

28F3 QVQLVESGGGVVQPGKSLRLSCAASGFTFSSYGMHWVRQA PGKGLEWVAVIWIYEGSNKYYADSVKGRFTI SRDNSKNTLYLQMNSLRAEDTAVYYCARGGSMVRGDYYYGMDVWGQGTITVTVSS
18E10
QVQLVESGGGVVQPGKSLRLSCAASGFTFSSYGMHWVRQA PGKGLEWVAVIWIYAGSNKYYADSVKGRFTI SRDNSKNTLYLQMNSLRAEDTAVYYCARGGRIA VAFYYSMVMDVWGQGTITVTVSS
19D3 QVQLVESGGGVVQPGKSLRLSCAASGFTFSSYGFHWVRQA PGKGLEWVAVIWIYAGSNKFYADSVKGRFTI SRDNSKNTLYLQMNSLRAEDTAVYYCARGGQLDYIYVMDVWGQGTITVTVSS
VL: 28F3 AIQLTQSPSSLSASVGDRVITTCRASQGISSALAWYQQKPKAPKLLIYDASSLESGVPSRFSGSGSGTDFTLTITISSSLQPEDFATYYCQQFNSEYPTFGQGTKLEIK
18E10 DIQMTQSPSSLSASVGDRVITTCRASQGISSWLAWYQQKPEKAPKSLIYAAASSLQSGVPSRFSGSGSGTDFTLTITISSSLQPEDFATYYCQQYNSYPTFGQGTKLEIK
19D3 DIQMTQSPSSLSASVGDRVITTCRASQGISSWLAWYQQKPEKAPKSLIYAAASSLQSGVPSRFSGSGSGTDFTLTITISSSLQPEDFATYYCQQYNSYPTFGQGTKLEIK

Figura 1

Anti-GITR 28F3 VH (hIgG2)

Segmento V: 3-33

Segmento D: 3-10

Segmento J: JH6b

1	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S
	CAG	GTG	CAG	CTG	GTG	GAG	TCT	GGG	GGA	GGC	GTG	GTC	CAG	CCT	GGG	AGG	TCC
52	L	R	L	S	C	A	A	S	G	F	T	F	S	_CDR1_			
	CTG	AGA	CTC	TCC	TGT	GCA	GCG	TCT	GGA	TTC	ACC	TTC	AGT	AGC	TAT	GGC	ATG
103	H	W	V	R	Q	A	P	G	K	G	L	E	W	V	A	_CDR2_	
	CAC	TGG	GTC	CGC	CAG	GCT	CCA	GGC	AAG	GGG	CTG	GAG	TGG	GTG	GCA	GTT	ATA
154	W	Y	E	G	S	N	K	Y	Y	A	D	S	V	K	G	R	F
	TGG	TAT	GAA	GGA	AGT	AAT	AAA	TAT	TAT	GCA	GAC	TCC	GTG	AAG	GGC	CGA	TTC
205	T	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S
	ACC	ATC	TCC	AGA	GAC	AAT	TCC	AAG	AAC	ACG	CTG	TAT	CTG	CAA	ATG	AAC	AGC
256	L	R	A	E	D	T	A	V	Y	Y	C	A	R	_CDR3_			
	CTG	AGA	GCC	GAG	GAC	ACG	GCT	GTG	TAT	TAC	TGT	GCG	AGA	GGG	GGG	AGT	ATG
307	V	R	G	D	Y	Y	Y	G	M	D	V	W	G	Q	G	T	T
	GTT	CGG	GGG	GAC	TAC	TAC	TAC	GGT	ATG	GAC	GTC	TGG	GGC	CAA	GGG	ACC	ACG
358	V	T	V	S	S												
	GTC	ACC	GTC	TCC	TCA												

Figura 2A

Anti-GITR 28F3 VK (hKappa)

Segmento V: L18

Segmento J: JK2

1	A	I	Q	L	T	Q	S	P	S	S	L	S	A	S	V	G	D
	GCC	ATC	CAG	TTG	ACC	CAG	TCT	CCA	TCC	TCC	CTG	TCT	GCA	TCT	GTA	GGA	GAC
							CDR1										
52	R	V	T	I	T	C	R	A	S	Q	G	I	S	S	A	L	A
	AGA	GTC	ACC	ATC	ACT	TGC	CGG	GCA	AGT	CAG	GGC	ATT	AGC	AGT	GCT	TTA	GCC
															CDR2		
103	W	Y	Q	Q	K	P	G	K	A	P	K	L	L	I	Y	D	A
	TGG	TAT	CAG	CAG	AAA	CCA	GGG	AAA	GCT	CCT	AAG	CTC	CTG	ATC	TAT	GAT	GCC
154	S	S	L	E	S	G	V	P	S	R	F	S	G	S	G	S	G
	TCC	AGT	TTG	GAA	AGT	GGG	GTC	CCA	TCA	AGG	TTC	AGC	GGC	AGT	GGA	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T
	ACA	GAT	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTG	CAG	CCT	GAA	GAT	TTT	GCA	ACT
							CDR3										
256	Y	Y	C	Q	Q	F	N	S	Y	P	Y	T	F	G	Q	G	T
	TAT	TAC	TGT	CAA	CAG	TTT	AAT	AGT	TAC	CCG	TAC	ACT	TTT	GGC	CAG	GGG	ACC
307	K	L	E	I	K												
	AAG	CTG	GAG	ATC	AAA												

Figura 2B

Anti-GITR 18E10 VH (hIgG2)

Segmento V: 3-33

Segmento D: 6-19

Segmento J: JH6b

1	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S
	CAG	GTG	CAG	CTG	GTG	GAG	TCT	GGG	GGA	GGC	GTG	GTC	CAG	CCT	GGG	AGG	TCC
52	L	R	L	S	C	A	A	S	G	F	T	F	S	_CDR1_			
	CTG	AGA	CTC	TCC	TGT	GCA	GCG	TCT	GGA	TTC	ACC	TTC	AGT	AGC	TAT	GGC	ATG
103	H	W	V	R	Q	A	P	G	K	G	L	E	W	V	A	V	I
	CAC	TGG	GTC	CGC	CAG	GCT	CCA	GGC	AAG	GGG	CTG	GAG	TGG	GTG	GCA	GTT	ATA
154	W	Y	A	G	S	N	K	Y	Y	A	D	S	V	K	G	R	F
	TGG	TAT	GCT	GGA	AGT	AAT	AAA	TAC	TAT	GCA	GAC	TCC	GTG	AAG	GGC	CGA	TTC
205	T	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S
	ACC	ATC	TCC	AGA	GAC	AAT	TCC	AAG	AAC	ACG	CTG	TAT	CTG	CAA	ATG	AAC	AGC
256	L	R	A	E	D	T	A	V	Y	Y	C	A	R	_CDR3_			
	CTG	AGA	GCC	GAG	GAC	ACG	GCT	GTG	TAT	TAC	TGT	GCG	AGA	GGG	GGG	CGT	ATA
307	A	V	A	F	Y	Y	S	M	D	V	W	G	Q	G	T	T	V
	GCA	GTG	GCC	TTC	TAC	TAC	AGT	ATG	GAC	GTC	TGG	GGC	CAA	GGG	ACC	ACG	GTC
358	T	V	S	S													
	ACC	GTC	TCC	TCA													

Figura 3A

Anti-GITR 18E10 VK (hKappa)

Segmento V: L15

Segmento J: JK2

1	D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D
	GAC	ATC	CAG	ATG	ACC	CAG	TCT	CCA	TCC	TCA	CTG	TCT	GCA	TCT	GTA	GGA	GAC
							CDR1										
52	R	V	T	I	T	C	R	A	S	Q	G	I	S	S	W	L	A
	AGA	GTC	ACC	ATC	ACT	TGT	CGG	GCG	AGT	CAG	GGT	ATT	AGC	AGC	TGG	TTA	GCC
																CDR2	
103	W	Y	Q	Q	K	P	E	K	A	P	K	S	L	I	Y	A	A
	TGG	TAT	CAG	CAG	AAA	CCA	GAG	AAA	GCC	CCT	AAG	TCC	CTG	ATC	TAT	GCT	GCA
154	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G
	TCC	AGT	TTG	CAA	AGT	GGG	GTC	CCA	TCA	AGG	TTC	AGC	GGC	AGT	GGA	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T
	ACA	GAT	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTG	CAG	CCT	GAA	GAT	TTT	GCA	ACT
				CDR3													
256	Y	Y	C	Q	Q	Y	N	S	Y	P	Y	T	F	G	Q	G	T
	TAT	TAC	TGC	CAA	CAG	TAT	AAT	AGT	TAC	CCG	TAC	ACT	TTT	GGC	CAG	GGG	ACC
307	K	L	E	I	K												
	AAG	CTG	GAG	ATC	AAA												

Figura 3B

Anti-GITR 19D3 VH (hIgG2)

Segmento V: 3-33

Segmento D: 3-16

Segmento J: JH6b

1	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S
	CAG	GTG	CAG	CTG	GTG	GAG	TCT	GGG	GGA	GGC	GTG	GTC	CAA	CCT	GGG	AGG	TCC
52	L	R	L	S	C	A	A	S	G	F	T	F	S	_CDR1_			
	CTG	AGA	CTC	TCC	TGT	GCA	GCG	TCT	GGA	TTC	ACC	TTC	AGT	AGC	TAT	GGC	TTC
103	H	W	V	R	Q	A	P	G	K	G	L	E	W	V	A	_CDR2_	
	CAC	TGG	GTC	CGC	CAG	GCT	CCA	GGC	AAG	GGG	CTG	GAG	TGG	GTG	GCA	GTT	ATA
154	W	Y	A	G	S	N	K	F	Y	A	D	S	V	K	G	R	F
	TGG	TAT	GCT	GGA	AGT	AAT	AAA	TTC	TAT	GCA	GAC	TCC	GTG	AAG	GGC	CGA	TTC
205	T	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S
	ACC	ATC	TCC	AGA	GAC	AAT	TCC	AAG	AAC	ACG	CTG	TAT	CTG	CAA	ATG	AAC	AGC
256	L	R	A	E	D	T	A	V	Y	Y	C	A	R	_CDR3_			
	CTA	AGA	GCC	GAG	GAC	ACG	GCT	GTG	TAT	TAC	TGT	GCG	AGA	GGG	GGA	CAG	TTG
307	D	Y	Y	Y	Y	Y	V	M	D	V	W	G	Q	G	T	T	V
	GAC	TAC	TAC	TAC	TAT	TAC	GTT	ATG	GAC	GTC	TGG	GGC	CAA	GGG	ACC	ACG	GTC
358	T	V	S	S													
	ACC	GTC	TCC	TCA													

Figura 4A

Anti-GITR 19D3 VK (hKappa)

Segmento V: L15

Segmento J: JK2

1	D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D
	GAC	ATC	CAG	ATG	ACC	CAG	TCT	CCA	TCC	TCA	CTG	TCT	GCA	TCT	GTA	GGA	GAC
							CDR1										
52	R	V	T	I	T	C	R	A	S	Q	G	I	S	S	W	L	A
	AGA	GTC	ACC	ATC	ACT	TGT	CGG	GCG	AGT	CAG	GGT	ATT	AGC	AGC	TGG	TTA	GCC
																CDR2	
103	W	Y	Q	Q	K	P	E	K	A	P	K	S	L	I	Y	A	A
	TGG	TAT	CAG	CAG	AAA	CCA	GAG	AAA	GCC	CCT	AAG	TCC	CTG	ATC	TAT	GCT	GCA
154	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G
	TCC	AGT	TTG	CAA	AGT	GGG	GTC	CCA	TCA	AGG	TTC	AGC	GGC	AGT	GGA	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T
	ACA	GAT	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTG	CAG	CCT	GAA	GAT	TTT	GCA	ACT
				CDR3													
256	Y	Y	C	Q	Q	Y	N	S	Y	P	Y	T	F	G	Q	G	T
	TAT	TAC	TGC	CAA	CAG	TAT	AAT	AGT	TAC	CCG	TAC	ACT	TTT	GGC	CAG	GGG	ACC
307	K	L	E	I	K												
	AAG	CTG	GAG	ATC	AAA												

Figura 4B

Anti-GITR 3C3 VH (hIgG1)

Segmento V: 4-34

Segmento J: JH3b

1	CAG GTG CAA CTA CAG CAG TGG GGC GCA GGA CTG TTG AAG CCT TCG GAG ACC
	LSLTCAVYGGSFS CTG TCC CTC ACC TGC GCT GTC TAT GGT GGG TCC TTC AGT CDR1 GYYW GGT TAC TAC TGG
52	
	TWIRQPPGKGLEWIG ACC TGG ATC CGC CAG CCC CCA GGG AAG GGG CTG GAG TGG ATT GGG CDR2 KI AAA ATC
103	
	NHSGNTNYNPSLKS AAT CAT AGT GGA AAC ACC AAC TAC AAC CCG TCC CTC AAG AGT RVT CGA GTC ACC
154	
	ISVDTSKNQFSLKLSSV ATA TCA GTA GAC ACG TCC AAG AAC CAG TTC TCC CTG AAG CTG AGC TCT GTG
205	
	TAADTAVYYCAR ACC GCC GCG GAC ACG GCT GTG TAT TAC TGT GCG AGA CDR3 LGAFD CTG GGG GCC TTT GAT
256	
	AFDI GCT TTT GAT ATC WGQGTMVTVSS TGG GGC CAA GGG ACA ATG GTC ACC GTC TCT TCA
307	

Figura 5A

Anti-GITR 3C3 VK1 (hKappa)

Segmento V: L15

Segmento J: JK2

1	D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D
	GAC	ATC	CAG	ATG	ACC	CAG	TCT	CCA	TCC	TCA	CTG	TCT	GCA	TCT	GTA	GGA	GAC
							CDR1										
52	R	V	T	I	T	C	R	A	S	Q	G	I	S	S	W	L	A
	AGA	GTC	ACC	ATC	ACT	TGT	CGG	GCG	AGT	CAG	GGT	ATT	AGC	AGC	TGG	TTA	GCC
															CDR2		
103	W	Y	Q	Q	K	P	E	K	A	P	K	S	L	I	Y	A	A
	TGG	TAT	CAG	CAG	AAA	CCA	GAG	AAA	GCC	CCT	AAG	TCC	CTG	ATC	TAT	GCT	GCA
154	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G
	TCC	AGT	TTG	CAA	AGT	GGG	GTC	CCA	TCA	AGG	TTC	AGC	GGC	AGT	GGA	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T
	ACA	GAT	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTG	CAG	CCT	GAA	GAT	TTT	GCA	ACT
							CDR3										
256	Y	Y	C	Q	Q	Y	N	S	Y	P	Y	T	F	G	Q	G	T
	TAT	TAC	TGC	CAA	CAG	TAT	AAT	AGT	TAC	CCG	TAC	ACT	TTT	GGC	CAG	GGG	ACC
307	K	L	E	I	K												
	AAG	CTG	GAG	ATC	AAA												

Figura 5B

Anti-GITR 3C3 VK2 (hKappa)

Segmento V: L20

Segmento J: JK2

1	E	I	V	L	T	Q	S	P	A	T	L	S	L	S	P	G	E
	GAA	ATT	GTG	TTG	ACA	CAG	TCT	CCA	GCC	ACC	CTG	TCT	TTG	TCT	CCA	GGG	GAA
							CDR1										
52	R	A	T	L	S	C	R	A	S	Q	G	V	S	S	Y	L	A
	AGA	GCC	ACC	CTC	TCC	TGC	AGG	GCC	AGT	CAG	GGT	GTT	AGC	AGC	TAC	TTA	GCC
															CDR2		
103	W	Y	Q	Q	K	P	G	Q	A	P	R	L	L	I	Y	D	A
	TGG	TAC	CAG	CAG	AAA	CCT	GGC	CAG	GCT	CCC	AGG	CTC	CTC	ATC	TAT	GAT	GCA
154	S	N	R	A	T	G	I	P	A	R	F	S	G	S	G	P	G
	TCC	AAC	AGG	GCC	ACT	GGC	ATC	CCA	GCC	AGG	TTC	AGT	GGC	AGT	GGG	CCT	GGG
205	T	D	F	T	L	T	I	S	S	L	E	P	E	D	F	A	V
	ACA	GAC	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTA	GAG	CCT	GAA	GAT	TTT	GCA	GTT
				CDR3													
256	Y	Y	C	Q	Q	R	S	N	W	H	T	F	G	Q	G	T	K
	TAT	TAC	TGT	CAG	CAG	CGT	AGC	AAC	TGG	CAC	ACT	TTT	GGC	CAG	GGG	ACC	AAG
307	L	E	I	K													
	CTG	GAG	ATC	AAA													

Figura 5C

Anti-GITR 2G6 VH (hIgG1)

