Figure 1A

IMGN853 Pharmacokinetic Results

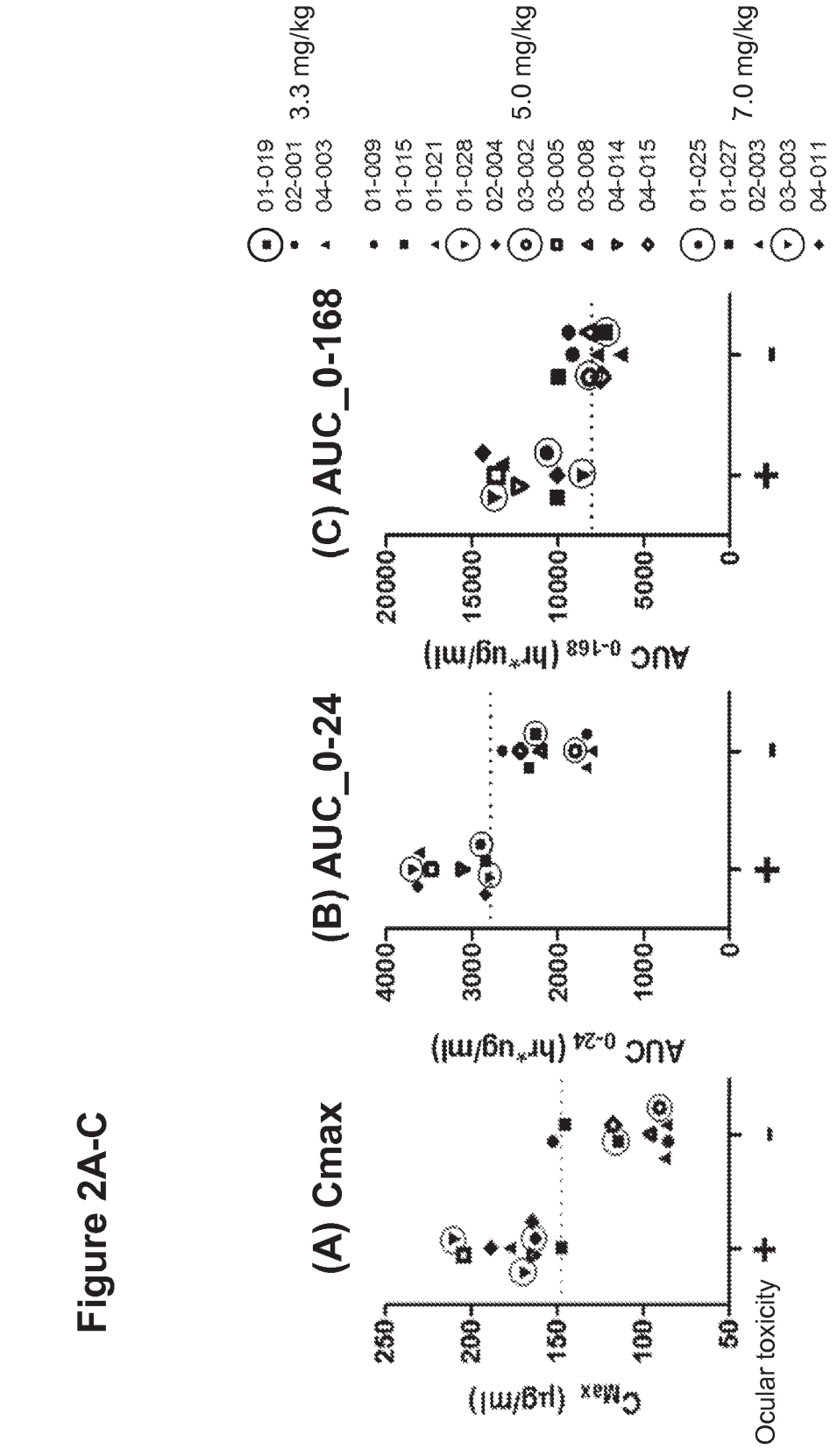
Dose (mg/kg)	0.15 (n=2)	0.5 (n=1)	1.0 (n=1)	2.0 (n=1)	3.3 (n=3)	5.0 (n=3)	7.0 (n=4)
Cmax (ug/mL)	2.9	10.5	22.1	65.7	96.6 (16.4)	108 (32.7)	179 (21.8)
Half-life (hr)	35.4	41.2	70.1	69.9	99.5 (15.7)	105 (4.4)	87.6 (11.5)
Half-life (d)	1.5	1.7	2.9	2.9	4.1 (0.65)	4.4 (1.0)	3.6 (0.5)
$AUC_{(0\text{-}inf)}$ (hr*ug/mL)	150.9	596.6	1779	6505	12188 (2581)	12708(2112)	17559(2850)
$AUC_{(0\text{-}168)}$(hr*ug/mL)	104.6	496.6	1678	5330	8178 (1129)	8254 (1771)	12177(1621)
CL (mL/hr/kg)	1.1	0.8	0.6	0.3	0.3 (0.06)	0.4 (0.07)	0.4 (0.06)
Vss (mL/kg)	51.8	46.7	54.9	28.2	38.6 (5.2)	61.2 (16.1)	52.8 (8.3)

