Figure 1

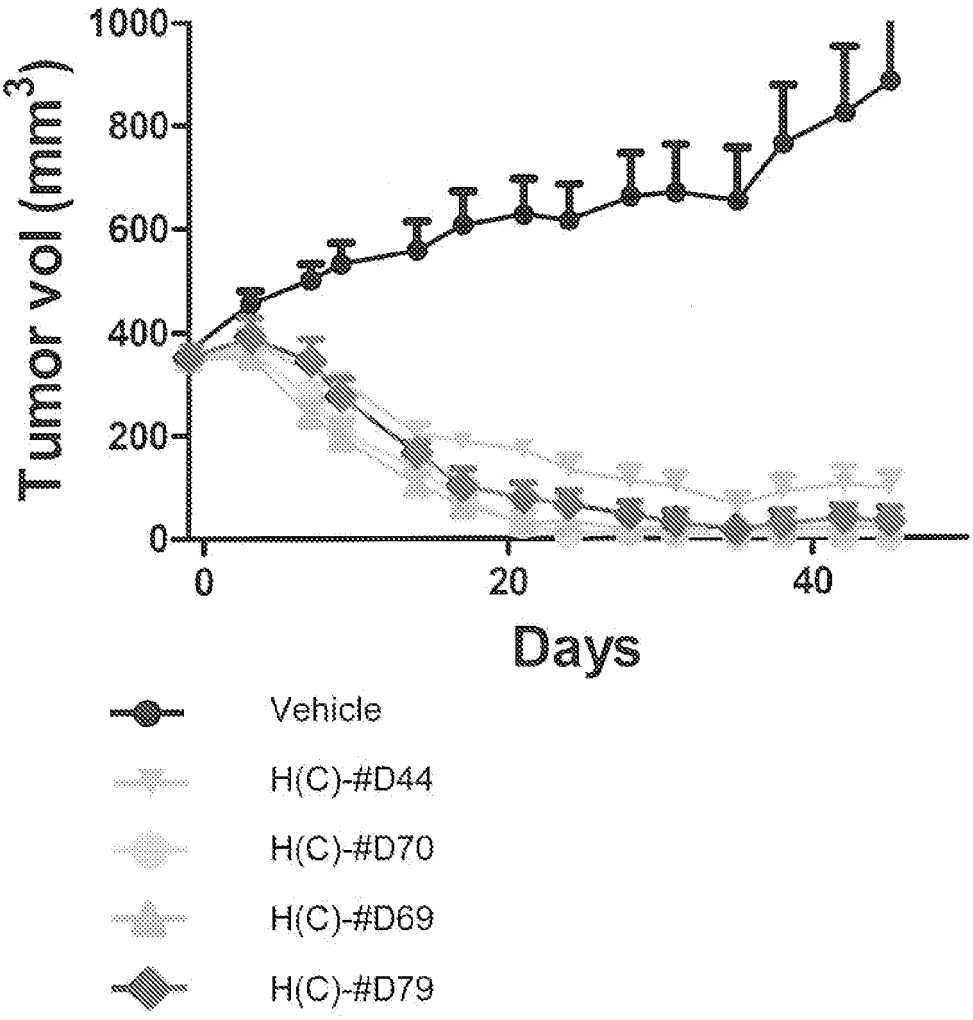

Tumor vol (mm3)
1000
800
600
400
200
0
0
20
40
Days
Vehicle
H(C)-#D44
H(C)-#D70
H(C)-#D69
H(C)-#D79

Figure 2

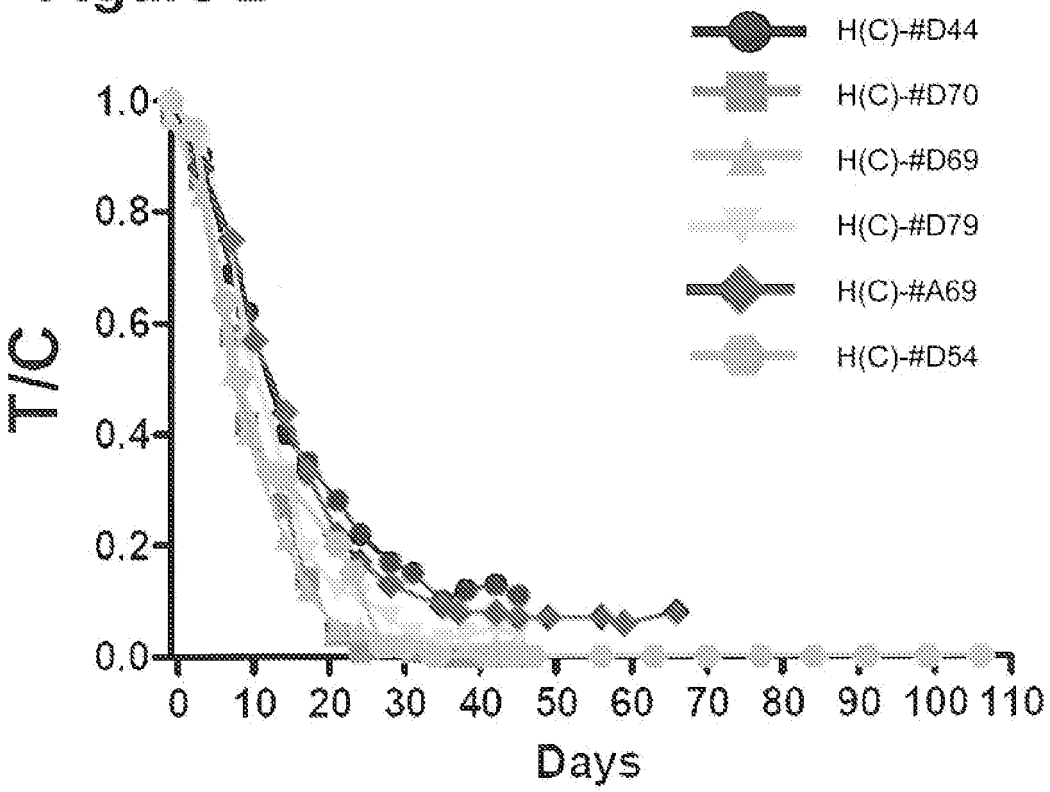

H(C)-#D44
H(C)-#D70
H(C)-#D69
H(C)-#D79
H(C)-#A69
H(C)-#D54
T/C
1.0
0.8
0.6
0.4
0.2
0.0
0
10
20
30
40
50
60
70
80
90
100
110
Days

