** | **2.40** | **0.34** | **6** | **2.51** | **0.26** | **0.219** | **-0.24** | **0.67** |
| **Log(SDH)** | **5** | **2.53** | **0.33** | **4** | **2.05** | **0.46** | **6** | **2.18** | **0.45** | **0.33252** | **-0.35** | **1.02** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S5. Prop-RA Results for Binary Mixture of BDCM and CDBM1,2**

**0.5:0.5 BDCM:CDBM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **7** | **4.69** | **0.63** | **7** | **5.24** | **0.27** | **7** | **5.05** | **0.42** | **-0.09003** | **-0.66** | **0.48** |
| **Log(ALT)** | **6** | **1.54** | **0.10** | **6** | **1.36** | **0.18** | **7** | **1.39** | **0.08** | **0.056** | **-0.10** | **0.22** |
| **Log(AST)** | **6** | **1.79** | **0.11** | **6** | **1.63** | **0.05** | **7** | **1.62** | **0.03** | **0.085** | **-0.01** | **0.18** |
| **Log(SDH)** | **6** | **1.52** | **0.13** | **6** | **1.42** | **0.08** | **7** | **1.44** | **0.06** | **0.0315** | **-0.09** | **0.15** |

**0.5:0.5 BDCM:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **6** | **5.94** | **0.44** | **7** | **6.09** | **0.48** | **2** | **6.48** | **0.24** | **-0.46903** | **-1.42** | **0.48** |
| **Log(ALT)** | **6** | **1.90** | **0.44** | **7** | **1.93** | **0.44** | **2** | **2.00** | **0.34** | **-0.085** | **-1.00** | **0.83** |
| **Log(AST)** | **6** | **1.92** | **0.29** | **7** | **1.93** | **0.23** | **2** | **1.90** | **0.31** | **0.023** | **-0.54** | **0.58** |
| **Log(SDH)** | **6** | **1.95** | **0.35** | **7** | **1.88** | **0.33** | **2** | **1.89** | **0.01** | **0.024** | **-0.66** | **0.71** |

**0.5:0.5 BDCM:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **5** | **8.78** | **0.59** | **2** | **5.86** | **0.51** | **3** | **8.05** | **1.11** | **-0.72649** | **-2.41** | **0.96** |
| **Log(ALT)** | **5** | **3.13** | **0.08** | **2** | **2.90** | **0.01** | **3** | **2.64** | **0.38** | **0.37** | **-0.09** | **0.83** |
| **Log(AST)** | **5** | **2.96** | **0.12** | **2** | **3.03** | **0.06** | **3** | **2.61** | **0.28** | **0.3885** | **-0.002** | **0.78** |
| **Log(SDH)** | **5** | **2.67** | **0.19** | **2** | **2.53** | **0.05** | **3** | **2.31** | **0.32** | **0.2885** | **-0.21** | **0.78** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S5 (cont). Prop-RA Results Binary Mixture of BDCM and CDBM1,2**

**0.706:0.294 BDCM:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **6** | **5.94** | **0.44** | **7** | **6.09** | **0.48** | **7** | **5.78** | **0.30** | **0.205249** | **-0.34** | **0.75** |
| **Log(ALT)** | **6** | **1.90** | **0.44** | **7** | **1.93** | **0.44** | **7** | **2.02** | **0.44** | **-0.10959** | **-0.68** | **0.46** |
| **Log(AST)** | **6** | **1.92** | **0.29** | **7** | **1.93** | **0.23** | **7** | **1.93** | **0.35** | **-0.00665** | **-0.39** | **0.38** |
| **Log(SDH)** | **6** | **1.95** | **0.35** | **7** | **1.88** | **0.33** | **7** | **1.94** | **0.31** | **-0.00693** | **-0.44** | **0.43** |

**0.706:0.294 BDCM:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **5** | **8.78** | **0.59** | **2** | **5.86** | **0.51** | **3** | **10.19** | **1.33** | **-2.27129** | **-4.11** | **-0.44** |
| **Log(ALT)** | **5** | **3.13** | **0.08** | **2** | **2.90** | **0.01** | **3** | **3.11** | **0.44** | **-0.05121** | **-0.57** | **0.47** |
| **Log(AST)** | **5** | **2.96** | **0.12** | **2** | **3.03** | **0.06** | **3** | **2.96** | **0.49** | **0.021286** | **-0.57** | **0.61** |
| **Log(SDH)** | **5** | **2.67** | **0.19** | **2** | **2.53** | **0.05** | **3** | **2.68** | **0.15** | **-0.04645** | **-0.40** | **0.31** |

|  |
| --- |
| 1PcLiv calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S6. Prop-RA Results for Binary Mixture of BDCM and CDBM-rep1,2**

**0.5:0.5 BDCM:CDBM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **8** | **5.27** | **0.48** | **11** | **5.01** | **0.44** | **7** | **5.07** | **0.48** | **0.072058** | **-0.47** | **0.61** |
| **Log(ALT)** | **8** | **1.43** | **0.10** | **11** | **1.28** | **0.14** | **7** | **1.36** | **0.14** | **-0.004** | **-0.15** | **0.14** |
| **Log(AST)** | **8** | **1.62** | **0.07** | **11** | **1.59** | **0.11** | **7** | **1.61** | **0.10** | **-0.008** | **-0.12** | **0.10** |
| **Log(SDH)** | **8** | **1.39** | **0.06** | **11** | **1.35** | **0.06** | **7** | **1.36** | **0.07** | **0.0065** | **-0.06** | **0.08** |

**0.5:0.5 BDCM:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **7** | **6.14** | **0.37** | **8** | **6.19** | **0.42** | **5** | **6.57** | **0.67** | **-0.40968** | **-1.06** | **0.24** |
| **Log(ALT)** | **7** | **1.89** | **0.26** | **8** | **1.82** | **0.22** | **5** | **1.91** | **0.33** | **-0.059** | **-0.42** | **0.31** |
| **Log(AST)** | **7** | **1.82** | **0.27** | **8** | **1.83** | **0.16** | **5** | **1.83** | **0.26** | **-0.004** | **-0.32** | **0.31** |
| **Log(SDH)** | **7** | **1.83** | **0.15** | **8** | **1.80** | **0.18** | **5** | **1.91** | **0.33** | **-0.097** | **-0.39** | **0.20** |

**0.5:0.5 BDCM:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **4** | **8.81** | **0.38** | **1** | **8.74** | **.** | **1** | **7.81** | **.** | **0.964414** | **N/A** | **N/A** |
| **Log(ALT)** | **4** | **2.00** | **0.78** | **1** | **2.82** | **.** | **1** | **2.17** | **.** | **0.2435** | **N/A** | **N/A** |
| **Log(AST)** | **4** | **2.32** | **0.29** | **1** | **2.72** | **.** | **1** | **2.11** | **.** | **0.407** | **N/A** | **N/A** |
| **Log(SDH)** | **4** | **2.58** | **0.03** | **1** | **2.57** | **.** | **1** | **2.02** | **.** | **0.5595** | **N/A** | **N/A** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S6 (cont). Prop-RA Results for Binary Mixture of BDCM and CDBM-rep1,2**