Figure 1B

IMGN853 Pharmacokinetic Results

Cohort Dose (mg/kg)	N	C_{Max} (µg/ml)	$T_{1/2}$ (days)	Cl (ml/hr/kg)	V_{ss} (ml/kg)	AUC_{0_∞} (hr*µg/ml)
0.15	2	2.90 (± 0.47)	1.5 (± 0.2)	1.08 (± 0.38)	53.2 (± 11.2)	149 (± 52)
0.5	1	10.55	3.2	0.58	59.2	865
1	1	22.09	3.4	0.56	57.0	1,777
2	1	65.73	5.0	0.30	32.4	6,757
3.3	3	96.64 (± 16.39)	4.7 (± 0.3)	0.26 (± 0.05)	36.82 (± 5.7)	12,915 (± 2,056)
5	10	141.90 (± 41.63)	4.9 (± 1.2)	0.35 (± 0.08)	51.9 (± 17.0)	15,079 (± 3,740)
7	5	172.98 (± 23.58)	6.2 (± 1.3)	0.36 (± 0.08)	65.7 (± 8.9)	20,146 (± 4,099)

Figure 2A-C


(A) Cmax
(B) AUC_0-24
(C) AUC_0-168
C_{Max} (µg/ml)
AUC_{0-24} (hr*ug/ml)
AUC_{0-168} (hr*ug/ml)
250
200
150
100
50
4000
3000
2000
1000
0
20000
15000
10000
5000
0
Ocular toxicity
+
-
01-019
02-001
04-003
3.3 mg/kg
01-009
01-015
01-021
01-028
02-004
03-002
03-005
03-008
04-014
04-015
5.0 mg/kg
01-025
01-027
02-003
03-003
04-011
7.0 mg/kg
= Objective Response

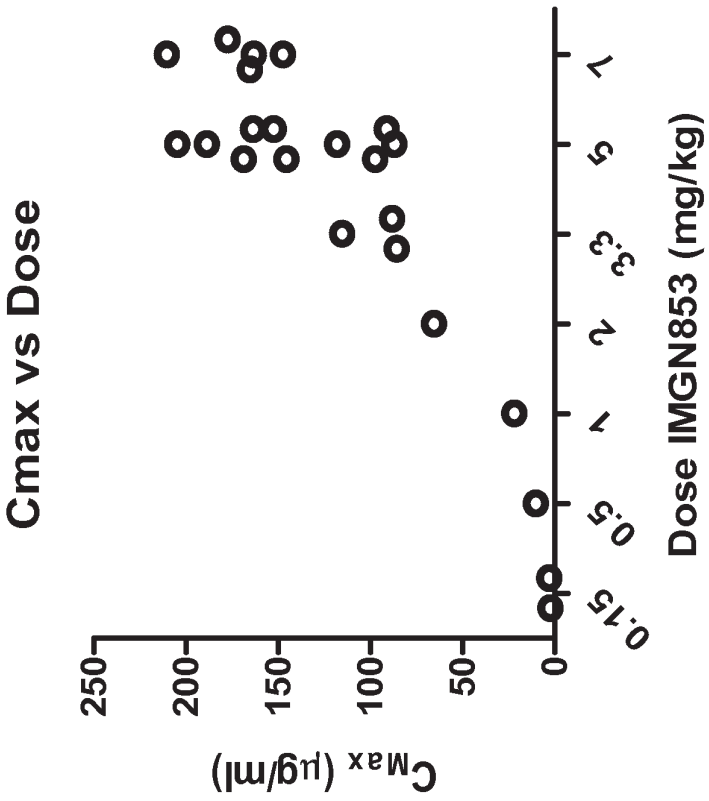

Cmax vs Dose
C_{max} (µg/ml)
250
200
150
100
50
0
0.15
0.5
1
2
3.3
5
1
Dose IMGN853 (mg/kg)