Segmento V: 3-33

Segmento J: JH6b

1	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	G	S
	CAG	GTT	CAG	CTG	GTG	GAG	TCT	GGG	GGA	GGC	GTG	GTC	CAG	CCT	GGG	GGG	TCC
52	L	R	L	S	C	A	A	S	G	F	I	L	S	CDR1			
	CTG	AGA	CTC	TCC	TGT	GCA	GCG	TCT	GGA	TTC	ATC	TTG	AGT	D	Y	G	M
103	H	W	V	R	Q	A	P	G	K	G	L	E	W	V	T	V	I
	CAC	TGG	GTC	CGC	CAG	GCT	CCA	GGC	AAG	GGA	CTG	GAG	TGG	GTG	ACA	GTT	ATC
154	W	Y	D	G	S	N	K	F	Y	V	D	S	V	K	G	R	F
	TGG	TAT	GAT	GGA	AGT	AAT	AAA	TTC	TAT	GTA	GAC	TCC	GTG	AAG	GGC	CGA	TTC
205	T	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S
	ACC	ATC	TCC	AGA	GAC	AAT	TCC	AAG	AAC	ACG	TTG	TAT	CTG	CAA	ATG	AAC	AGC
256	L	R	V	E	D	T	A	V	Y	Y	C	A	R	CDR3			
	CTG	AGA	GTC	GAG	GAC	ACG	GCT	GTG	TAT	TAC	TGT	GCG	AGA	G	G	R	L
307	A	T	G	H	F	Y	Y	V	M	D	V	W	G	Q	G	T	T
	GCA	ACA	GGT	CAC	TTC	TAC	TAC	GTT	ATG	GAC	GTC	TGG	GGC	CAA	GGG	ACC	ACG
358	V	T	V	S	S												
	GTC	ACC	GTC	TCC	TCA												

Figura 6A

Anti-GITR 2G6 VK (hKappa)

Segmento V: L15

Segmento J: JK2

1	D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D
	GAC	ATC	CAG	ATG	ACC	CAG	TCT	CCA	TCC	TCA	CTG	TCT	GCA	TCT	GTA	GGA	GAC
							CDR1										
52	R	V	T	I	T	C	R	A	S	Q	G	I	S	S	W	L	A
	AGA	GTC	ACC	ATC	ACT	TGT	CGG	GCG	AGT	CAG	GGT	ATT	AGC	AGC	TGG	TTA	GCC
															CDR2		
103	W	Y	Q	Q	K	P	E	K	A	P	K	S	L	I	Y	A	A
	TGG	TAT	CAG	CAG	AAA	CCA	GAG	AAA	GCC	CCT	AAG	TCC	CTG	ATC	TAT	GCT	GCA
154	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G
	TCC	AGT	TTG	CAA	AGT	GGG	GTC	CCA	TCA	AGG	TTC	AGC	GGC	AGT	GGA	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T
	ACA	GAT	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTG	CAG	CCT	GAA	GAT	TTT	GCA	ACT
							CDR3										
256	Y	Y	C	Q	Q	Y	N	S	Y	P	Y	T	F	G	Q	G	T
	TAT	TAC	TGC	CAA	CAG	TAT	AAT	AGT	TAC	CCG	TAC	ACT	TTT	GGC	CAG	GGG	ACC
307	K	L	E	I	K												
	AAG	CTG	GAG	ATC	AAA												

Figura 6B

Anti-GITR 8A6 VH (hIgG2)

Segmento V: 3-33

Segmento D: 3-10

Segmento J: JH6b

1	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S
	CAG	GTG	CAG	CTG	GTG	GAG	TCT	GGG	GGA	GGC	GTG	GTC	CAG	CCT	GGG	AGG	TCC
52	L	R	L	S	C	T	A	S	G	F	T	F	S	CDR1			
	CTG	AGA	CTC	TCC	TGT	ACA	GCG	TCT	GGA	TTC	ACC	TTC	AGT	AGC	TAT	GGC	ATG
103	Q	W	V	R	Q	A	P	G	K	G	L	E	W	V	A	V	I
	CAG	TGG	GTC	CGC	CAG	GCT	CCA	GGC	AAG	GGG	CTG	GAG	TGG	GTG	GCA	GTT	ATA
154	W	Y	E	G	S	N	K	Y	Y	A	D	S	V	K	G	R	F
	TGG	TAT	GAA	GGA	AGT	AAT	AAA	TAC	TAT	GCA	GAC	TCC	GTG	AAG	GGC	CGA	TTC
205	T	I	S	R	E	N	S	K	N	T	L	Y	L	Q	M	N	S
	ACC	ATC	TCC	AGA	GAA	AAT	TCC	AAG	AAC	ACG	CTG	TAT	CTG	CAA	ATG	AAC	AGC
256	L	R	A	E	D	T	A	V	Y	Y	C	A	R	CDR3			
	CTG	AGA	GCC	GAG	GAC	ACG	GCT	GTG	TAT	TAC	TGT	GCG	AGA	GGC	GGT	CTT	ATG
307	V	R	G	L	F	Y	Y	G	M	D	V	W	G	Q	G	T	T
	GTT	CGG	GGT	CTC	TTC	TAC	TAC	GGT	ATG	GAC	GTC	TGG	GGC	CAA	GGG	ACC	ACG
358	V	T	V	S	S												
	GTC	ACC	GTC	TCC	TCA												

Figura 7A

Anti-GITR 8A6 VK (hKappa)

Segmento V: L18

Segmento J: JK2

1	A	I	Q	L	T	Q	S	P	S	S	L	S	A	S	V	G	D
	GCC	ATC	CAG	TTG	ACC	CAG	TCT	CCA	TCC	TCC	CTG	TCT	GCA	TCT	GTA	GGA	GAC
							CDR1										
52	R	V	T	I	T	C	R	A	S	Q	G	I	S	S	A	L	A
	AGA	GTC	ACC	ATC	ACT	TGC	CGG	GCA	AGT	CAG	GGC	ATT	AGC	AGT	GCT	TTA	GCC
																CDR2	
103	W	Y	Q	Q	K	P	G	K	A	P	K	F	L	I	Y	D	A
	TGG	TAT	CAG	CAG	AAA	CCA	GGG	AAA	GCT	CCT	AAG	TTC	CTG	ATC	TAT	GAT	GCC
154	S	S	L	E	S	G	V	P	S	R	F	S	G	S	G	S	G
	TCC	AGT	TTG	GAA	AGT	GGG	GTC	CCA	TCA	AGG	TTC	AGC	GGC	AGT	GGA	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T
	ACA	GAT	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTG	CAG	CCT	GAA	GAT	TTT	GCA	ACT
							CDR3										
256	Y	Y	C	Q	Q	F	N	S	Y	P	Y	T	F	G	Q	G	T
	TAT	TAC	TGT	CAA	CAG	TTT	AAT	AGT	TAC	CCG	TAC	ACT	TTT	GGC	CAG	GGG	ACC
307	K	L	E	I	K												
	AAG	CTG	GAG	ATC	AAA												

Figura 7B

Anti-GITR 9G7 VH (hIgG4)

Segmento V: 3-15

Segmento D: 3-10

Segmento J: JH6b

1	E	V	Q	L	V	E	S	G	G	G	L	V	K	P	G	G	S
	GAG	GTG	CAG	CTG	GTG	GAG	TCT	GGG	GGA	GGC	TTA	GTA	AAG	CCT	GGG	GGG	TCC
52	L	R	L	S	C	A	A	S	G	F	T	F	S	CDR1			
	CTT	AGA	CTC	TCC	TGT	GCA	GCC	TCT	GGA	TTC	ACT	TTC	AGT	ACC	GTC	TGG	ATG
103	S	W	V	R	Q	A	P	G	K	G	L	E	W	V	G	CDR2	
	AGC	TGG	GTC	CGC	CAG	GCT	CCA	GGG	AAG	GGG	CTG	GAG	TGG	GTT	GGC	CGT	ATT
154	K	S	K	T	D	G	G	T	T	D	Y	A	A	P	V	K	G
	AAA	AGC	AAA	ACT	GAT	GGT	GGG	ACA	ACA	GAC	TAC	GCT	GCA	CCC	GTG	AAA	GGC
205	R	F	T	I	S	R	D	D	S	K	N	T	L	Y	L	Q	M
	AGA	TTC	ACC	ATC	TCA	AGA	GAT	GAT	TCA	AAA	AAC	ACG	CTG	TAT	CTG	CAA	ATG
256	N	S	L	H	T	E	D	T	A	V	Y	Y	C	T	T	CDR3	
	AAC	AGC	CTG	CAC	ACC	GAG	GAC	ACA	GCC	GTG	TAT	TAC	TGT	ACC	ACA	GGG	CAG
307	L	I	P	Y	S	Y	Y	Y	G	M	D	V	W	G	Q	G	T
	CTG	ATC	CCT	TAC	TCC	TAC	TAC	TAC	GGT	ATG	GAC	GTC	TGG	GGC	CAA	GGG	ACC
358	S	V	T	V	S	S											
	TCG	GTC	ACC	GTC	TCC	TCA											

Figura 8A

Anti-GITR 9G7 VK1 (hKappa)

Segmento V: A27

Segmento J: JK1

E	I	V	L	T	Q	S	P	G	T	L	S	L	S	P	G	E									
1		GAA	ATT	GTG	TTG	ACG	CAG	TCT	CCA	GGC	ACC	CTG	TCT	TTG	TCT	CCA	GGG	GAA							
								CDR1																	
52		R	A	T	L	S	C	R	A	S	Q	S	V	S	S	S	Y	L							
		AGA	GCC	ACC	CTC	TCC	TGC	AGG	GCC	AGT	CAG	AGT	GTT	AGC	AGC	AGC	TAC	TTA							
																	CDR2								
103		A	W	Y	Q	Q	K	P	G	Q	A	P	R	L	L	I	Y	G							
		GCC	TGG	TAC	CAG	CAG	AAA	CCT	GGC	CAG	GCT	CCC	AGG	CTC	CTC	ATC	TAT	GGT							
									A	S	S	R	A	T	G	I	P	D	R	F	S	G	S	G	S
154		GCA	TCC	AGC	AGG	GCC	ACT	GGC	ATC	CCA	GAC	AGG	TTC	AGT	GGC	AGT	GGG	TCT							
		G	T	D	F	T	L	T	I	S	R	L	E	P	E	D	F	A							
205		GGG	ACA	GAC	TTC	ACT	CTC	ACC	ATC	AGC	AGA	CTG	GAG	CCT	GAA	GAT	TTT	GCA							
									CDR3																
256		V	Y	Y	C	Q	Q	Y	G	S	S	P	W	T	F	G	Q	G							
		GTG	TAT	TAC	TGT	CAG	CAG	TAT	GGT	AGC	TCA	CCG	TGG	ACG	TTC	GGC	CAA	GGG							
		T	K	V	E	I	K																		
307		ACC	AAG	GTG	GAA	ATC	AAA																		

Figura 8B

Anti-GITR 9G7 VK2 (hKappa)

Segmento V: A27

Segmento J: JK5

1	E	I	V	L	T	Q	S	P	G	T	L	S	L	S	P	G	E
	GAA	ATT	GTG	TTG	ACG	CAG	TCT	CCA	GGC	ACC	CTG	TCT	TTG	TCT	CCA	GGG	GAA
	CDR1																
52	R	A	T	L	S	C	R	A	S	Q	S	V	T	S	S	Y	L
	AGA	GCC	ACC	CTC	TCC	TGC	AGG	GCC	AGT	CAG	AGT	GTT	ACC	AGC	AGC	TAC	TTA
	CDR2																
103	A	W	Y	Q	Q	K	P	G	Q	A	P	R	L	L	I	Y	G
	GCC	TGG	TAC	CAG	CAG	AAA	CCT	GGC	CAG	GCT	CCC	AGG	CTC	CTC	ATC	TAT	GGT
	CDR3																
154	A	S	S	R	A	T	G	I	P	E	R	F	S	G	S	G	S
	GCA	TCC	AGC	AGG	GCC	ACT	GGC	ATC	CCA	GAG	AGG	TTC	AGT	GGC	AGT	GGG	TCT
205	G	T	D	F	T	L	T	I	S	R	L	E	P	E	D	F	A
	GGG	ACA	GAC	TTC	ACT	CTC	ACC	ATC	AGC	AGA	CTG	GAG	CCT	GAA	GAT	TTT	GCA
	CDR3																
256	V	Y	Y	C	Q	Q	Y	G	S	S	P	I	T	F	G	Q	G
	GTG	TAT	TAC	TGT	CAG	CAG	TAT	GGT	AGC	TCA	CCG	ATC	ACC	TTC	GGC	CAA	GGG
307	T	R	L	E	I	K											
	ACA	CGA	CTG	GAG	ATT	AAA											

Figura 8C

Anti-GITR 14E3 VH (hIgG1)

Segmento V: 4-34

Segmento J: JH3b

1	Q	V	Q	L	Q	Q	W	G	A	G	L	L	K	P	S	E	T
	CAG	GTG	CAG	CTA	CAG	CAG	TGG	GGC	GCA	GGA	CTG	TTG	AAG	CCT	TCG	GAG	ACC
52	L	S	L	T	C	A	V	Y	G	G	S	F	S	CDR1			
	CTG	TCC	CTC	ACC	TGC	GCT	GTC	TAT	GGT	GGG	TCC	TTC	AGT	G	Y	Y	W
103	S	W	I	R	Q	P	P	G	K	G	L	E	W	I	G	CDR2	
	AGC	TGG	ATC	CGC	CAG	CCC	CCA	GGG	AAG	GGG	CTG	GAG	TGG	ATT	GGA	E	I
154	N	H	S	G	N	T	Y	Y	N	P	S	L	K	S	R	V	T
	AAT	CAT	AGT	GGA	AAC	ACC	TAC	TAC	AAC	CCG	TCC	CTC	AAG	AGT	CGC	GTC	ACC
205	I	S	V	D	T	S	K	N	Q	L	S	L	K	L	S	S	V
	ATA	TCA	GTA	GAC	ACG	TCC	AAG	AAC	CAG	TTA	TCC	CTG	AAG	CTG	AGC	TCT	GTG
256	T	A	A	D	T	A	V	Y	Y	C	A	R	CDR3				
	ACC	GCC	GCG	GAC	ACG	GCT	GTG	TAT	TAC	TGT	GCG	AGA	TTT	G	S	N	D
307	A	F	D	I	W	G	Q	G	T	M	V	T	V	S	S		
	GCT	TTT	GAT	ATC	TGG	GGC	CAA	GGG	ACA	ATG	GTC	ACC	GTC	TCT	TCA		

Figura 9A

Anti-GITR 14E3 VK (hKappa)

Segmento V: L15

Segmento J: JK1

1	D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D
	GAC	ATC	CAG	ATG	ACC	CAG	TCT	CCA	TCC	TCA	CTG	TCT	GCA	TCT	GTA	GGA	GAC
							<u>CDR1</u>										
52	R	V	T	I	T	C	R	A	S	Q	G	I	S	S	W	L	A
	AGA	GTC	ACC	ATC	ACT	TGT	CGG	GCG	AGT	CAG	GGT	ATT	AGC	AGC	TGG	TTA	GCC
															<u>CDR2</u>		
103	W	Y	Q	Q	K	P	E	K	A	P	K	S	L	I	Y	A	A
	TGG	TAT	CAG	CAG	AAA	CCA	GAG	AAA	GCC	CCT	AAG	TCC	CTG	ATC	TAT	GCT	GCA
154	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G
	TCC	AGT	TTG	CAA	AGT	GGG	GTC	CCA	TCA	AGG	TTC	AGC	GGC	AGT	GGA	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T
	ACA	GAT	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTG	CAG	CCT	GAA	GAT	TTT	GCA	ACT
				<u>CDR3</u>													
256	Y	Y	C	Q	Q	Y	N	S	Y	P	P	T	F	G	Q	G	T
	TAT	TAC	TGC	CAA	CAG	TAT	AAT	AGT	TAC	CCT	CCG	ACG	TTC	GGC	CAA	GGG	ACC
307	K	V	E	I	K												
	AAG	GTG	GAA	ATC	AAA												

Figura 9B

Anti-GITR 19H8 VH (hIgG2)

Segmento V: 3-33

Segmento D: 3-10

Segmento J: JH6b

1	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S
	CAG	GTG	CAG	CTG	GTG	GAG	TCT	GGG	GGA	GGC	GTG	GTC	CAG	CCT	GGG	AGG	TCC
52	L	R	L	S	C	A	A	S	G	F	T	F	S	CDR1			
	CTG	AGA	CTC	TCC	TGT	GCA	GCG	TCT	GGA	TTC	ACC	TTC	AGT	AAC	TAT	GGC	ATG
103	H	W	V	R	Q	A	P	G	K	G	L	E	W	M	A	CDR2	
	CAC	TGG	GTC	CGC	CAG	GCT	CCA	GGC	AAG	GGG	CTG	GAG	TGG	ATG	GCA	GTT	ATA
154	W	Y	G	G	S	N	K	F	Y	A	D	S	V	K	G	R	F
	TGG	TAT	GGT	GGA	AGT	AAT	AAA	TTC	TAT	GCA	GAC	TCC	GTG	AAG	GGC	CGA	TTC
205	T	I	S	R	D	N	S	K	N	S	L	S	L	Q	M	N	S
	ACC	ATC	TCC	AGA	GAC	AAT	TCC	AAG	AAC	TCG	CTG	TCT	CTG	CAA	ATG	AAC	AGC
256	L	R	A	E	D	T	A	V	Y	Y	C	A	R	CDR3			
	CTG	AGA	GCC	GAG	GAC	ACG	GCT	GTG	TAT	TAC	TGT	GCG	AGA	GGG	GGG	GCT	ATG
307	V	R	G	V	Y	Y	Y	G	M	D	V	W	G	Q	G	T	T
	GTT	CGG	GGA	GTC	TAC	TAC	TAC	GGT	ATG	GAC	GTC	TGG	GGC	CAA	GGG	ACC	ACG
358	V	T	V	S	S												
	GTC	ACC	GTC	TCC	TCA												

Figura 10A

Anti-GITR 19H8 VK1 (hKappa)

