Figure 1

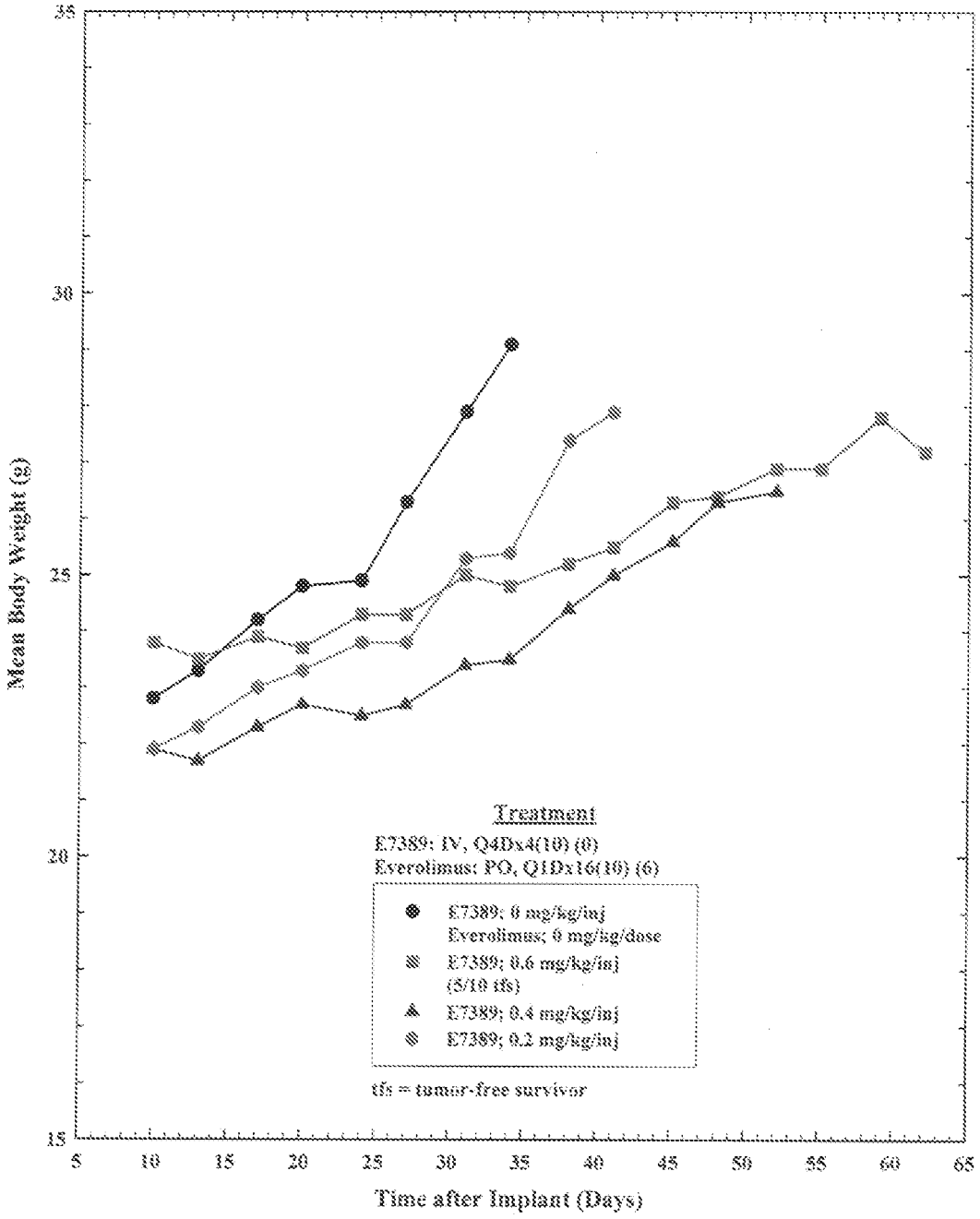

Mean Body Weight (g)
35
30
25
20
15
5
10
15
20
25
30
35
40
45
50
55
60
65
Time after Implant (Days)
Treatment
E7389: IV, Q4Dx4(10) (0)
Everolimus: PO, Q1Dx16(10) (6)
E7389: 0 mg/kg/inj
Everolimus: 0 mg/kg/dose
E7389: 0.6 mg/kg/inj
(5/10 tfs)
E7389: 0.4 mg/kg/inj
E7389: 0.2 mg/kg/inj
tfs = tumor-free survivor