Figure 3

Figure 4

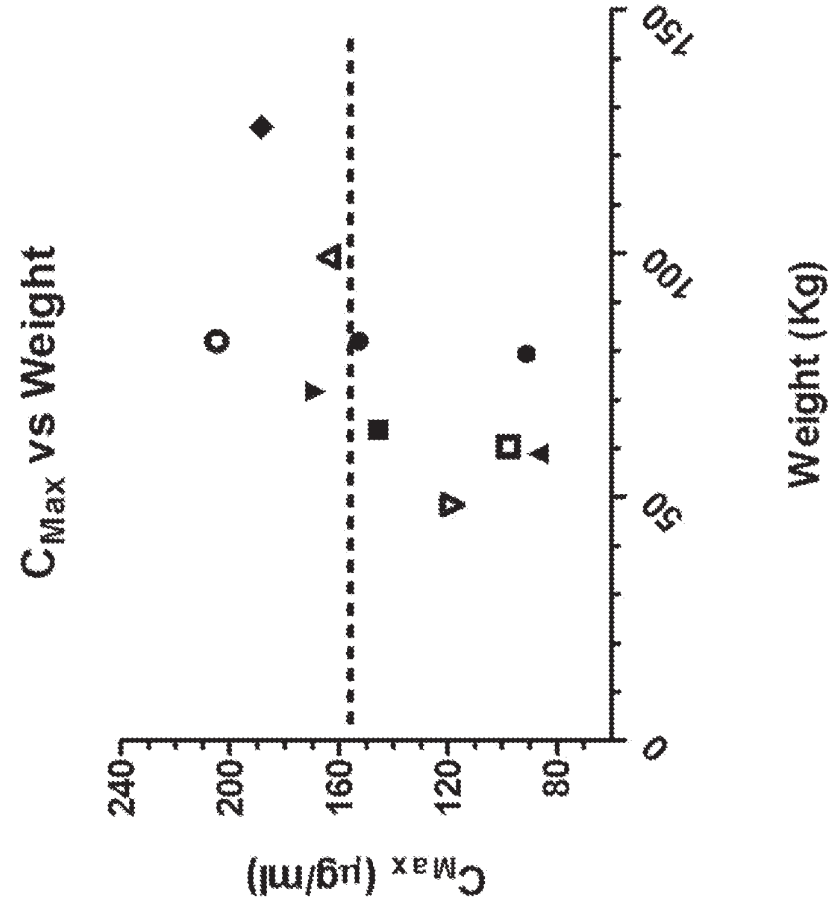
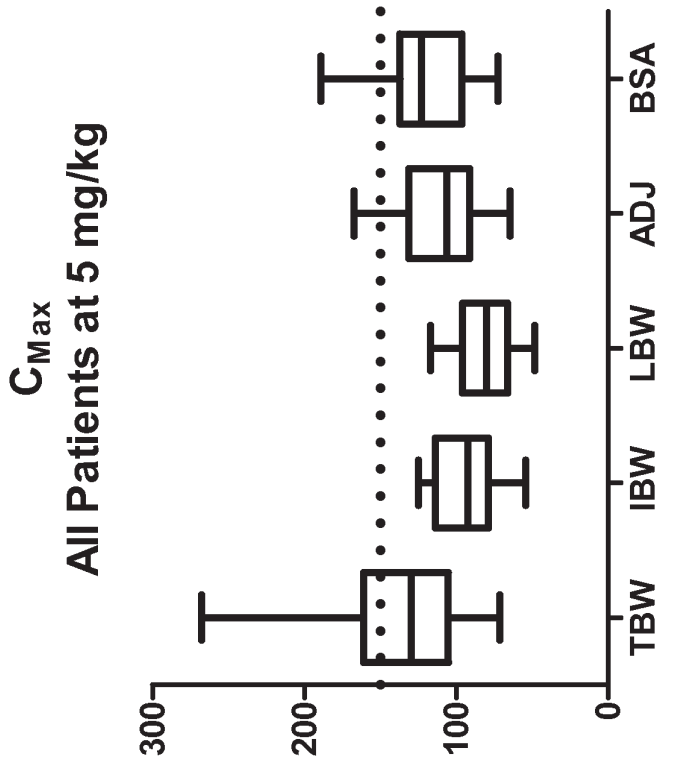

C_{Max} vs Weight
C_{Max} (µg/ml)
240
200
160
120
80
0
50
100
150
Weight (Kg)

Figure 5


C_{Max}
All Patients at 5 mg/kg
C_{Max} (µg/ml)
300
200
100
0
TBW
IBW
LBW
ADJ
BSA

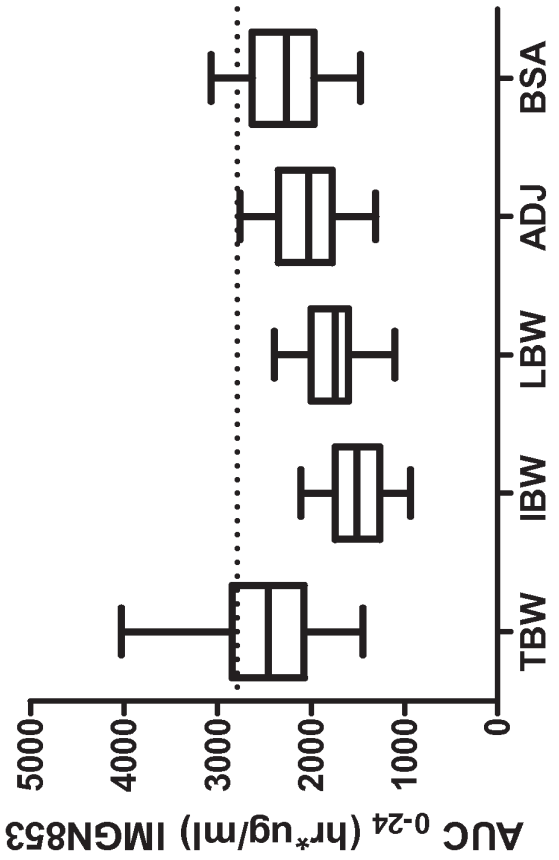

AUC_{0-24}
All Patients at 5 mg/kg
AUC_{0-24} (hr*ug/ml) IMGN853
5000
4000
3000
2000
1000
0
TBW
IBW
LBW
ADJ
BSA