Segmento V: L18

Segmento J: JK1

1	A	I	Q	L	T	Q	S	P	S	S	L	S	A	S	V	G	D
	GCC	ATC	CAG	TTG	ACC	CAG	TCT	CCA	TCC	TCC	CTG	TCT	GCA	TCT	GTA	GGA	GAC
	CDR1																
52	R	V	T	I	T	C	R	A	S	Q	G	I	S	S	A	L	A
	AGA	GTC	ACC	ATC	ACT	TGC	CGG	GCA	AGT	CAG	GGC	ATT	AGC	AGT	GCT	TTA	GCC
	CDR2																
103	W	Y	Q	Q	K	P	G	K	A	P	K	F	L	I	Y	D	A
	TGG	TAT	CAG	CAG	AAA	CCA	GGG	AAA	GCT	CCT	AAG	TTC	CTG	ATC	TAT	GAT	GCC
	CDR3																
154	S	S	L	E	S	G	V	P	S	R	F	S	G	S	G	S	G
	TCC	AGT	TTG	GAA	AGT	GGG	GTC	CCA	TCA	AGG	TTC	AGC	GGC	AGT	GGA	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T
	ACA	GAT	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTG	CAG	CCT	GAA	GAT	TTT	GCA	ACT
	CDR3																
256	Y	Y	C	Q	Q	F	N	S	Y	P	Q	T	F	G	Q	G	T
	TAT	TAC	TGT	CAA	CAG	TTT	AAT	AGT	TAC	CCT	CAG	ACG	TTC	GGC	CAA	GGG	ACC
307	K	V	E	I	K												
	AAG	GTG	GAA	ATC	AAA												

Figura 10B

Anti-GITR 19H8 VK2 (hKappa)

Segmento V: L6

Segmento J: JK4

1	E	I	V	L	T	Q	S	P	A	T	L	S	L	S	P	G	E
	GAA	ATT	GTG	TTG	ACA	CAG	TCT	CCA	GCC	ACC	CTG	TCT	TTG	TCT	CCA	GGG	GAA
							CDR1										
52	R	A	T	L	S	C	R	A	S	Q	S	V	S	S	Y	L	A
	AGA	GCC	ACC	CTC	TCC	TGC	AGG	GCC	AGT	CAG	AGT	GTT	AGC	AGC	TAC	TTA	GCC
																CDR2	
103	W	Y	Q	Q	K	P	G	Q	A	P	R	L	L	I	Y	D	A
	TGG	TAC	CAA	CAG	AAA	CCT	GGC	CAG	GCT	CCC	AGG	CTC	CTC	ATC	TAT	GAT	GCA
154	S	N	R	A	T	G	I	P	A	R	F	S	G	S	G	S	G
	TCC	AAC	AGG	GCC	ACT	GGC	ATC	CCA	GCC	AGG	TTC	AGT	GGC	AGT	GGG	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	E	P	E	D	F	A	V
	ACA	GAC	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTA	GAG	CCT	GAA	GAT	TTT	GCA	GTT
				CDR3													
256	Y	Y	C	Q	Q	R	S	N	W	P	L	T	F	G	G	G	T
	TAT	TAC	TGT	CAG	CAG	CGT	AGC	AAC	TGG	CCG	CTC	ACT	TTC	GGC	GGA	GGG	ACC
307	K	V	E	I	K												
	AAG	GTG	GAG	ATC	AAA												

Figura 10C

Anti-GITR 6G10 VH (hIgG2)

Segmento V: 3-33

Segmento D: 3-10

Segmento J: JH6b

1	Q	V	Q	L	V	E	S	G	G	D	V	V	Q	P	G	R	S
	CAG	GTG	CAG	CTG	GTG	GAG	TCT	GGG	GGA	GAC	GTG	GTC	CAG	CCT	GGG	AGG	TCC
52	L	R	L	S	C	A	A	S	G	F	T	F	S	CDR1			
	CTG	AGA	CTC	TCC	TGT	GCA	GCG	TCT	GGA	TTC	ACC	TTC	AGT	T	Y	G	M
103	H	W	V	R	Q	A	P	G	K	G	L	E	W	V	A	V	T
	CAC	TGG	GTC	CGC	CAG	GCT	CCA	GGC	AAG	GGG	CTG	GAG	TGG	GTG	GCA	GTT	ACA
154	W	Y	A	G	S	N	K	F	Y	A	D	S	V	K	G	R	F
	TGG	TAT	GCT	GGA	AGT	AAT	AAA	TTT	TAT	GCA	GAC	TCC	GTG	AAG	GGC	CGA	TTC
205	T	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S
	ACC	ATC	TCC	AGA	GAC	AAT	TCC	AAG	AAC	ACG	CTG	TAT	CTG	CAA	ATG	AAC	AGC
256	L	R	A	E	D	T	A	V	Y	Y	C	A	R	CDR3			
	CTG	AGA	GCC	GAG	GAC	ACG	GCT	GTG	TAT	TAC	TGT	GCG	AGA	G	G	S	M
307	V	R	G	L	Y	Y	Y	G	M	D	V	W	G	Q	G	T	T
	GTT	CGG	GGA	CTT	TAT	TAT	TAC	GGT	ATG	GAC	GTC	TGG	GGC	CAA	GGG	ACC	ACG
358	V	T	V	S	S												
	GTC	ACC	GTC	TCC	TCA												

Figura 11A

Anti-GITR 6G10 VK1 (hKappa)

Segmento V: L18

Segmento J: JK2

1	A	I	Q	L	T	Q	S	P	S	S	L	S	A	S	V	G	D
	GCC	ATC	CAG	TTG	ACC	CAG	TCT	CCA	TCC	TCC	CTG	TCT	GCA	TCT	GTA	GGA	GAC
	CDR1																
52	R	V	T	I	T	C	R	A	S	Q	G	I	S	S	A	L	A
	AGA	GTC	ACC	ATC	ACT	TGC	CGG	GCA	AGT	CAG	GGC	ATT	AGC	AGT	GCT	TTA	GCC
	CDR2																
103	W	Y	Q	Q	K	P	G	K	A	P	K	L	L	I	Y	D	A
	TGG	TAT	CAG	CAG	AAA	CCA	GGG	AAA	GCT	CCT	AAG	CTC	CTG	ATC	TAT	GAT	GCC
	CDR3																
154	S	S	L	E	S	G	V	P	S	R	F	S	G	S	G	S	G
	TCC	AGT	TTG	GAA	AGT	GGG	GTC	CCA	TCA	AGG	TTC	AGC	GGC	AGT	GGA	TCT	GGG
205	T	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T
	ACA	GAT	TTC	ACT	CTC	ACC	ATC	AGC	AGC	CTG	CAG	CCT	GAA	GAT	TTT	GCA	ACT
	CDR3																
256	Y	Y	C	Q	Q	F	N	S	Y	P	Y	T	F	G	Q	G	T
	TAT	TAC	TGT	CAA	CAG	TTT	AAT	AGT	TAC	CCG	TAC	ACT	TTT	GGC	CAG	GGG	ACC
307	K	L	E	I	K												
	AAG	CTG	GAG	ATC	AAA												

Figura 11B

Anti-GITR 28F3 VH (hIgG2)

3-33	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S	L	R	L	S	C	A	
28F3 VH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
3-33	A	S	G	F	T	F	S	<u>CDR1</u>	S	Y	G	M	H	W	V	R	Q	A	P	G	K	G	L	E
28F3 VH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3-33	W	V	A	<u>CDR2</u>	V	I	W	Y	D	G	S	N	K	Y	Y	A	D	S	V	K	G	R	F	T
28F3 VH	-	-	-	-	-	-	-	E	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3-33	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S	L	R	A	E	D	T	A	
28F3 VH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3-33	V	Y	Y	C	A	R	<u>CDR3</u>																	
3-10											M	V	R	G										
JH6																Y	Y	Y	G	M	D	V	W	G
28F3 VH	-	-	-	-	-	-	G	G	S	-	-	-	-	D	-	-	-	-	-	-	-	-	-	-
JH6	Q	G	T	T	V	T	V	S	S															
28F3 VH	-	-	-	-	-	-	-	-	-															

Figura 12

Anti-GITR 28F3 VK (hKappa)

L18	A	I	Q	L	T	Q	S	P	S	S	L	S	A	S	V	G	D	R	V	T	I	T	C
28F3 VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
		CDR1																					
L18	R	A	S	Q	G	I	S	S	A	L	A	W	Y	Q	Q	K	P	G	K	A	P	K	L
28F3 VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
		CDR2																					
L18	L	I	Y	D	A	S	S	L	E	S	G	V	P	S	R	F	S	G	S	G	S	G	T
28F3 VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
		CDR3																					
L18	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T	Y	Y	C	Q	Q	F	N
28F3 VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
L18	N	Y																					
JK2			Y	T	F	G	Q	G	T	K	L	E	I	K									
28F3 VK	S	-	P	-	-	-	-	-	-	-	-	-	-	-									

Figura 13

Anti-GITR 18E10 VH (hIgG2)

3-33	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S	L	R	L	S	C	A																					
18E10	VH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-																					
3-33	A	S	G	F	T	F	S	CDR1															S	Y	G	M	H	W	V	R	Q	A	P	G	K	G	L	E						
18E10	VH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-													
3-33	W	V	A	CDR2																		V	I	W	Y	D	G	S	N	K	Y	Y	A	D	S	V	K	G	R	F	T			
18E10	VH	-	-	-	-	-	-	-	-	A	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-											
3-33	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S	L	R	A	E	D	T	A																					
18E10	VH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-											
3-33	V	Y	Y	C	A	R	CDR3																I	A	V	A	Y	Y	G	M	D	V	W	G	Q									
6-19																							I	A	V	A																		
JH6																																				Y	Y	G	M	D	V	W	G	Q
18E10	VH	-	-	-	-	-	-	G	G	R	-	-	-	-	F	-	-	S	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-											
JH6	G	T	T	V	T	V	S	S																																				
18E10	VH	-	-	-	-	-	-	-																																				

Figura 14

Anti-GITR 18E10 VK (hKappa)

L15		D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D	R	V	T	I	T	C										
18E10	VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-										
		CDR1																																
L15		R	A	S	Q	G	I	S	S	W	L	A	W	Y	Q	Q	K	P	E	K	A	P	K	S										
18E10	VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-										
		CDR2																																
L15		L	I	Y	A	A	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G	T										
18E10	VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-										
		CDR3																																
L15		D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T	Y	Y	C	Q	Q	Y	N										
18E10	VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-										
L15		S										Y																						
JK2												Y												T	F	G	Q	G	T	K	L	E	I	K
18E10	VK	-	-	P	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-										

Figura 15

Anti-GITR 19D3 VH (hIgG2)

3-33	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S	L	R	L	S	C	A	
19D3	VH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
		CDR1																						
3-33	A	S	G	F	T	F	S	S	Y	G	M	H	W	V	R	Q	A	P	G	K	G	L	E	
19D3	VH	-	-	-	-	-	-	-	-	-	F	-	-	-	-	-	-	-	-	-	-	-	-	
		CDR2																						
3-33	W	V	A	V	I	W	Y	D	G	S	N	K	Y	Y	A	D	S	V	K	G	R	F	T	
19D3	VH	-	-	-	-	-	-	A	-	-	-	-	F	-	-	-	-	-	-	-	-	-	-	
3-33	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S	L	R	A	E	D	T	A	
19D3	VH	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
		CDR3																						
3-33	V	Y	Y	C	A	R																		
3-16																								
JH6													D	Y										
JH6													Y	Y	Y	Y	Y	G	M	D	V	W	G	Q
19D3	VH	-	-	-	-	-	G	G	Q	L	-	-	-	-	-	-	V	-	-	-	-	-	-	
JH6	G	T	T	V	T	V	S	S																
19D3	VH	-	-	-	-	-	-	-																

Figura 16

Anti-GITR 19D3 VK (hKappa)

L15	D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D	R	V	T	I	T	C
19D3 VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
		CDR1																					
L15	R	A	S	Q	G	I	S	S	W	L	A	W	Y	Q	Q	K	P	E	K	A	P	K	S
19D3 VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
		CDR2																					
L15	L	I	Y	A	A	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G	T
19D3 VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
		CDR3																					
L15	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T	Y	Y	C	Q	Q	Y	N
19D3 VK	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
L15	S	Y																					
JK2			Y	T	F	G	Q	G	T	K	L	E	I	K									
19D3 VK	-	-	P	-	-	-	-	-	-	-	-	-	-	-									

Figura 17

Anti-GITR 3C3 VH (hIgG1)

4-34	Q	V	Q	L	Q	Q	W	G	A	G	L	L	K	P	S	E	T	L	S	L	T	C	A
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4-34	V	Y	G	G	S	F	S	CDR1					W	I	R	Q	P	P	G	K	G	L	E
ENTRADA	-	-	-	-	-	-	-	G	Y	Y	W	S	T	-	-	-	-	-	-	-	-	-	-
4-34	W	I	G	CDR2										S	L	K	S	R	V	T	I		
ENTRADA	-	-	-	E	I	N	H	S	G	S	T	N	Y	N	P	S	L	K	S	-	-	-	-
4-34	S	V	D	T	S	K	N	Q	F	S	L	K	L	S	S	V	T	A	A	D	T	A	V
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4-34	Y	Y	C	A	R	CDR3																	
JH3																							
ENTRADA	-	-	-	-	-	L	G	A	F	D	A	F	D	I	W	G	Q	G	T	M	V	T	V
JH3	S	S																					
ENTRADA	-	-																					

Figura 18

Anti-GITR 3C3 VK1 (hKappa)

L15	D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D	R	V	T	I	T	C											
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-											
L15	CDR1											R	A	S	Q	G	I	S	S	W	L	A	W	Y	Q	Q	K	P	E	K	A	P	K	S
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
L15	L	I	Y	CDR2			A	A	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G	S	G	T						
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
L15	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T	Y	Y	C	CDR3					Q	Q	Y	N						
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
L15						S	Y																											
JK2				Y	T	F	G	Q	G	T	K	L	E	I	K																			
ENTRADA	-	-	P	-	-	-	-	-	-	-	-	-	-	-	-																			

Figura 19A

Anti-GITR 3C3 VK2 (hKappa)

L20	E	I	V	L	T	Q	S	P	A	T	L	S	L	S	P	G	E	R	A	T	L	S	C	
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
L20	CDR1																							
ENTRADA	R	A	S	Q	G	V	S	S	Y	L	A	W	Y	Q	Q	K	P	G	Q	A	P	R	L	
L20	L	I	Y	CDR2							G	I	P	A	R	F	S	G	S	G	P	G	T	
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
L20	D	F	T	L	T	I	S	S	L	E	P	E	D	F	A	V	Y	Y	C	CDR3				
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	Q	Q	R	S	
L20																								
JK2	N	W																						
ENTRADA	-	-	H	T	F	G	Q	G	T	K	L	E	I	K										

Figura 19B

Anti-GITR 2G6 VH (hIgG2)

3-33	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S	L	R	L	S	C	A		
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	G	-	-	-	-	-	-	-		
3-33	A	S	G	F	T	F	S	CDR1					W	V	R	Q	A	P	G	K	G	L	E		
ENTRADA	-	-	-	-	I	L	-	S	Y	G	M	H	-	-	-	-	-	-	-	-	-	-	-		
3-33	W	V	A	CDR2													R	F	T						
ENTRADA	-	-	T	V	I	W	Y	D	G	S	N	K	Y	Y	A	D	S	V	K	G	-	-			
3-33	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S	L	R	A	E	D	T	A		
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	V	-	-	-	-		
3-33	V	Y	Y	C	A	R	CDR3																	W	G
JH6																	Y	Y	G	M	D	V			
ENTRADA	-	-	-	-	-	-	G	G	R	L	A	T	G	H	F	-	-	V	-	-	-	-	-		
JH6	Q	G	T	T	V	T	V	S	S																
ENTRADA	-	-	-	-	-	-	-	-	-																

Figura 20

Anti-GITR 2G6 VK (hKappa)

L15	D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D	R	V	T	I	T	C
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
	<u>CDR1</u>																						
L15	R	A	S	Q	G	I	S	S	W	L	A	W	Y	Q	Q	K	P	E	K	A	P	K	S
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
	<u>CDR2</u>																						
L15	L	I	Y	A	A	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G	T
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
	<u>CDR3</u>																						
L15	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T	Y	Y	C	Q	Q	Y	N
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
	<u>CDR4</u>																						
L15	S	Y																					
JK2												Y	T	F	G	Q	G	T	K	L	E	I	K
ENTRADA	-	-	P	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

Figura 21

Anti-GITR 8A6 VH (hIgG2)

3-33	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S	L	R	L	S	C	A
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	T
3-33	A	S	G	F	T	F	S	S	Y	G	M	H	W	V	R	Q	A	P	G	K	G	L	E
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	Q	-	-	-	-	-	-	-	-	-	-	-
3-33	W	V	A	V	I	W	Y	D	G	S	N	K	Y	Y	A	D	S	V	K	G	R	F	T
ENTRADA	-	-	-	-	-	-	-	E	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3-33	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S	L	R	A	E	D	T	A
ENTRADA	-	-	-	E	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3-33	V	Y	Y	C	A	R																	
3-10																							
JH6																							
ENTRADA	-	-	-	-	-	-	G	G	L	-	-	-	-	L	F	-	-	-	-	-	-	-	-
JH6	Q	G	T	T	V	T	V	S	S														
ENTRADA	-	-	-	-	-	-	-	-	-														

Figura 22

Anti-GITR 8A6 VK (hKappa)