**0.706:0.294 BDCM:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **7** | **6.14** | **0.37** | **8** | **6.19** | **0.42** | **8** | **6.17** | **0.35** | **-0.01415** | **-0.47** | **0.44** |
| **Log(ALT)** | **7** | **1.89** | **0.26** | **8** | **1.82** | **0.22** | **8** | **1.76** | **0.11** | **0.109244** | **-0.13** | **0.35** |
| **Log(AST)** | **7** | **1.82** | **0.27** | **8** | **1.83** | **0.16** | **8** | **1.74** | **0.06** | **0.081176** | **-0.13** | **0.30** |
| **Log(SDH)** | **7** | **1.83** | **0.15** | **8** | **1.80** | **0.18** | **8** | **1.80** | **0.10** | **0.022592** | **-0.15** | **0.19** |

**0.706:0.294 BDCM:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **BDCM** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **4** | **8.81** | **0.38** | **1** | **8.74** | **.** | **6** | **8.02** | **0.64** | **0.771884** | **N/A** | **N/A** |
| **Log(ALT)** | **4** | **2.00** | **0.78** | **1** | **2.82** | **.** | **6** | **2.51** | **0.18** | **-0.26239** | **N/A** | **N/A** |
| **Log(AST)** | **4** | **2.32** | **0.29** | **1** | **2.72** | **.** | **6** | **2.38** | **0.16** | **0.051776** | **N/A** | **N/A** |
| **Log(SDH)** | **4** | **2.58** | **0.03** | **1** | **2.57** | **.** | **6** | **2.28** | **0.31** | **0.303178** | **N/A** | **N/A** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S7. Prop-RA Results for Binary Mixture of CHCl3 and CDBM1,2**

**0.5:0.5 CHCl3:CDBM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **8** | **4.50** | **0.27** | **8** | **4.88** | **0.52** | **8** | **4.76** | **0.89** | **-0.06814** | **-0.77** | **0.63** |
| **Log(ALT)** | **8** | **1.49** | **0.22** | **8** | **1.41** | **0.07** | **7** | **1.47** | **0.10** | **-0.019** | **-0.20** | **0.16** |
| **Log(AST)** | **8** | **1.57** | **0.05** | **8** | **1.56** | **0.09** | **7** | **1.69** | **0.06** | **-0.126** | **-0.21** | **-0.04** |
| **Log(SDH)** | **8** | **1.26** | **0.07** | **8** | **1.25** | **0.06** | **7** | **1.37** | **0.15** | **-0.114** | **-0.23** | **0.01** |

**0.5:0.5 CHCl3:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **8** | **6.09** | **0.29** | **10** | **6.31** | **0.43** | **9** | **6.01** | **0.58** | **0.189974** | **-0.29** | **0.67** |
| **Log(ALT)** | **8** | **1.84** | **0.29** | **10** | **1.74** | **0.26** | **9** | **1.83** | **0.19** | **-0.045** | **-0.31** | **0.22** |
| **Log(AST)** | **8** | **1.77** | **0.22** | **10** | **1.75** | **0.24** | **9** | **1.86** | **0.27** | **-0.105** | **-0.37** | **0.16** |
| **Log(SDH)** | **8** | **1.61** | **0.28** | **10** | **1.68** | **0.16** | **8** | **1.73** | **0.19** | **-0.085** | **-0.32** | **0.15** |

**0.5:0.5 CHCl3:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **4** | **6.74** | **1.26** | **3** | **8.05** | **0.76** | **5** | **7.77** | **1.00** | **-0.37062** | **-2.17** | **1.43** |
| **Log(ALT)** | **4** | **1.68** | **0.73** | **3** | **2.25** | **0.18** | **5** | **2.29** | **0.81** | **-0.33** | **-1.51** | **0.85** |
| **Log(AST)** | **4** | **2.29** | **0.49** | **3** | **2.58** | **0.15** | **5** | **2.70** | **0.36** | **-0.267** | **-0.91** | **0.38** |
| **Log(SDH)** | **4** | **2.27** | **0.64** | **3** | **2.37** | **0.29** | **5** | **2.50** | **0.34** | **-0.1845** | **-0.96** | **0.59** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S7 (cont). Prop-RA Results for Binary Mixture of CHCl3 and CDBM1,2**

**0.867:0.133 CHCl3:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **8** | **6.09** | **0.29** | **10** | **6.31** | **0.43** | **8** | **6.01** | **0.42** | **0.110483** | **-0.37** | **0.59** |
| **Log(ALT)** | **8** | **1.84** | **0.29** | **10** | **1.74** | **0.26** | **8** | **1.77** | **0.39** | **0.0517** | **-0.34** | **0.44** |
| **Log(AST)** | **8** | **1.77** | **0.22** | **10** | **1.75** | **0.24** | **8** | **1.73** | **0.25** | **0.034074** | **-0.26** | **0.32** |
| **Log(SDH)** | **8** | **1.61** | **0.28** | **10** | **1.68** | **0.16** | **8** | **1.61** | **0.25** | **0.010044** | **-0.27** | **0.29** |

**0.867:0.133 CHCl3:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **CDBM** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **4** | **6.74** | **1.26** | **3** | **8.05** | **0.76** | **5** | **7.71** | **0.51** | **-0.78663** | **-2.40** | **0.82** |
| **Log(ALT)** | **4** | **1.68** | **0.73** | **3** | **2.25** | **0.18** | **5** | **2.19** | **0.13** | **-0.42899** | **-1.23** | **0.37** |
| **Log(AST)** | **4** | **2.29** | **0.49** | **3** | **2.58** | **0.15** | **5** | **2.27** | **0.26** | **0.061102** | **-0.56** | **0.68** |
| **Log(SDH)** | **4** | **2.27** | **0.64** | **3** | **2.37** | **0.29** | **5** | **2.32** | **0.32** | **-0.0443** | **-0.86** | **0.77** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S8. Prop-RA Results for Binary Mixture of CDBM and CHBr31,2**

**0.5:0.5 CDBM and CHBr3**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CDBM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **11** | **5.02** | **0.39** | **11** | **5.08** | **0.25** | **8** | **5.27** | **0.36** | **-0.21684** | **-0.58** | **0.14** |
| **Log(ALT)** | **11** | **1.29** | **0.04** | **11** | **1.33** | **0.12** | **9** | **1.29** | **0.08** | **0.027** | **-0.06** | **0.12** |
| **Log(AST)** | **11** | **1.64** | **0.14** | **11** | **1.61** | **0.18** | **9** | **1.55** | **0.07** | **0.078** | **-0.07** | **0.23** |
| **Log(SDH)** | **11** | **1.14** | **0.05** | **11** | **1.14** | **0.07** | **9** | **1.14** | **0.06** | **0** | **-0.06** | **0.06** |

**0.5:0.5 CDBM and CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CDBM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **10** | **6.40** | **0.24** | **11** | **6.43** | **0.33** | **7** | **6.58** | **0.46** | **-0.16748** | **-0.55** | **0.22** |
| **Log(ALT)** | **10** | **1.70** | **0.26** | **11** | **1.57** | **0.20** | **7** | **1.54** | **0.12** | **0.093** | **-0.15** | **0.33** |
| **Log(AST)** | **10** | **1.76** | **0.21** | **11** | **1.68** | **0.13** | **7** | **1.68** | **0.06** | **0.038** | **-0.14** | **0.22** |
| **Log(SDH)** | **10** | **1.44** | **0.16** | **11** | **1.35** | **0.17** | **7** | **1.33** | **0.08** | **0.067** | **-0.10** | **0.23** |