Figure 2

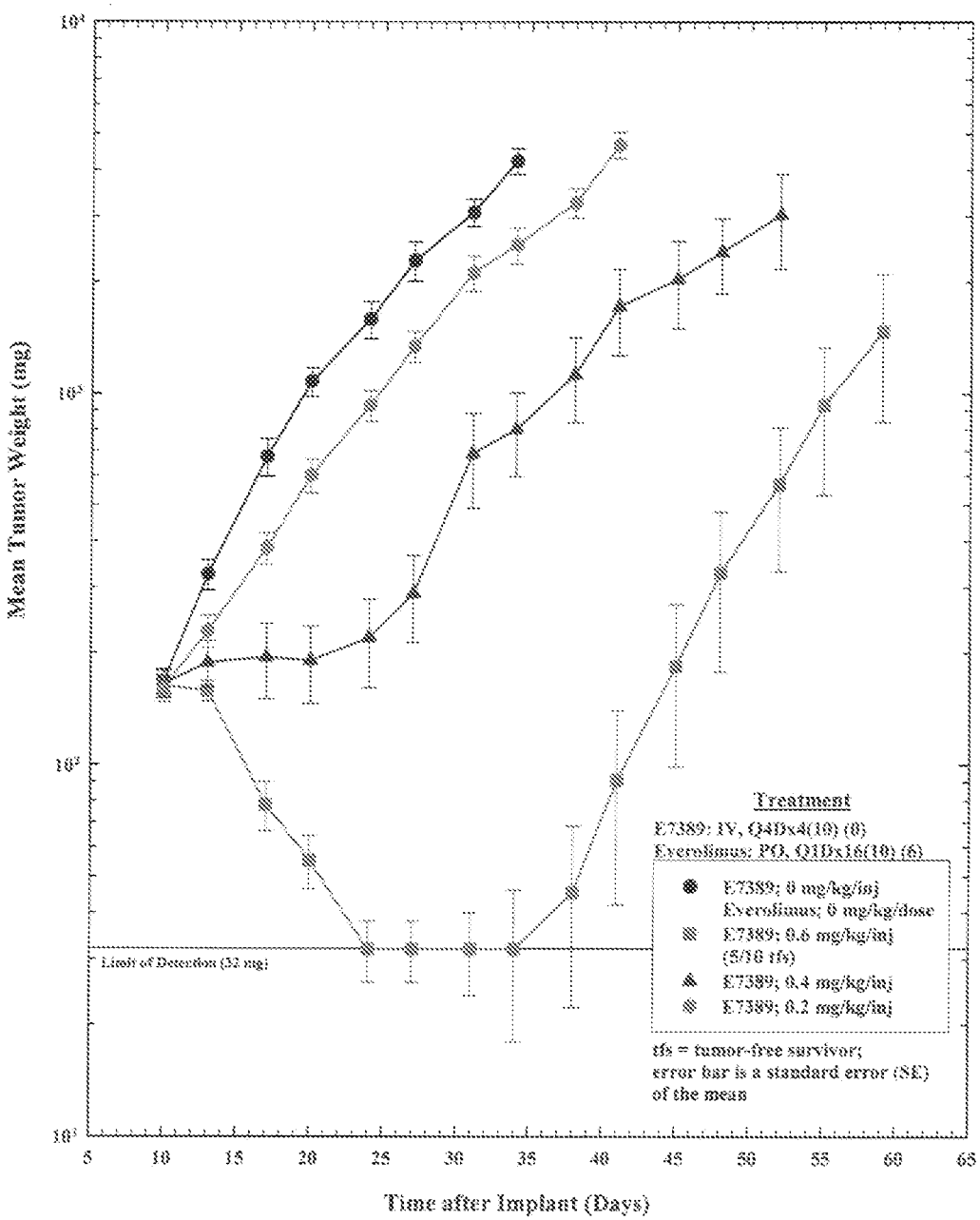

Mean Tumor Weight (mg)
Time after Implant (Days)
Limit of Detection (32 mg)
Treatment
E7389: IV, Q4Dx4(10) (8)
Everolimus: PO, Q1Dx16(10) (6)
E7389; 0 mg/kg/inj
Everolimus; 0 mg/kg/dose
E7389; 0.6 mg/kg/inj
(5/16 tfs)
E7389; 0.4 mg/kg/inj
E7389; 0.2 mg/kg/inj
tfs = tumor-free survivor;
error bar is a standard error (SE)
of the mean

Figure 3

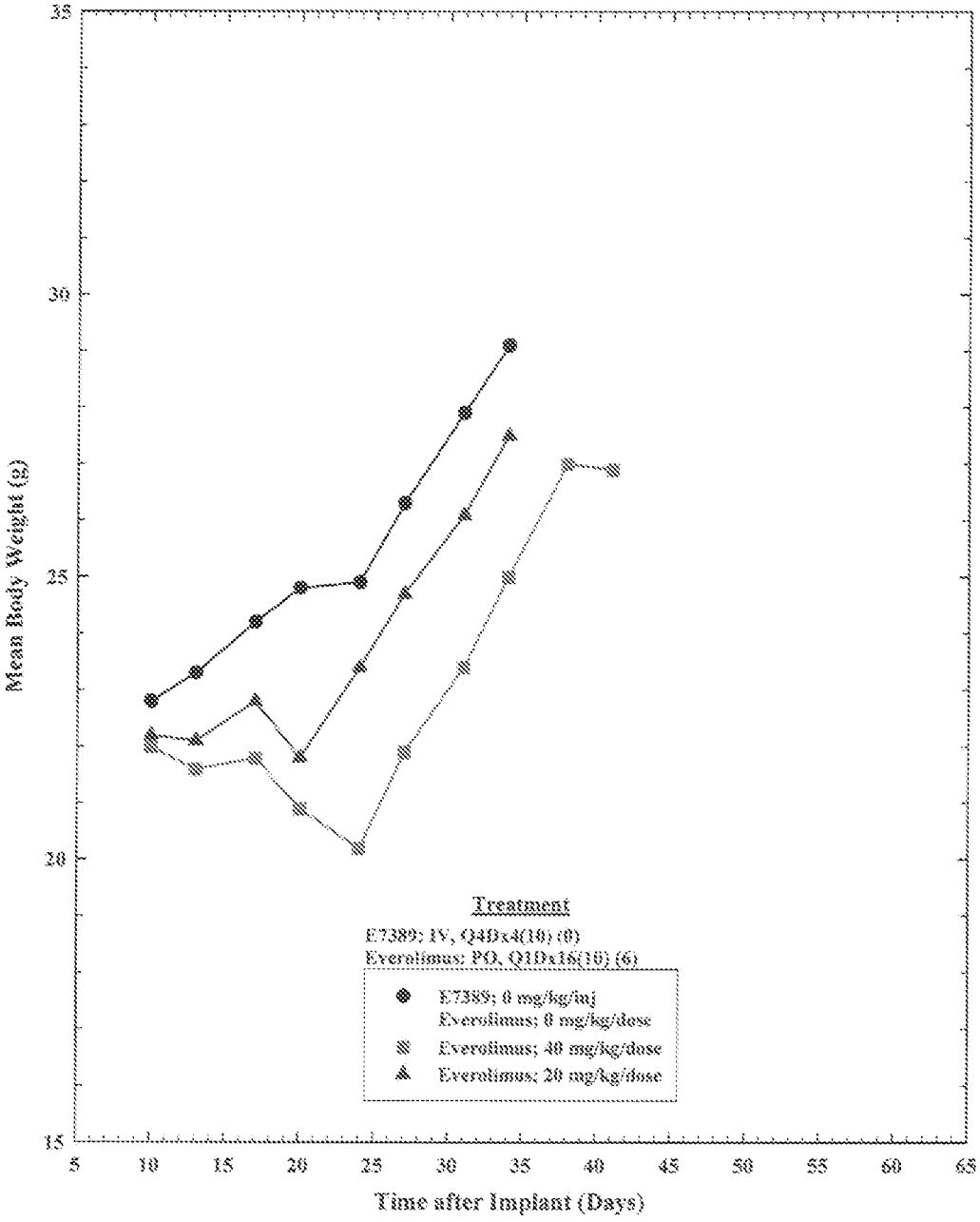

Mean Body Weight (g)
35
30
25
20
15
5
10
15
20
25
30
35
40
45
50
55
60
65
Time after Implant (Days)
Treatment
E7389; IV, Q4Dx4(10) (0)
Everolimus; PO, Q1Dx15(10) (6)
E7389; 0 mg/kg/inj
Everolimus; 0 mg/kg/dose
Everolimus; 40 mg/kg/dose
Everolimus; 20 mg/kg/dose

