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Table C (Table 6 of manuscript). C_{max} values after each dose from Elcombe et al. (2013)

Patient	Daily Dose mg/kg-day*	C _{max} after each weekly dose in μM					
	week>	1	2	3	4	5	6
1	0.67	25.72	na	na	na	na	na
2	0.67	29.79	na	na	na	na	na
3	0.67	24.64	na	na	na	na	na
4	0.10	19.95	40.37	40.6	52.28	77.49	81.07
	Avg	25	40	41	52	77	81
5	0.19	23.66	50.82	80.2	87.35	100.84	109.1
6	0.19	32.32	47.47	70.55	97	89.54	179.07
7	0.19	30.91	-	55.78	73.03	-	-
	Avg	29	49	69	86	95	144
8	0.38	114.25	171.02	276.84	368.27	426.16	414.33
9	0.38	93.43	170.29	253.19	362.32	471.59	373.31
10	0.38	58.6	119.44	181.86	276.15	256.06	232.44
	Avg	89	154	237	336	385	340
11	0.57	111.65	178.42	237.26	288.21	326.13	386.77
12	0.57	122.9	182.32	240.93	303.06	372.99	-
13	0.57	85.32	-	-	-	-	-
14	0.57	131.24	179.97	297.35	420.49	478.38	562.63
	Avg	113	180	259	337	393	475
15	0.86	231.36	324.96	463.43	578.86	707.8	800.55
16	0.86	164.05	348.41	545.74	721.48	906.59	-
17	0.86	163.18	276.16	341.96	427.08	497.22	525.98
	Avg	186	317	450	576	704	663
18	1.1	338.52	406.73	590.95	-	-	-

20	1.1	413.39	327.38	474.01	562.88	651.85	770.32
21	1.1	203.29	504.5	652.79	734.36	847.13	995.39
22	1.1	198.74	309.8	433.41	595.95	-	-
23	1.1	236.13	400.07	635.73	-	-	-
24	1.1	282.55	488.31	691.46	858.92	813.92	966.13
25	1.1	230	360	480	640	750	780
	Avg	272	400	565	678	766	878
26	1.4	200.07	397.76	624.63	625.39	732.46	823.68
27	1.4	240.51	410.69	569.22	719.7	811.16	-
28	1.4	206.86	321.26	472.99	654.6	757.67	853.05
	Avg	216	377	556	667	767	838
29	1.8	352.58	606.03	896.3	896.9	971.71	1043.2
30	1.8	332.61	-	-	-	-	-
31	1.8	347.52	554.28	799.77	998.35	1031.14	-
32	1.8	291.69	516.7	-	-	-	-
40	1.9	189.71	367.81	487.42	554.18	697.26	826.44
41	1.9	232.54	412.52	558.23	748.03	802.5	1209.31
42	1.9	358.73	585.96	764.91	1231.51	1281.13	1251.9
	Avg	301	507	701	886	957	1083
33	2.3	441.43	734.84	925.6	1172.58	1231.36	1317.84
34	2.3	559.64	893.14	1115.82	1440.82	1448.79	-
35	2.3	316.74	592.29	704.4	1172.95	-	-
36	2.3	708.42	679.68	968.95	1143.19	-	1293.03
37	2.3	418.44	841.24	1135.41	1393.91	1530.33	-
38	2.3	314.43	538.47	808.36	787.75	931.5	958.1
	Avg	460	713	943	1185	1285	1190

* Doses given in mg/week. Mg/kg-day doses are determined from average body weight of 75 kg as stated by Convertino et al. (2018) and further dividing by seven days/week.