Figure 6

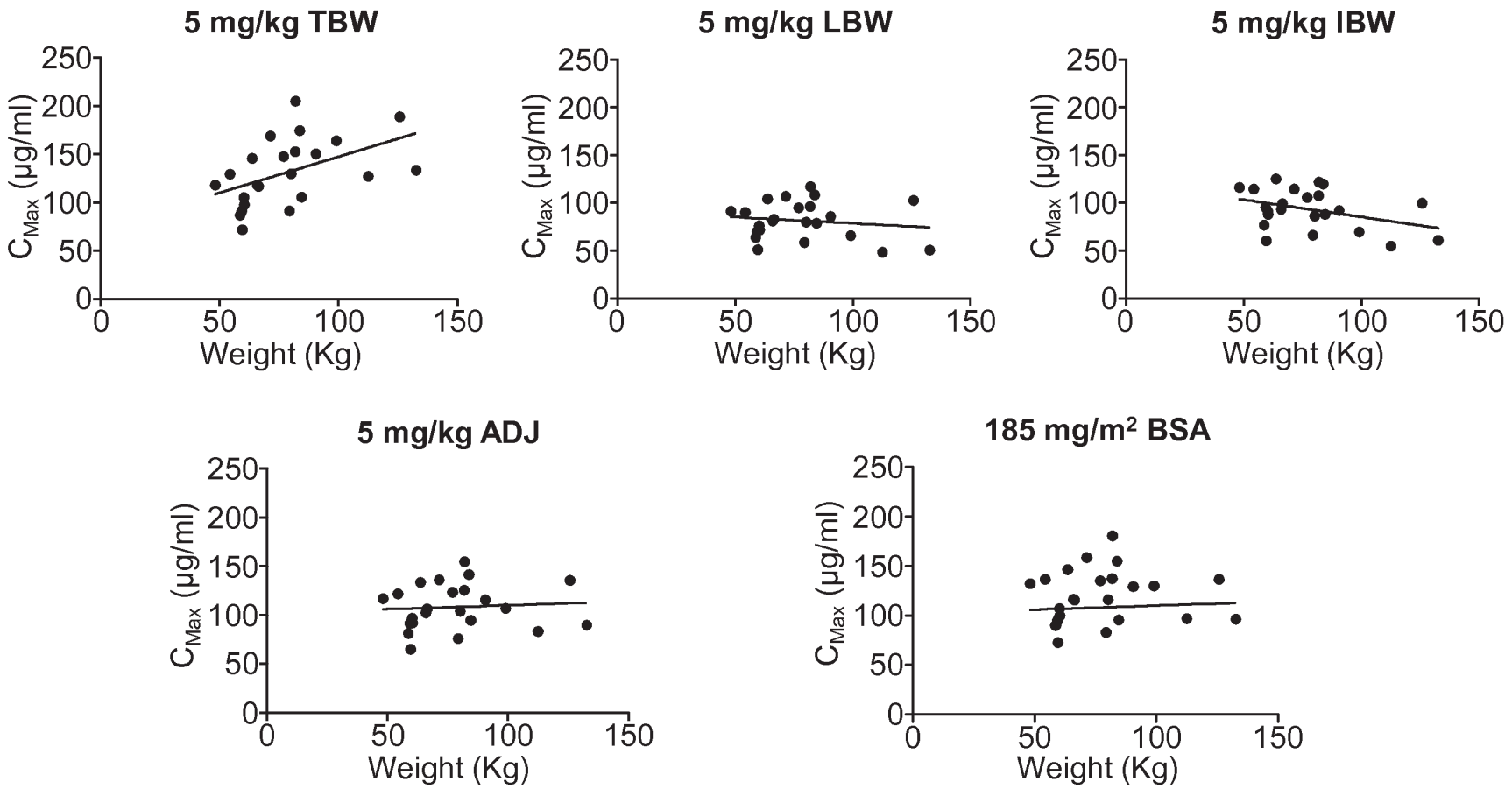
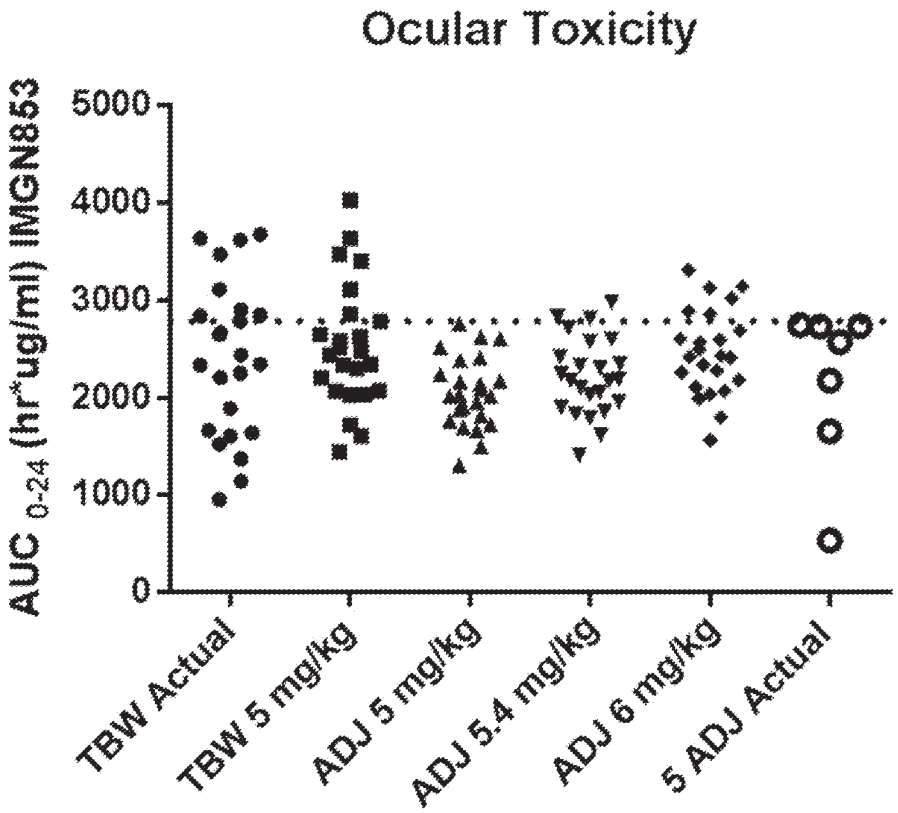