L18	A	I	Q	L	T	Q	S	P	S	S	L	S	A	S	V	G	D	R	V	T	I	T	C											
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-											
L18	CDR1											R	A	S	Q	G	I	S	S	A	L	A	W	Y	Q	Q	K	P	G	K	A	P	K	L
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	F	
L18	L	I	Y	CDR2			D	A	S	S	L	E	S	G	V	P	S	R	F	S	G	S	G	S	G	T								
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-								
L18	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T	Y	Y	C	CDR3				Q	Q	F	N							
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-								
L18	N			Y																														
JK2							Y	T	F	G	Q	G	T	K	L	E	I	K																
ENTRADA	S	-	P	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-																

Figura 23

Anti-GITR 9G7 VH (hIgG4)

3-15	E	V	Q	L	V	E	S	G	G	G	L	V	K	P	G	G	S	L	R	L	S	C	A
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3-15	A	S	G	F	T	F	S	CDR1				W	V	R	Q	A	P	G	K	G	L	E	
ENTRADA	-	-	-	-	-	-	-	N	A	T	V	-	-	-	-	-	-	-	-	-	-	-	-
3-15	W	V	G	CDR2																			
ENTRADA	-	-	-	R	I	K	S	K	T	D	G	G	T	T	D	Y	A	A	P	V	K	G	R
3-15	F	T	I	S	R	D	D	S	K	N	T	L	Y	L	Q	M	N	S	L	K	T	E	D
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	H	-	-	-
3-15	T	A	V	Y	Y	C	T	T	CDR3														
3-10																	Y	Y	Y	G			
JH6																Y	Y	Y	Y	G	M	D	V
ENTRADA	-	-	-	-	-	-	-	-	G	Q	L	I	P	-	S	-	-	-	-	-	-	-	-
JH6	G	Q	G	T	T	V	T	V	S	S													
ENTRADA	-	-	-	-	S	-	-	-	-	-													

Figura 24

Anti-GITR 9G7 VK1 (hKappa)

A27	E	I	V	L	T	Q	S	P	G	T	L	S	L	S	P	G	E	R	A	T	L	S	C	
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
A27	CDR1																							
A27	R	A	S	Q	S	V	S	S	S	Y	L	A	W	Y	Q	Q	K	P	G	Q	A	P	R	
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
A27	L	L	I	Y	CDR2								G	I	P	D	R	F	S	G	S	G	S	G
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
A27	T	D	F	T	L	T	I	S	R	L	E	P	E	D	F	A	V	Y	Y	C	CDR3			
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	Q	Q	Y	
A27																								
JK1	G	S	S																					
ENTRADA	-	-	-	P	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	

Figura 25A

Anti-GITR 9G7 VK2 (hKappa)

A27	E	I	V	L	T	Q	S	P	G	T	L	S	L	S	P	G	E	R	A	T	L	S	C
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
A27	R	A	S	Q	S	V	S	S	S	Y	L	A	W	Y	Q	Q	K	P	G	Q	A	P	R
ENTRADA	-	-	-	-	-	-	T	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
A27	L	L	I	Y	G	A	S	S	R	A	T	G	I	P	D	R	F	S	G	S	G	S	G
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	E	-	-	-	-	-	-	-	-
A27	T	D	F	T	L	T	I	S	R	L	E	P	E	D	F	A	V	Y	Y	C	Q	Q	Y
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
A27	G	S	S																				
JK5					I	T	F	G	Q	G	T	R	L	E	I	K							
ENTRADA	-	-	-	P	-	-	-	-	-	-	-	-	-	-	-	-							

Figura 25B

Anti-GITR 14E3 VH (hIgG1)

4-34	Q	V	Q	L	Q	Q	W	G	A	G	L	L	K	P	S	E	T	L	S	L	T	C	A
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4-34	V	Y	G	G	S	F	S	CDR1				W	I	R	Q	P	P	G	K	G	L	E	
ENTRADA	-	-	-	-	-	-	-	G	Y	Y	W	S	-	-	-	-	-	-	-	-	-	-	-
4-34	W	I	G	CDR2				S	G	S	T	N	Y	N	P	S	L	K	S	R	V	T	I
ENTRADA	-	-	-	E	I	N	H	-	-	N	-	Y	-	-	-	-	-	-	-	-	-	-	-
4-34	S	V	D	T	S	K	N	Q	F	S	L	K	L	S	S	V	T	A	A	D	T	A	V
ENTRADA	-	-	-	-	-	-	-	-	L	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4-34	Y	Y	C	A	R	CDR3				D	A	F	D	I	W	G	Q	G	T	M	V	T	V
JH3																							
ENTRADA	-	-	-	-	-	F	G	S	N	-	-	-	-	-	-	-	-	-	-	-	-	-	-
JH3	S	S																					
ENTRADA	-	-																					

Figura 26

Anti-GITR 14E3 VK (hKappa)

L15	D	I	Q	M	T	Q	S	P	S	S	L	S	A	S	V	G	D	R	V	T	I	T	C																
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-																
L15	CDR1										R	A	S	Q	G	I	S	S	W	L	A	W	Y	Q	Q	K	P	E	K	A	P	K	S						
ENTRADA											-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-					
L15	L	I	Y	CDR2							A	A	S	S	L	Q	S	G	V	P	S	R	F	S	G	S	G	S	G	T									
ENTRADA	-	-	-								-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-						
L15	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T	Y	Y	C	CDR3					Q	Q	Y	N											
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-						-	-	-	-	-	-	-	-	-						
L15					S	Y																																	
JK1							T	F	G	Q	G	T	K	V	E	I	K																						
ENTRADA	-	-	P	P	-	-	-	-	-	-	-	-	-	-	-	-																							

Figura 27

Anti-GITR 19H8 VH (hIgG2)

3-33	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S	L	R	L	S	C	A
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3-33	A	S	G	F	T	F	S	S	Y	G	M	H	W	V	R	Q	A	P	G	K	G	L	E
ENTRADA	-	-	-	-	-	-	-	N	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3-33	W	V	A	V	I	W	Y	D	G	S	N	K	Y	Y	A	D	S	V	K	G	R	F	T
ENTRADA	-	M	-	-	-	-	-	G	-	-	-	-	F	-	-	-	-	-	-	-	-	-	-
3-33	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S	L	R	A	E	D	T	A
ENTRADA	-	-	-	-	-	-	-	-	S	-	S	-	-	-	-	-	-	-	-	-	-	-	-
3-33	V	Y	Y	C	A	R																	
3-10														Y	Y	Y							
JH6														Y	Y	Y	G	M	D	V	W	G	
ENTRADA	-	-	-	-	-	-	G	G	A	M	V	R	G	V	-	-	-	-	-	-	-	-	-
JH6	Q	G	T	T	V	T	V	S	S														
ENTRADA	-	-	-	-	-	-	-	-	-														

Figura 28

Anti-GITR 19H8 VK1 (hKappa)

L18	A	I	Q	L	T	Q	S	P	S	S	L	S	A	S	V	G	D	R	V	T	I	T	C	
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
	CDR1																							
L18	R	A	S	Q	G	I	S	S	A	L	A	W	Y	Q	Q	K	P	G	K	A	P	K	L	
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	F	
	CDR2																							
L18	L	I	Y	D	A	S	S	L	E	S	G	V	P	S	R	F	S	G	S	G	S	G	T	
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
	CDR3																							
L18	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T	Y	Y	C	Q	Q	F	N	
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
L18	N				Y																			
JK1					T				F	G	Q	G	T	K	V	E	I	K						
ENTRADA	S	-	P	Q	-	-	-	-	-	-	-	-	-	-	-	-								

Figura 29A

Anti-GITR 19H8 VK2 (hKappa)

L6	E	I	V	L	T	Q	S	P	A	T	L	S	L	S	P	G	E	R	A	T	L	S	C
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
	CDR1																						
L6	R	A	S	Q	S	V	S	S	Y	L	A	W	Y	Q	Q	K	P	G	Q	A	P	R	L
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
	CDR2																						
L6	L	I	Y	D	A	S	N	R	A	T	G	I	P	A	R	F	S	G	S	G	S	G	T
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
	CDR3																						
L6	D	F	T	L	T	I	S	S	L	E	P	E	D	F	A	V	Y	Y	C	Q	Q	R	S
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
L6	N	W																					
JK4				L	T	F	G	G	G	T	K	V	E	I	K								
ENTRADA	-	-	P	-	-	-	-	-	-	-	-	-	-	-	-								

Figura 29B

Anti-GITR 6G10 VH (hIgG2)

3-33	Q	V	Q	L	V	E	S	G	G	G	V	V	Q	P	G	R	S	L	R	L	S	C	A	
ENTRADA	-	-	-	-	-	-	-	-	-	D	-	-	-	-	-	-	-	-	-	-	-	-	-	
3-33	A	S	G	F	T	F	S	<u>CDR1</u> S	Y	G	M	H	W	V	R	Q	A	P	G	K	G	L	E	
ENTRADA	-	-	-	-	-	-	-	T	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
3-33	W	V	A	<u>CDR2</u> V	I	W	Y	D	G	S	N	K	Y	Y	A	D	S	V	K	G	R	F	T	
ENTRADA	-	-	-	-	T	-	-	A	-	-	-	-	F	-	-	-	-	-	-	-	-	-	-	
3-33	I	S	R	D	N	S	K	N	T	L	Y	L	Q	M	N	S	L	R	A	E	D	T	A	
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
3-33	V	Y	Y	C	A	R	<u>CDR3</u>																	
3-10								M V R G																
JH6								Y Y Y G M D V W G																
ENTRADA	-	-	-	-	-	-	G	G	S	-	-	-	-	L	-	-	-	-	-	-	-	-	-	
JH6	Q	G	T	T	V	T	V	S	S															
ENTRADA	-	-	-	-	-	-	-	-	-															

Figura 30

Anti-GITR 6G10 VK1 (hKappa)

L18	A	I	Q	L	T	Q	S	P	S	S	L	S	A	S	V	G	D	R	V	T	I	T	C											
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-											
L18	CDR1											R	A	S	Q	G	I	S	S	A	L	A	W	Y	Q	Q	K	P	G	K	A	P	K	L
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
L18	L	I	Y	CDR2							D	A	S	S	L	E	S	G	V	P	S	R	F	S	G	S	G	S	G	T				
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
L18	D	F	T	L	T	I	S	S	L	Q	P	E	D	F	A	T	Y	Y	C	CDR3					Q	Q	F	N						
ENTRADA	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
L18				N		Y																												
JK2						Y	T	F	G	Q	G	T	K	L	E	I	K																	
ENTRADA	S	-	P	-	-	-	-	-	-	-	-	-	-	-	-	-	-																	