Figure 3

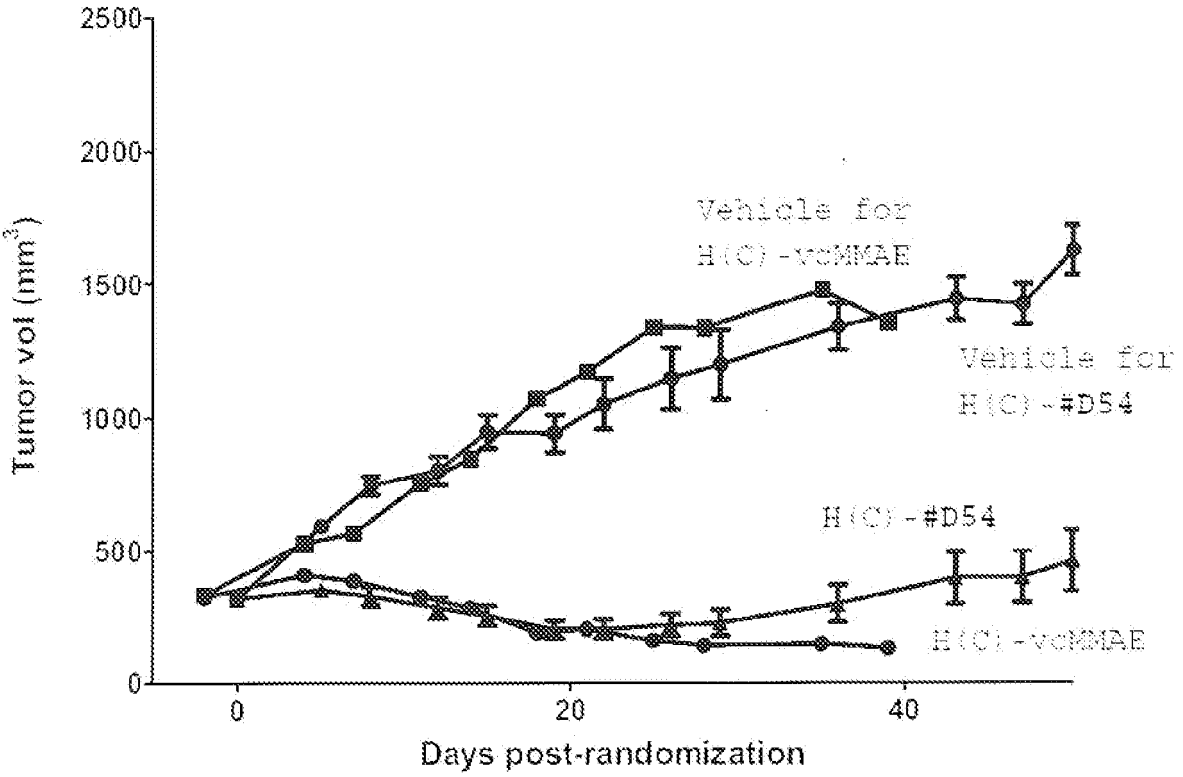

2500
2000
1500
1000
500
0
Tumor vol (mm3)
0
20
40
Days post-randomization
Vehicle for H(C)-vcMMAE
Vehicle for H(C)-#D54
H(C)-#D54
H(C)-vcMMAE

Figure 4A

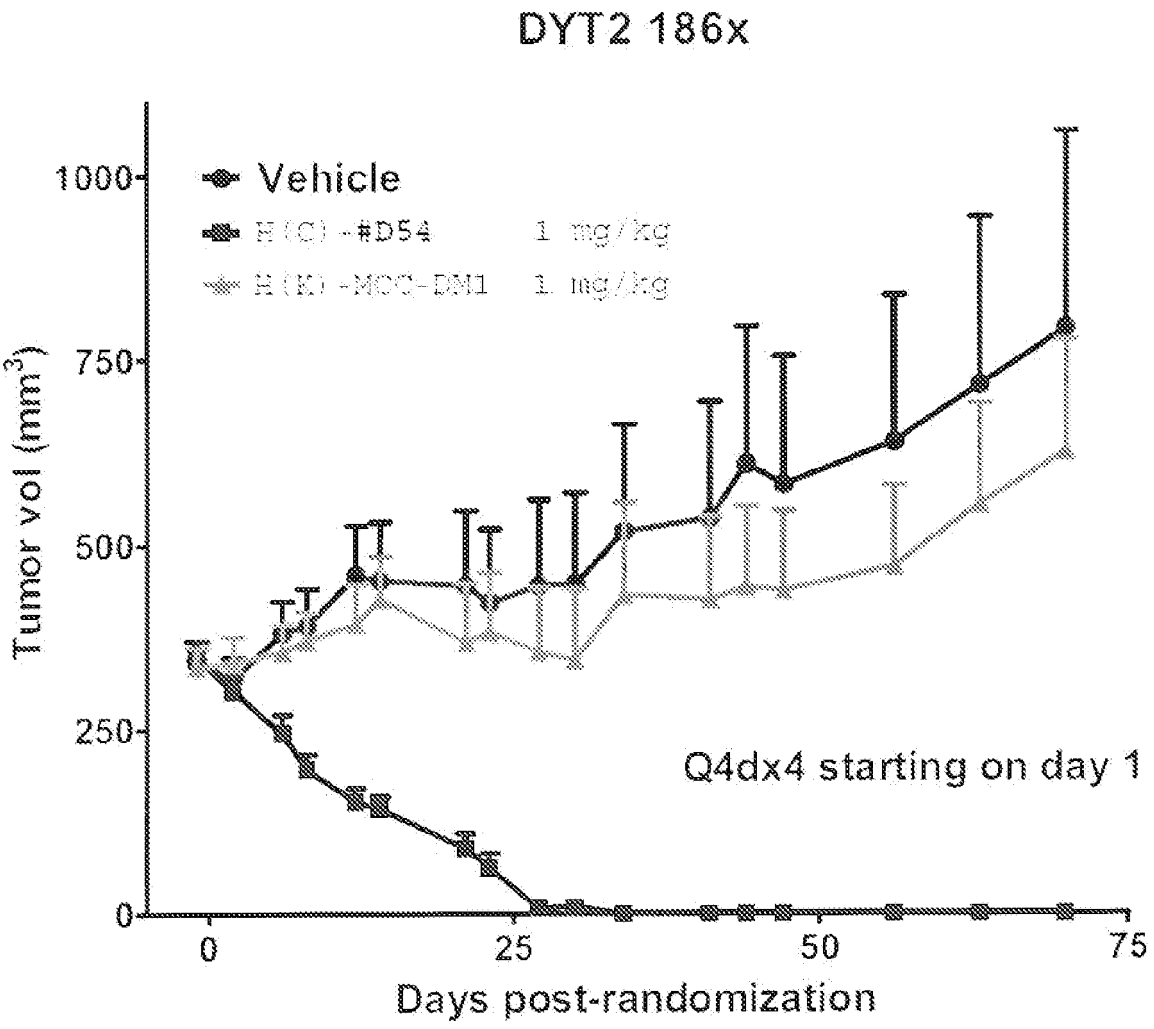

DYT2 186x
Vehicle
H(C)-#D54 1 mg/kg
H(K)-MCC-DM1 1 mg/kg
Tumor vol (mm3)
1000
750
500
250
0
0
25
50
75
Days post-randomization
Q4dx4 starting on day 1