**0.5:0.5 CDBM and CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CDBM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **5** | **7.59** | **0.46** | **4** | **7.55** | **0.58** | **5** | **6.96** | **0.52** | **0.611068** | **-0.20** | **1.42** |
| **Log(ALT)** | **5** | **2.82** | **0.20** | **4** | **2.19** | **0.23** | **5** | **2.23** | **0.35** | **0.2755** | **-0.15** | **0.70** |
| **Log(AST)** | **5** | **2.74** | **0.22** | **4** | **2.30** | **0.19** | **5** | **2.28** | **0.25** | **0.239** | **-0.11** | **0.59** |
| **Log(SDH)** | **5** | **2.08** | **0.28** | **4** | **1.76** | **0.17** | **5** | **1.85** | **0.34** | **0.0675** | **-0.38** | **0.51** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S8 (cont). Prop-RA Results for Binary Mixture of CDBM and CHBr31,2**

**0.909:0.091 CDBM:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CDBM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **10** | **6.40** | **0.24** | **11** | **6.43** | **0.33** | **9** | **6.30** | **0.41** | **0.100385** | **-0.28** | **0.48** |
| **Log(ALT)** | **10** | **1.70** | **0.26** | **11** | **1.57** | **0.20** | **9** | **1.64** | **0.25** | **0.04917** | **-0.22** | **0.32** |
| **Log(AST)** | **10** | **1.76** | **0.21** | **11** | **1.68** | **0.13** | **9** | **1.74** | **0.20** | **0.014356** | **-0.20** | **0.22** |
| **Log(SDH)** | **10** | **1.44** | **0.16** | **11** | **1.35** | **0.17** | **9** | **1.38** | **0.19** | **0.049174** | **-0.15** | **0.24** |

**0.909:0.091 CDBM:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CDBM** | **CHBr3** | **Mixture** | **L** | **Scheffé 95% CI** |
| **N** | **Mean** | **Std Dev** | **N** | **Mean** | **St Dev** | **N** | **Mean** | **St Dev** | **Low CL** | **Upp CL** |
| **PcLiv** | **5** | **7.59** | **0.46** | **4** | **7.55** | **0.58** | **2** | **8.52** | **0.36** | **-0.93904** | **-2.15** | **0.27** |
| **Log(ALT)** | **5** | **2.82** | **0.20** | **4** | **2.19** | **0.23** | **2** | **2.19** | **0.10** | **0.570307** | **0.08** | **1.06** |
| **Log(AST)** | **5** | **2.74** | **0.22** | **4** | **2.30** | **0.19** | **2** | **2.11** | **0.17** | **0.590324** | **0.10** | **1.08** |
| **Log(SDH)** | **5** | **2.08** | **0.28** | **4** | **1.76** | **0.17** | **2** | **1.77** | **0.16** | **0.277335** | **-0.29** | **0.85** |

|  |
| --- |
| 1Relative Liver Weight calculations use untransformed data. ALT, AST and SDH calculations use Log10 transformed data. |
| 2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05. |

**Table S9. Prop-RA Untransformed Data Results for Binary Mixture of CHCl3 and BDCM1,2**

**0.5:0.5 CHCl3:BDCM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Scheffe 95% Confidence** |
| **N Mean Std Dev****10 4.66 0.48****10 26.30 5.76****10 41.20 8.59****10 14.41 1.16** | **N Mean St Dev****10 4.72 0.40****10 26.10 7.56****10 45.80 12.61****10 12.66 3.45** | **N Mean St Dev****11 4.98 0.49****11 25.82 6.08****11 43.36 10.64****11 14.50 1.90** | **Lower CL****-0.74****-5.92****-10.28****-3.25** | **Upper CL****0.16****6.69****10.55****1.32** |
| **Rel Liver Wt****ALT AST SDH** | **-0.28717** |
| **0.381818** |
| **0.136364** |
| **-0.965** |

**0.5:0.5 CHCl3:BDCM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Scheffe 95% Confidence** |
| **N Mean Std Dev****9 5.40 0.31****9 43.44 14.56****9 44.00 8.09****9 21.43 3.57** | **N Mean St Dev****7 6.15 0.83****7 94.86 60.82****7 64.43 25.86****7 34.14 20.69** | **N Mean St Dev****10 5.76 0.37****11 45.00 14.01****11 42.82 7.17****11 24.06 6.04** | **Lower CL****-0.53****-9.49****-3.47****-7.81** | **Upper CL****0.56****57.79****26.26****15.25** |
| **Rel Liver Wt****ALT AST SDH** | **0.018606** |
| **24.15079** |
| **11.3961** |
| **3.724459** |

**0.5:0.5 CHCl3:BDCM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Scheffe 95% Confidence** |
| **N Mean Std Dev****7 7.72 0.95****7 630.86 461.50****7 418.43 436.43****7 104.73 68.50** | **N Mean St Dev****7 8.16 1.59****7 989.00 749.65****7 941.14 810.13****7 133.59 158.15** | **N Mean St Dev****9 8.42 0.93****9 948.44 737.60****9 847.22 885.45****9 155.68 144.59** | **Lower CL****-1.80****-896.12****-1018.22****-184.93** | **Upper CL****0.85****619.08****683.35****111.89** |
| **Rel Liver Wt****ALT AST SDH** | **-0.47626** |
| **-138.516** |
| **-167.437** |
| **-36.5206** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S9 (cont). Prop-RA Untransformed Data Results for Binary Mixture of CHCl3 and BDCM1,2**

**0.730:0.270 CHCl3:BDCM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Scheffe 95% Confidence** |
| **N Mean Std Dev****9 5.40 0.31****9 43.44 14.56****9 44.00 8.09****9 21.43 3.57** | **N Mean St Dev****7 6.15 0.83****7 94.86 60.82****7 64.43 25.86****7 34.14 20.69** | **N Mean St Dev****11 5.68 0.44****11 64.73 23.72****11 64.82 9.87****11 26.38 8.75** | **Lower CL****-0.63****-44.06****-31.14****-14.03** | **Upper CL****0.48****29.25****0.53****10.99** |
| **Rel Liver Wt****ALT AST SDH** | **-0.07441** |
| **-7.4014** |
| **-15.3025** |
| **-1.51691** |

**0.730:0.270 CHCl3:BDCM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Scheffe 95% Confidence** |
| **N Mean Std Dev****7 7.72 0.95****7 630.86 461.50****7 418.43 436.43****7 104.73 68.50** | **N Mean St Dev****7 8.16 1.59****7 989.00 749.65****7 941.14 810.13****7 133.59 158.15** | **N Mean St Dev****11 8.11 0.85****11 1128.55 869.60****11 718.27 536.39****11 141.90 101.32** | **Lower CL****-1.51****-1224.61****-823.57****-154.29** | **Upper CL****0.98****422.62****506.15****95.53** |
| **Rel Liver Wt****ALT AST SDH** | **-0.26488** |
| **-400.994** |
| **-158.711** |
| **-29.38** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S10. Prop-RA Untransformed Data Results for Binary Mixture of CHCl3 and BDCM-rep1,2**