Figure 4

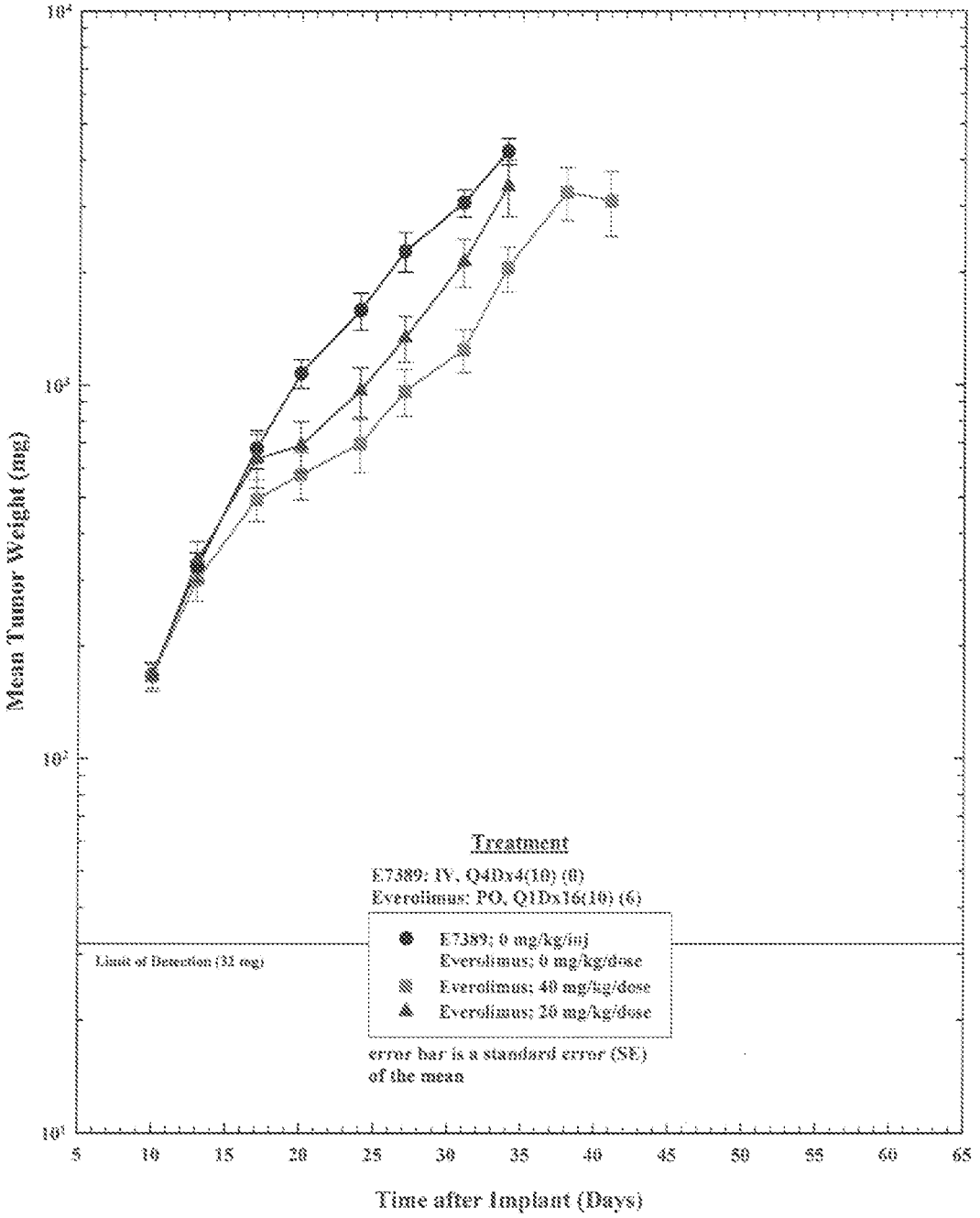

Mean Tumor Weight (mg)
Time after Implant (Days)
Treatment
E7389: IV, Q4Dx4(10) (8)
Everolimus: PO, Q1Dx16(10) (6)
E7389; 0 mg/kg/inj
Everolimus; 0 mg/kg/dose
Everolimus; 40 mg/kg/dose
Everolimus; 20 mg/kg/dose
Limit of Detection (32 mg)
error bar is a standard error (SE) of the mean