Figure 4B

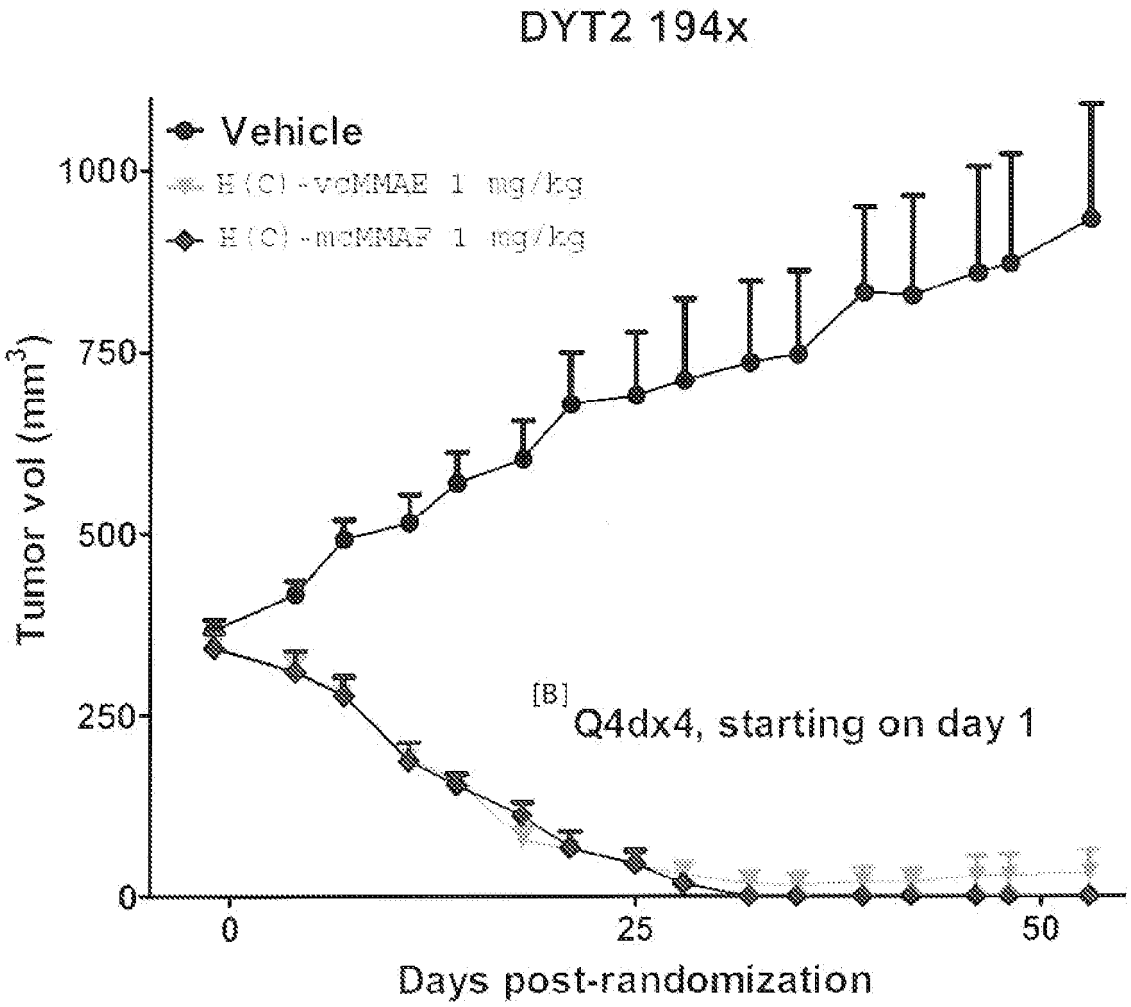

DYT2 194x
Vehicle
H(C)-vcMMAE 1 mg/kg
H(C)-mcMMAF 1 mg/kg
Tumor vol (mm3)
1000
750
500
250
0
0
25
50
Days post-randomization
[B] Q4dx4, starting on day 1

Figure 4C

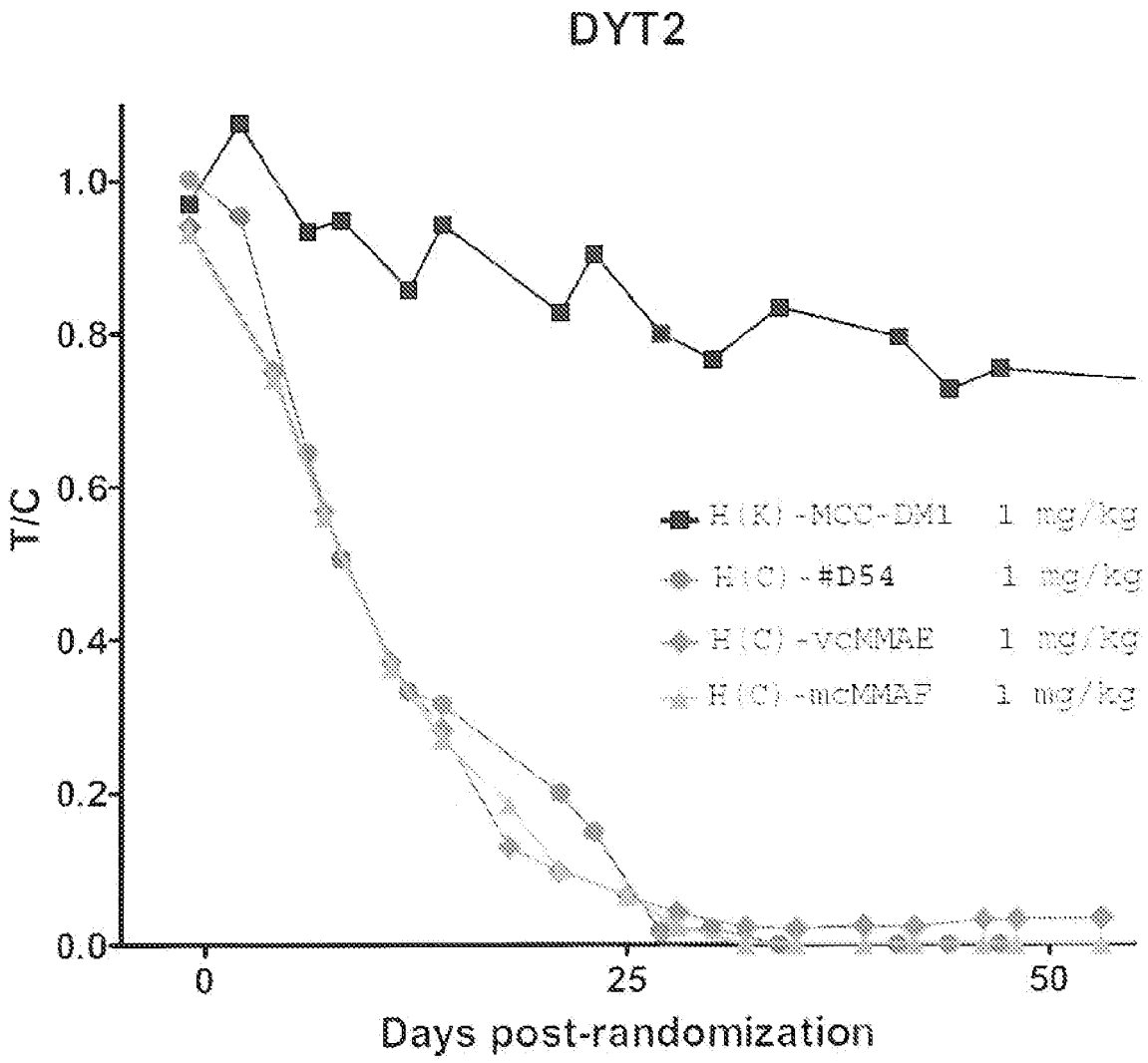

DYT2
T/C
1.0
0.8
0.6
0.4
0.2
0.0
0
25
50
Days post-randomization
H(K)-MCC-DM1 1 mg/kg
H(C)-#D54 1 mg/kg
H(C)-vcMMAE 1 mg/kg
H(C)-mcMMAF 1 mg/kg