**0.5:0.5 CHCl3:BDCM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****8 5.18 0.39****8 25.63 9.43****8 40.38 7.84****8 13.83 3.12** | **N Mean St Dev****10 5.38 0.39****10 23.30 8.37****10 41.50 10.64****10 12.52 2.18** | **N Mean St Dev****10 5.23 0.37****10 25.30 7.12****10 38.70 7.06****10 13.35 2.08** | **Lower CL****-0.34****-9.34****-6.72****-2.69** | **Upper CL****0.45****7.66****11.20****2.34** |
| **Wt****ALT AST SDH** | **0.053491** |
| **-0.8375** |
| **2.2375** |
| **-0.1775** |

**0.5:0.5 CHCl3:BDCM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****9 6.34 0.54****9 52.56 16.44****9 45.44 5.59****9 28.04 10.52** | **N Mean St Dev****8 6.22 0.50****7 77.57 39.09****7 59.29 17.92****7 43.97 18.10** | **N Mean St Dev****10 6.30 0.63****9 61.22 35.15****9 56.67 18.94****9 32.69 14.97** | **Lower CL****-0.61****-30.21****-20.91****-12.61** | **Upper CL****0.57****37.89****12.31****19.25** |
| **Wt****ALT AST SDH** | **-0.01859** |
| **3.84127** |
| **-4.30159** |
| **3.319048** |

**0.5:0.5 CHCl3:BDCM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****5 8.29 1.78****3 1205.00 1057.60****3 633.33 596.96****3 301.27 253.79** | **N Mean St Dev****7 8.41 0.55****7 641.86 264.05****7 478.71 215.89****7 188.46 82.55** | **N Mean St Dev****9 9.22 1.77****10 1508.30 1258.59****10 1141.00 1324.22****10 333.01 341.88** | **Lower CL****-2.62****-1835.98****-1830.99****-424.40** | **Upper CL****0.88****666.24****661.04****248.11** |
| **Wt****ALT AST SDH** | **-0.86813** |
| **-584.871** |
| **-584.976** |
| **-88.1481** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S10 (cont). Prop-RA Untransformed Data Results for Binary Mixture of CHCl3 and BDCM-rep1,2**

**0.730:0.270 CHCl3:BDCM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****9 6.34 0.54****9 52.56 16.44****9 45.44 5.59****9 28.04 10.52** | **N Mean St Dev****8 6.22 0.50****7 77.57 39.09****7 59.29 17.92****7 43.97 18.10** | **N Mean St Dev****10 6.40 0.72****10 44.20 14.73****10 46.00 10.28****10 23.86 7.60** | **Lower CL****-0.74****-10.78****-9.39****-4.56** | **Upper CL****0.55****41.00****15.75****21.53** |
| **Wt****ALT AST SDH** | **-0.0947** |
| **15.10984** |
| **3.181587** |
| **8.48473** |

**0.730:0.270 CHCl3:BDCM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **CHCl3** | **BDCM** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****5 8.29 1.78****3 1205.00 1057.60****3 633.33 596.96****3 301.27 253.79** | **N Mean St Dev****7 8.41 0.55****7 641.86 264.05****7 478.71 215.89****7 188.46 82.55** | **N Mean St Dev****7 8.43 0.98****7 855.57 769.66****7 657.00 627.80****7 240.03 216.22** | **Lower CL****-1.66****-850.25****-835.72****-251.30** | **Upper CL****1.43****1245.01****704.89****312.86** |
| **Wt****ALT AST SDH** | **-0.11181** |
| **197.38** |
| **-65.4138** |
| **30.77952** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S11. Prop-RA Untransformed Data Results for Binary Mixture of CHCl3 and CHBr31,2**

**0.5:0.5 CHCl3:CHBr3**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****10 4.97 0.62****10 27.10 10.72****10 43.50 12.29****9 19.34 3.28** | **N Mean St Dev****4 5.06 0.33****17 29.44 5.77****34 45.33 13.59****12 19.94 3.33** | **N Mean St Dev****11 4.75 0.40****10 28.90 9.55****10 42.50 7.93****10 20.91 3.74** | **CL****-0.28****-8.65****-9.59****-4.70** | **Upper CL****0.81****7.39****13.42****2.17** |
| **Rel Liver Wt****ALT AST SDH** | **0.268194** |
| **-0.62778** |
| **1.916667** |
| **-1.26556** |

**0.5:0.5 CHCl3:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****9 5.70 0.56****9 64.11 27.72****9 47.44 11.92****9 44.77 13.52** | **N Mean St Dev****9 6.42 0.63****9 54.00 28.57****9 55.00 13.96****9 39.71 14.71** | **N Mean St Dev****9 6.08 0.39****9 52.89 26.49****9 51.78 15.43****9 34.32 5.94** | **CL****-0.58****-23.24****-15.30****-4.90** | **Upper CL****0.56****35.57****14.19****20.73** |
| **Rel Liver Wt****ALT AST SDH** | **-0.01351** |
| **6.166667** |
| **-0.55556** |
| **7.916667** |

**0.5:0.5 CHCl3:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****8 7.69 1.14****8 728.13 616.68****8 490.50 499.45****8 208.94 187.35** | **N Mean St Dev****8 7.47 0.70****8 272.75 118.97****8 268.75 88.78****8 70.13 8.55** | **N Mean St Dev****4 7.12 0.29****4 138.75 63.84****4 116.50 43.88****4 63.53 9.09** | **CL****-0.84****-243.45****-225.36****-104.39** | **Upper CL****1.76****966.83****751.61****256.40** |
| **Rel Liver Wt****ALT AST SDH** | **0.461538** |
| **361.6875** |
| **263.125** |
| **76.00625** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S11 (cont). Prop-RA Untransformed Data Results for Binary Mixture of CHCl3 and CHBr31,2**

**0.985:0.015 CHCl3:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****9 5.70 0.56****9 64.11 27.72****9 47.44 11.92****9 44.77 13.52** | **N Mean St Dev****9 6.42 0.63****9 54.00 28.57****9 55.00 13.96****9 39.71 14.71** | **N Mean St Dev****10 6.09 0.49****10 56.40 30.96****10 42.40 14.95****10 36.45 12.71** | **CL****-1.05****-27.07****-11.12****-7.93** | **Upper CL****0.28****42.19****21.44****24.41** |
| **Rel Liver Wt****ALT AST SDH** | **-0.38271** |
| **7.559444** |
| **5.157778** |
| **8.240833** |

**0.985:0.015 CHCl3:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****8 7.69 1.14****8 728.13 616.68****8 490.50 499.45****8 208.94 187.35** | **N Mean St Dev****8 7.47 0.70****8 272.75 118.97****8 268.75 88.78****8 70.13 8.55** | **N Mean St Dev****6 7.23 1.77****5 836.00 588.73****5 689.60 821.54****5 210.74 188.71** | **CL****-1.28****-840.19****-958.19****-225.96** | **Upper CL****2.19****610.78****553.34****218.20** |
| **Rel Liver Wt****ALT AST SDH** | **0.456933** |
| **-114.706** |
| **-202.426** |
| **-3.88469** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S12. Prop-RA Untransformed Data Results for Binary Mixture of BDCM and CHBr31,2**

