

Table S1 Pharmacokinetic parameters of metformin

	C _{max} (μ M)	T _{max} (h)	AUC _{inf} ^a (μ M \times h)	f _e ^b	T _{1/2} (h)	CL/F ^c (ml/min)	CL _{r,p} ^d (ml/min)	F ^e	V _d /F ^f (L)
Control	12.0 \pm 2.8	2.67 \pm 0.99	75.0 \pm 13.6	0.683 \pm 0.110	4.83 \pm 0.56	692 \pm 130	471 \pm 52.7	0.542 \pm 0.086	266 \pm 55
+PYR10mg	11.5 \pm 2.6	2.00 \pm 0.74	78.7 \pm 12.0	0.619 \pm 0.125	6.98 \pm 1.45	654 \pm 102	416 \pm 61.7	0.505 \pm 0.097	269 \pm 46
+PYR25mg	15.3 \pm 6.9	2.83 \pm 1.03	97.2 \pm 26.7	0.566 \pm 0.232	6.92 \pm 1.56	552 \pm 143	317 \pm 74.7	0.461 \pm 0.094	233 \pm 68
+PYR75mg	18.5 \pm 7.7	3.00 \pm 1.04	132 \pm 35	0.530 \pm 0.146	6.23 \pm 0.92	406 \pm 113	213 \pm 31.7	0.430 \pm 0.113	180 \pm 61

^a. AUC_{inf} was calculated as $AUC_{0-24h} + C_{24h}/(\ln 2/T_{1/2})$

^b. f_e was calculated as $X_{urine}(0-24h)/Dose$

^c. CL/F was calculated as $Dose/AUC_{inf}$

^d. CL_{r,p} was calculated as $X_{urine}(0-24h)/AUC_{0-24h}$

^e. F was calculated as $CL_r/(CL/F)$

^f. V_d/F was calculated as $CL/F \times MRT$

Each parameter was shown as arithmetic mean \pm SEM. Statistical significance in the difference in the pharmacokinetic parameters between control and pyrimethamine-treated conditions was shown in Supplemental TableS2.

Parameters: C_{max}, maximum plasma concentration; T_{max}, time to achieve the maximum concentration; AUC_{inf}, AUC from time 0 to infinity; f_e, fraction recovered in the urine; T_{1/2}, half-life at the terminal phase; CL/F, oral clearance; CL_r, renal clearance; F, bioavailability; V_d/F, apparent volume of distribution; MRT, mean residence time