Figure 5A

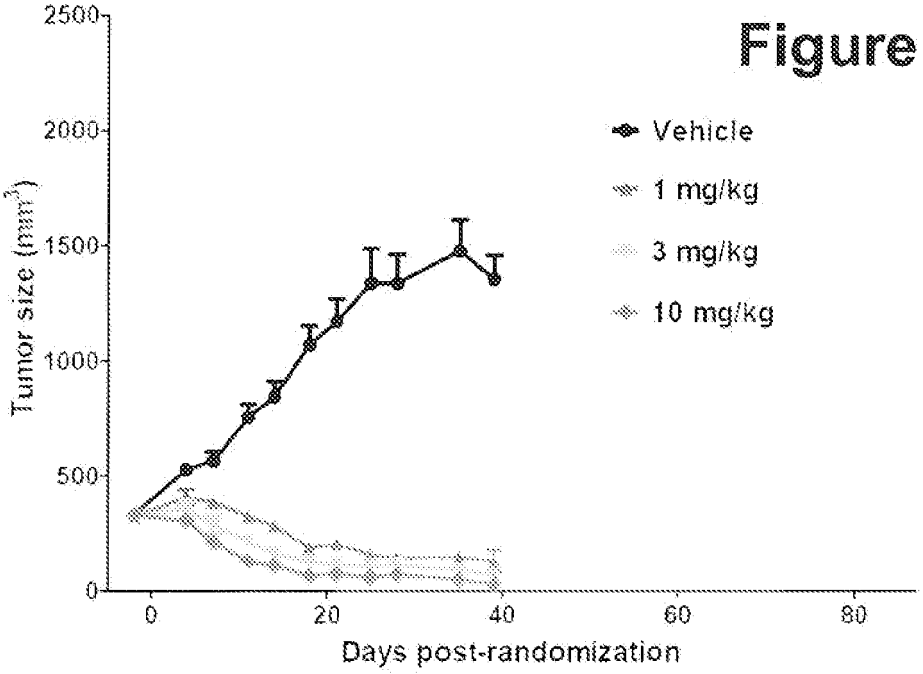


Figure 5B

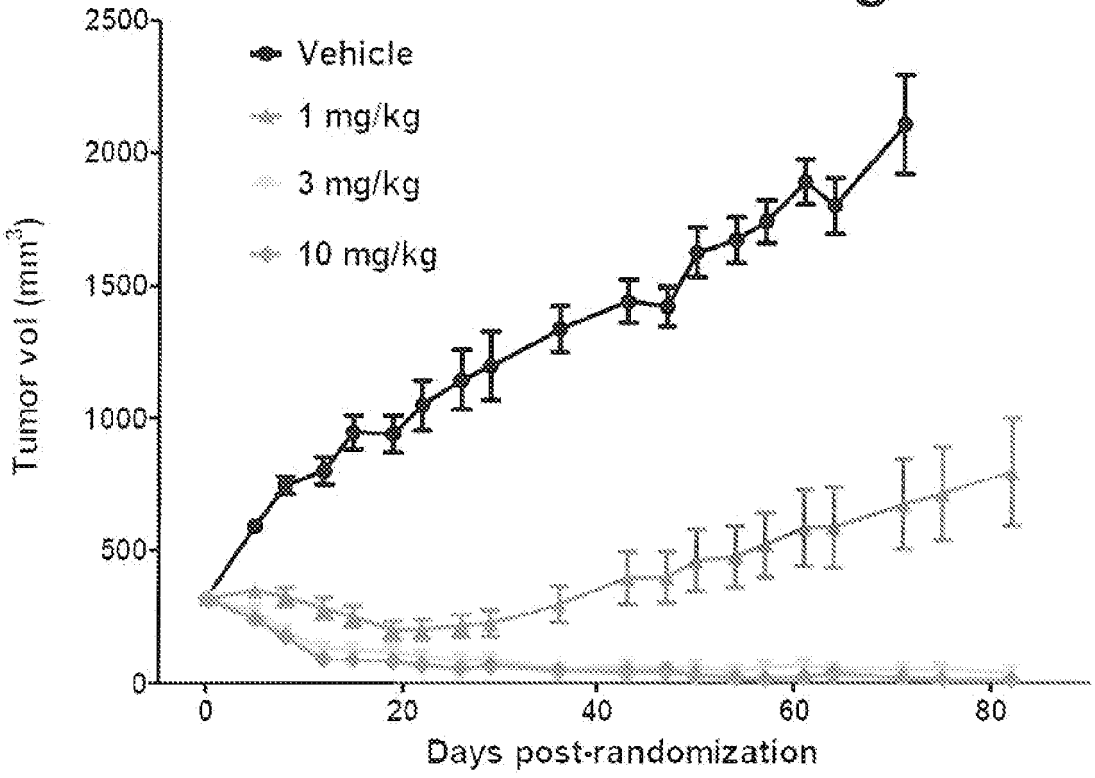


Figure 5C

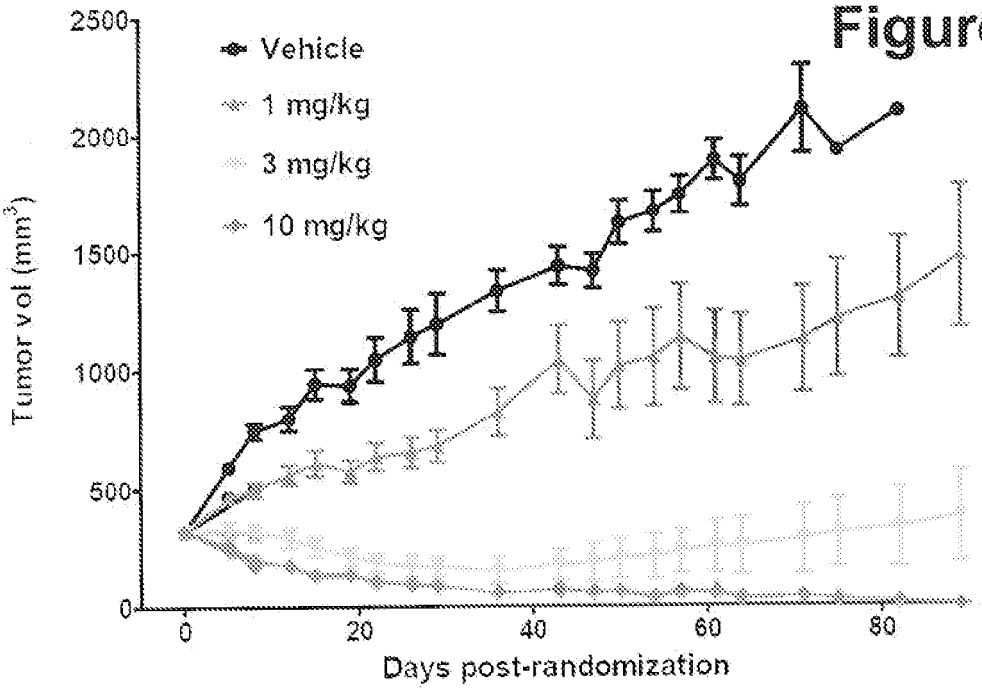


Figure 5D

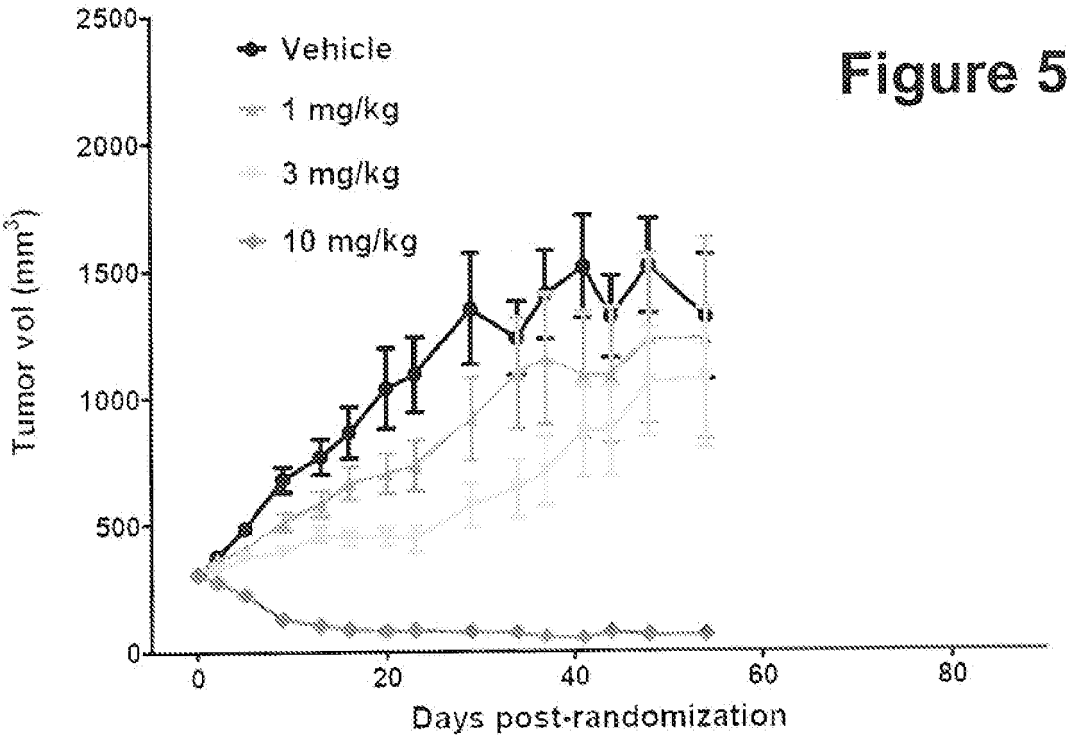


Figure 5E

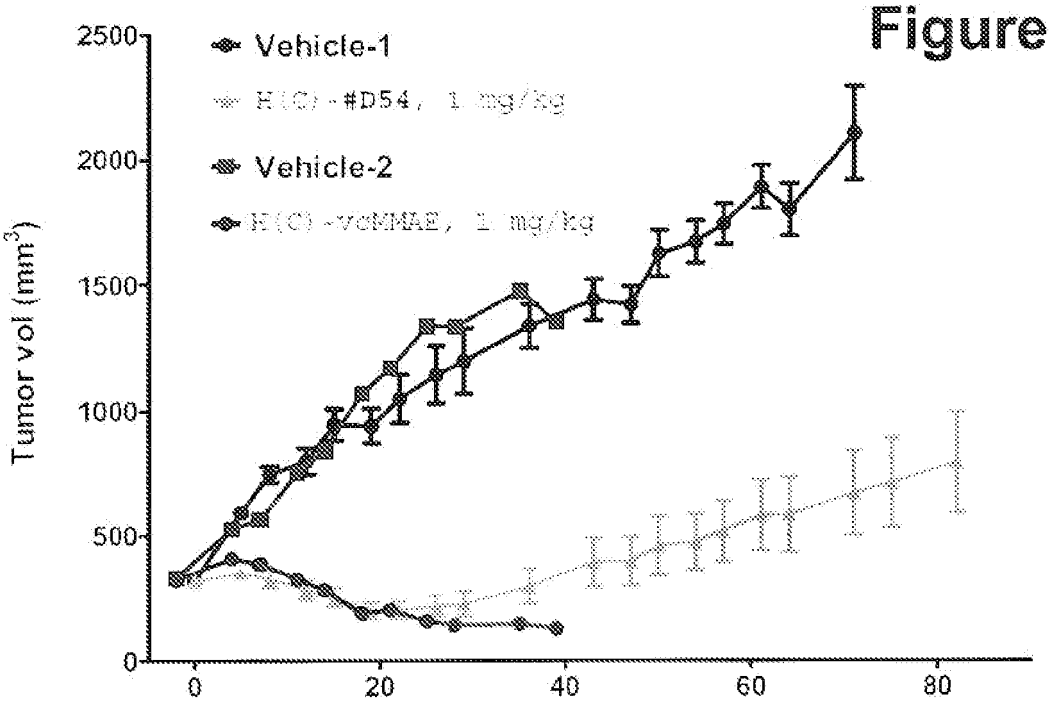