**0.5:0.5 BDCM:CHBr3**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****9 4.98 0.32****9 22.22 7.74****9 34.00 5.50****9 16.06 2.34** | **N Mean St Dev****11 5.23 1.29****11 39.27 45.02****11 51.45 37.21****11 20.20 14.38** | **N Mean St Dev****10 5.00 0.65****10 24.30 7.07****10 55.80 39.80****10 16.11 2.70** | **CL****-0.79****-21.71****-45.63****-7.01** | **Upper CL****1.00****34.60****19.48****11.04** |
| **Rel Liver Wt****ALT AST SDH** | **0.10586** |
| **6.447475** |
| **-13.0727** |
| **2.017778** |

**0.5:0.5 BDCM:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****10 5.99 0.45****10 110.80 72.03****10 69.00 40.90****10 56.28 18.33** | **N Mean St Dev****9 6.39 0.37****8 73.13 56.93****8 61.88 34.08****8 37.48 15.38** | **N Mean St Dev****7 6.87 0.92****6 111.17 99.36****6 86.67 53.57****6 43.07 24.74** | **CL****-1.36****-112.69****-73.79****-20.06** | **Upper CL****0.00****74.28****31.33****27.68** |
| **Rel Liver Wt****ALT AST SDH** | **-0.68016** |
| **-19.2042** |
| **-21.2292** |
| **3.810833** |

**0.5:0.5 BDCM:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****5 8.34 1.16****5 754.80 333.44****5 622.00 303.18****5 404.30 193.86** | **N Mean St Dev****4 7.69 1.13****4 346.50 251.46****4 309.75 212.74****4 183.28 233.98** | **N Mean St Dev****6 7.69 0.49****6 259.83 132.67****6 199.83 76.30****6 62.72 13.00** | **CL****-1.05****-70.57****-44.23****-7.71** | **Upper CL****1.69****652.20****576.32****469.85** |
| **Rel Liver Wt****ALT AST SDH** | **0.323436** |
| **290.8167** |
| **266.0417** |
| **231.0708** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S12 (cont). Prop-RA Untransformed Data Results for Binary Mixture of BDCM and CHBr31,2**

**0.96:0.040 BDCM:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****10 5.99 0.45****10 110.80 72.03****10 69.00 40.90****10 56.28 18.33** | **N Mean St Dev****9 6.39 0.37****8 73.13 56.93****8 61.88 34.08****8 37.48 15.38** | **N Mean St Dev****8 5.98 0.33****8 141.13 107.57****8 77.63 43.16****8 58.89 26.09** | **CL****-0.45****-130.46****-57.31****-28.08** | **Upper CL****0.50****66.79****39.49****21.36** |
| **Rel Liver Wt****ALT AST SDH** | **0.026164** |
| **-31.832** |
| **-8.91** |
| **-3.3597** |

**0.96:0.040 BDCM:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****5 8.34 1.16****5 754.80 333.44****5 622.00 303.18****5 404.30 193.86** | **N Mean St Dev****4 7.69 1.13****4 346.50 251.46****4 309.75 212.74****4 183.28 233.98** | **N Mean St Dev****6 8.75 0.97****6 651.83 605.20****6 381.33 241.75****6 241.70 254.94** | **CL****-2.21****-662.41****-197.31****-227.71** | **Upper CL****1.34****835.68****653.67****535.23** |
| **Rel Liver Wt****ALT AST SDH** | **-0.43673** |
| **86.63467** |
| **228.1767** |
| **153.759** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S13. Prop-RA Untransformed Data Results for Binary Mixture of BDCM and CDBM1,2**

**0.5:0.5 BDCM:CDBM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****7 4.69 0.63****6 35.17 7.11****6 62.67 16.97****6 34.15 10.87** | **N Mean St Dev****7 5.24 0.27****6 24.33 8.52****6 42.83 5.08****6 26.92 5.20** | **N Mean St Dev****7 5.05 0.42****7 25.00 4.43****7 42.00 3.06****7 27.77 4.39** | **CL****-0.66****-3.93****-2.17****-6.54** | **Upper CL****0.48****13.43****23.67****12.06** |
| **Rel Liver Wt****ALT AST SDH** | **-0.09003** |
| **4.75** |
| **10.75** |
| **2.761905** |

**0.5:0.5 BDCM:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****6 5.94 0.44****6 125.83 148.05****6 101.50 71.89****6 125.23 134.38** | **N Mean St Dev****7 6.09 0.48****7 130.71 119.52****7 95.29 46.42****7 102.61 109.75** | **N Mean St Dev****2 6.48 0.24****2 116.00 82.02****2 90.50 60.10****2 77.80 0.99** | **CL****-1.42****-262.56****-117.98****-210.41** | **Upper CL****0.48****287.10****133.77****282.66** |
| **Rel Liver Wt****ALT AST SDH** | **-0.46903** |
| **12.27381** |
| **7.892857** |
| **36.12381** |

**0.5:0.5 BDCM:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****5 8.78 0.59****5 1359.00 237.47****5 950.40 264.84****5 509.58 231.20** | **N Mean St Dev****2 5.86 0.51****2 786.00 16.97****2 1084.50 142.13****2 340.65 39.67** | **N Mean St Dev****3 8.05 1.11****3 572.33 524.68****3 469.00 306.26****3 239.60 135.00** | **CL****-2.41****-230.75****-31.21****-230.74** | **Upper CL****0.96****1231.08****1128.11****601.77** |
| **Rel Liver Wt****ALT AST SDH** | **-0.72649** |
| **500.1667** |
| **548.45** |
| **185.515** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S13 (cont). Prop-RA Untransformed Data Results for Binary Mixture of BDCM and CDBM1,2**

**0.706:0.294 BDCM:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****6 5.94 0.44****6 125.83 148.05****6 101.50 71.89****6 125.23 134.38** | **N Mean St Dev****7 6.09 0.48****7 130.71 119.52****7 95.29 46.42****7 102.61 109.75** | **N Mean St Dev****7 5.78 0.30****7 158.86 160.51****7 114.57 105.38****7 109.47 88.45** | **CL****-0.34****-219.27****-117.91****-136.12** | **Upper CL****0.75****156.09****88.11****154.35** |
| **Rel Liver Wt****ALT AST SDH** | **0.205249** |
| **-31.5888** |
| **-14.8984** |
| **9.111905** |

**0.706:0.294 BDCM:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****5 8.78 0.59****5 1359.00 237.47****5 950.40 264.84****5 509.58 231.20** | **N Mean St Dev****2 5.86 0.51****2 786.00 16.97****2 1084.50 142.13****2 340.65 39.67** | **N Mean St Dev****3 10.19 1.33****3 1714.33 1310.64****3 1288.33 1047.27****3 494.13 177.96** | **CL** | **Upper CL** |
| **Rel Liver Wt****ALT AST SDH** | **-2.27129** | **-4.11** | **-0.44** |
| **-523.792** | **-2060.08****-1566.52****-458.09** | **1012.50****969.51****389.65** |
| **-298.505** |
| **-34.2188** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S14. Prop-RA Untransformed Data Results for Binary Mixture of BDCM and CDBM-rep1,2**

**0.5:0.5 BDCM:CDBM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****8 5.27 0.48****8 27.38 6.25****8 42.13 6.45****8 24.86 3.30** | **N Mean St Dev****11 5.01 0.44****11 19.91 5.45****11 40.36 12.42****11 22.54 2.99** | **N Mean St Dev****7 5.07 0.48****7 24.00 9.17****7 42.14 10.87****7 23.43 4.18** | **CL****-0.47****-8.29****-13.10****-3.71** | **Upper CL****0.61****7.58****11.30****4.25** |
| **Rel Liver Wt****ALT AST SDH** | **0.072058** |
| **-0.35795** |
| **-0.89854** |
| **0.27086** |