5 mg/kg TBW
5 mg/kg LBW
5 mg/kg IBW
5 mg/kg ADJ
185 mg/m² BSA
C_Max (µg/ml)
Weight (Kg)
0
50
100
150
200
250

Figure 7



			Percent of Patients Above Threshold					
Readout	PK Parameter	Threshold (hr*ug/ml)	TBW Actual	TBW 5 mg/kg	ADJ 5 mg/kg	ADJ 5.4 mg/kg	ADJ 6 mg/kg	5 ADJ Actual
Ocular Toxicity	AUC_0-24	>2741	38%	29%	0%	13%	25%	14%*

* - Patient @ 2749 hr*ug/ml

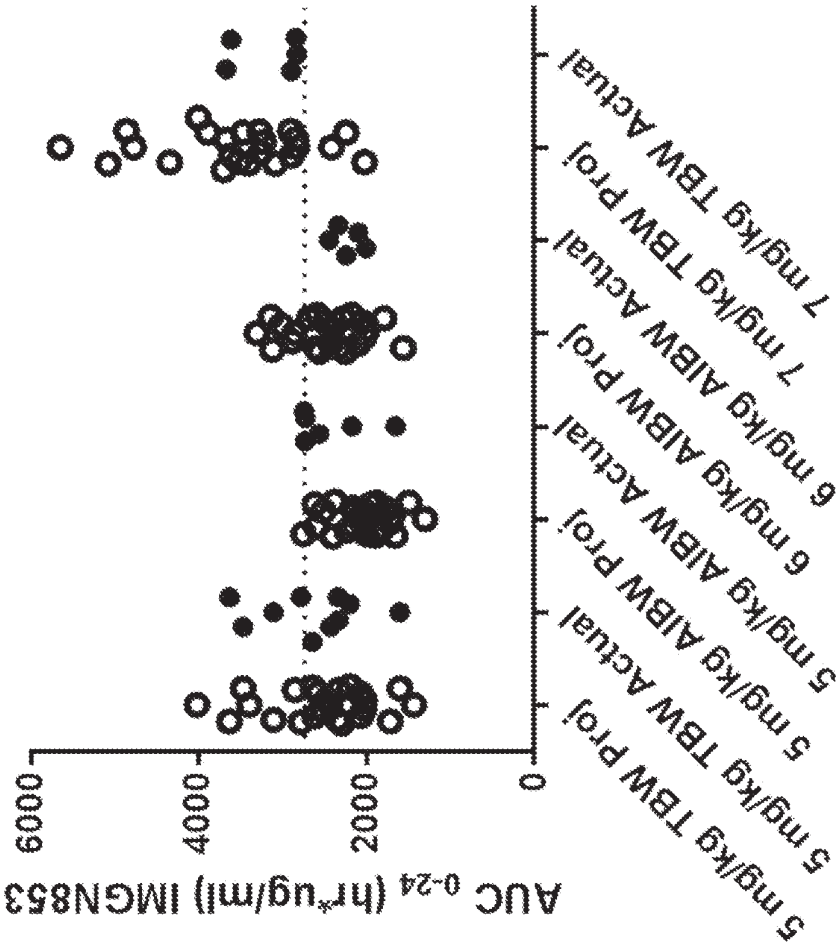
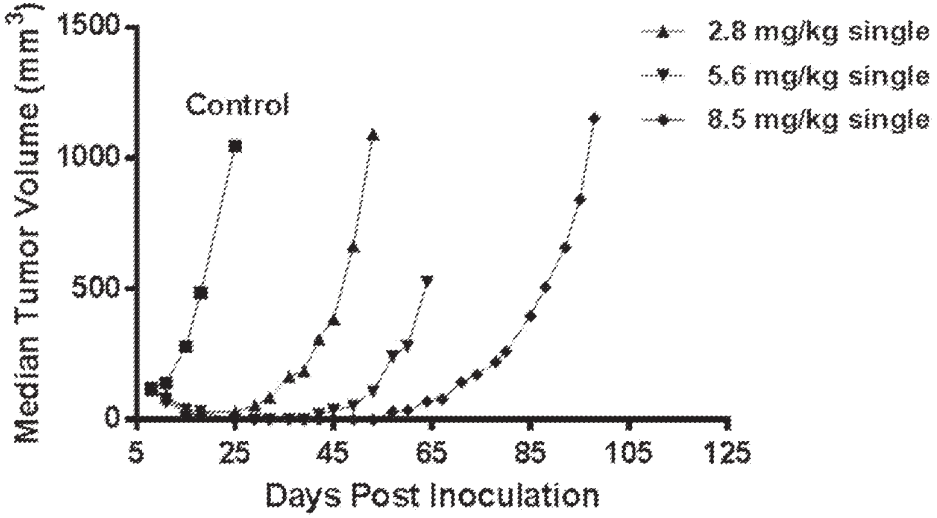
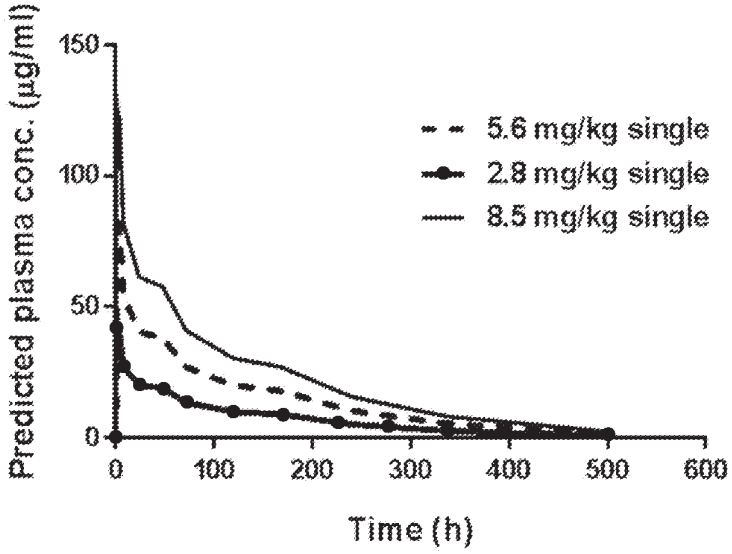

AUC0-24 Projected vs Actual Data
AUC 0-24 (hr*ug/ml) IMGN853
6000
4000
2000
0
5 mg/kg TBW Proj
5 mg/kg TBW Actual
5 mg/kg AIBW Proj
5 mg/kg AIBW Actual
6 mg/kg AIBW Proj
6 mg/kg AIBW Actual
7 mg/kg TBW Proj
7 mg/kg TBW Actual