Figure 5F

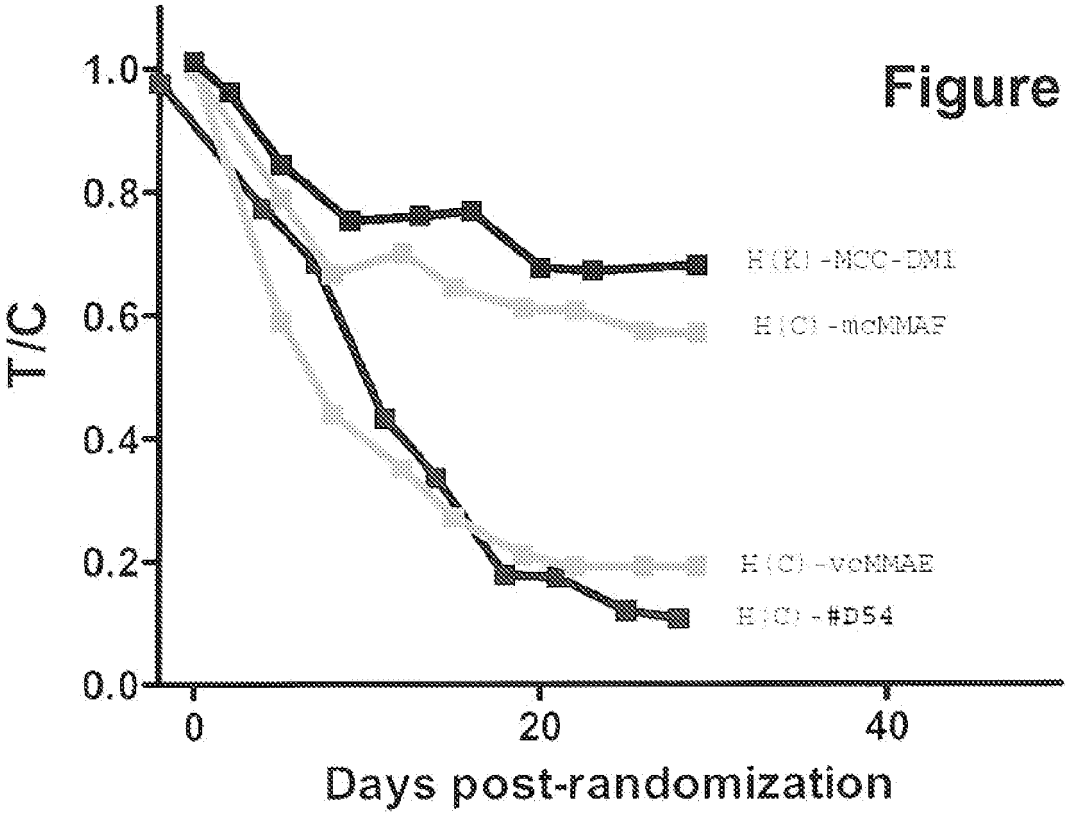


Figure 6

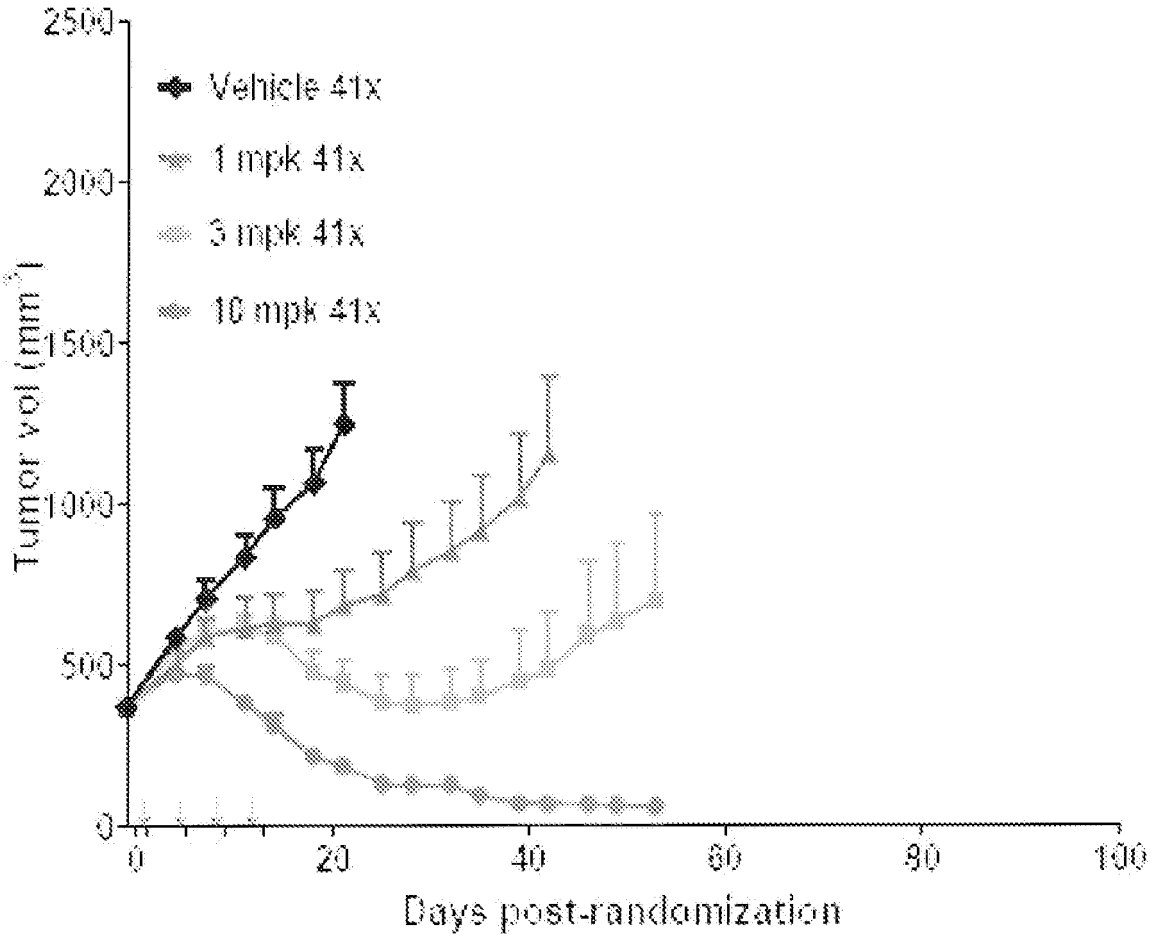

2500
2000
1500
1000
500
0
Tumor vol (mm3)
Vehicle 41x
1 mpk 41x
3 mpk 41x
10 mpk 41x
0
20
40
60
80
100
Days post-randomization