**0.5:0.5 BDCM:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****7 6.14 0.37****7 91.71 62.56****7 81.00 67.15****7 70.70 23.73** | **N Mean St Dev****8 6.19 0.42****8 72.75 33.67****8 71.25 26.40****8 67.13 27.24** | **N Mean St Dev****5 6.57 0.67****5 108.20 104.37****5 78.80 54.86****5 109.08 113.63** | **CL****-1.06****-117.94****-73.10****-122.59** | **Upper CL****0.24****66.00****67.75****42.25** |
| **Rel Liver Wt****ALT AST SDH** | **-0.40968** |
| **-25.9679** |
| **-2.675** |
| **-40.1675** |

**0.5:0.5 BDCM:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****4 8.81 0.38****4 244.25 258.37****4 243.50 156.92****4 383.70 27.46** | **N Mean St Dev****1 8.74 .****1 657.00 .****1 525.00 .****1 371.70 .** | **N Mean St Dev****1 7.81 .****1 147.00 .****1 129.00 .****1 104.00 .** | **CL****N/A N/A N/A N/A** | **Upper CL****N/A N/A N/A N/A** |
| **Rel Liver Wt****ALT AST SDH** | **0.964414** |
| **303.625** |
| **255.25** |
| **273.7** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S14 (cont). Prop-RA Untransformed Data Results for Binary Mixture of BDCM and CDBM-rep1,2**

**0.706:0.294 BDCM:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****7 6.14 0.37****7 91.71 62.56****7 81.00 67.15****7 70.70 23.73** | **N Mean St Dev****8 6.19 0.42****8 72.75 33.67****8 71.25 26.40****8 67.13 27.24** | **N Mean St Dev****8 6.17 0.35****8 59.38 17.05****8 55.75 7.40****8 63.94 14.44** | **CL****-0.47****-22.41****-25.95****-21.22** | **Upper CL****0.44****75.94****70.72****32.64** |
| **Rel Liver Wt****ALT AST SDH** | **-0.01415** |
| **26.76379** |
| **22.3835** |
| **5.71145** |

**0.706:0.294 BDCM:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****4 8.81 0.38****4 244.25 258.37****4 243.50 156.92****4 383.70 27.46** | **N Mean St Dev****1 8.74 .****1 657.00 .****1 525.00 .****1 371.70 .** | **N Mean St Dev****6 8.02 0.64****6 346.17 157.91****6 256.00 96.94****6 229.92 143.26** | **CL****N/A N/A N/A N/A** | **Upper CL****N/A N/A N/A N/A** |
| **Rel Liver Wt****ALT AST SDH** | **0.771884** |
| **19.43183** |
| **70.261** |
| **150.2553** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S15. Prop-RA Untransformed Data Results for Binary Mixture of CHCl3 and CDBM1,2**

**0.5:0.5 CHCl3:CDBM**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****8 4.50 0.27****8 35.50 26.59****8 37.50 4.24****8 18.53 2.64** | **N Mean St Dev****8 4.88 0.52****8 25.88 4.29****8 36.63 7.50****8 17.85 2.63** | **N Mean St Dev****8 4.76 0.89****7 29.86 6.28****7 49.43 6.92****7 24.79 9.20** | **CL****-0.77****-18.69** | **Upper CL****0.63****20.35** |
| **Rel Liver Wt****ALT AST SDH** | **-0.06814** |
| **0.830357** |
| **-12.3661** | **-19.98****-13.19** | **-4.76****-0.01** |
| **-6.59821** |

**0.5:0.5 CHCl3:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****8 6.09 0.29****8 82.13 49.37****8 66.88 44.34****8 48.43 27.99** | **N Mean St Dev****10 6.31 0.43****10 67.70 62.65****10 66.80 55.82****10 51.32 22.07** | **N Mean St Dev****9 6.01 0.58****9 74.11 35.36****9 93.89 99.11****8 57.93 21.20** | **CL****-0.29****-53.61****-102.64****-34.54** | **Upper CL****0.67****55.21****48.54****18.44** |
| **Rel Liver Wt****ALT AST SDH** | **0.189974** |
| **0.801389** |
| **-27.0514** |
| **-8.0525** |

**0.5:0.5 CHCl3:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****4 6.74 1.26****4 107.75 120.52****4 269.25 181.73****4 296.95 199.21** | **N Mean St Dev****3 8.05 0.76****3 186.33 74.14****3 397.33 130.59****3 265.33 137.20** | **N Mean St Dev****5 7.77 1.00****5 544.60 812.77****5 698.20 737.64****5 386.74 234.70** | **CL****-2.17****-1336.83****-1234.12****-456.79** | **Upper CL****1.43****541.71****504.30****245.60** |
| **Rel Liver Wt****ALT AST SDH** | **-0.37062** |
| **-397.558** |
| **-364.908** |
| **-105.598** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S15 (cont). Prop-RA Untransformed Data Results for Binary Mixture of CHCl3 and CDBM1,2**

**0.867:0.133 CHCl3:CDBM**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****8 6.09 0.29****8 82.13 49.37****8 66.88 44.34****8 48.43 27.99** | **N Mean St Dev****10 6.31 0.43****10 67.70 62.65****10 66.80 55.82****10 51.32 22.07** | **N Mean St Dev****8 6.01 0.42****8 88.50 93.06****8 63.13 42.23****8 47.66 29.79** | **CL****-0.37****-94.45****-55.98****-31.35** | **Upper CL****0.59****77.86****63.46****33.65** |
| **Rel Liver Wt****ALT AST SDH** | **0.110483** |
| **-8.29352** |
| **3.740025** |
| **1.147535** |

**0.867:0.133 CHCl3:CDBM**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****4 6.74 1.26****4 107.75 120.52****4 269.25 181.73****4 296.95 199.21** | **N Mean St Dev****3 8.05 0.76****3 186.33 74.14****3 397.33 130.59****3 265.33 137.20** | **N Mean St Dev****5 7.71 0.51****5 159.00 45.74****5 211.00 115.23****5 255.14 148.92** | **CL****-2.40****-193.92****-188.15****-264.75** | **Upper CL****0.82****112.32****338.72****339.96** |
| **Rel Liver Wt****ALT AST SDH** | **-0.78663** |
| **-40.7984** |
| **75.28508** |
| **37.60498** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S16. Prop-RA Untransformed Data Results for Binary Mixture of CDBM and CHBr31,2**

**0.5:0.5 CDBM and CHBr3**

**Dose = 0.1 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****11 5.02 0.39****11 19.64 1.96****11 46.36 22.67****11 13.97 1.70** | **N Mean St Dev****11 5.08 0.25****11 22.27 5.88****11 45.45 26.95****11 13.86 2.36** | **N Mean St Dev****8 5.27 0.36****9 19.56 3.64****9 35.78 6.18****9 13.88 1.93** | **CL****-0.58****-2.88****-11.65****-2.03** | **Upper CL****0.14****5.68****31.92****2.11** |
| **Rel Liver Wt****ALT AST SDH** | **-0.21684** |
| **1.39899** |
| **10.13131** |
| **0.040404** |