Figure 8

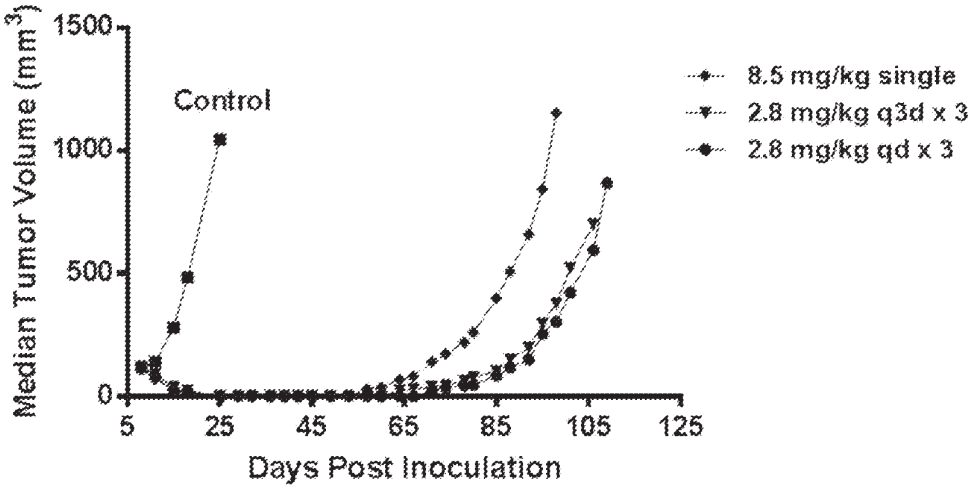
Figure 9

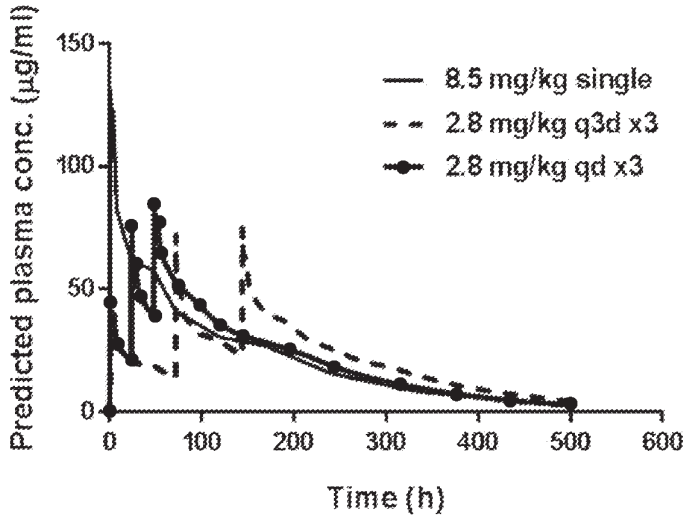




Dose (mg/kg)	Schedule	T/C (%)	Log cell kill	CR	Cmax (µg/ml)	Cavg (µg/ml)	AUC 0-540 (hr*µg/ml)
2.8	Single	3	1.5	2/6	43	7	3740
5.6	Single	0	2.7	4/6	86	15	7481
8.5	Single	0	3.9	6/6	131	23	11354