Figure 5

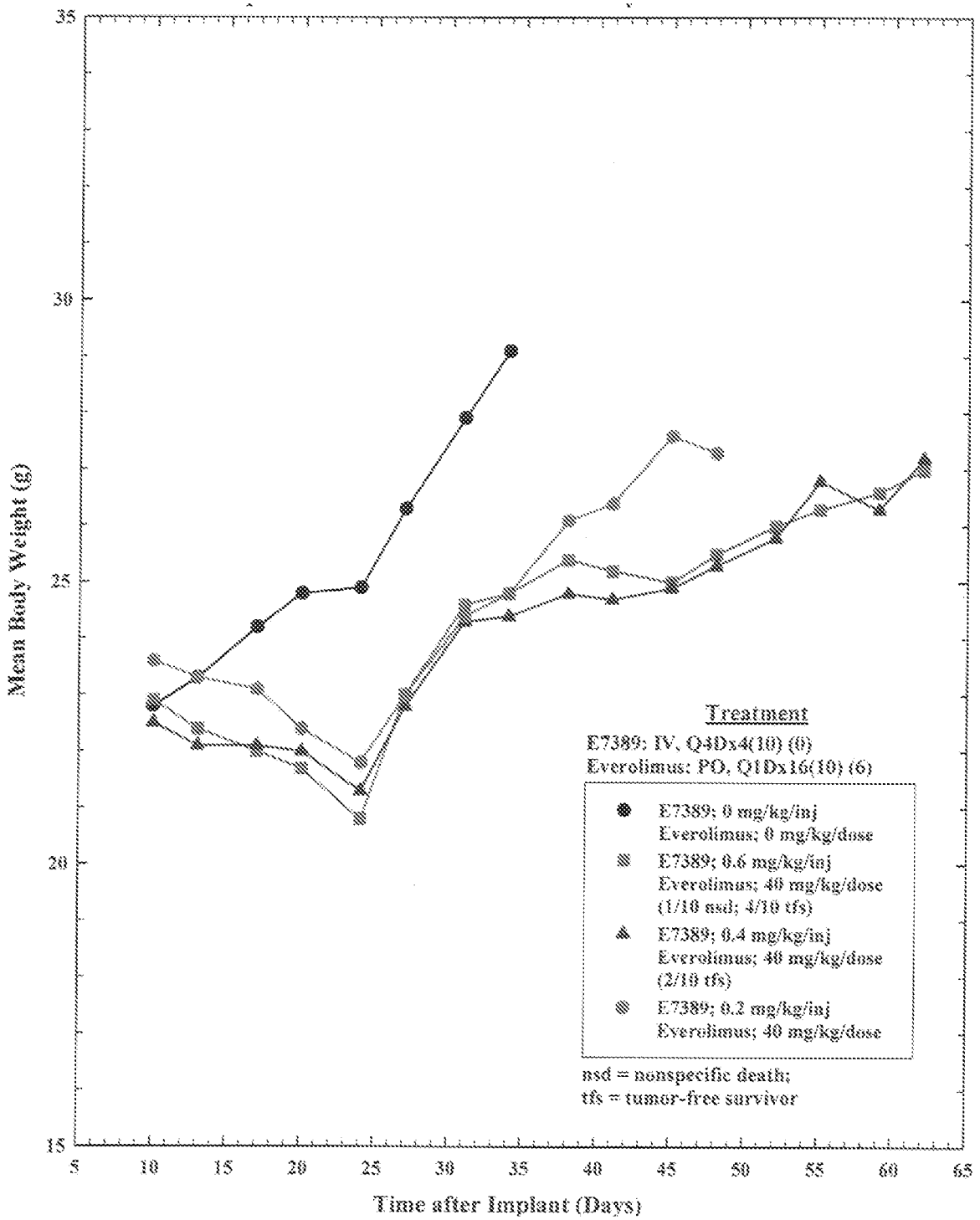

Mean Body Weight (g)
Time after Implant (Days)
35
30
25
20
15
5
10
15
20
25
30
35
40
45
50
55
60
65
Treatment
E7389: IV, Q4Dx4(10) (0)
Everolimus: PO, Q1Dx16(10) (6)
E7389; 0 mg/kg/inj
Everolimus; 0 mg/kg/dose
E7389; 0.6 mg/kg/inj
Everolimus; 40 mg/kg/dose
(1/10 nsd; 4/10 tfs)
E7389; 0.4 mg/kg/inj
Everolimus; 40 mg/kg/dose
(2/10 tfs)
E7389; 0.2 mg/kg/inj
Everolimus; 40 mg/kg/dose
nsd = nonspecific death;
tfs = tumor-free survivor

Figure 6

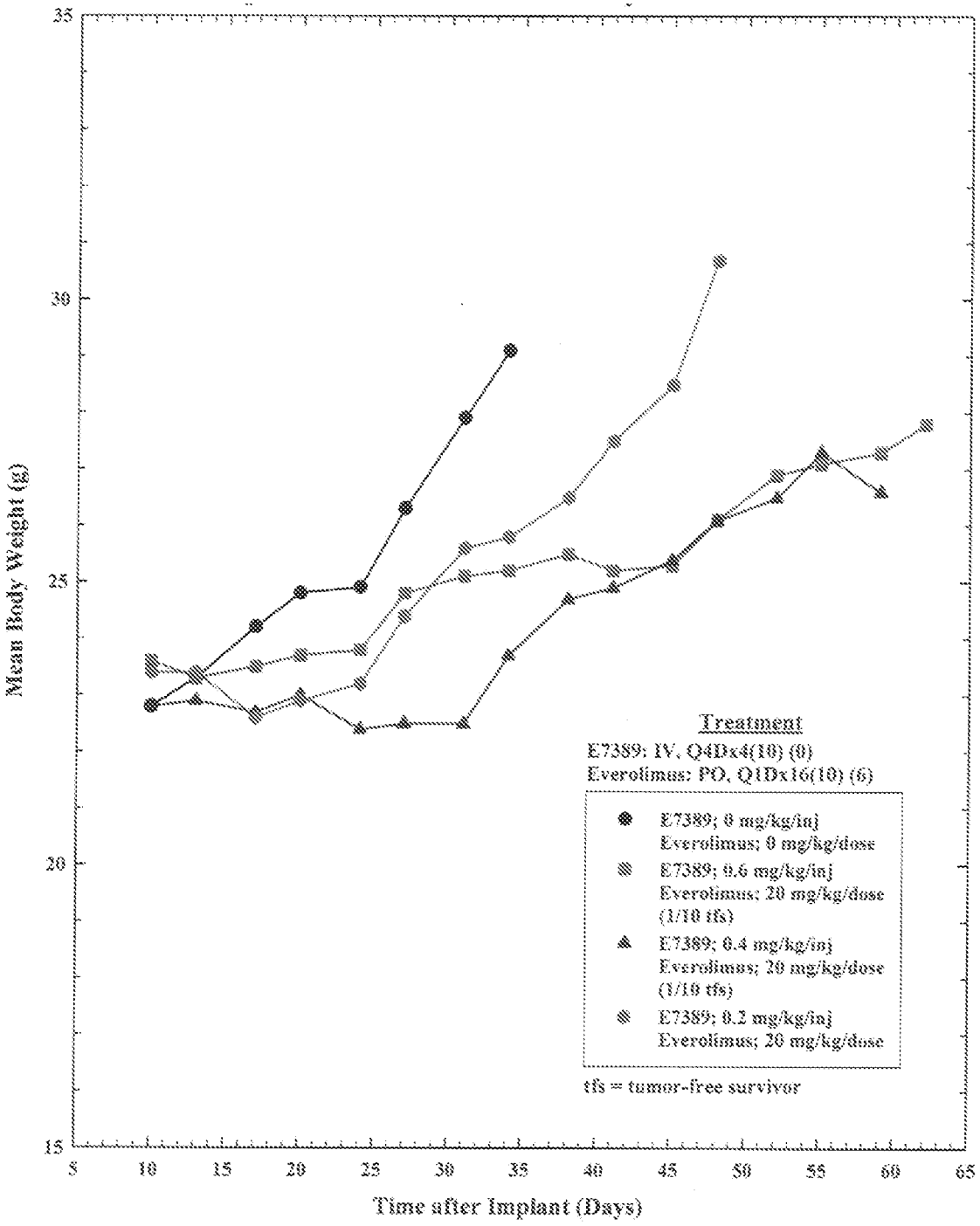

Mean Body Weight (g)
Time after Implant (Days)
35
30
25
20
15
5
10
15
20
25
30
35
40
45
50
55
60
65
Treatment
E7389: IV, Q4Dx4(10) (0)
Everolimus: PO, Q1Dx16(10) (6)
E7389; 0 mg/kg/inj
Everolimus; 0 mg/kg/dose
E7389; 0.6 mg/kg/inj
Everolimus; 20 mg/kg/dose
(1/10 tfs)
E7389; 0.4 mg/kg/inj
Everolimus; 20 mg/kg/dose
(1/10 tfs)
E7389; 0.2 mg/kg/inj
Everolimus; 20 mg/kg/dose
tfs = tumor-free survivor