**0.5:0.5 CDBM and CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****10 6.40 0.24****10 60.60 44.74****10 64.50 32.56****10 28.94 11.00** | **N Mean St Dev****11 6.43 0.33****11 42.45 30.23****11 50.00 19.13****11 24.41 14.34** | **N Mean St Dev****7 6.58 0.46****7 36.00 9.61****7 48.43 6.00****7 21.47 4.19** | **CL****-0.55****-22.29****-17.49****-7.75** | **Upper CL****0.22****53.34****35.14****18.16** |
| **Rel Liver Wt****ALT AST SDH** | **-0.16748** |
| **15.52727** |
| **8.821429** |
| **5.203117** |

**0.5:0.5 CDBM and CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****5 7.59 0.46****5 705.40 278.77****5 600.60 256.70****5 143.38 99.96** | **N Mean St Dev****4 7.55 0.58****4 171.75 82.43****4 214.75 81.06****4 61.08 20.67** | **N Mean St Dev****5 6.96 0.52****5 221.60 191.59****5 225.60 167.87****5 96.72 101.32** | **CL****-0.20****-111.91****-117.27****-130.97** | **Upper CL****1.42****545.86****481.42****141.98** |
| **Rel Liver Wt****ALT AST SDH** | **0.611068** |
| **216.975** |
| **182.075** |
| **5.5075** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

**Table S16 (cont). Prop-RA Untransformed Data Results for Binary Mixture of CDBM and CHBr31,2**

**0.909:0.091 CDBM:CHBr3**

**Dose = 1.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****10 6.40 0.24****10 60.60 44.74****10 64.50 32.56****10 28.94 11.00** | **N Mean St Dev****11 6.43 0.33****11 42.45 30.23****11 50.00 19.13****11 24.41 14.34** | **N Mean St Dev****9 6.30 0.41****9 51.67 37.56****9 61.22 34.30****9 26.31 14.46** | **CL****-0.28****-35.82****-31.10****-13.05** | **Upper CL****0.48****50.38****35.01****17.48** |
| **Rel Liver Wt****ALT AST SDH** | **0.100385** |
| **7.282097** |
| **1.958278** |
| **2.216576** |

**0.909:0.091 CDBM:CHBr3**

**Dose = 3.0 mmol/kg/day**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
| **End Point** | **Chem A** | **Chem B** | **Mixture** | **L** | **Confidence Interval** |
| **N Mean Std Dev****5 7.59 0.46****5 705.40 278.77****5 600.60 256.70****5 143.38 99.96** | **N Mean St Dev****4 7.55 0.58****4 171.75 82.43****4 214.75 81.06****4 61.08 20.67** | **N Mean St Dev****2 8.52 0.36****2 156.50 36.06****2 133.50 50.20****2 61.25 22.42** | **CL****-2.15** | **Upper CL****0.27** |
| **Rel Liver Wt****ALT AST SDH** | **-0.93904** |
| **500.3379** | **2.98** | **997.70** |
| **431.9877** | **-29.12****-101.60** | **893.09****250.88** |
| **74.64025** |

1All calculations use original untransformed data.

2Shaded areas are statistically significant. Overall type I error rate for this study using Scheffé’s method is α=0.05.

# Homogeneity of Variances Test Results for Binary Mixtures For Analysis of Departures from Proportional Response Addition

To test for departures from proportional response addition (Prop-RA), a linear contrast is constructed for a given experiment for each dose/ratio/endpoint combination, and homogeneity of variance (HOV) is assumed across the two single chemical and the binary mixture datasets. To calculate a 95% Scheffé confidence interval for the linear contrast estimate used to analyze departures from Prop-RA, we generate an unbiased estimate of the variance of the linear contrast estimate. This calculation includes a value for the mean square error, which is a pooled weighted average of the three individual dataset variances.

To satisfy the HOV assumption, we used the O’Brien test for HOV. Based on published research, the O’Brien test was chosen for several reasons that are generally characteristic of our THM datasets. The O’Brien test is: the only HOV method that has adequate control of type I error rates for average sample sizes < 10; not sensitive to normality assumptions of the data; and, controls for type I error rate across all population shapes (Wang et al., 2017). In general, it is also not sensitive to skewed data (i.e., data points out in the tail of the distribution as compared with other values in the dataset), although this can sometimes be an issue with small sample sizes.

Supporting HOV results are shown below in Tables S17-S24 and Figures S1-S4. "Table 1" in footnote a refers to Table 1 of the main manuscript. In the figures, the lower and upper ends of the box represent the 1st and 3rd quartiles of the data distribution. Yellow-highlights identify HOV p-values that were higher with the log transform, indicating more homogeneity of variance, and blue-highlights identify HOV p-values that were lower.

HOV testing results were helpful in ascertaining the appropriateness of applying our method for determining departures from Prop-RA to the relative liver weight and serum enzyme data for the THMs. Because HOV testing showed general consistency (nonsignificant differences) across all but 2 experiments for dose/ratio/endpoint combinations for PcLiv, we chose to use the original untransformed data for testing departures from Prop-RA. In general, HOV testing results for the serum enzyme data showed improvements in similarity of variances using the transformed data across all experiments for dose/ratio/endpoint combinations. Evidence included O’Brien test results and visual inspection of changes in the similarities of the variances. Thus, for AST, ALT and SDH, we chose to use the Log10 transformed data for testing departures from Prop-RA.

# Table S17. Testing Equality of Variances- Relative Liver Weight

|  |  |  |
| --- | --- | --- |
| **Dose (mmol/ kg-d)** | **Ratio Studieda** | **Experimentb,c** |
| **1** | **1A** | **2** | **3** | **4** | **4A** | **5** | **6** |
| 0.1 | 1:1 | ns | ns | ns | ns | ns | ns | ns | ns |
| 1.0 | 1:1 | 0.01 | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | 1:1 | ns | ns | ns | ns | ns | d | ns | ns |
| 1.0 | envir. | 0.02 | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | envir. | ns | ns | ns | ns | ns\* | e | ns | ns |

1. See table 1 for the chemicals and environmentally relevant ratios used in each experiment
2. ns=not statistically significant (p<0.05) for O'Brien variance test. When shown, value is the significance level for that dose-ratio combination.
3. Areas with an \* represent data that were statistically different from Prop-RA.
4. No calculation, n=1 for mixture data and for one component's data.
5. ns but n=1 for one component's data.

# Table S18. Testing Equality of Variances- Log(Relative Liver Weight)

|  |  |  |
| --- | --- | --- |
| **Dose (mmol/ kg-d)** | **Ratio Studieda** | **Experimentb,c** |
| **1** | **1A** | **2** | **3** | **4** | **4A** | **5** | **6** |
| 0.1 | 1:1 | ns | ns | ns | ns | ns | ns | ns | ns |
| 1.0 | 1:1 | 0.02 | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | 1:1 | ns | ns | ns | ns | ns | d | ns | ns |
| 1.0 | envir. | 0.04 | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | envir. | ns | ns | ns | ns | ns\* | e | ns | ns |

1. See table 1 for the chemicals and environmentally relevant ratios used in each experiment
2. ns=not statistically significant (p<0.05) for O'Brien variance test. When shown, value is the significance level for that dose-ratio combination.
3. Areas with an \* represent data that were statistically different from Prop-RA.
4. No calculation, n=1 for mixture data and for one component's data.
5. ns but n=1 for one component's data.

