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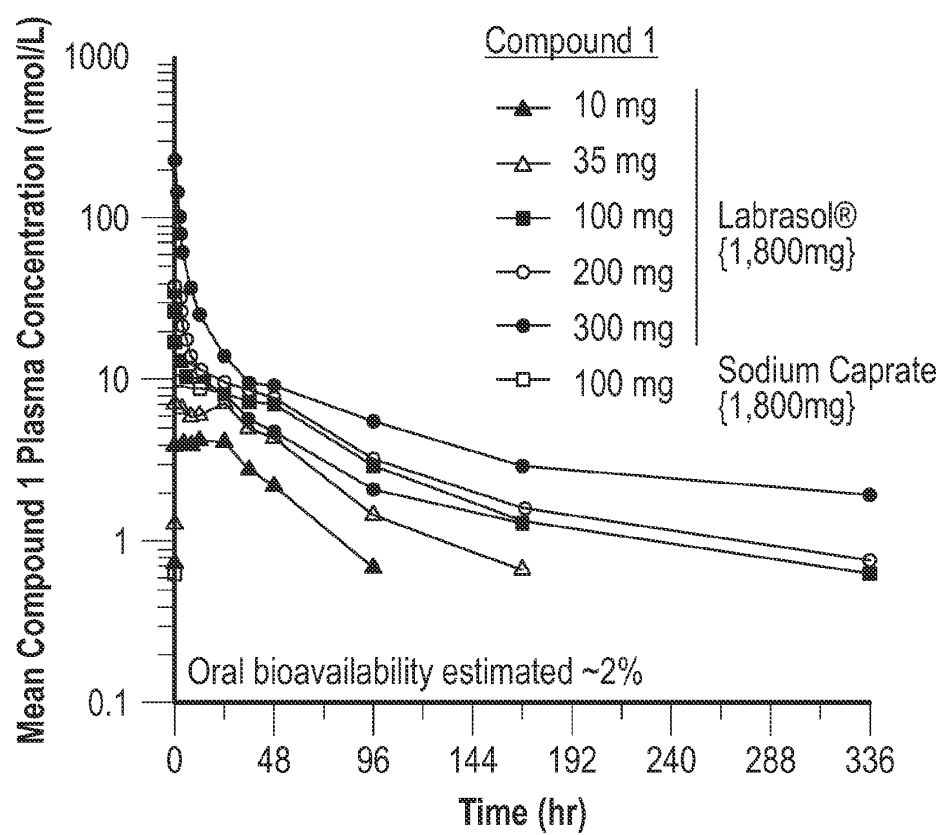


FIG. 1

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Pharmacokinetic parameters	Panel A 10 mg Period 1 N=9			Panel B 35 mg Period 1 N=9			Panel C 100 mg Period 1 N=9			Panel D 20 mg Period 1 N=9			Panel F 40 mg Period 1 N=8		
	GM	95% CI		GM	95% CI		GM	95% CI		GM	95% CI		GM	95% CI	
AUC _{0-inf} (h*nmol/L) ^a	259	(205, 328)		540	(427, 683)		1080	(852, 1360)		465	(367, 588)		439	(342, 564)	
AUC ₀₋₂₄ (h*nmol/L) ^a	93.0	(72.1, 120)		165	(128, 213)		309	(239, 398)		139	(108, 179)		133	(102, 174)	
AUC _{last} (h*nmol/L) ^a	235	(185, 297)		501	(396, 634)		1020	(805, 1290)		429	(339, 543)		402	(313, 516)	
C _{max} (nmol/L) ^a	5.21	(3.48, 7.80)		17.8	(11.9, 26.7)		46.2	(30.9, 69.0)		9.78	(6.55, 14.6)		9.86	(6.44, 15.1)	
C ₂₄ (nmol/L) ^a	3.90	(3.28, 4.65)		6.98	(5.86, 8.32)		8.47	(7.11, 10.1)		6.11	(5.13, 7.28)		5.58	(4.64, 6.73)	
T _{max} (h) ^b	1.50	(0.50, 24.08)		1.50	(1.50, 1.50)		1.50	(1.00, 3.00)		1.50	(1.00, 12.00)		1.25	(1.00, 12.00)	
t _{1/2} (h) ^c	35.14	17.5		42.86	5.8		81.52	16.9		45.71	36.6		43.33	15.2	
CL/F (L/h) ^c	24.98	40.5		42.96	26.7		59.83	29.9		27.74	47.1		58.71	53.7	
Vz/F (L) ^c	1266.52	39.4		2656.39	26.0		7037.04	24.0		1829.27	25.9		3669.46	37.9	
Pharmacokinetic parameters	Panel A 200 mg Period 2 N=8			Panel B 300 mg Period 2 N=9			Panel C 100 mg Period 2 N=9			Panel D 40 mg Period 2 N=9			Panel F 40 mg Period 2 N=8		
	GM	95% CI		GM	95% CI		GM	95% CI		GM	95% CI		GM	95% CI	
AUC _{0-inf} (h*nmol/L) ^a	1260	(984, 1620)		2260	(1790, 2850)		979	(774, 1240)		758	(599, 959)		505	(394, 649)	
AUC ₀₋₂₄ (h*nmol/L) ^a	339	(259, 443)		778	(603, 1000)		281	(218, 363)		171	(132, 220)		138	(106, 181)	
AUC _{last} (h*nmol/L) ^a	1170	(910, 1500)		2000	(1580, 2540)		855	(675, 1080)		694	(548, 879)		454	(353, 583)	
C _{max} (nmol/L) ^a	45.3	(29.5, 69.5)		149	(99.5, 223)		41.3	(27.6, 61.6)		14.3	(9.57, 21.4)		9.21	(6.02, 14.1)	
C ₂₄ (nmol/L) ^a	9.91	(8.23, 11.9)		13.3	(11.2, 15.9)		8.22	(6.90, 9.80)		7.80	(6.55, 9.30)		6.03	(5.01, 7.27)	
T _{max} (h) ^b	2.02	(1.07, 5.00)		1.50	(1.00, 1.98)		1.50	(1.00, 3.00)		1.50	(1.00, 2.00)		0.75	(0.50, 2.00)	
t _{1/2} (h) ^c	95.47	27.7		129.95	40.5		56.80	10.2		97.20	31.8		46.37	23.7	
CL/F (L/h) ^c	103.17	34.3		84.64	52.2		65.87	30.0		34.03	29.8		51.03	31.9	
Vz/F (L) ^c	14209.96	42.3		15868.31	65.9		5397.58	26.4		4771.73	18.0		3414.17	27.0	