Figure 7

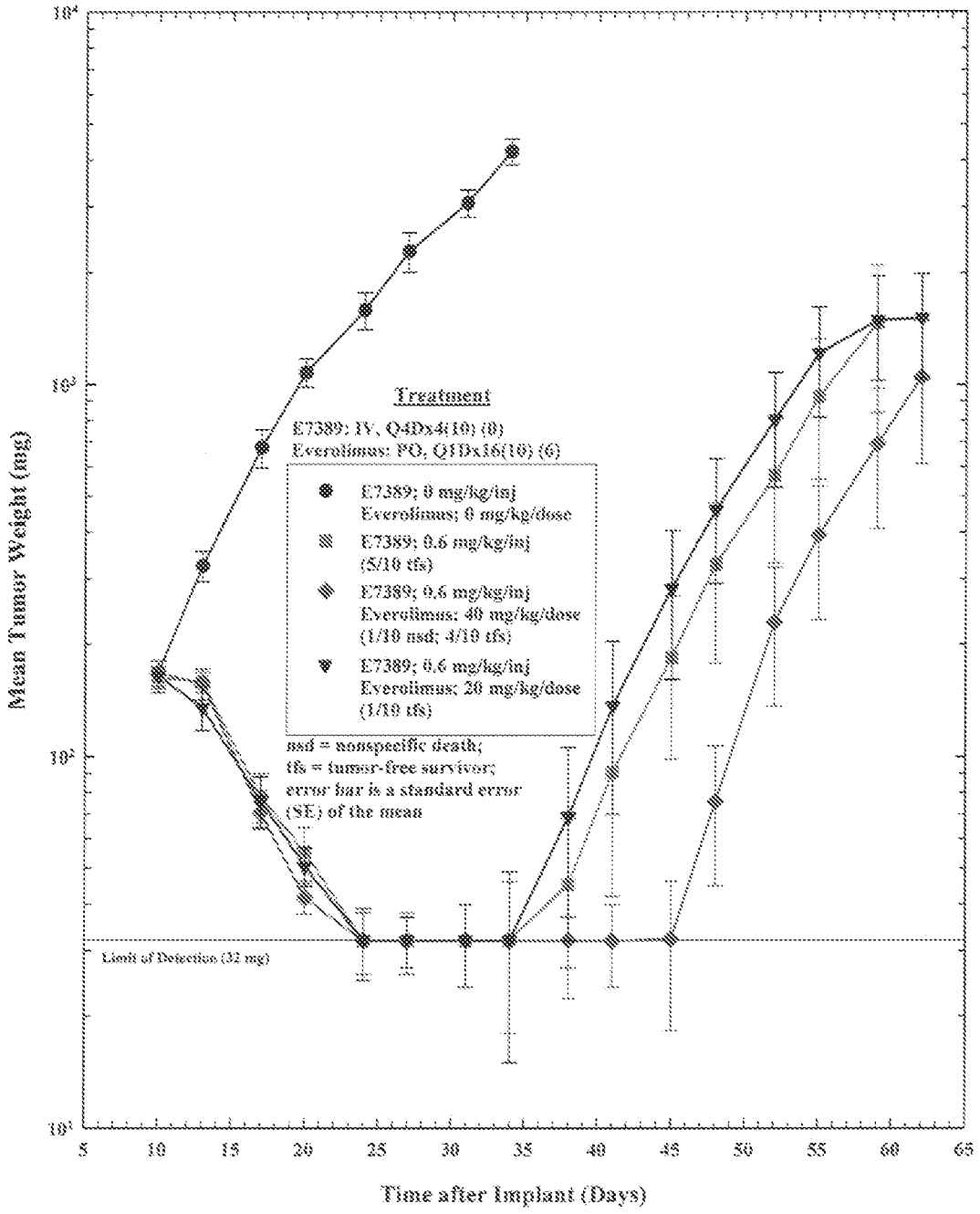

Mean Tumor Weight (mg)
Time after Implant (Days)
Treatment
E7389: IV, Q4Dx4(10) (0)
Everolimus: PO, QDx16(16) (6)
E7389; 0 mg/kg/inj
Everolimus; 0 mg/kg/dose
E7389; 0.6 mg/kg/inj
(5/10 tfs)
E7389; 0.6 mg/kg/inj
Everolimus; 40 mg/kg/dose
(1/10 nsd; 4/10 tfs)
E7389; 0.6 mg/kg/inj
Everolimus; 20 mg/kg/dose
(1/10 tfs)
nsd = nonspecific death;
tfs = tumor-free survivor;
error bar is a standard error
(SE) of the mean
Limit of Detection (32 mg)

Figure 8

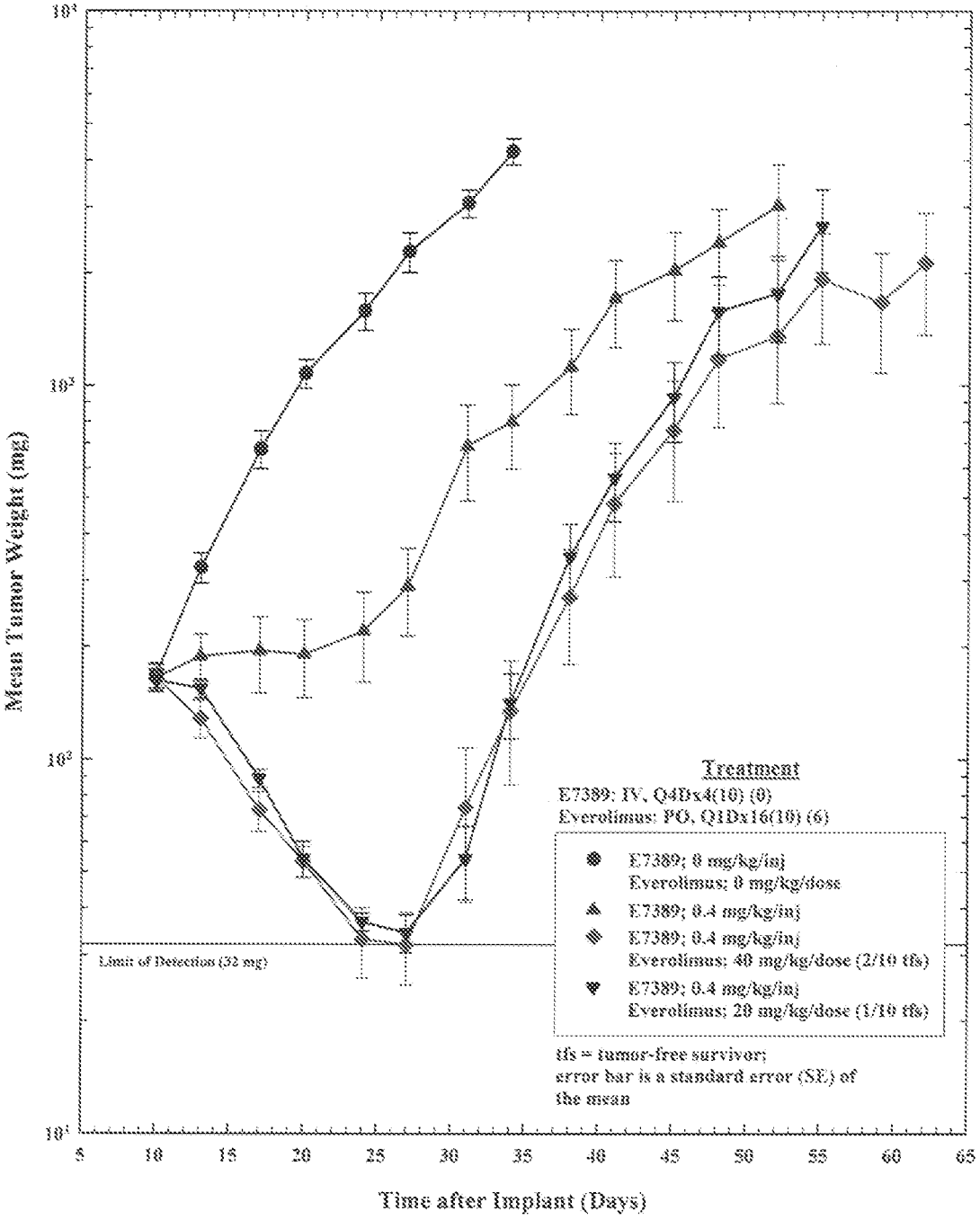

Mean Tumor Weight (mg)
Time after Implant (Days)
Limit of Detection (32 mg)
Treatment
E7389: IV, Q4Dx4(10) (0)
Everolimus: PO, Q1Dx16(10) (6)
E7389; 0 mg/kg/inj
Everolimus; 0 mg/kg/dose
E7389; 0.4 mg/kg/inj
E7389; 0.4 mg/kg/inj
Everolimus; 40 mg/kg/dose (2/10 tfs)
E7389; 0.4 mg/kg/inj
Everolimus; 20 mg/kg/dose (1/10 tfs)
tfs = tumor-free survivor;
error bar is a standard error (SE) of the mean