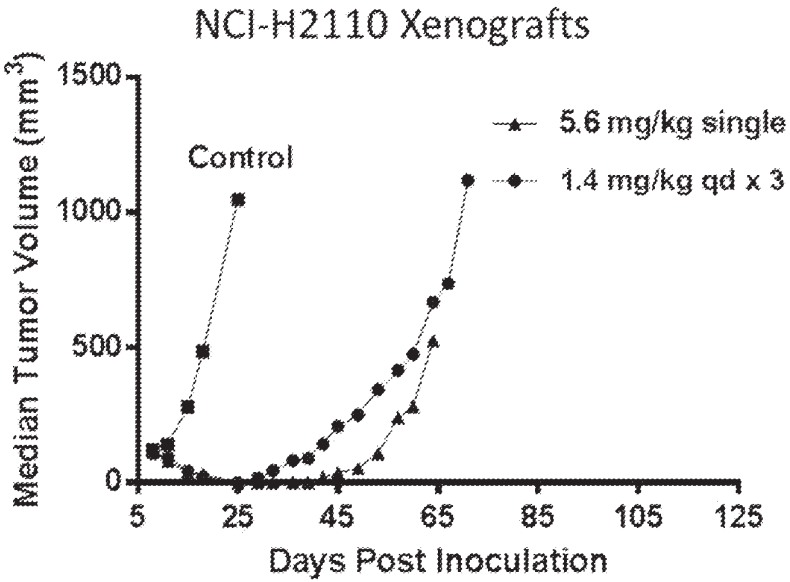
Figure 10

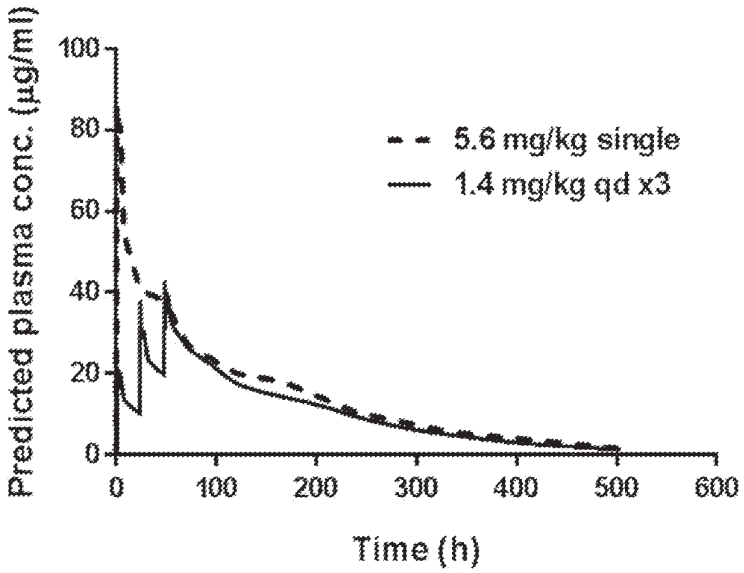




Dose (mg/kg)	Schedule	Total dose (mg/kg)	T/C (%)	Log cell kill	CR	Tumor-free (Day 120)	Cmax (µg/ml)	Cavg (µg/ml)	AUC 0-540 (hr*µg/ml)
8.5	Single	8.5	0	3.9	6/6	0/6	131	23	11354
2.8	qd x 3	8.4	0	4.6	6/6	1/6	85	28	10961
2.8	q3d x 3	8.4	0	4.1	6/6	2/6	75	21	10762

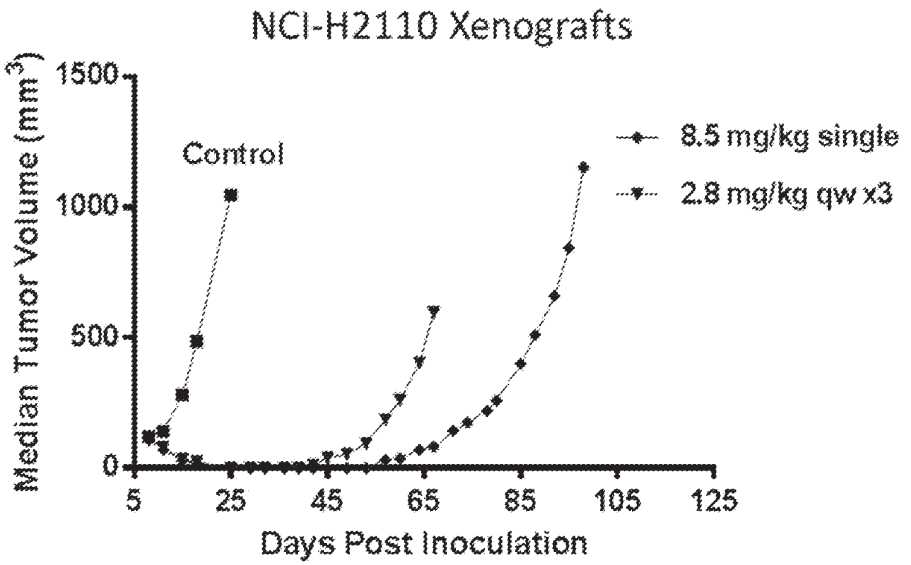
Figure 11

