

Supplementary Table 1: Pharmacokinetic Parameters in WT and TKO Mice after Oral Dosing with 15 mg/kg Dasatinib (Data presented as Mean \pm S.E. of Estimate)

Mouse Genotype	Tissue	T _{max} (hr)	C _{max} (μg/mL)	AUC _{last} ¹ (μg.hr/mL)	Kp ²	Kp Ratio ³
WT	Plasma	2	0.417 \pm 0.189	1.31 \pm 0.318	0.12	7.6
WT	Brain	4	0.052 \pm 0.039	0.160 \pm 0.078		
TKO	Plasma	0.5	0.451 \pm 0.053	1.78 \pm 0.240 [#]	0.93	
TKO	Brain	6	0.161 \pm 0.048	1.63 \pm 0.241 [*]		

1. Area under the curve from time zero to 12 hour post dose
2. Kp = AUC_{brain}/AUC_{plasma}
3. Kp Ratio = Kp in KO Mice / Kp in WT Mice
4. [#], p > 0.05 compared to WT plasma
5. ^{*}, p = 0.001 compared to WT brain

Pharmacokinetic Parameters in WT and TKO Mice after Oral Dosing with 15 mg/kg Dasatinib (Data presented as Mean \pm S.E. of Estimate). The total plasma exposure in wild-type mice is not significantly different as compared to the *Mdr1a/b(-/-)Bcrp1(-/-)* mice. However, the total brain exposure in *Mdr1a/b(-/-)Bcrp1(-/-)* mice is ~10 fold higher than the wild-type mice, resulting in a drug targeting index of 7.6.