Figure 7

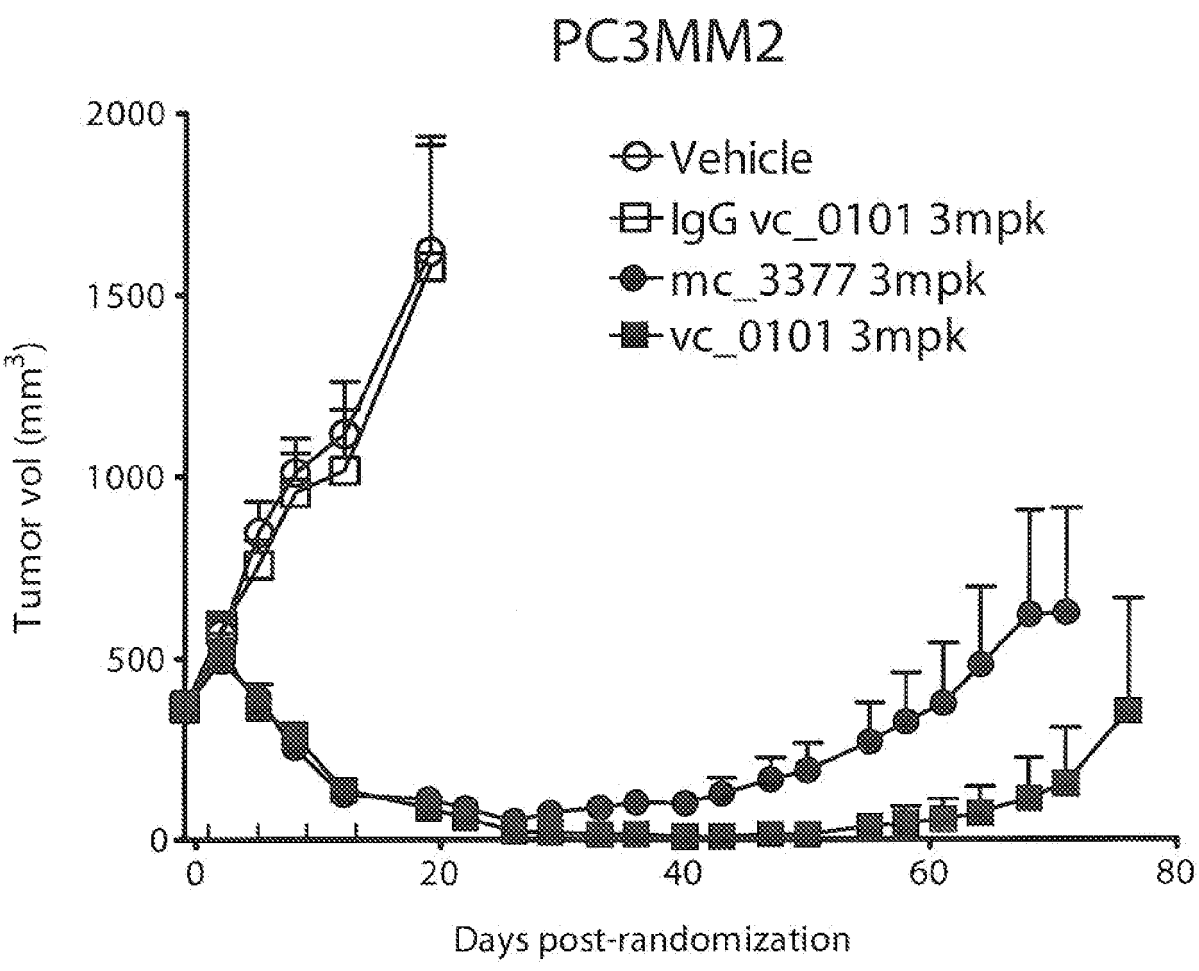

PC3MM2
Vehicle
IgG vc_0101 3mpk
mc_3377 3mpk
vc_0101 3mpk
Tumor vol (mm3)
2000
1500
1000
500
0
0
20
40
60
80
Days post-randomization

Figure 8A

Efficacy of rat-human chimeric anti-Notch ADC's dosed at 5mg/kg in HCC2429 lung xenografts

HCC2429 Lung xenografts, tumor volume (mm^3 +/- SEM)

Day	PBS	ch28-mc0101	ch28-mc3377	ch28-me0131	ch28-MALPEG6-C2-0131	ch75-me0131	ch28-vc0101	ch28-vc6780	ch75-vc0101	ch75-vc6780	huNeg8.8-mc0131	huNeg8.8-mc3377	huNeg8.8-me0131	huNeg8.9-vc0101	huNeg8.8-vc6780
-2	136 ±44	195 ±37	195 ±38	195 ±39	196 ±47	195 ±35	195 ±26	195 ±33	195 ±25	205 ±35	196 ±43	195 ±45	195 ±42	209 ±45	195 ±43
2	503 ±106	457 ±73	525 ±101	458 ±90	505 ±128	483 ±96	395 ±54	378 ±77	450 ±49	420 ±66	603 ±119	500 ±127	543 ±115	550 ±135	516 ±117
4	838 ±187	506 ±66	654 ±136	504 ±112	638 ±192	622 ±125	596 ±46	528 ±73	366 ±47	430 ±92	845 ±149	637 ±175	746 ±188	845 ±201	758 ±177
6	1008 ±229	562 ±126	783 ±156	600 ±152	750 ±207	634 ±158	559 ±33	519 ±79	515 ±48	474 ±108	1117 ±171	939 ±243	1016 ±225	1118 ±270	968 ±229
9	1618 ±265	652 ±133	1186 ±253	787 ±177	1020 ±213	828 ±129	777 ±24	780 ±72	717 ±30	502 ±125	1773 ±234	1348 ±284	1508 ±387	1549 ±394	1449 ±304
11	1605± 228	761 ±166	1458 ±314	941 ±199	1362 ±257	737 ±162	1169 ±22	956 ±77	1196 ±30	801 ±155	1965 ±170	1608 ±259	1475 ±283	1532 ±407	1582 ±316
13	2033± 293	865 ±176	1534 ±339	1096 ±264	1598 ±274	802 ±184	1222 ±15	993 ±88	1166 ±33	735 ±166	2535 ±228	2257 ±350	1831 ±400	-	-
16	-	917 ±207	-	949 ±190	1556 ±300	878 ±200	988 ±13	866 ±105	970 ±26	781 ±210	-	-	-	-	-
19	-	1222 ±275	-	1322 ±276	1986 ±414	1051 ±247	884 ±14	484 ±171	1022 ±28	770 ±153	-	-	-	-	-
23	-	1694 ±461	-	-	-	1354 ±340	794 ±25	496 ±238	724 ±25	855 ±156	-	-	-	-	-
26	-	-	-	-	-	-	774 ±47	659 ±329	644 ±28	1159 ±253	-	-	-	-	-
30	-	-	-	-	-	-	1099 ±86	533 ±290	364 ±20	1270 ±205	-	-	-	-	-
33	-	-	-	-	-	-	1367 ±128	775 ±411	364 ±34	1628 ±309	-	-	-	-	-
37	-	-	-	-	-	-	2107 ±199	-	464 ±45	-	-	-	-	-	-
40	-	-	-	-	-	-	3235 ±311	-	587 ±57	-	-	-	-	-	-
44	-	-	-	-	-	-	1767 ±17	-	1087 ±107	-	-	-	-	-	-
51	-	-	-	-	-	-	4333 ±43	-	2017 ±200	-	-	-	-	-	-
58	-	-	-	-	-	-	8727 ±82	-	0±0	-	-	-	-	-	-
65	-	-	-	-	-	-	1398 ±138	-	0±0	-	-	-	-	-	-
73	-	-	-	-	-	-	3011 ±301	-	0±0	-	-	-	-	-	-
79	-	-	-	-	-	-	-	-	0±0	-	-	-	-	-	-
89	-	-	-	-	-	-	-	-	0±0	-	-	-	-	-	-
96	-	-	-	-	-	-	-	-	0±0	-	-	-	-	-	-

Figure 8B

Efficacy of rat-human chimeric anti-Notch ADCs dosed at 5mg/kg in MDA-MB-468 breast xenografts