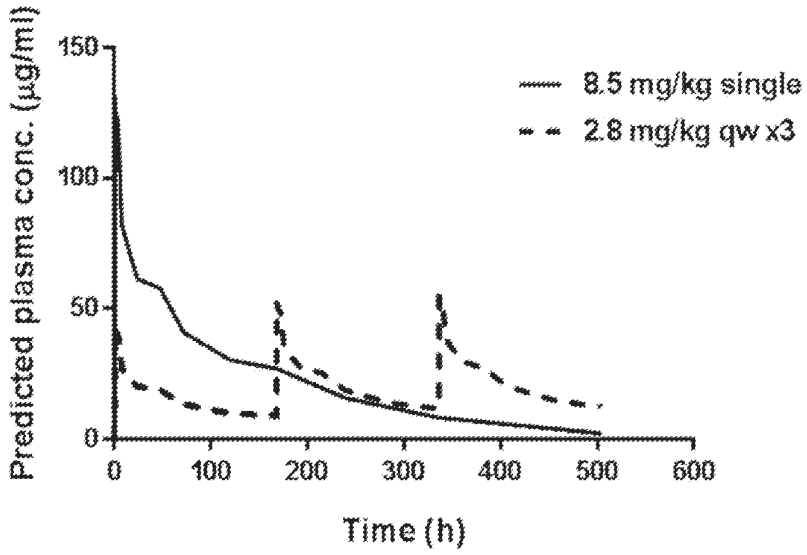




Dose (mg/kg)	Schedule	Total Dose (mg/kg)	T/C (%)	Log cell kill	CR	Cmax (µg/ml)	Cavg (µg/ml)	AUC 0-540 (hr*µg/ml)
5.6	Single	5.6	0	2.7	4/6	86	15	7481
1.4	qd x 3	4.2	0	2.5	4/6	43	14	5480

Figure 12





Dose (mg/kg)	Schedule	Total Dose (mg/kg)	T/C (%)	Log cell kill	CR	Cmax (µg/ml)	Cavg (µg/ml)	AUC 0-540 (hr*µg/ml)
8.5	Single	8.5	0	3.9	6/6	131	23	11354
2.8	qw x 3	8.4	0	3.0	6/6	55	15	9614