Figura 31

Fijación de mAbs GITR a linfocitos T humanos activados

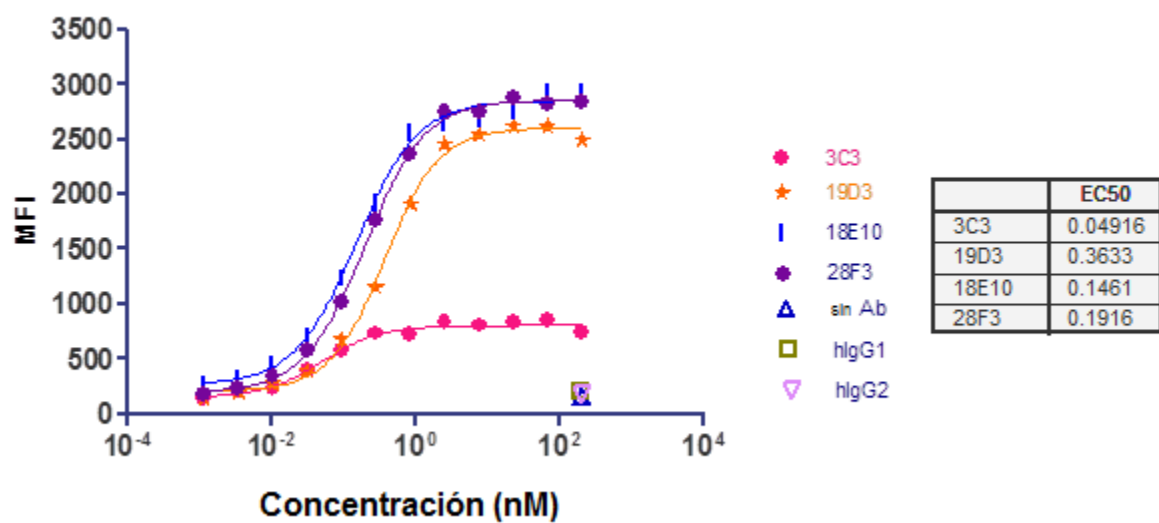


Figura 32

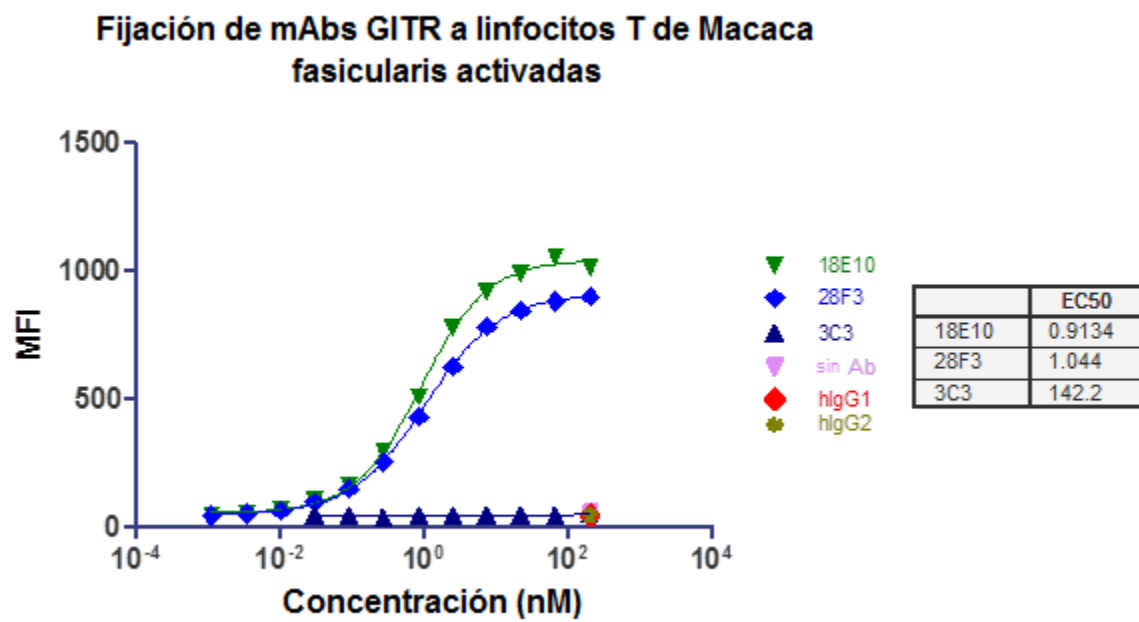


Figura 33

HuMabs GITR bloquean la fijación de GITRL a células GITR-3A9

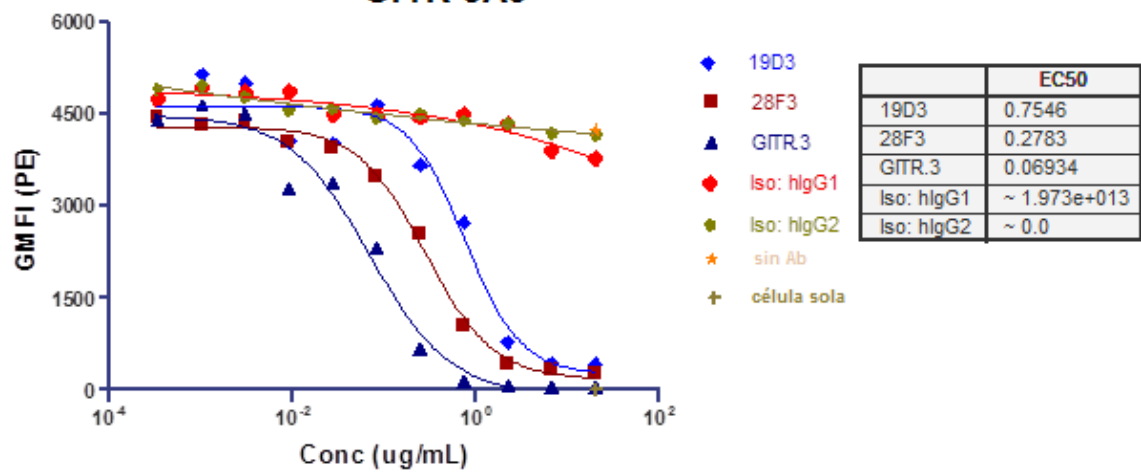


Figura 34A

HuMabs GITR inhiben la fijación de GITRL a células GITR-3A9

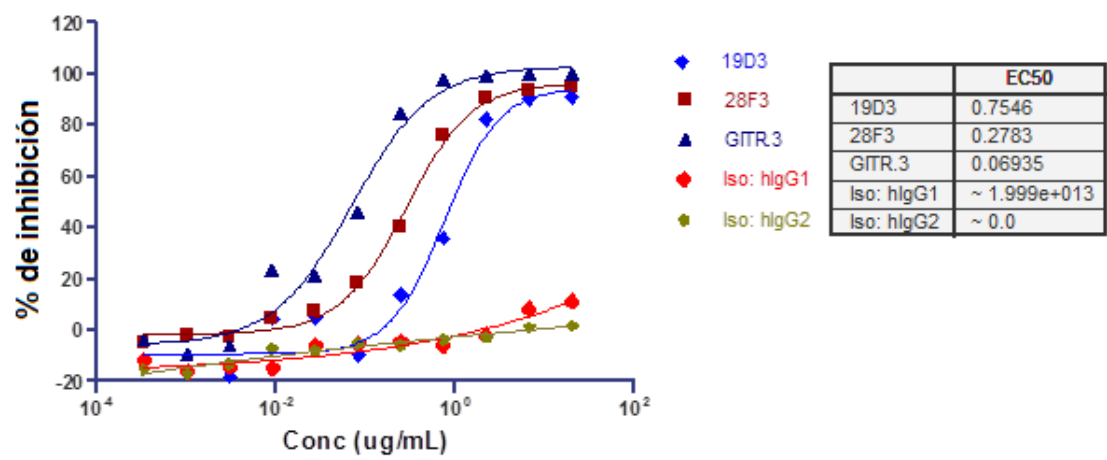


Figura 34B

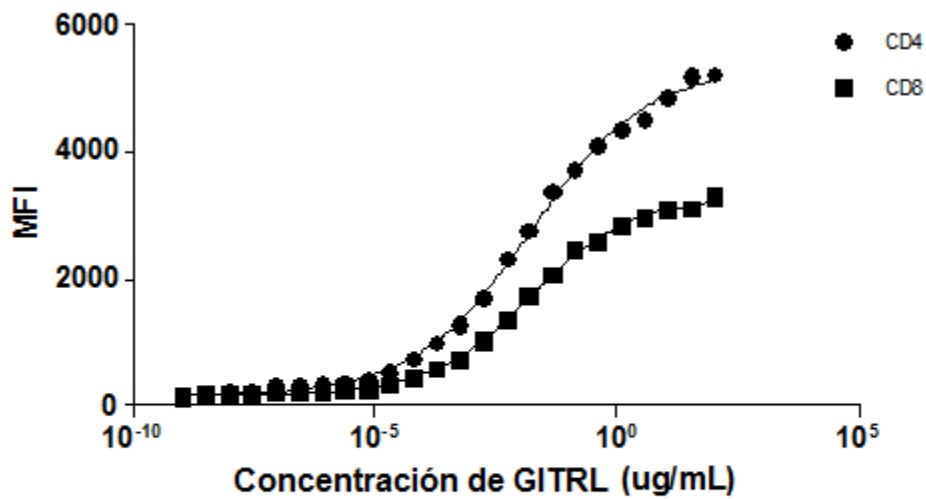
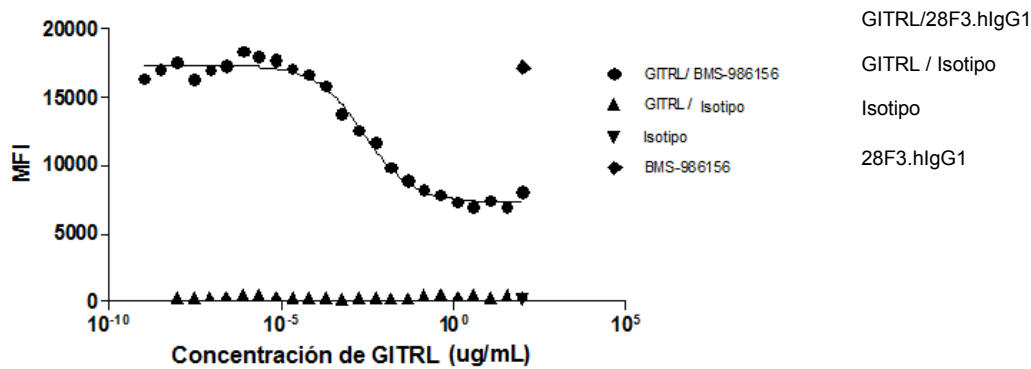


Figura 34C



	GITRL/28F3.hlgG1
IC50	0.002440

Figura 34D

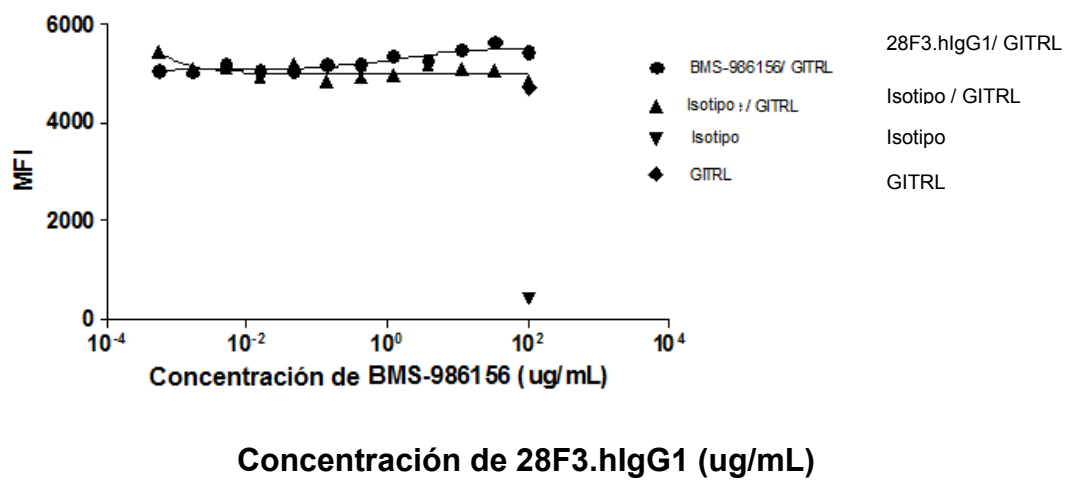
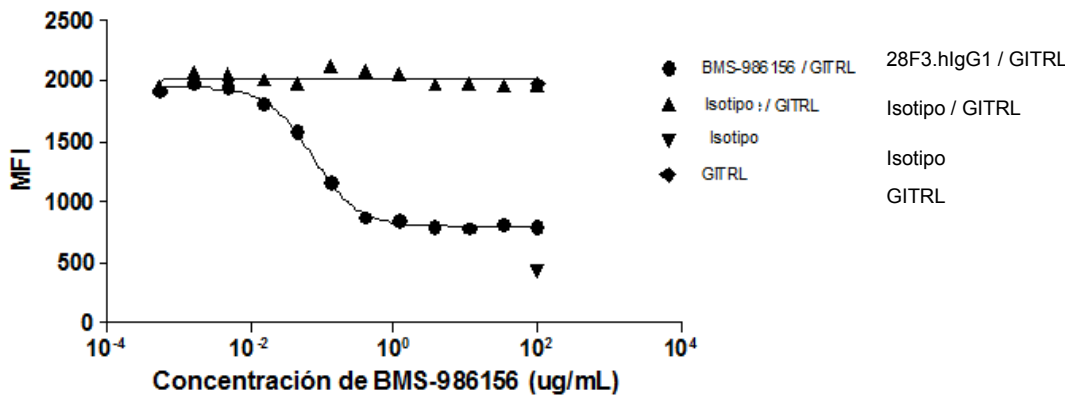


Figura 34E

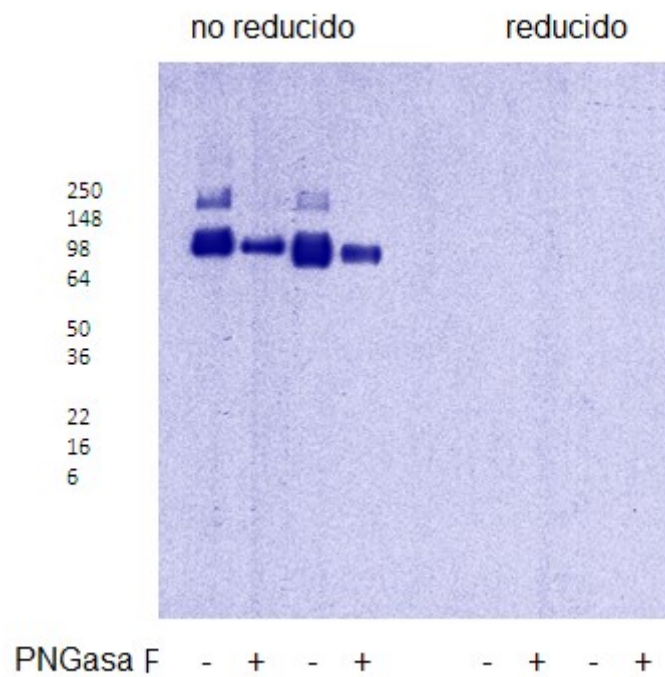


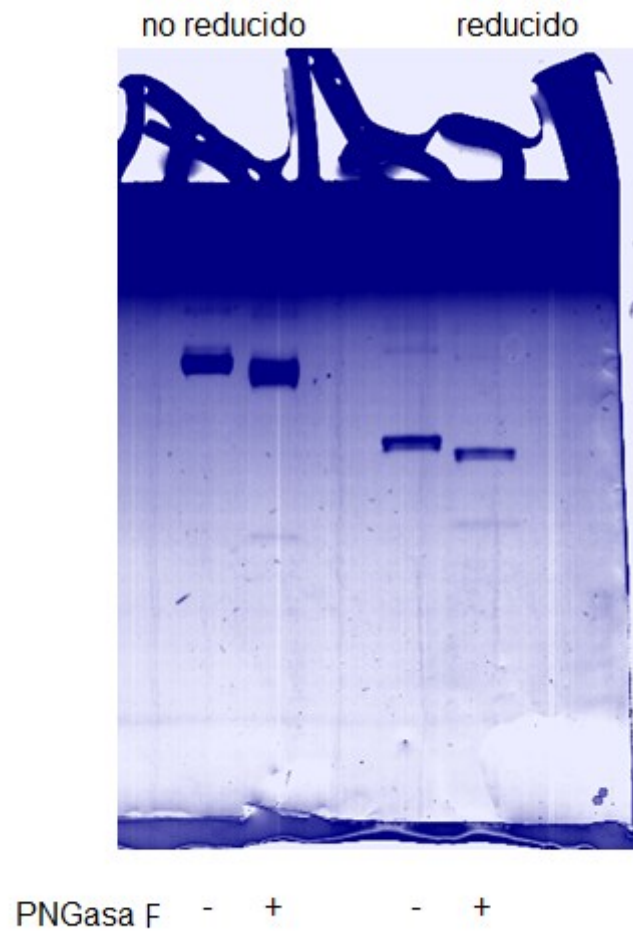
	BMS-986156 / GITRL
IC50	0.07575

Concentración de 28F3.hlgG1 (ug/mL)

	28F3.hlgG1 / GITRL
IC50	0.07575

Figura 34F

**Figura 35A**

**Figura 35B**

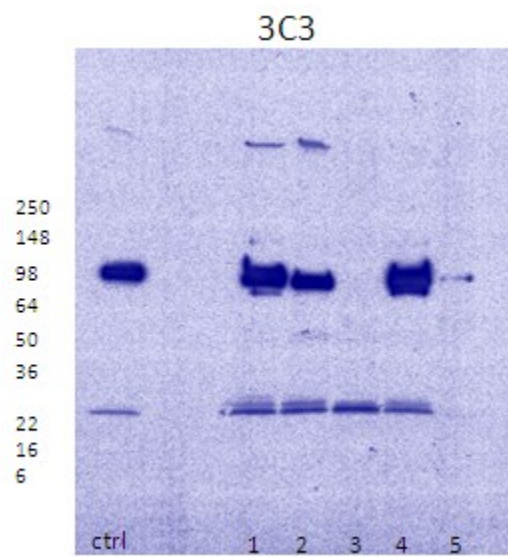


Figura 36A

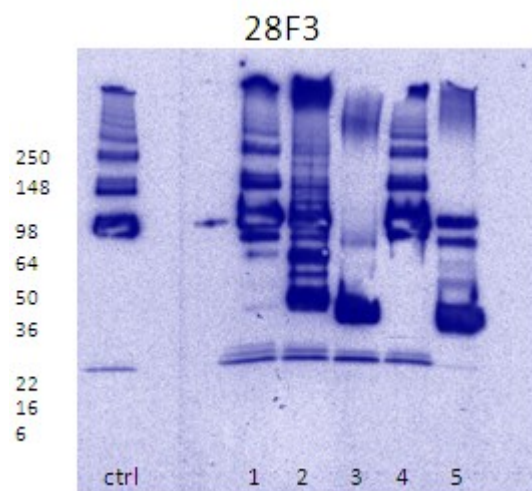
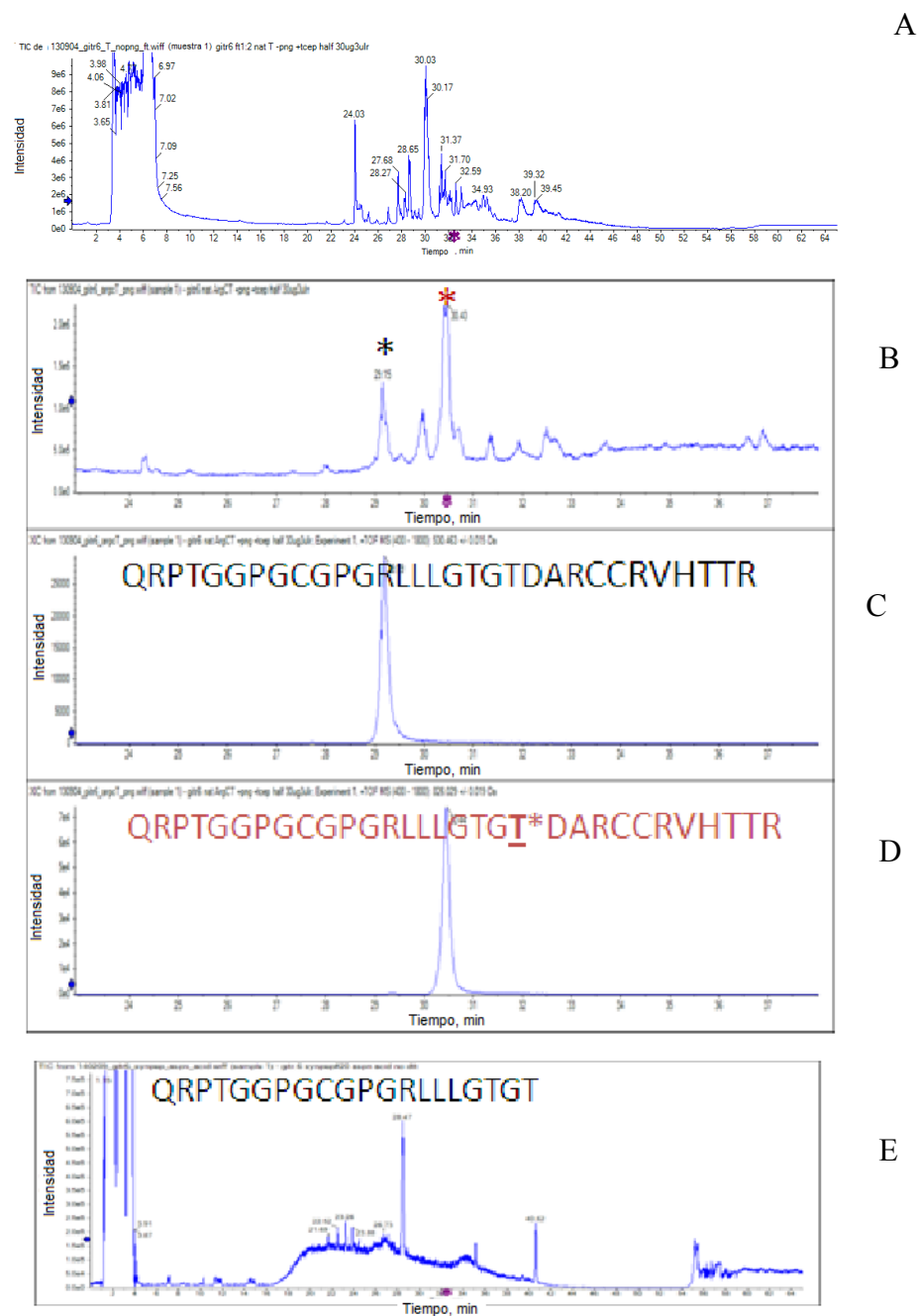