TABLE D. Average Cmax values after each dose (data not used in manuscript)

Daily Dose mg/kg-day*	Average Cmax after each weekly dose in μM					
	1	2	3	4	5	6
0.1	25	40	41	50	77	80
0.19	29	49	67	86	95	144
0.38	89	154	201	336	385	340
0.57	113	180	259	329	393	475
0.86	186	317	425	576	704	663
1.1	279	398	572	688	770	911
1.4	216	377	556	667	767	838
1.8 & 1.9	301	487	673	877	957	1083
2.3	460	713	937	1185	1285	1190

* Doses given in mg/week. Mg/kg-day doses are determined from average body weight of 75 kg as stated by Convertino et al. (2018) and further dividing by seven days/week.

Figure of Table D (data not used in manuscript)

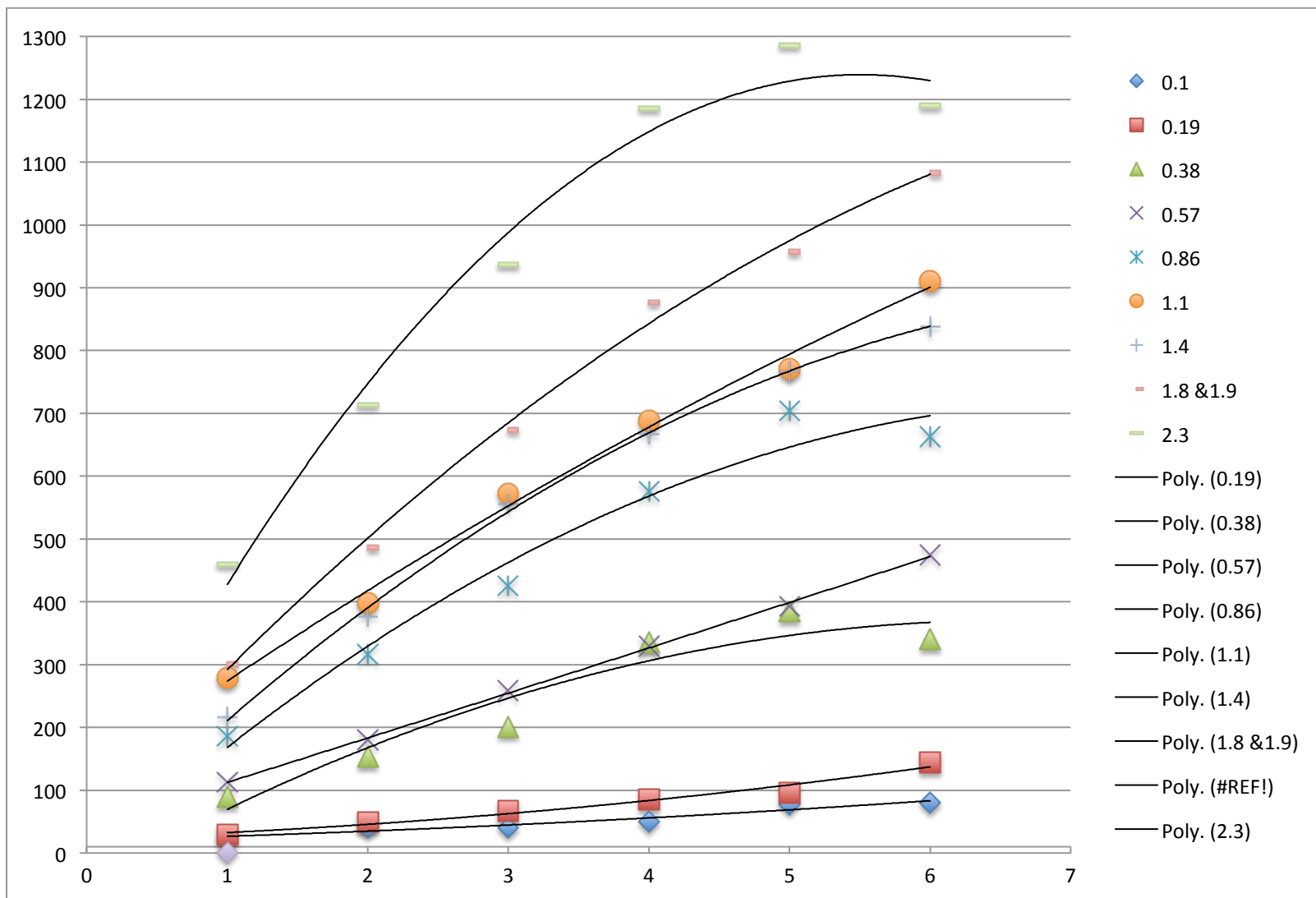
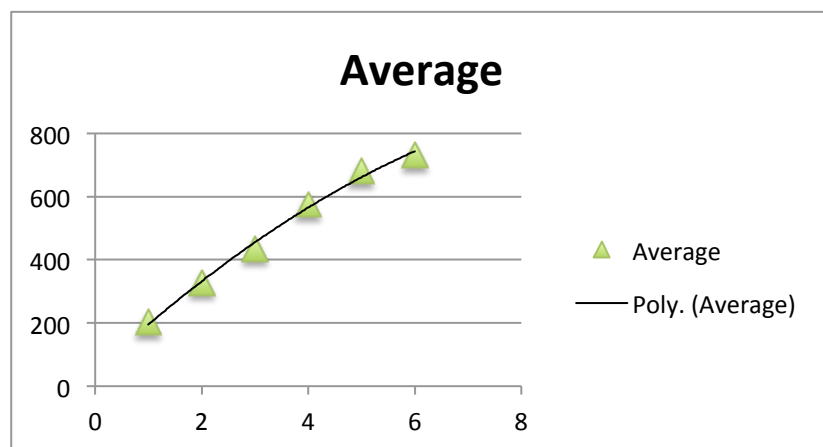