	MDA-MB-468 Breast xenografts, tumor volume (mm^3 ± SEM)							
Day	PBS	ch75-mc3377	ch75-me0131	ch75-MALPEG6 CZ-0131	ch75-mc0131	huNeg8.8-mc0131	huNeg8.8-mc3377	huNeg8.8-me0131
-1	303 ±10	304 ±19	303 ±13	308 ±22	305 ±14	306 ±14	302 ±15	306 ±21
2	344 ±14	315 ±26	348 ±21	375 ±16	354 ±7	340 ±16	363 ±25	343 ±26
5	360 ±17	339 ±25	339 ±16	325 ±23	346 ±14	349 ±19	366 ±19	349 ±31
8	439 ±28	340 ±20	352 ±21	346 ±27	368 ±15	437 ±35	431 ±25	421 ±36
12	533 ±41	337 ±22	311 ±26	312 ±31	350 ±18	492 ±27	449 ±30	466 ±41
15	542 ±40	304 ±23	297 ±22	271 ±18	322 ±22	543 ±42	484 ±48	523 ±35
19	608 ±46	234 ±20	225 ±25	204 ±20	283 ±19	552 ±37	516 ±26	542 ±45
22	644 ±34	240 ±20	210 ±26	156 ±21	261 ±22	602 ±38	561 ±38	586 ±47
26	707 ±58	256 ±22	169 ±14	127 ±14	252 ±25	615 ±65	604 ±34	664 ±62
29	768 ±72	278 ±33	191 ±19	119 ±16	285 ±27	761 ±59	641 ±42	695 ±57
33	822 ±78	320 ±45	203 ±23	133 ±22	315 ±31	765 ±80	737 ±57	759 ±66
36	840 ±87	308 ±53	220 ±18	119 ±20	320 ±30	748 ±55	865 ±54	788 ±81
40	894 ±80	346 ±51	276 ±39	143 ±20	367 ±41	882 ±90	946 ±69	853 ±77
44	1040 ±104	373 ±49	312 ±34	162 ±29	421 ±29	1030 ±109	1055 ±54	1002 ±82
48	1174 ±146	459 ±69	382 ±58	193 ±31	455 ±42	1138 ±131	1148 ±36	1066 ±92
51	1266 ±144	539 ±83	418 ±49	237 ±31	563 ±48	1301 ±127	1368 ±57	1164 ±98
54	1244 ±155	543 ±80	422 ±45	220 ±37	592 ±52	1268 ±134	1321 ±56	1206 ±139
57	1366 ±163	594 ±97	474 ±85	260 ±36	665 ±56	1380 ±151	1530 ±23	1132 ±102
61	1492 ±129	659 ±100	497 ±80	275 ±36	690 ±52	1380 ±149	1607 ±39	-
64	-	718 ±117	519 ±93	363 ±45	758 ±64	1420 ±160	1580 ±47	-
68	-	872 ±152	636 ±117	369 ±48	-	1552 ±198	1560 ±71	-
71	-	897 ±151	705 ±102	377 ±65	-	-	-	-
78	-	-	906 ±137	511 ±74	-	-	-	-

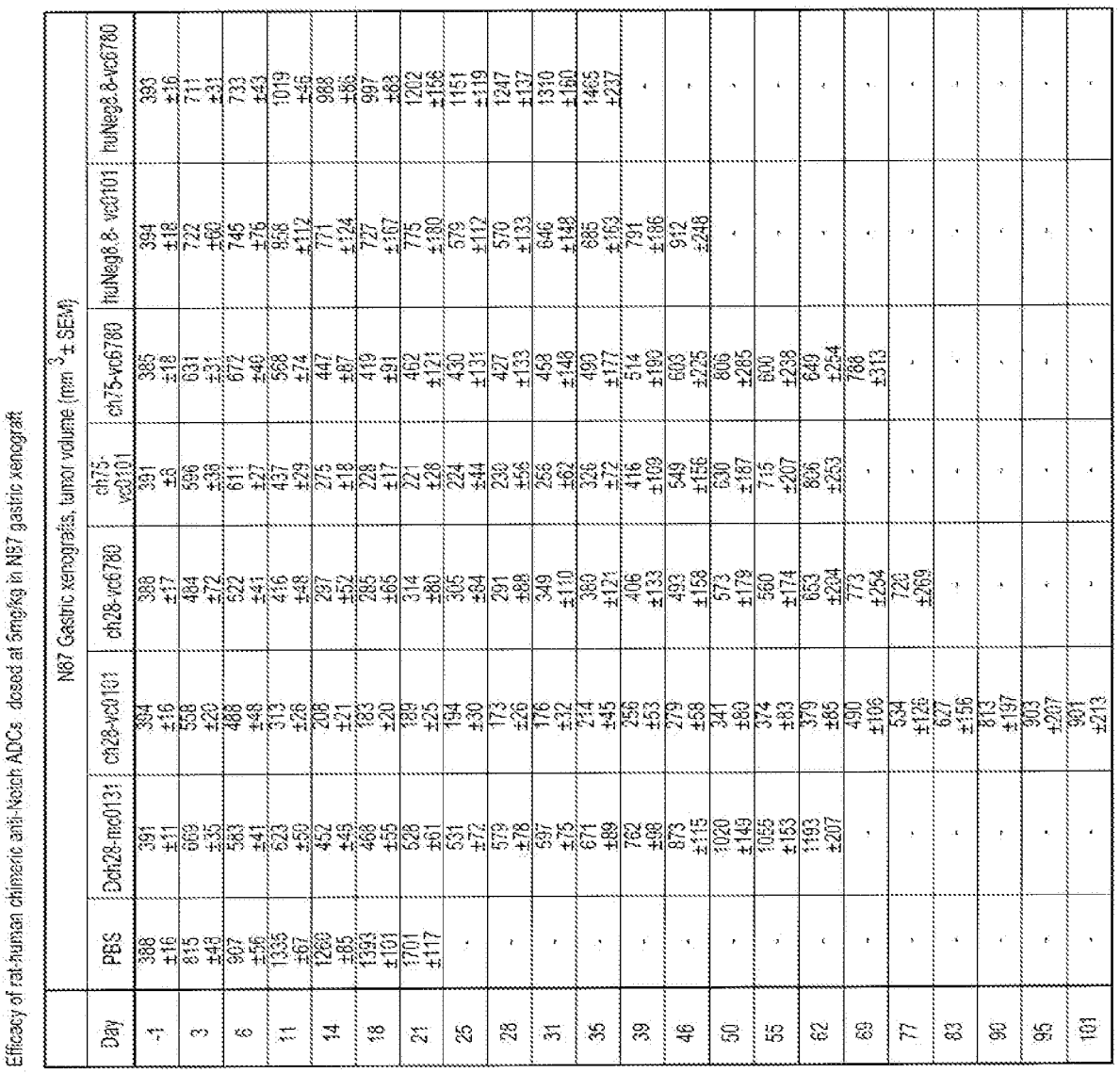
Figure 8C

Efficacy of rat-human chimeric anti-Notch ADCs dosed at 5mg/kg in MDA-MB-468 breast xenograft

MDA-MB-468 Breast xenografts, tumor volume (mm ^{3} ± SEM)