FIG. 2

Pharmacokinetic parameters	Panel A 200 mg Period 3 N=8		Panel B 120 mg Period 3 N=9		Panel C 40 mg Period 3 N=8			
	GM	95% CI	GM	95% CI	GM	95% CI		
AUC _{0-inf} (h*nmol/L) ^a	562	(438, 721)	994	(787, 1260)	538	(420, 690)		
AUC ₀₋₂₄ (h*nmol/L) ^a	131	(100, 172)	333	(258, 429)	156	(119, 204)		
AUC _{last} (h*nmol/L) ^a	487	(379, 625)	861	(680, 1090)	489	(381, 629)		
C _{max} (nmol/L) ^a	7.84	(5.11, 12.0)	57.7	(38.5, 86.4)	13.5	(8.81, 20.7)		
C ₂₄ (nmol/L) ^a	6.33	(5.26, 7.63)	8.35	(7.00, 9.95)	6.55	(5.44, 7.89)		
T _{max} (h) ^b	14.53	(0.50, 36.00)	1.48	(1.00, 1.50)	1.28	(1.00, 2.00)		
t _{1/2} (h) ^c	56.41	12.0	61.78	13.9	47.59	9.7		
CL/F (L/h) ^c	231.82	26.3	75.88	33.4	47.62	28.5		
V _z /F (L) ^c	18866.57	22.1	6763.21	29.7	3269.38	32.1		

FIG. 2 (cont.)