TABLE E (Table 7 of manuscript). Average Cmax values after each dose in μM per mg/kg-day.

Daily Dose mg/kg-day*	Average Cmax after each weekly dose in μM per mg/kg-day						Data not used		
	week>	1	2	3	4	5	6	Weighted Analysis	
0.1	250	404	406	504	775	801	# of patient:	Week 1	week 6
0.19	152	259	353	452	501	758	1	250	801
0.38	234	404	530	883	1012	895	2	305	1517
0.57	198	316	454	577	689	833	3	701	2684
0.86	217	368	495	670	818	771	2	396	1666
1.1	253	362	520	625	700	828	2	433	1542
1.4	154	269	397	476	548	599	4	1014	3311
1.85	163	263	364	474	517	585	2	308	1198
2.3	200	310	407	515	559	517	4	651	2341
Average	202	328	436	575	680	732	3	600	1552
							23	4657	16612
							avg. Cmax	202	722

Figure of Table E



Values were approximated from
Figure 78 of Elcombe et al. (2013)

TABLE F

Patient	Dose	6 week level	SS level	ratio SS/6
5	100	109	280	2.6
9	200	373	520	1.4
10	200	232	600	2.6
11	300	387	530	1.4
14	300	563	950	1.7
15	450	801	1050	1.3
20	600	770	1000	1.3
25	600	780	950	1.2
26	750	824	750	0.9

Value used to adjust 6 week DDEF in TERA report 1.6

TABLE G. Information from Elcombe et al. (2013) on patients beyond 6 weeks of exposure

Patient	Dose	6 week level	Measurements Past 6 weeks				Average of Ratio past/6	
			1st	2nd	3rd	4th	Past 6 wk	
5	100	109	276				276	2.5
9	200	399	471	617	501	540	532	1.3
10	200	232	570	650	660	570	613	2.6
11	300	387	574				574	1.5
14	300	562	890	980	940		937	1.7
15	450	801	1175	1000			1088	1.4
20	600	770	950				950	1.2
25	600	780	1050				1050	1.3
26	750	823	670				670	0.8

Values in yellow were read from Figure 78; other values were read from patient tables Average 1.6