Day	PBS	ch28-me0131	ch28-mc3377	ch28-me0131	ch28-MALPEG 6C2-0131	ch75-me0131	ch28-vc0101	ch28-vc6780	ch75-vc0101	ch75-vc6780	huNeg8.8-me0131	huNeg8.8-mc3377	huNeg8.8-me0131	huNeg8.8-vc0101	huNeg8.8-vc6780
-1	309 ±11	314 ±12	313 ±20	314 ±15	311 ±22	307 ±19	313 ±22	309 ±23	312 ±7	305 ±8	313 ±21	310 ±15	308 ±22	312 ±18	314 ±21
4	396 ±20	358 ±19	321 ±20	329 ±18	331 ±26	337 ±16	303 ±21	323 ±20	328 ±21	341 ±17	353 ±27	333 ±19	358 ±27	333 ±27	379 ±18
8	414 ±25	315 ±16	297 ±17	306 ±14	295 ±25	315 ±14	241 ±24	293 ±21	297 ±23	323 ±21	384 ±33	391 ±26	398 ±32	349 ±22	387 ±18
11	452 ±31	262 ±15	237 ±15	256 ±12	238 ±22	324 ±22	149 ±17	237 ±25	219 ±16	308 ±25	422 ±33	408 ±31	422 ±40	360 ±35	426 ±21
18	578 ±48	143 ±11	121 ±18	173 ±21	115 ±9	257 ±28	38 ±8	136 ±19	108 ±12	257 ±35	496 ±45	479 ±40	537 ±56	310 ±61	485 ±22
21	615 ±46	107 ±9	119 ±10	145 ±19	68 ±6	272 ±27	24 ±7	138 ±23	70 ±10	232 ±43	499 ±47	569 ±49	572 ±55	275 ±66	545 ±43
28	691 ±89	113 ±16	132 ±16	95 ±10	26 ±10	171 ±24	14 ±7	163 ±28	37 ±11	259 ±50	584 ±51	638 ±72	638 ±61	251 ±66	614 ±31
35	835 ±87	137 ±25	191 ±20	127 ±19	15 ±8	236 ±28	7 ±5	176 ±33	26 ±9	340 ±58	662 ±64	758 ±101	772 ±86	279 ±72	737 ±64
39	968 ±103	179 ±37	234 ±19	145 ±33	32 ±12	267 ±36	0 ±0	229 ±41	13 ±9	400 ±74	702 ±56	882 ±110	881 ±114	361 ±99	860 ±65
42	1004 ±112	168 ±35	256 ±22	174 ±37	32 ±13	240 ±42	0 ±0	242 ±41	10 ±8	420 ±72	753 ±75	945 ±123	926 ±115	362 ±90	888 ±49
46	1140 ±126	191 ±43	290 ±27	188 ±48	44 ±16	293 ±35	0 ±0	243 ±41	11 ±8	442 ±77	834 ±77	1016 ±148	930 ±162	377 ±101	965 ±102
49	1211 ±92	233 ±56	333 ±33	207 ±45	37 ±17	334 ±46	0 ±0	295 ±52	7 ±7	435 ±86	896 ±90	1114 ±150	1133 ±171	439 ±115	1038 ±128
57	1402 ±112	298 ±67	447 ±37	272 ±59	72 ±28	400 ±53	0 ±0	369 ±65	13 ±13	652 ±110	1093 ±106	1355 ±189	1328 ±175	577 ±155	1275 ±143
60	-	328 ±69	513 ±42	335 ±67	89 ±28	472 ±59	0 ±0	373 ±61	25 ±16	688 ±118	1108 ±97	1298 ±138	1187 ±161	483 ±38	1161 ±77
63	-	320 ±70	516 ±44	354 ±67	67 ±29	498 ±58	0 ±0	415 ±76	21 ±14	709 ±114	1092 ±113	1379 ±137	1332 ±160	579 ±148	1270 ±93
67	-	379 ±85	538 ±49	377 ±77	85 ±37	563 ±63	0 ±0	462 ±94	21 ±15	708 ±136	1203 ±121	-	-	655 ±166	-
74	-	434 ±98	582 ±57	382 ±47	114 ±47	585 ±74	0 ±0	499 ±91	42 ±27	925 ±143	-	-	-	715 ±206	-
77	-	470 ±113	-	423 ±42	122 ±51	-	0 ±0	-	45 ±26	-	-	-	-	-	-
84	-	-	-	-	177 ±72	-	0 ±0	-	67 ±37	-	-	-	-	-	-
98	-	-	-	-	162 ±75	-	0 ±0	-	55 ±37	-	-	-	-	-	-
104	-	-	-	-	-	-	0 ±0	-	-	-	-	-	-	-	-
112	-	-	-	-	-	-	0 ±0	-	-	-	-	-	-	-	-
117	-	-	-	-	-	-	0 ±0	-	-	-	-	-	-	-	-
126	-	-	-	-	-	-	0 ±0	-	-	-	-	-	-	-	-

Figure 8D



Efficacy of rat-human chimeric anti-Notch ADCs dosed at 5mg/kg in N87 gastric xenograft

	N87 Gastric xenografts, tumor volume (mm ^{3} ± SEM)								
Day	PBS	Dch28-mc0131	ch28-vc0131	ch28-vc6780	ch75-vc0101	ch75-vc6780	huNeg8.8- vc0101	huNeg8.8-vc6780
-1	388 ±16	391 ±11	394 ±16	388 ±17	391 ±6	385 ±18	394 ±18	393 ±16
3	815 ±48	669 ±35	558 ±28	484 ±72	596 ±36	631 ±31	722 ±60	711 ±31
6	907 ±56	583 ±41	488 ±48	522 ±41	611 ±27	672 ±40	745 ±76	733 ±43
11	1035 ±67	623 ±58	313 ±28	416 ±48	437 ±29	568 ±74	956 ±112	1019 ±46
14	1268 ±85	452 ±48	208 ±21	297 ±52	275 ±18	447 ±87	771 ±124	968 ±86
18	1393 ±101	468 ±55	183 ±20	285 ±65	228 ±17	419 ±91	727 ±107	997 ±83
21	1701 ±117	528 ±61	189 ±25	314 ±80	221 ±28	462 ±121	775 ±130	1202 ±158
25	-	531 ±72	164 ±30	305 ±84	224 ±44	430 ±131	579 ±112	1151 ±119
28	-	579 ±78	173 ±26	291 ±88	238 ±56	427 ±133	570 ±133	1247 ±137
31	-	597 ±75	176 ±32	349 ±110	256 ±62	458 ±148	646 ±148	1310 ±160
35	-	671 ±89	214 ±45	360 ±121	326 ±72	493 ±177	685 ±163	1465 ±237
39	-	762 ±98	256 ±53	406 ±133	416 ±109	514 ±190	791 ±196	-
46	-	873 ±115	279 ±58	493 ±158	549 ±156	603 ±225	912 ±248	-
50	-	1020 ±149	341 ±80	573 ±179	630 ±187	805 ±285	-	-
55	-	1055 ±153	374 ±83	560 ±174	715 ±207	800 ±238	-	-
62	-	1193 ±207	379 ±85	653 ±234	806 ±253	849 ±254	-	-
69	-	-	490 ±106	773 ±254	-	788 ±313	-	-
77	-	-	534 ±128	726 ±269	-	-	-	-
83	-	-	627 ±156	-	-	-	-	-
90	-	-	813 ±197	-	-	-	-	-
95	-	-	903 ±267	-	-	-	-	-
101	-	-	991 ±213	-	-	-	-	-

Figure 8E

Efficacy of rat-human chimeric anti-Notch ADCs dosed at 5mg/kg in N87 gastric xenografts

	N87 Gastric xenografts, tumor volume (mm^3 ± SEM)				
Day	PBS	ch28-mc0131	ch28-m(H2O)c-0131	ch75-mc0131	ch75-m(H2O)c-0131
-1	372 ±8	371 ±18	371 ±17	364 ±12	373 ±15
4	589 ±28	473 ±41	469 ±41	461 ±33	497 ±34
7	706 ±56	479 ±39	446 ±34	466 ±28	484 ±35
11	837 ±71	442 ±37	391 ±29	468 ±33	414 ±28
14	959 ±98	395 ±43	303 ±24	430 ±33	337 ±20
18	1069 ±106	329 ±36	231 ±27	424 ±41	279 ±14
21	1252 ±123	357 ±48	232 ±29	448 ±42	268 ±13
25	1230 ±112	324 ±55	230 ±38	422 ±45	271 ±11
28	1352 ±132	363 ±62	256 ±41	487 ±52	266 ±17
32		381 ±66	255 ±49	530 ±56	298 ±22
35		419 ±85	283 ±52	541 ±56	307 ±21
39		453 ±87	315 ±65	570 ±55	358 ±23
42		522 ±108	347 ±71	619 ±66	388 ±34
46		559 ±119	381 ±87	629 ±68	432 ±41
49		537 ±139	418 ±96	697 ±87	447 ±48
53		681 ±149	441 ±107	755 ±91	484 ±52
56		713 ±167	479 ±113	753 ±80	512 ±55
63		783 ±192	542 ±139	868 ±112	563 ±70
70		847 ±219	535 ±148	827 ±107	579 ±86
77		1081 ±273	726 ±191	1041 ±139	788 ±110
84			855 ±230	1172 ±165	872 ±131
91			722 ±194	1332 ±194	1015 ±169
98			757 ±204		1125 ±198