Figura 36B

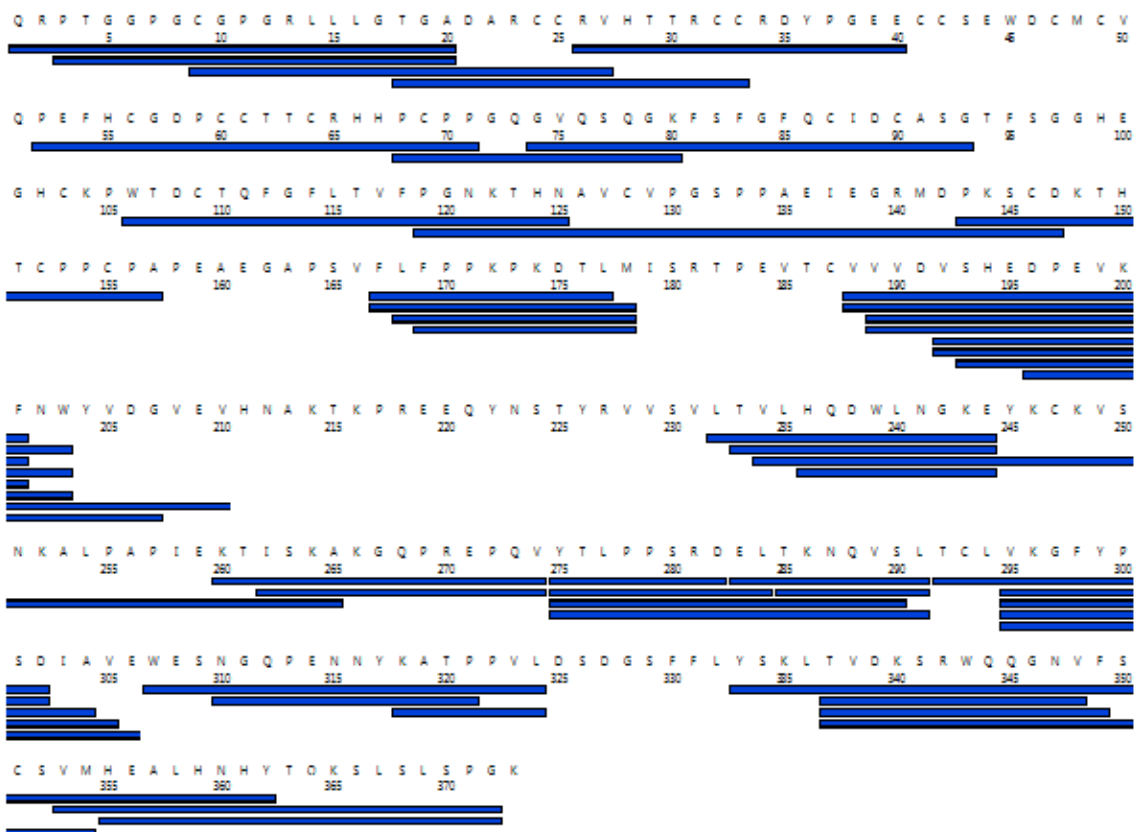


Figura 37



Figuras 38A-38E

GITR



Total: 81.0% de cobertura, 2.38 redundancia

cobertura de
secuencia de 86 %

Figura 39A

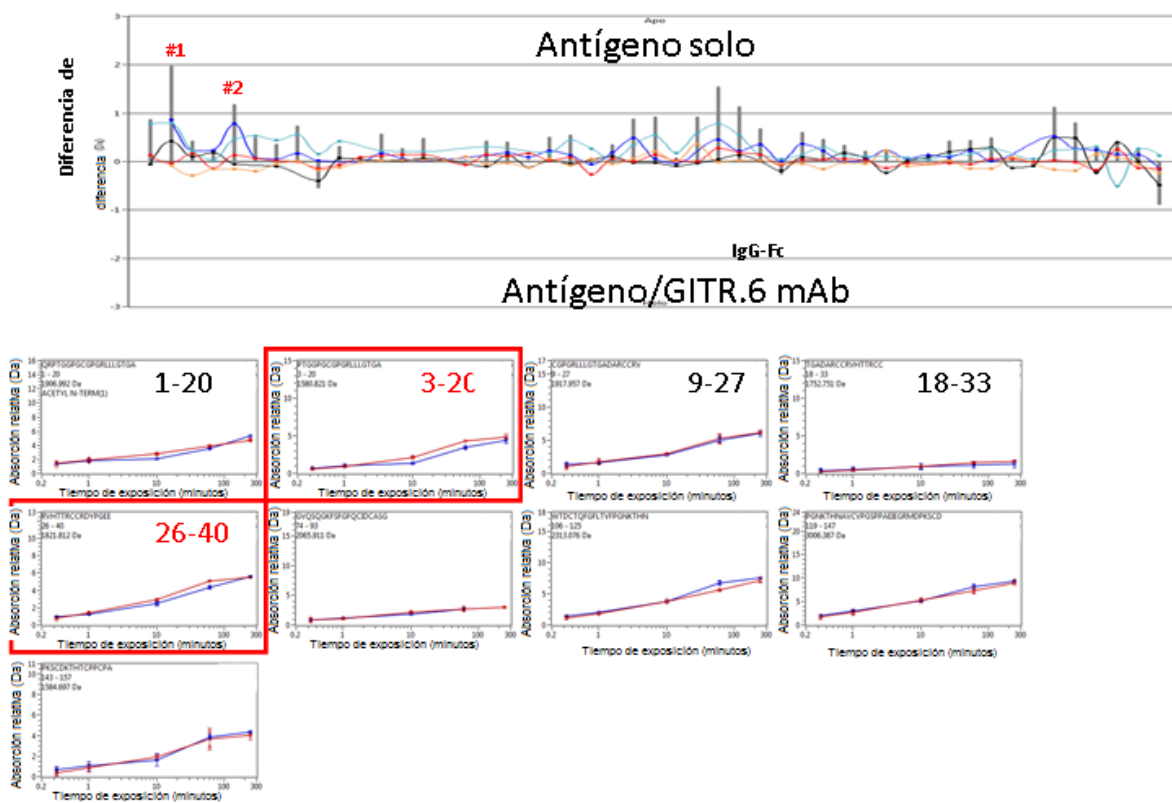


Figura 39B

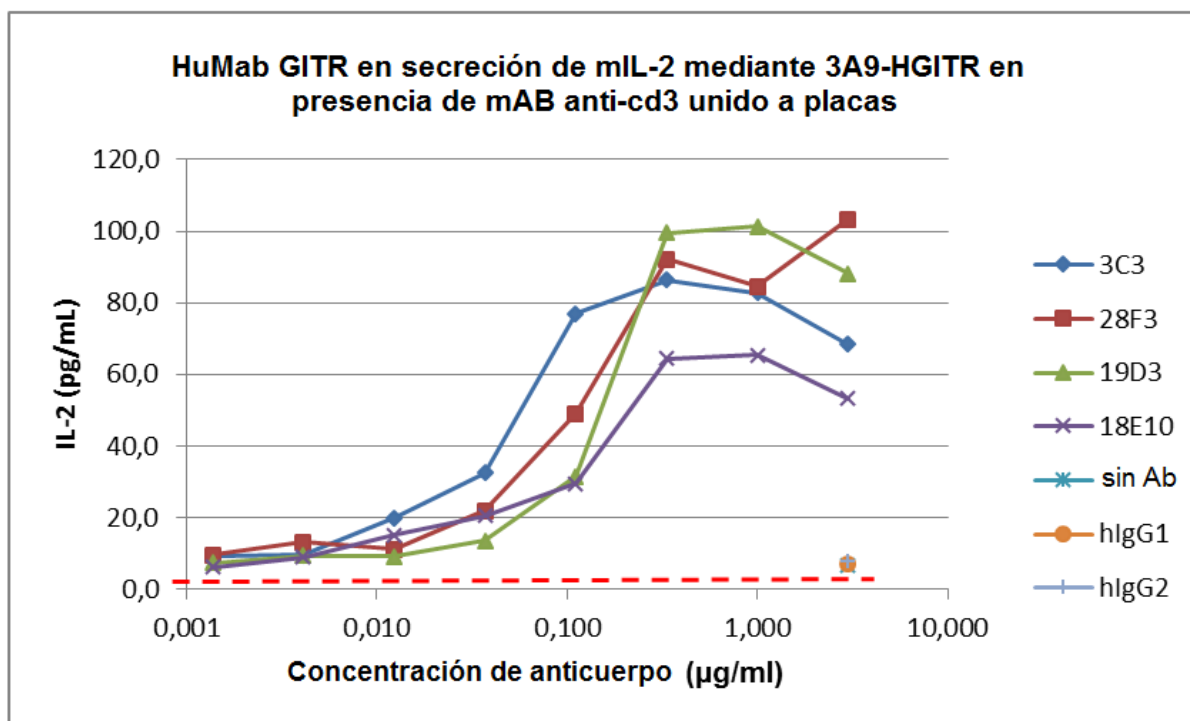
¹QRPTGGPGCG ¹¹PGRLLLGTGA ²¹DARCCRVHTT ³¹RCCRDYPGEE
 
Clasificación #1 Clasificación #2

⁴¹CCSEWDCMCV ⁵¹QPEFHCGDPC ⁶¹CTTCRHHPCP ⁷¹PGQGVQSQK

⁸¹FSFGFQCIDC ⁹¹ASGTFSGGHE ¹⁰¹GHCKPWT DCT ¹¹¹QFGFLT V FPG

¹²¹NKTHNAV CVP ¹³¹GSPPAEIEGR ¹⁴¹MD

Figura 39C

**Figura 40**

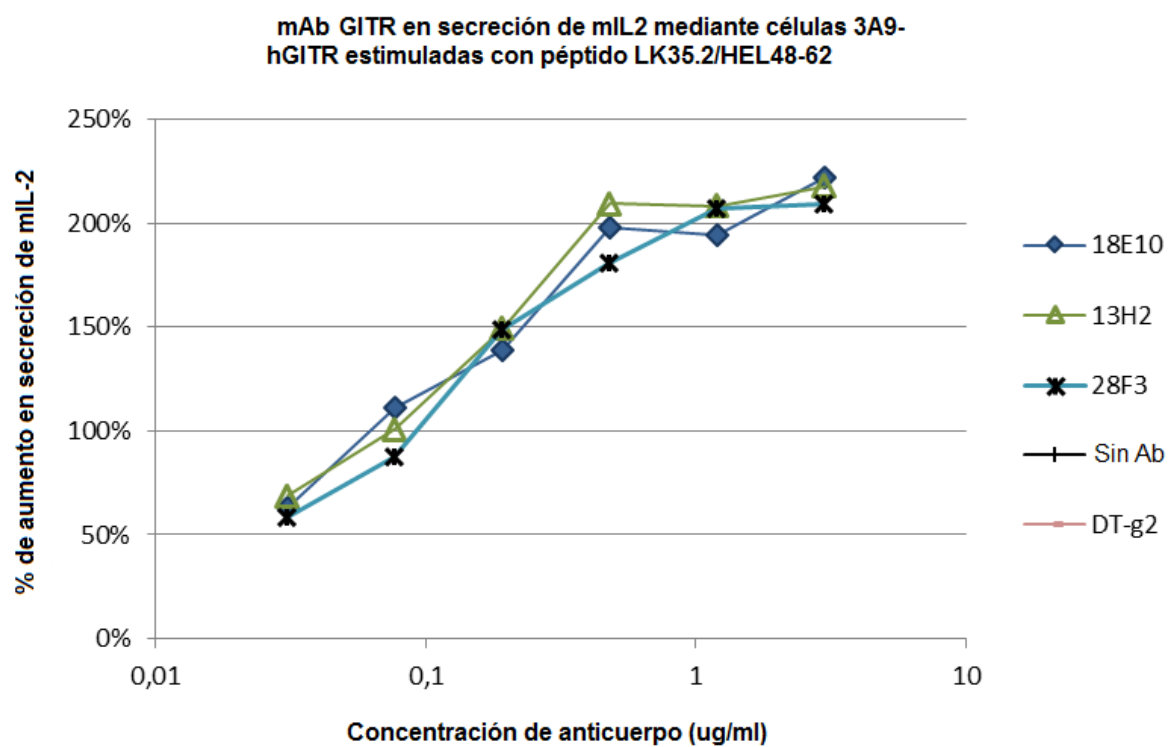


Figura 41A

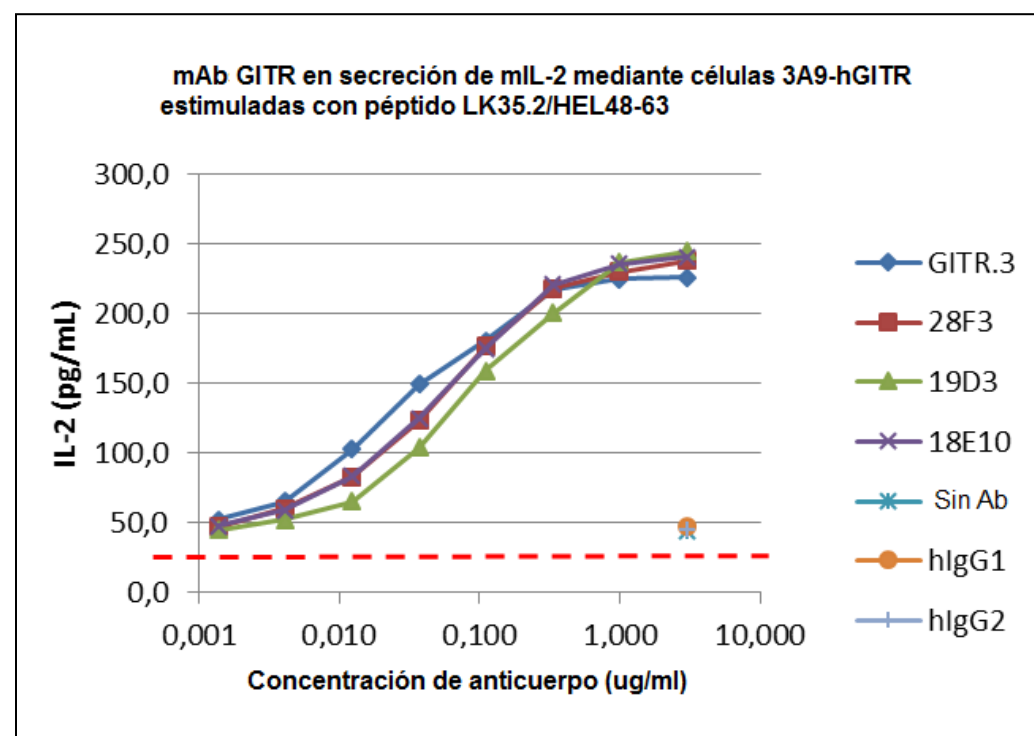


Figura 41B

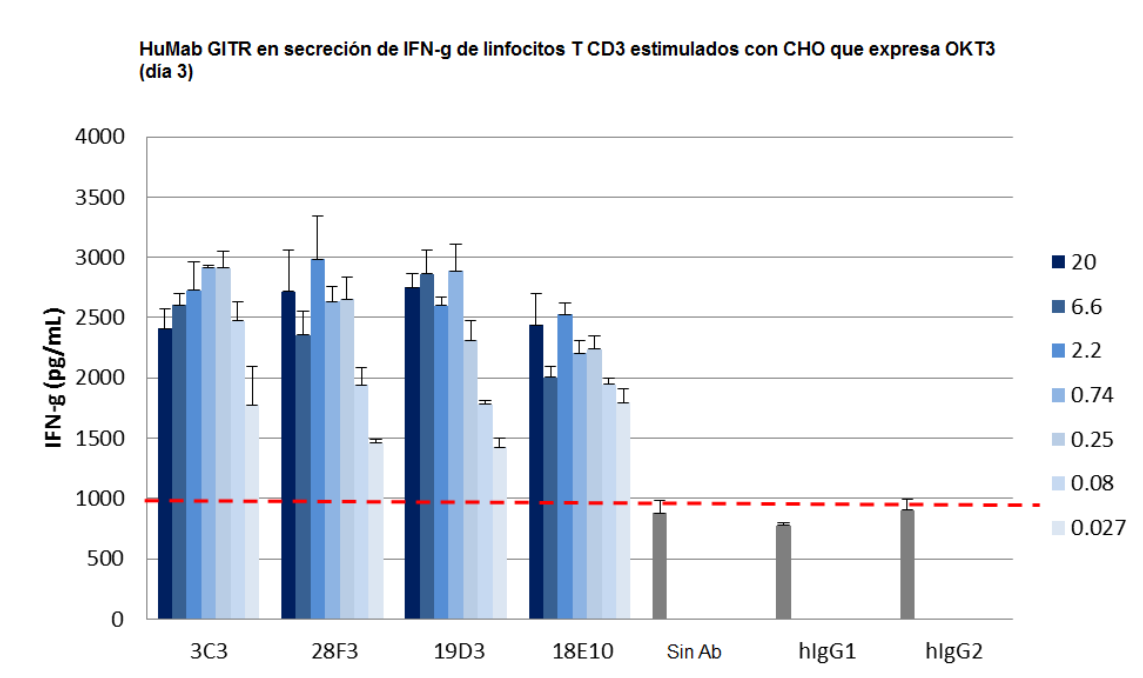


Figura 42A

mAb GITR en secreción de IL-2 mediante linfocitos T CD4 estimulados con células CHO que expresan OKT3 (día 2) - Donante #1

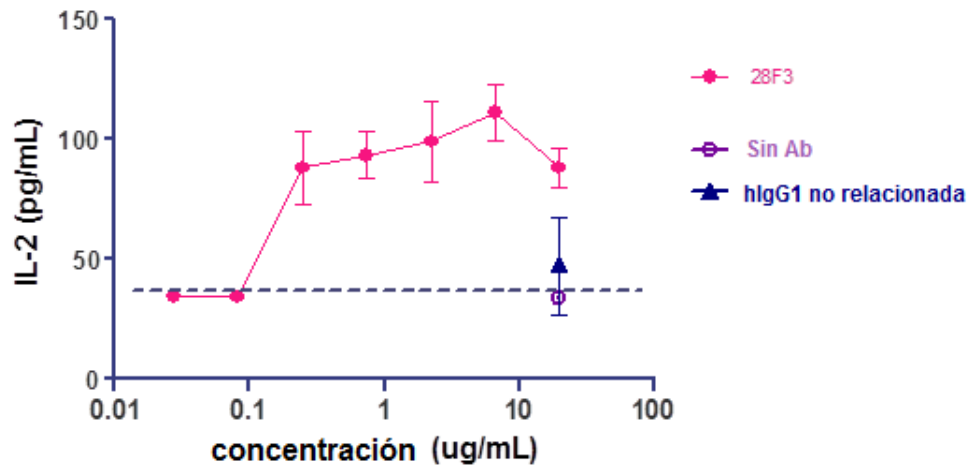


Figura 42B

mAb GITR en secreción de IFN-g mediante linfocitos T CD4 estimulados con células CHO que expresan OKT3 (día 3)- Donante #1