Figure 9

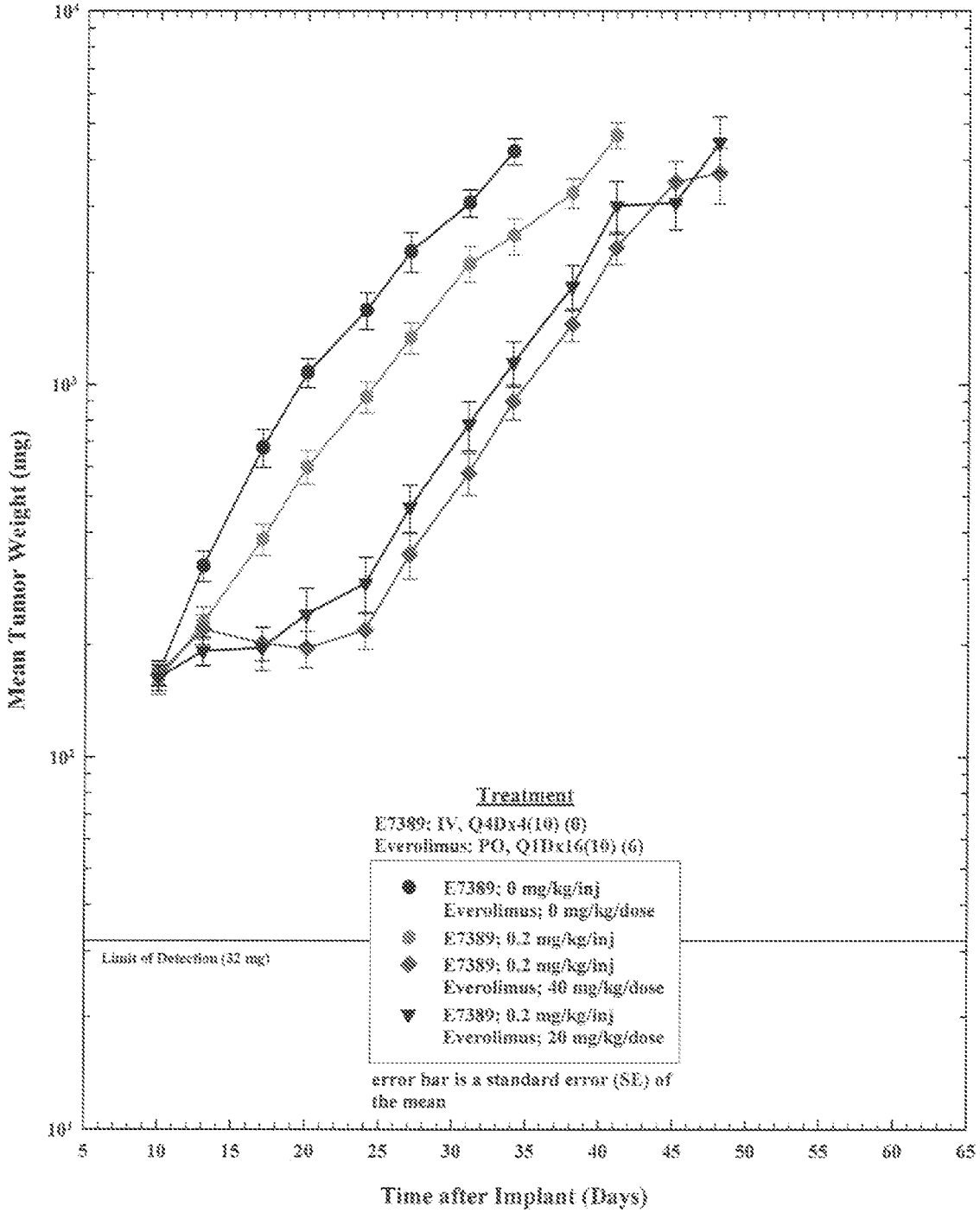

Mean Tumor Weight (mg)
Time after Implant (Days)
Treatment
E7389: IV, Q4Dx4(10) (6)
Everolimus: PO, Q1Dx16(10) (6)
E7389; 0 mg/kg/inj
Everolimus; 0 mg/kg/dose
E7389; 0.2 mg/kg/inj
E7389; 0.2 mg/kg/inj
Everolimus; 40 mg/kg/dose
E7389; 0.2 mg/kg/inj
Everolimus; 20 mg/kg/dose
Limit of Detection (32 mg)
error bar is a standard error (SE) of the mean

