

SUPPLEMENTAL TABLE 1. The effect of cyclosporine A (CsA) on [¹¹C]-radioactivity in the plasma present as verapamil or its metabolites. *, P<0.05.

		9 Minutes					
Subject	CsA dose	Before CsA			During CsA		
Identifier	(mg/kg/hr)	Verapamil	D-617/D-717	Polar metabolites	Verapamil	D-617/D-717	Polar metabolites
02088	12	33	30	37	41	37	22
02054	12	47	24	29	28	51	21
04052	24	59	17	24	61	23	16
02091	24	55	15	30	48	34	19
Mean		48	21	30	44	36	20*
SD		12	7	5	13	11	3

20 Minutes							
Subject	CsA dose	Before CsA			During CsA		
Identifier	(mg/kg/hr)	Verapamil	D-617/D-717	Polar metabolites	Verapamil	D-617/D-717	Polar metabolites
02088	12	21	41	38	14	59	27
02054	12	25	38	36	17	62	21
04052	24	34	33	33	32	39	28
02091	24	32	29	39	23	52	24
Mean		28	35	37	21.8	53	25*
SD		6	5	3	8.1	10	3

40 Minutes							
Subject	CsA dose	Before CsA			During CsA		
Identifier	(mg/kg/hr)	Verapamil	D-617/D-717	Polar metabolites	Verapamil	D-617/D-717	Polar metabolites
02088	12	10	51	40	9	66	25
02054	12	17	44	39	11	65	24
04052	24	21	44	35	17	41	42
02091	24	15	61	24	13	63	24
Mean		16	50	34	13	59	29
SD		5	8	7	4	12	8

SUPPLEMENTAL TABLE 2. The effect of cyclosporine A (CsA) on plasma and tissue [¹¹C]-radioactivity. Data are expressed as mean \pm SD (n=4) and were analyzed by Mann-Whitney-test. High variability in spinal and uterine wall radioactivity concentrations at early time points prevented accurate determination of t_{max} and C_{max} of this tissue. *, P<0.05.

ID-normalized AUC₍₀₋₉₎ (min/L)

Tissue	Before CsA	During CsA	% Change
Brain	1.02 \pm 0.16	3.45 \pm 0.69	238 \pm 49*
Fetal liver	1.02 \pm 0.28	1.97 \pm 0.57	108 \pm 90*
Maternal liver	6.03 \pm 0.87	4.92 \pm 1.00	-19 \pm 7
Gallbladder	13.69 \pm 3.97	14.21 \pm 4.24	6 \pm 27
Kidneys	7.13 \pm 1.29	7.70 \pm 1.26	9 \pm 12
Heart	4.82 \pm 1.12	4.98 \pm 0.71	5 \pm 13
Lungs	3.30 \pm 1.03	3.49 \pm 0.84	8 \pm 9
Vertebrae	5.37 \pm 2.08	4.75 \pm 1.44	-7 \pm 16
Spleen	6.82 \pm 1.43	7.30 \pm 1.86	2 \pm 2
Uterine wall	2.55 \pm 1.16	2.49 \pm 0.75	7 \pm 22
Placenta	1.07 \pm 0.28	1.27 \pm 0.24	20 \pm 9
Plasma	1.10 \pm 0.08	0.99 \pm 0.12	-8 \pm 14

Tmax (min)

Tissue	Before CsA	During CsA	% change
Brain	1.01 \pm 0.45	8.25 \pm 2.06	833 \pm 410*
Fetal liver	9.50 \pm 2.94	9.50 \pm 3.65	0 \pm 25
Maternal liver	10.50 \pm 2.16	8.75 \pm 3.20	-18 \pm 15
Gallbladder	35.50 \pm 14.00	27.75 \pm 14.86	-22 \pm 36
Kidneys	1.42 \pm 0.64	1.59 \pm 0.56	20 \pm 28
Heart	0.80 \pm 0.41	1.03 \pm 0.43	38 \pm 25
Lungs	0.76 \pm 0.38	1.05 \pm 0.44	45 \pm 23
Spleen	1.63 \pm 0.48	1.79 \pm 0.70	9 \pm 19
Placenta	1.54 \pm 0.42	4.54 \pm 2.76	214 \pm 186
Plasma	0.86 \pm 0.29	0.90 \pm 0.27	6 \pm 18

ID-normalized Cmax (L⁻¹)

Tissue	Before CsA	During CsA	% change
Brain	0.14 \pm 0.02	0.45 \pm 0.10	209 \pm 42*
Fetal liver	0.18 \pm 0.07	0.35 \pm 0.08	106 \pm 72
Maternal liver	0.88 \pm 0.10	0.71 \pm 0.10	-19 \pm 3
Gallbladder	14.64 \pm 9.18	3.07 \pm 1.25	-61 \pm 42
Kidneys	1.26 \pm 0.14	1.46 \pm 0.16	16 \pm 18
Heart	0.86 \pm 0.08	0.87 \pm 0.07	3 \pm 13
Lungs	0.84 \pm 0.22	0.88 \pm 0.15	8 \pm 15
Spleen	1.14 \pm 0.27	1.17 \pm 0.39	-2 \pm 7
Placenta	0.18 \pm 0.07	0.18 \pm 0.03	8 \pm 21
Plasma	0.73 \pm 0.27	0.61 \pm 0.22	-9 \pm 46