

Supplementary Materials: Dose-Dependent Pharmacokinetics of Tofacitinib in Rats: Influence of Hepatic and Intestinal First-Pass Metabolism

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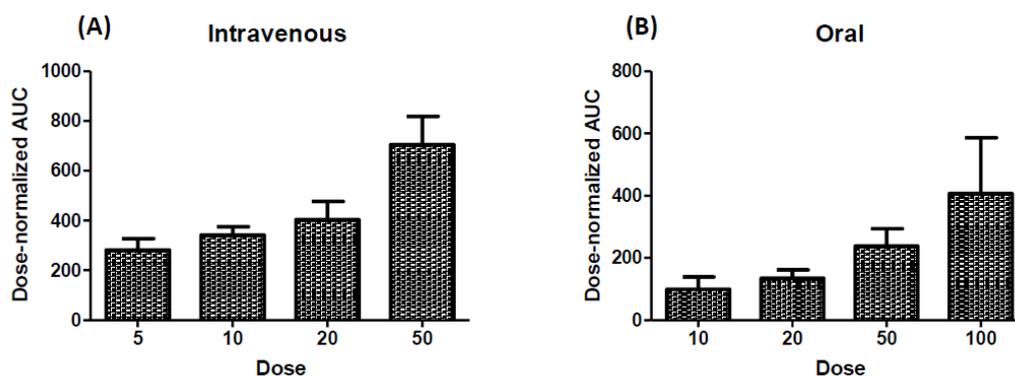


Figure S1. Mean dose (mg/kg) versus dose-normalized AUC (μg·min/mL) of tofacitinib in Sprague-Dawley rats after (A) 1-min intravenous infusion of 5 ($n = 9$), 10 ($n = 8$), 20 ($n = 7$), and 50 ($n = 7$) mg/kg tofacitinib and (B) oral administration of 10 ($n = 7$), 20 ($n = 8$), 50 ($n = 9$), and 100 ($n = 7$) mg/kg tofacitinib. Bars represent standard deviations (SD). (A) 20 mg/kg was significantly different ($p < 0.05$) from 5 mg/kg. 50 mg/kg was significantly different ($p < 0.001$) from 5, 10 and 20 mg/kg. (B) 10 mg/kg was significantly different ($p < 0.05$) from 50 mg/kg. 100 mg/kg was significantly different from 10 ($p < 0.001$), 20 ($p < 0.001$) and 50 ($p < 0.01$) mg/kg, respectively.

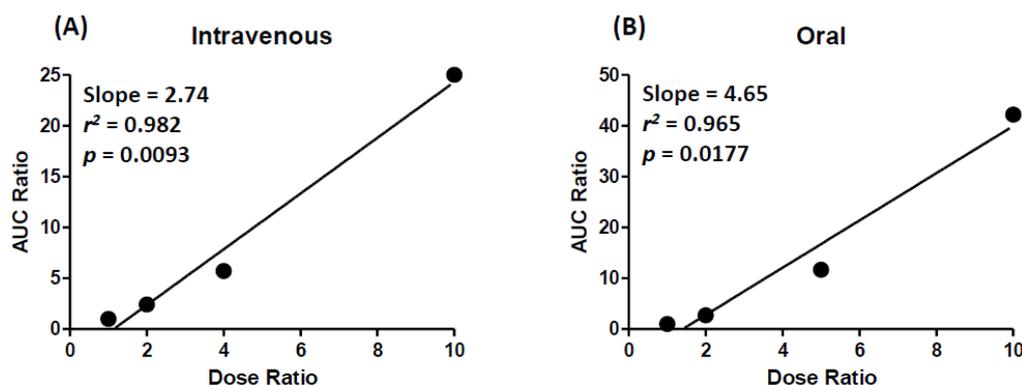


Figure S2. Plots of dose versus AUC of tofacitinib in Sprague-Dawley rats after (A) 1-min intravenous infusion of 5, 10, 20, and 50 mg/kg tofacitinib and (B) oral administration of 10, 20, 50, and 100 mg/kg tofacitinib. Dose and AUC ratios were calculated based on 5 and 10 mg/kg dose and respective AUC for intravenous and oral administration, respectively.