Figure 10

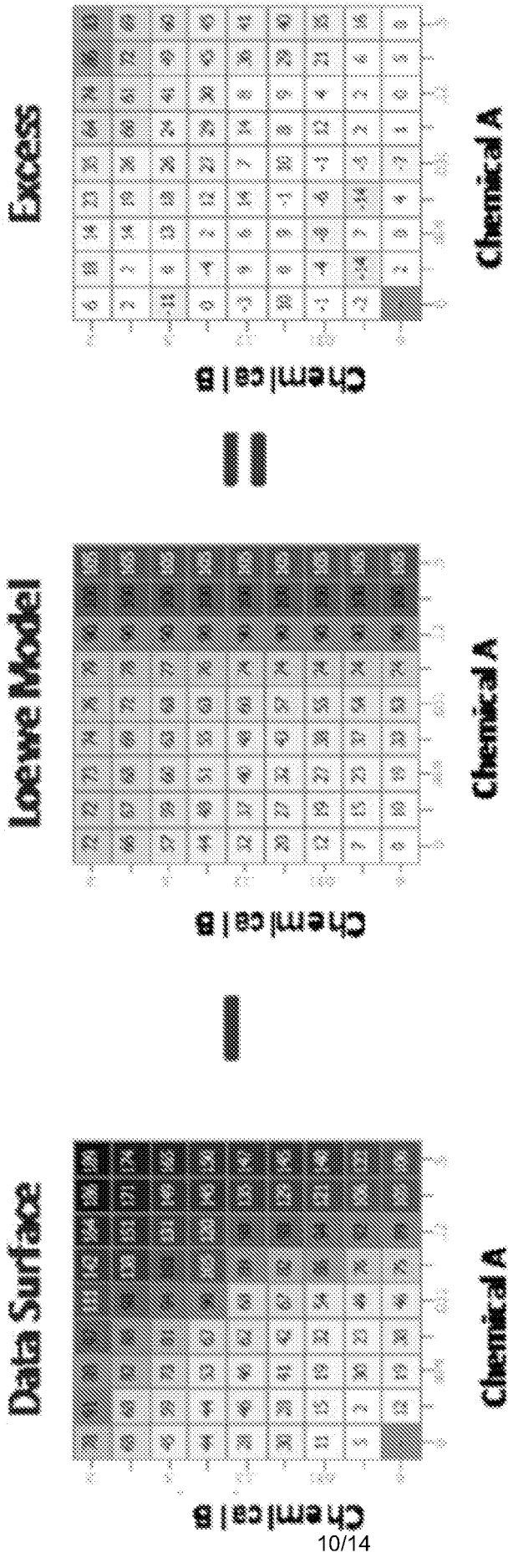
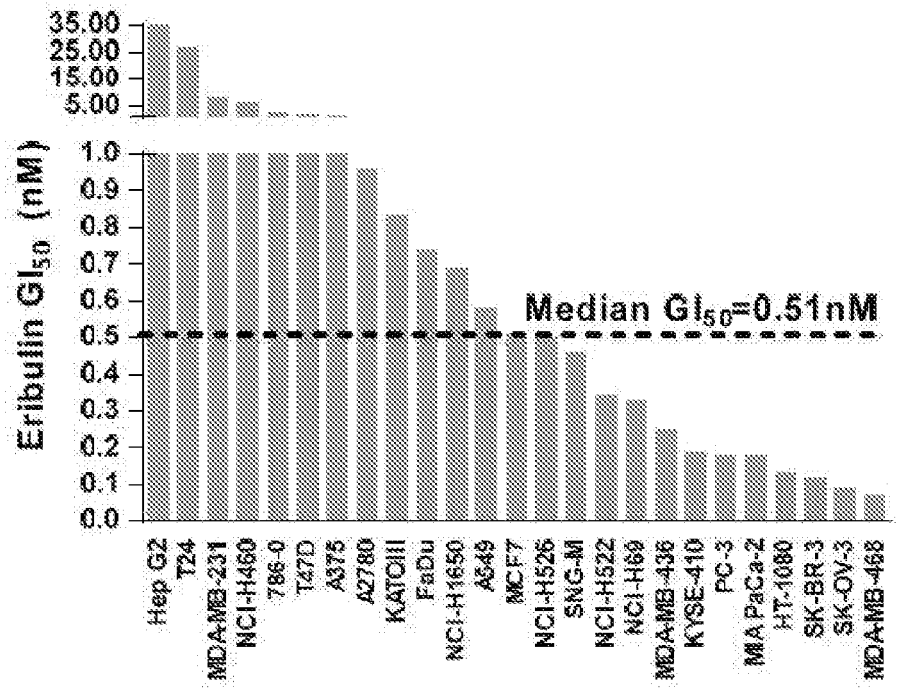


Figure 11


Eribulin GI50 (nM)
Median GI50=0.51nM
Hep G2
T24
MDA-MB-231
NCI-H460
786-0
T47D
A375
A2780
KATOIII
FaDu
NCI-H1650
A549
MCF7
NCI-H526
SNG-M
NCI-H522
NCI-H69
MDA-MB-436
KYSE-410
PC-3
MIA PaCa-2
HT-1080
SK-BR-3
SK-OV-3
MDA-MB-468

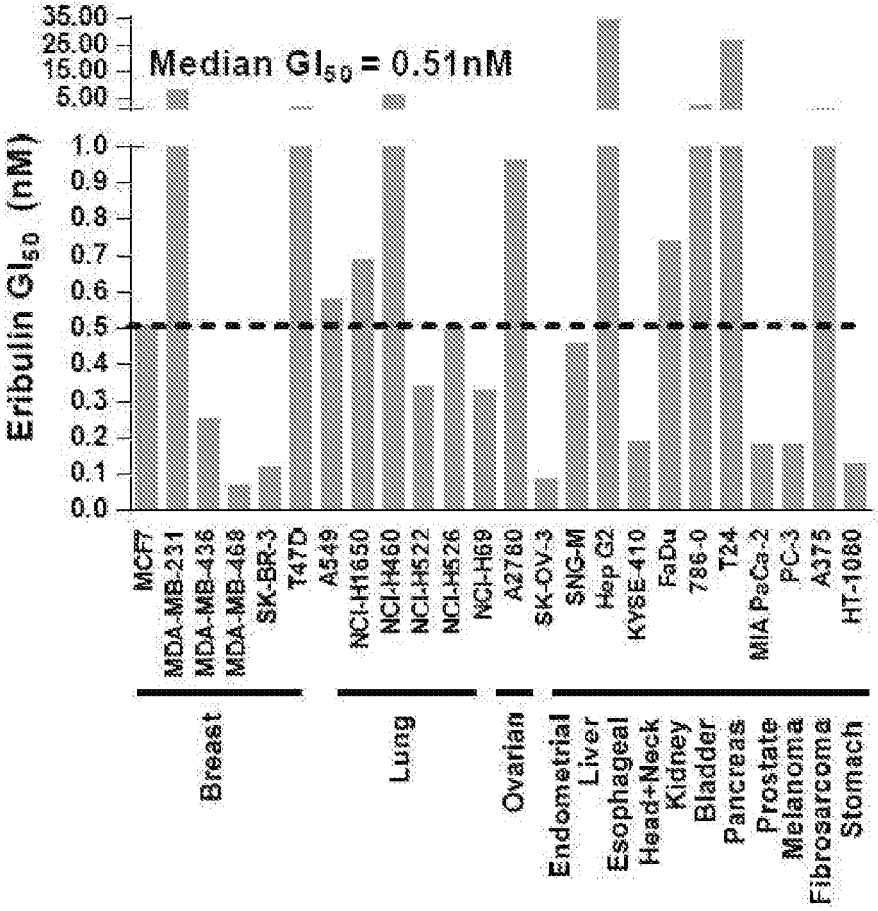

Eribulin GI50 (nM)
Median GI50 = 0.51nM
MCF7
MDA-MB-231
MDA-MB-436
MDA-MB-468
SK-BR-3
T47D
A549
NCI-H1650
NCI-H460
NCI-H522
NCI-H526
NCI-H69
A2780
SK-OV-3
SNG-M
Hep G2
KYSE-410
FaDu
786-0
T24
MIA PaCa-2
PC-3
A375
HT-1080
Breast
Lung
Ovarian
Endometrial
Liver
Esophageal
Head+Neck
Kidney
Bladder
Pancreas
Prostate
Melanoma
Fibrosarcoma
Stomach