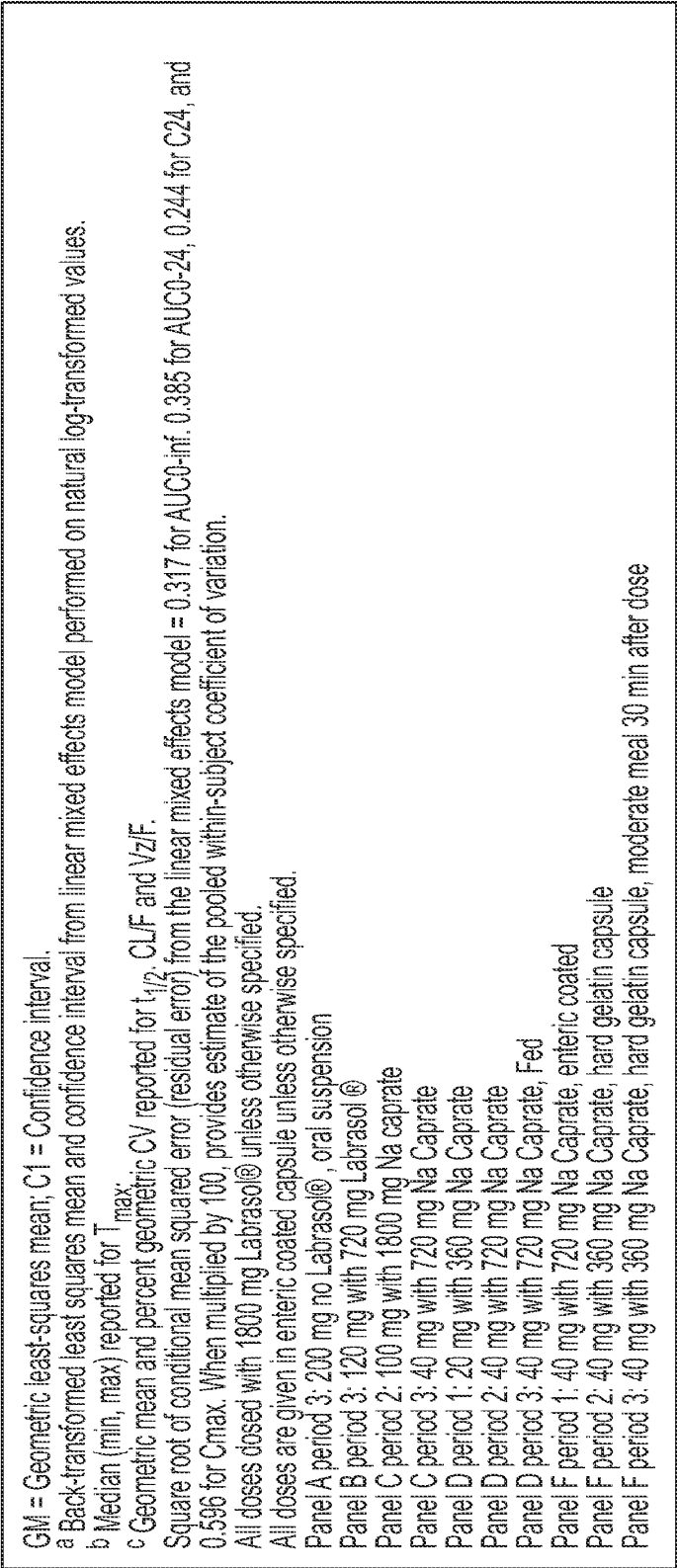


FIG. 2 (cont.)

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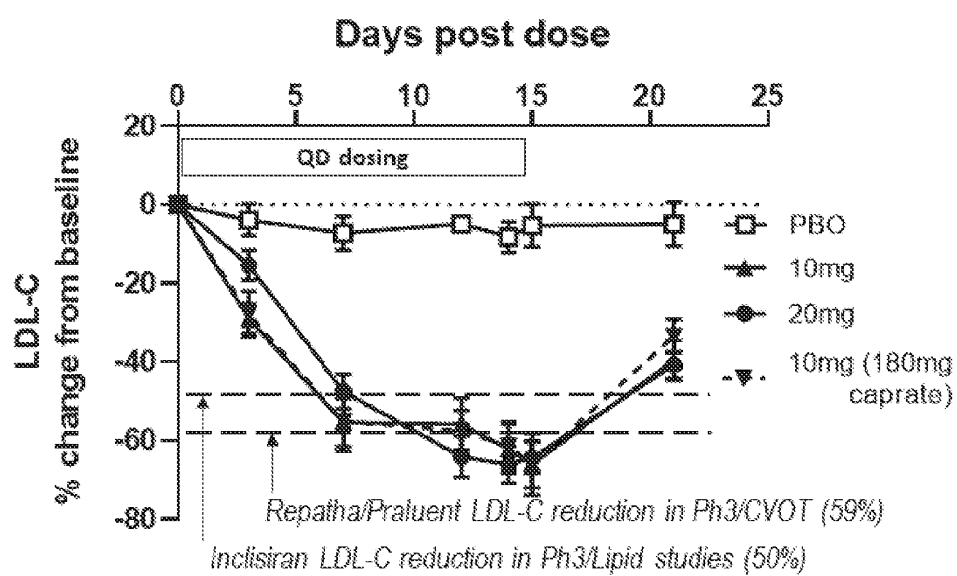