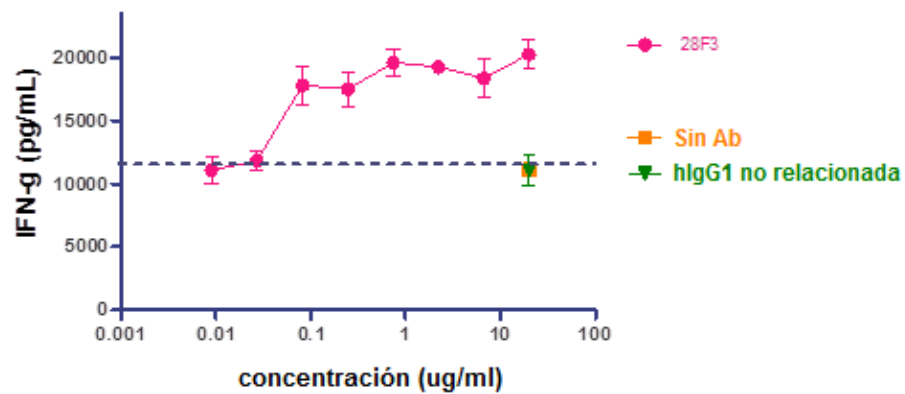


Figura 42C

mAb GITR en secreción de IL-2 mediante linfocitos T CD4 estimulados con células CHO que expresan OKT3 (día 2) - Donante #2

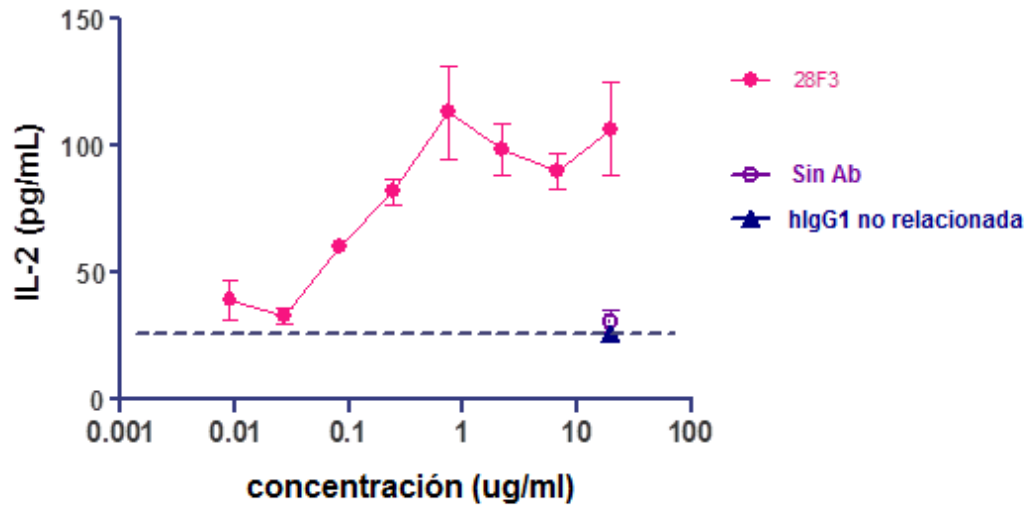


Figura 42D

mAb GITR en secreción de IFN-g mediante linfocitos T CD4 estimulados con células CHO que expresan OKT3 (día 3) - Donante #2

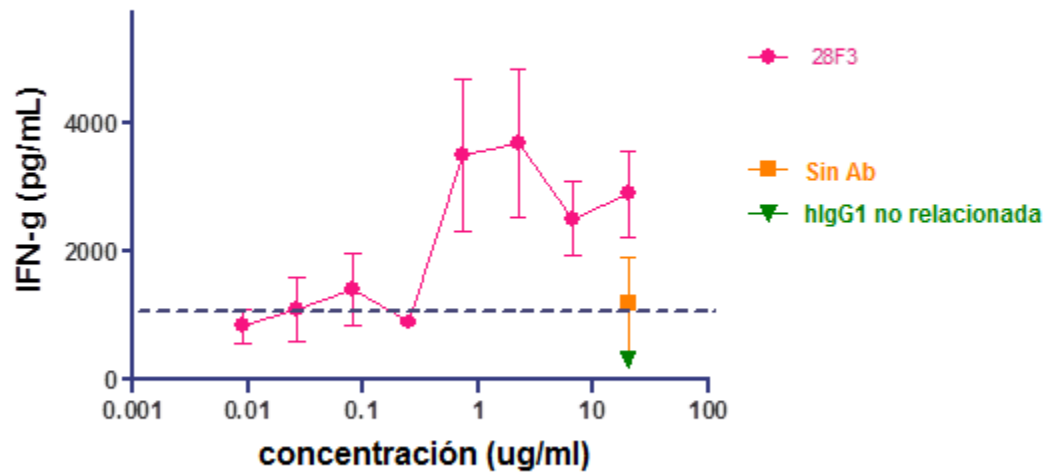


Figura 42E

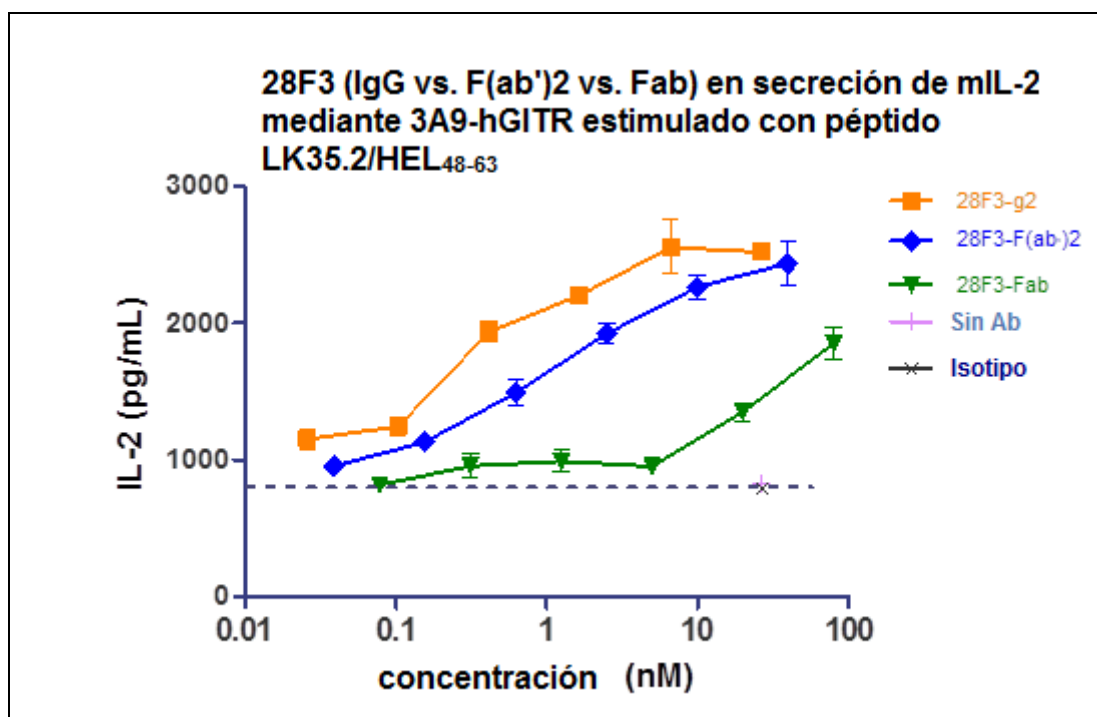


Figura 43

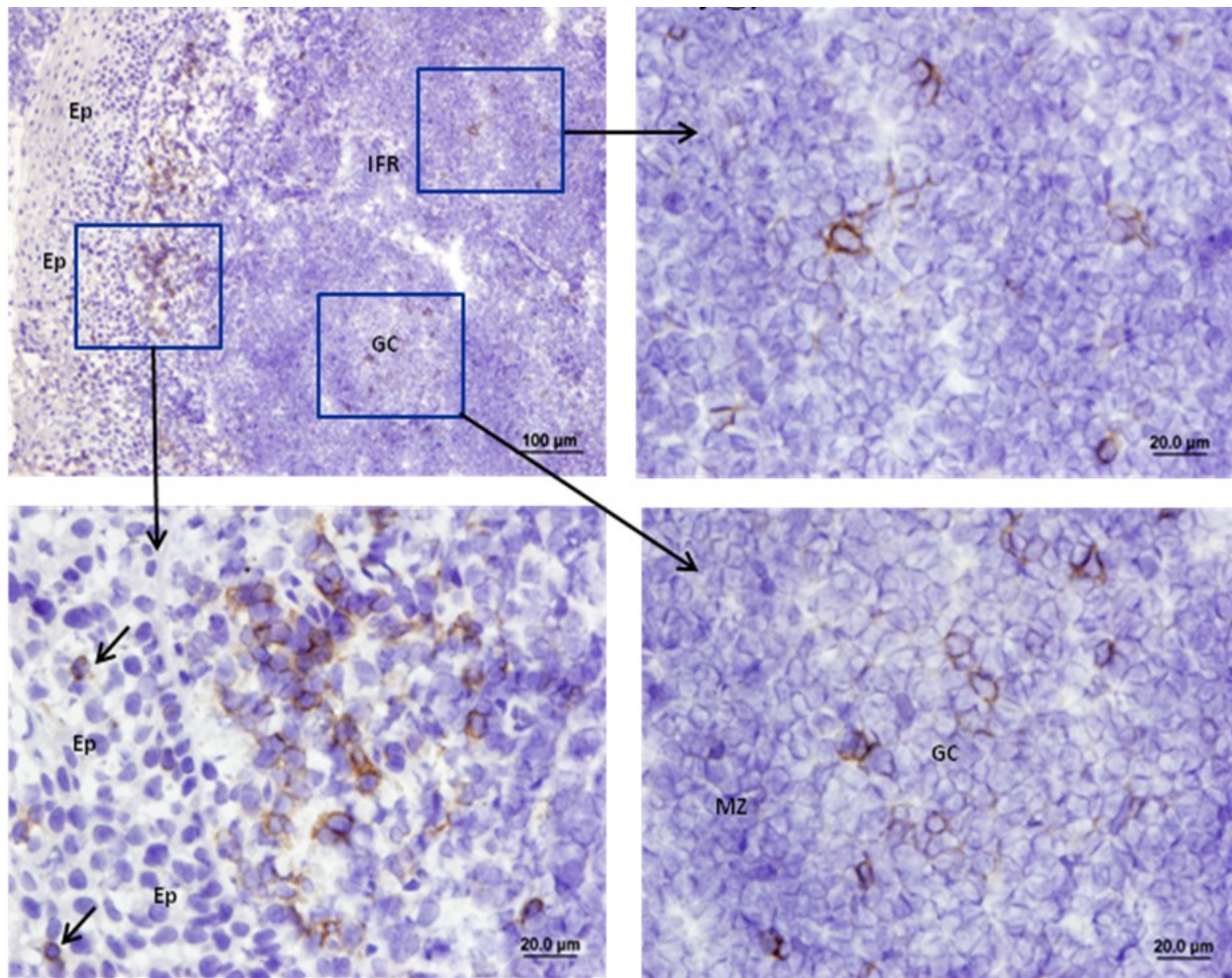


Figura 44

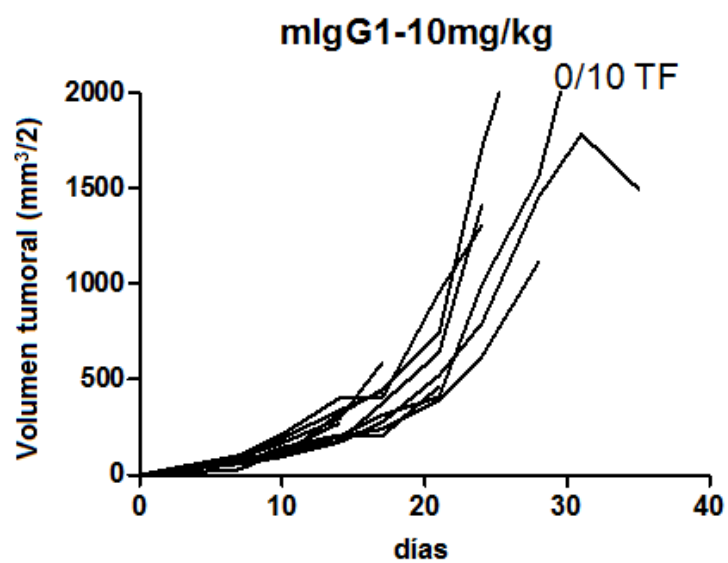


Figura 45A

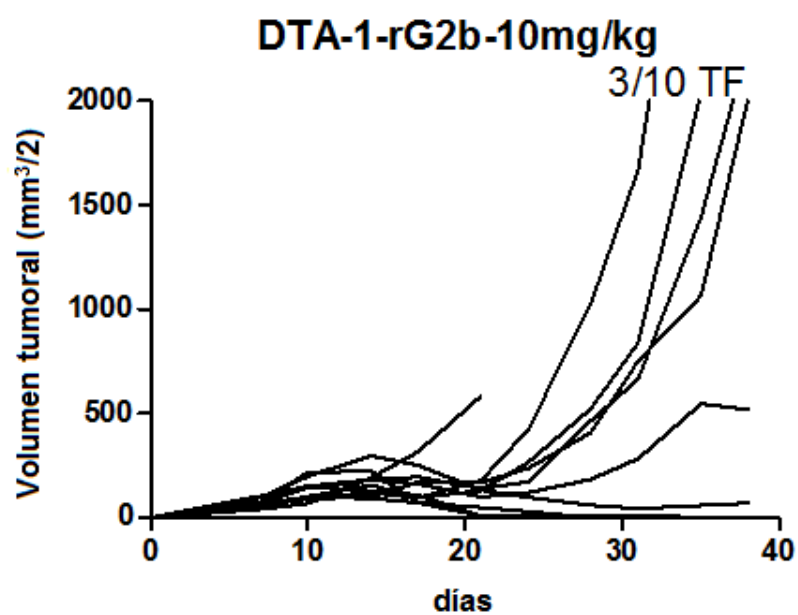


Figura 45B

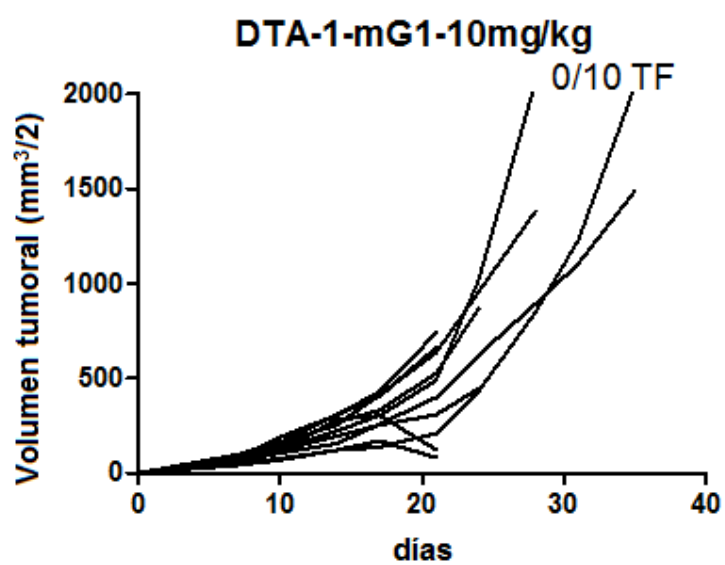


Figura 45C

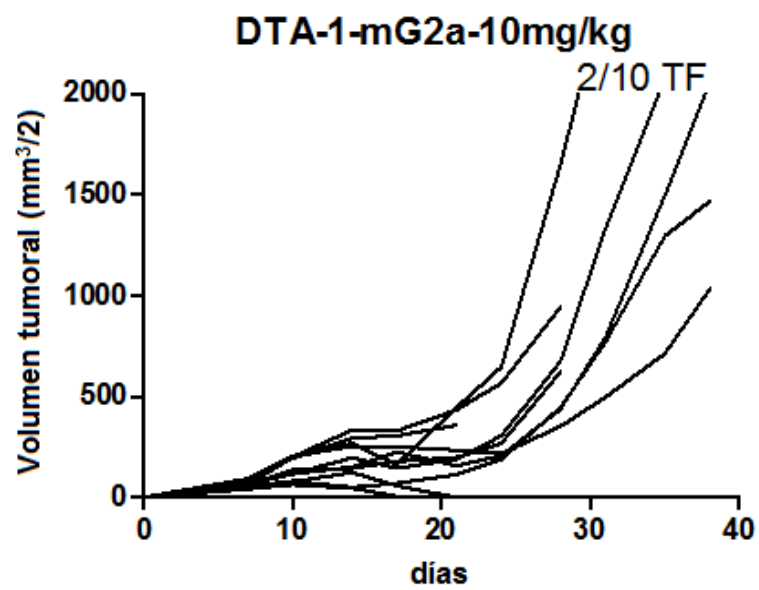


Figura 45D

A.