Figure 12

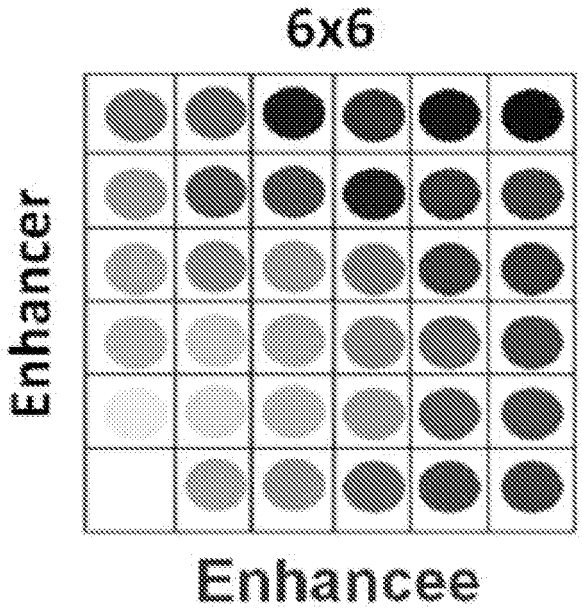


Figure 13

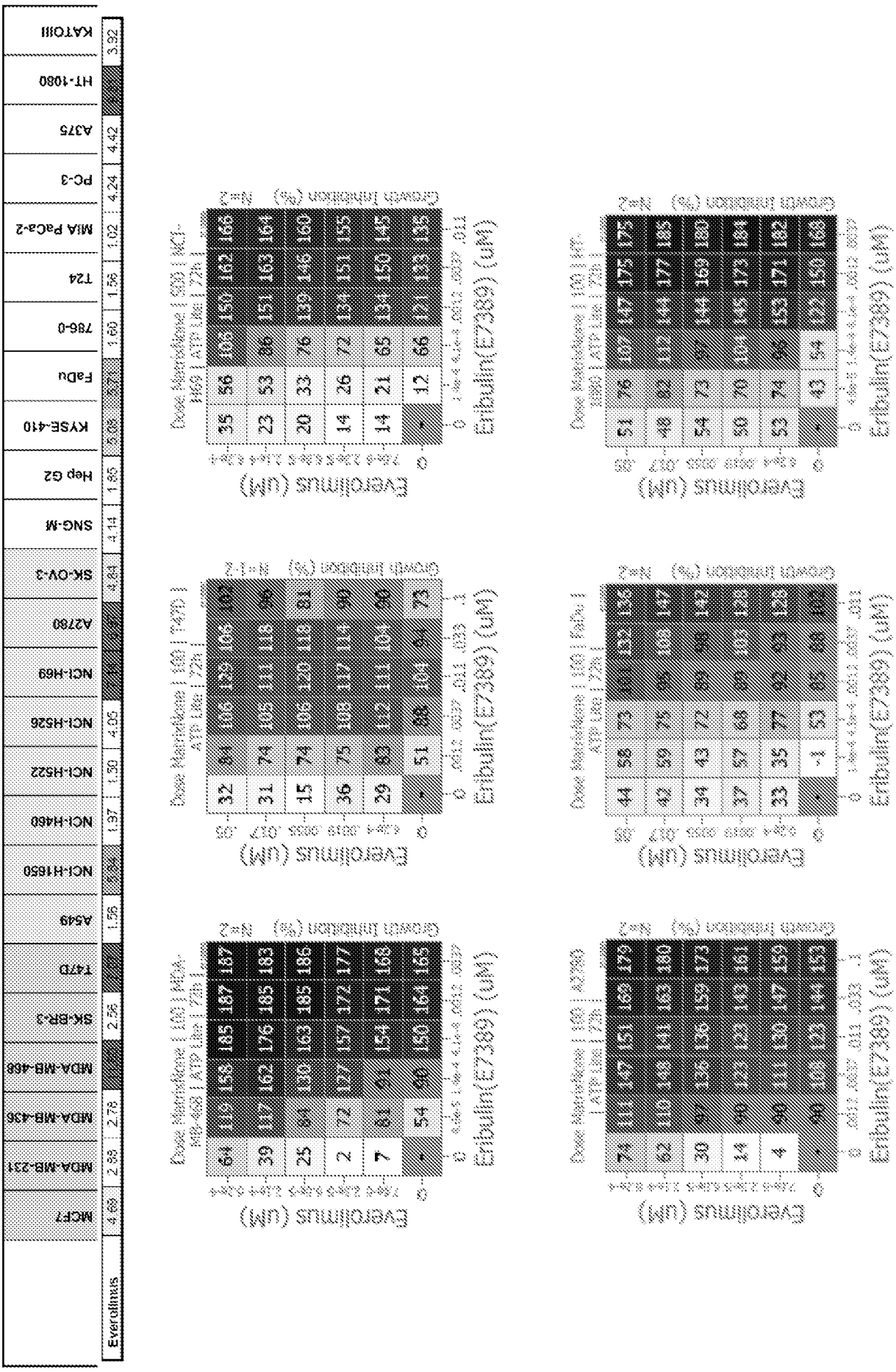

Everolimus
MCF7
MDA-MB-231
MDA-MB-436
MDA-MB-468
SK-BR-3
T47D
A549
NCI-H1650
NCI-H460
NCI-H522
NCI-H526
NCI-H69
A2780
SK-OV-3
SNG-M
Hep G2
KYSE-410
FaDu
786-0
T24
MIA PaCa-2
PC-3
A375
HT-1080
KATOIII
Everolimus (uM)
Eribulin(E7389) (uM)

Figure 14

Synergy Score Values

	MCF7	MDA-MB-231	MDA-MB-436	MDA-MB-468	SK-BR-3	T47D	A549	NCI-H1650	NCI-H460	NCI-H522	NCI-H526	NCI-H69	A2780	SK-OV-3	SNG-M	Hep G2	KYSE-410	FaDu	786-0	T24	RA PaCa-2	PC-3	A375	HT-1080	KATOIII
Everolimus	4.69	2.66	1.78	21.60	2.56	2.07	1.56	5.64	1.92	1.50	4.05	7.24	6.97	4.84	4.14	1.65	3.08	5.73	1.60	1.56	1.02	4.24	4.42	2.61	3.92

Loewe Volume Scores

	MCF7	MDA-MB-231	MDA-MB-436	MDA-MB-468	SK-BR-3	T47D	A549	NCI-H1650	NCI-H460	NCI-H522	NCI-H526	NCI-H69	A2780	SK-OV-3	SNG-M	Hep G2	KYSE-410	FaDu	786-0	T24	RA PaCa-2	PC-3	A375	HT-1080	KATOIII
Everolimus	2.71	2.93	2.45	6.68	1.43	5.59	1.85	5.86	1.40	-0.11	2.71	4.37	3.42	3.19	3.52	2.03	3.86	4.22	1.30	1.06	0.21	3.18	4.25	6.27	4.84