FIG. 3

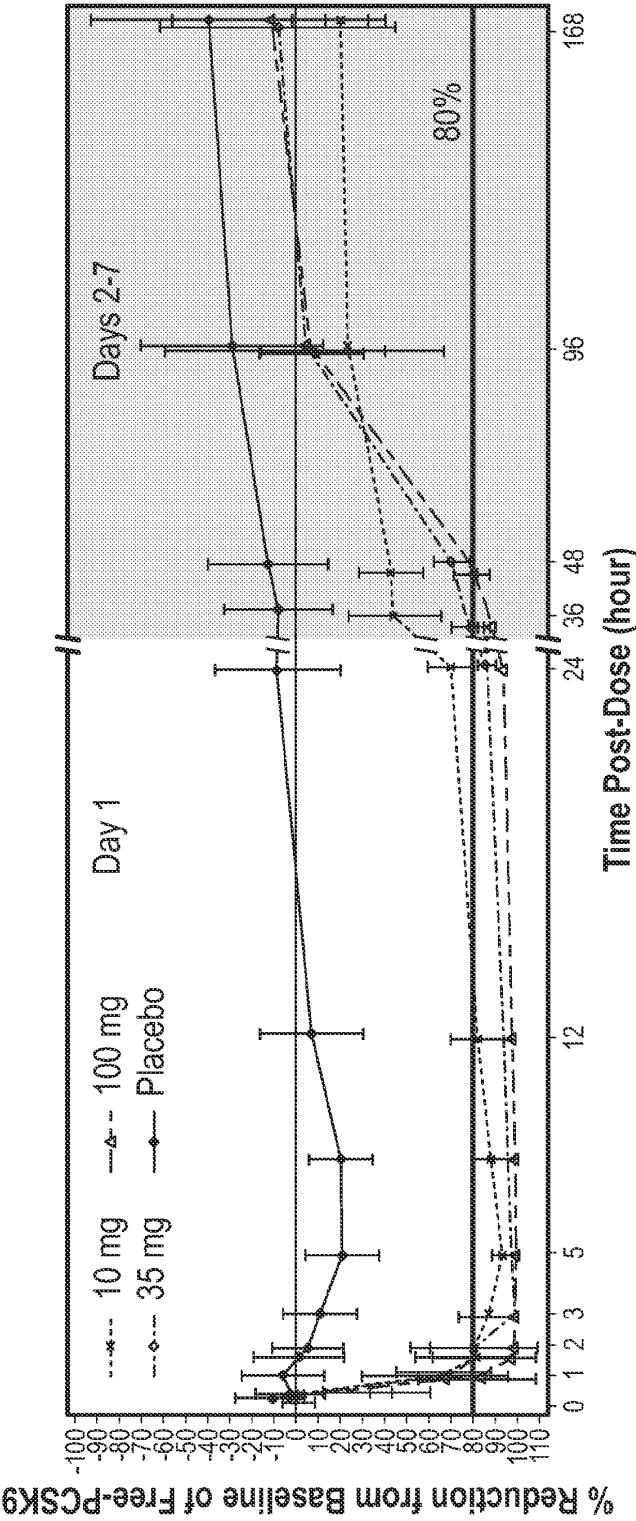


FIG. 4

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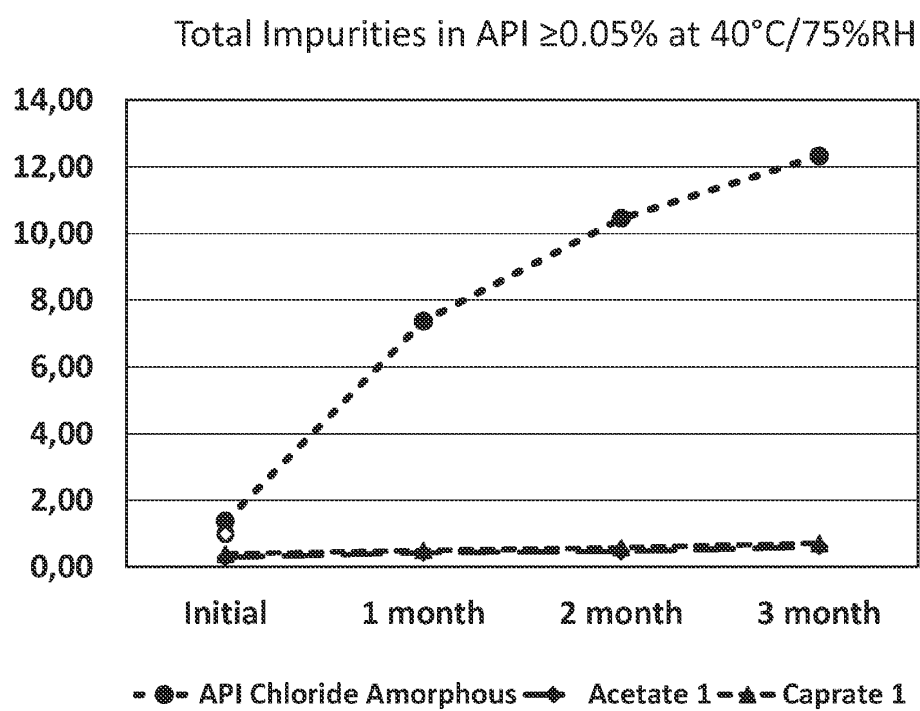


FIG. 5