Figure S1. HOV Analysis for (left) percent liver weight or (right) log10(percent liver weight), experiment #4, environmental mix ratio and dose of 3.0 mmol/kg/d

# Table S19. Testing Equality of Variances- ALT

|  |  |  |
| --- | --- | --- |
| **Dose (mmol/ kg-d)** | **Ratio Studieda** | **Experimentb,c** |
| **1** | **1A** | **2** | **3** | **4** | **4A** | **5** | **6** |
| 0.1 | 1:1 | ns | ns | ns | ns | ns | ns | ns | 0.01 |
| 1.0 | 1:1 | <0.01 | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | 1:1 | ns | ns | <0.01 | 0.03 | ns | d | ns | ns |
| 1.0 | envir. | 0.01 | 0.02 | ns | ns | ns | ns | ns | ns |
| 3.0 | envir. | ns | ns | ns | ns | ns | e | ns | ns\* |

1. See table 1 for the chemicals and environmentally relevant ratios used in each experiment
2. ns=not statistically significant (p<0.05) for O'Brien variance test. When shown, value is the significance level for that dose-ratio combination.
3. Areas with an \* represent data that were statistically different from Prop-RA.
4. No calculation, n=1 for mixture data and for one component's data.
5. p=0.04 but n=1 for one component's data.

# Table S20. Testing Equality of Variances- log10(ALT)

|  |  |  |
| --- | --- | --- |
| **Dose (mmol/ kg-d)** | **Ratio Studieda** | **Experimentb,c** |
| **1** | **1A** | **2** | **3** | **4** | **4A** | **5** | **6** |
| 0.1 | 1:1 | ns | ns | ns | ns | ns | ns | ns | 0.02 |
| 1.0 | 1:1 | 0.04 | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | 1:1 | ns | ns | 0.04\* | 0.02 | ns | d | ns | ns |
| 1.0 | envir. | ns | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | envir. | ns | ns | ns | ns | ns | e | 0.04 | ns\* |

1. See table 1 for the chemicals and environmentally relevant ratios used in each experiment
2. ns=not statistically significant (p<0.05) for O'Brien variance test. When shown, value is the significance level for that dose-ratio combination.
3. Areas with an \* represent data that were statistically different from Prop-RA.
4. No calculation, n=1 for mixture data and for one component's data.
5. p<0.01 but n=1 for one component's data.

Figure S2. HOV Analysis for (left) ALT and (right) Log(ALT), experiment #2, 1:1 ratio and dose of 3.0 mmol/kg/d

# Table S21. Testing Equality of Variances- AST

|  |  |  |
| --- | --- | --- |
| **Dose (mmol/ kg-d)** | **Ratio Studieda** | **Experimentbcd** |
| **1** | **1A** | **2** | **3** | **4** | **4A** | **5** | **6** |
| 0.1 | 1:1 | ns | ns | ns | ns | ns | ns | ns\* | ns |
| 1.0 | 1:1 | <0.01 | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | 1:1 | ns | ns | 0.01 | 0.01 | ns | d | ns | ns |
| 1.0 | envir. | <0.01 | 0.01 | ns | ns | ns | ns | ns | ns |
| 3.0 | envir. | ns | ns | ns | ns | ns | e | ns | ns |

1. See table 1 for the chemicals and environmentally relevant ratios used in each experiment
2. ns=not statistically significant (p<0.05) for O'Brien variance test. When shown, value is the significance level for that dose-ratio combination.
3. Areas with an \* represent data that were statistically different from Prop-RA.
4. No calculation, n=1 for mixture data and for one component's data.
5. ns but n=1 for one component's data.

# Table S22. Testing Equality of Variances- log10(AST)

|  |  |  |
| --- | --- | --- |
| **Dose (mmol/ kg-d)** | **Ratio Studieda** | **Experimentb,c** |
| **1** | **1A** | **2** | **3** | **4** | **4A** | **5** | **6** |
| 0.1 | 1:1 | ns | ns | ns | ns | ns | ns | ns\* | ns |
| 1.0 | 1:1 | <0.01 | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | 1:1 | ns | ns | <0.01 | ns | ns | d | ns | ns |
| 1.0 | envir. | <0.01\* | 0.046 | ns | ns | ns | ns | ns | ns |
| 3.0 | envir. | ns | 0.04 | ns | ns | ns | e | ns | ns\* |

1. See table 1 for the chemicals and environmentally relevant ratios used in each experiment
2. ns=not statistically significant (p<0.05) for O'Brien variance test. When shown, value is the significance level for that dose-ratio combination.
3. Areas with an \* represent data that were statistically different from Prop-RA.
4. No calculation, n=1 for mixture data and for one component's data.
5. ns but n=1 for one component's data.

Figure S3. HOV Analysis for (left) AST and (right) Log(AST), experiment #1, environmental mix ratio and dose of 1.0 mmol/kg/d

# Table S23. Testing Equality of Variances- SDH

|  |  |  |
| --- | --- | --- |
| **Dose (mmol/ kg-d)** | **Ratio Studieda** | **Experimentb,c** |
| **1** | **1A** | **2** | **3** | **4** | **4A** | **5** | **6** |
| 0.1 | 1:1 | ns | ns | ns | ns | ns | ns | ns\* | ns |
| 1.0 | 1:1 | ns | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | 1:1 | ns | ns | ns | ns | ns | d | ns | ns |
| 1.0 | envir. | ns | 0.04 | ns | ns | ns | ns | ns | ns |
| 3.0 | envir. | ns | ns | ns | ns | ns | e | ns | ns |

1. See table 1 for the chemicals and environmentally relevant ratios used in each experiment
2. ns=not statistically significant (p<0.05) for O'Brien variance test. When shown, value is the significance level for that dose-ratio combination.
3. Areas with an \* represent data that were statistically different from Prop-RA.
4. No calculation, n=1 for mixture data and for one component's data.
5. p<0.01 but n=1 for one component's data.

# Table S24. Testing Equality of Variances- log10(SDH)

|  |  |  |
| --- | --- | --- |
| **Dose (mmol/ kg-d)** | **Ratio Studieda** | **Experimentb,c** |
| **1** | **1A** | **2** | **3** | **4** | **4A** | **5** | **6** |
| 0.1 | 1:1 | ns | ns | ns | ns | ns | ns | ns | ns |
| 1.0 | 1:1 | ns | ns | ns | 0.04 | ns | ns | ns | ns |
| 3.0 | 1:1 | ns | ns | 0.02 | ns\* | ns | d | ns | ns |
| 1.0 | envir. | ns | ns | ns | ns | ns | ns | ns | ns |
| 3.0 | envir. | ns | ns | <0.01 | ns | ns | e | ns | ns |

1. See table 1 for the chemicals and environmentally relevant ratios used in each experiment
2. ns=not statistically significant (p<0.05) for O'Brien variance test. When shown, value is the significance level for that dose-ratio combination.
3. Areas with an \* represent data that were statistically different from Prop-RA.
4. No calculation, n=1 for mixture data and for one component's data.
5. p<0.01 but n=1 for one component's data.

Figure S4. HOV analysis for (left) SDH and (right) Log(SDH), experiment #5, 1:1 mix ratio and dose of 0.1 mmol/kg/d