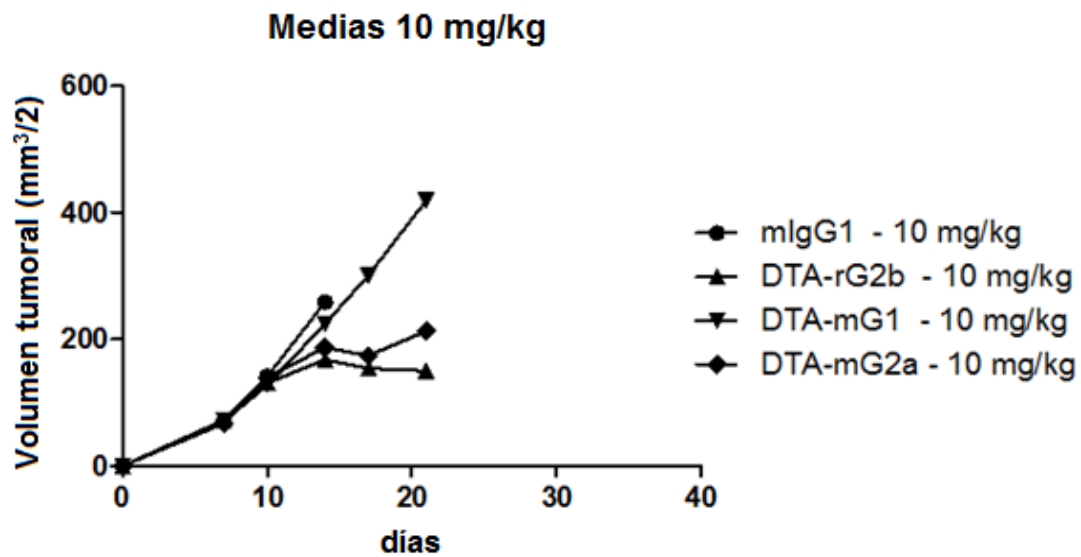


Figura 46A

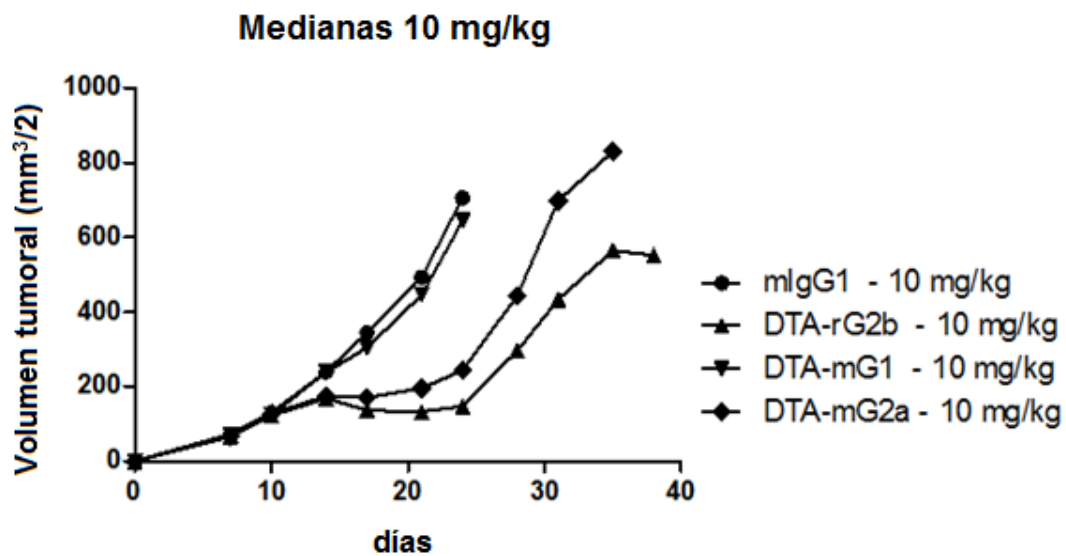


Figura 46B

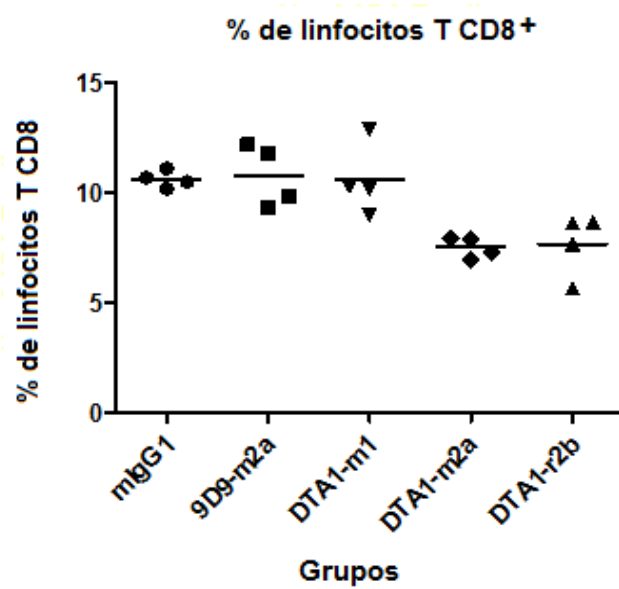


Figura 47A

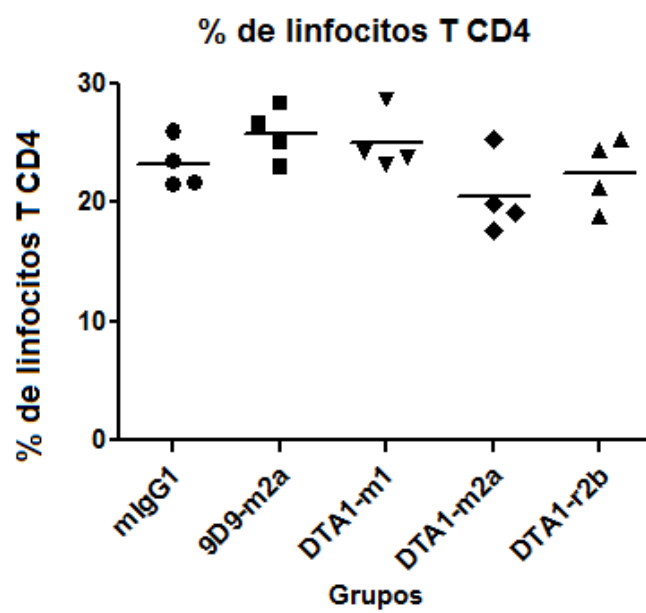


Figura 47B

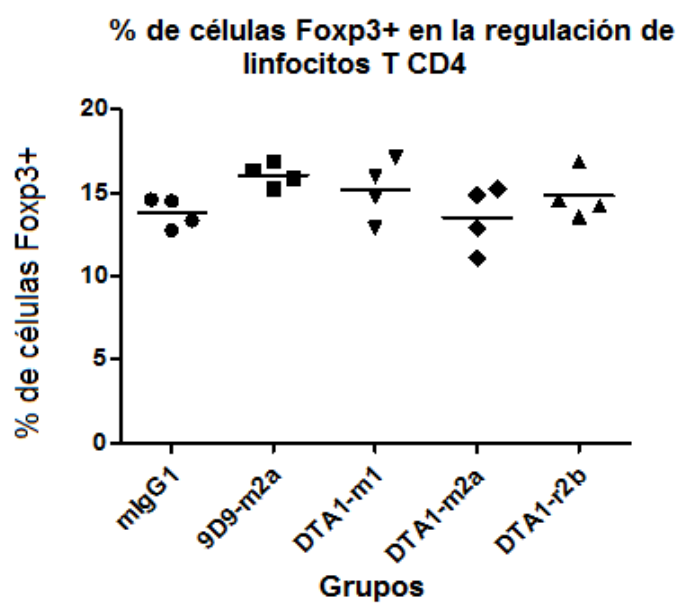


Figura 47C

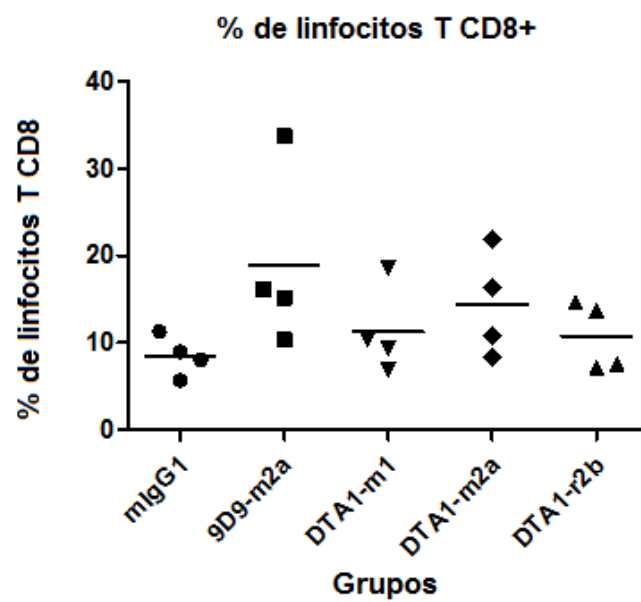


Figura 47D

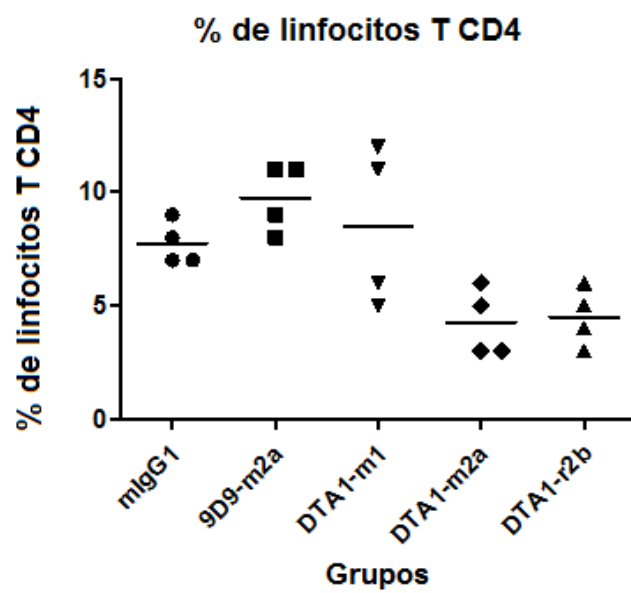


Figura 47E

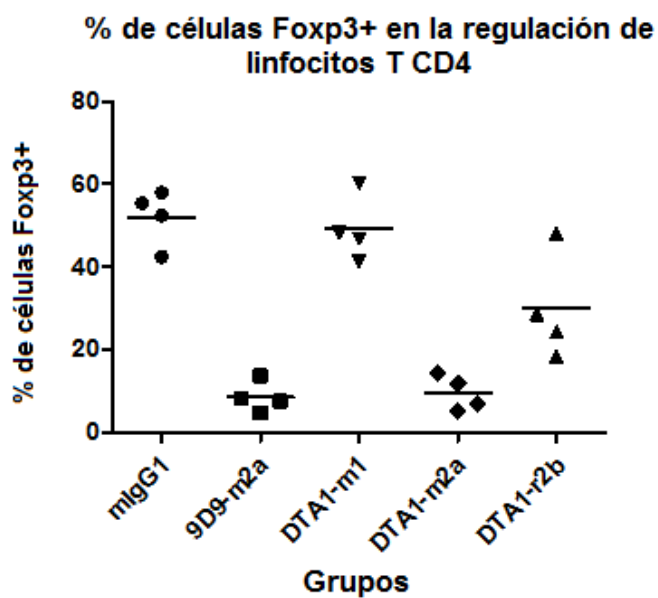
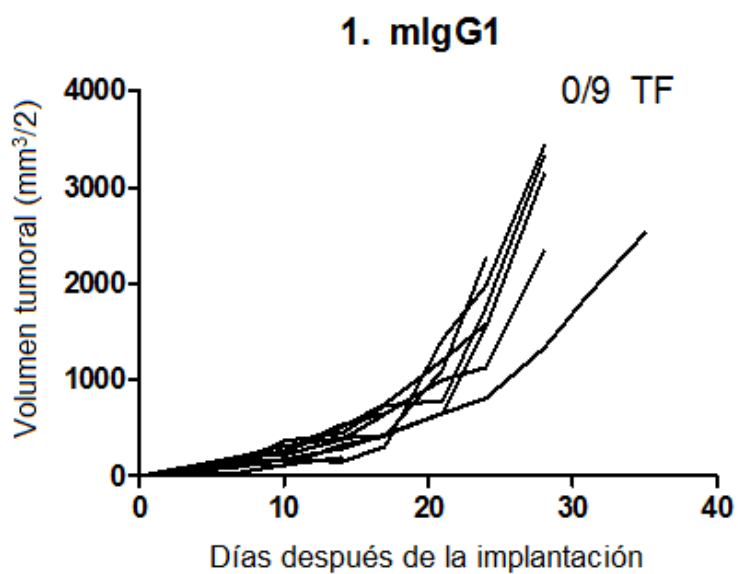
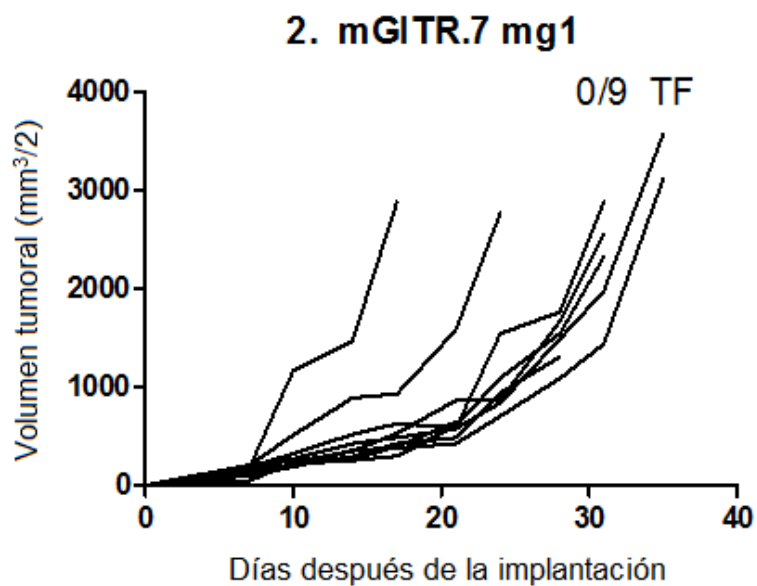
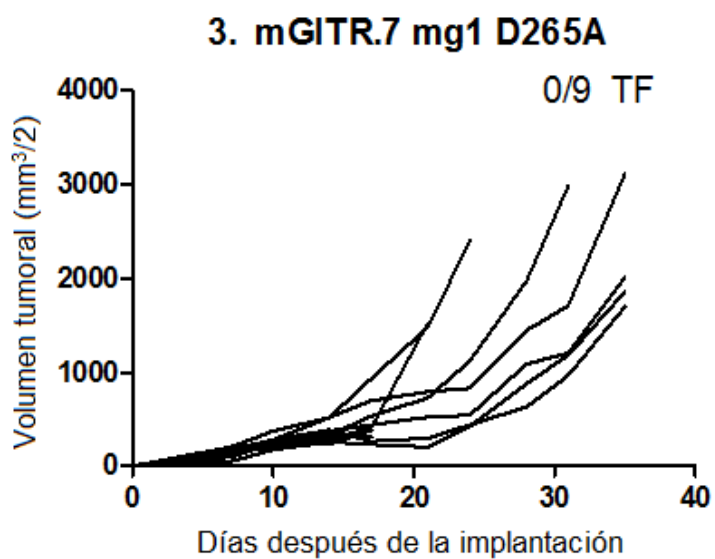
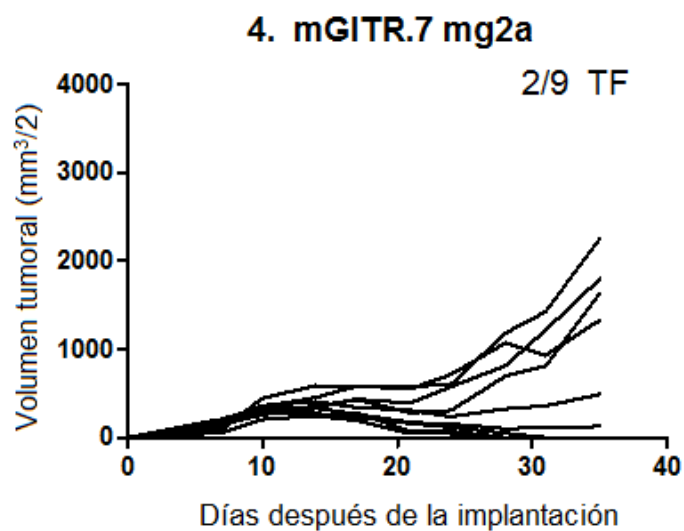
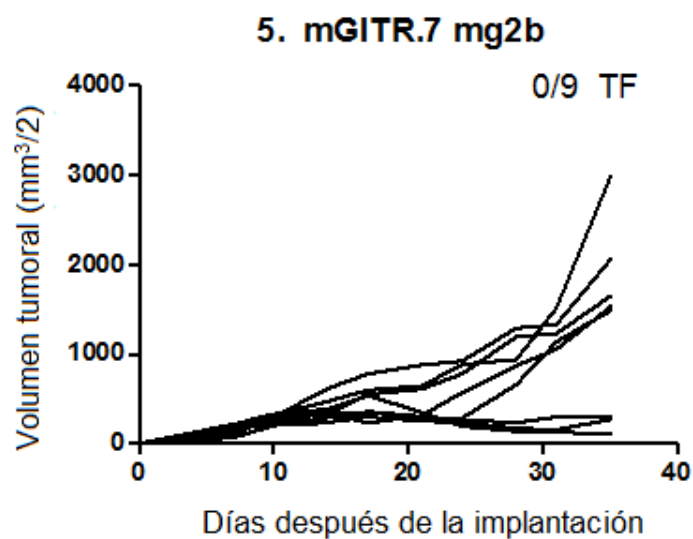
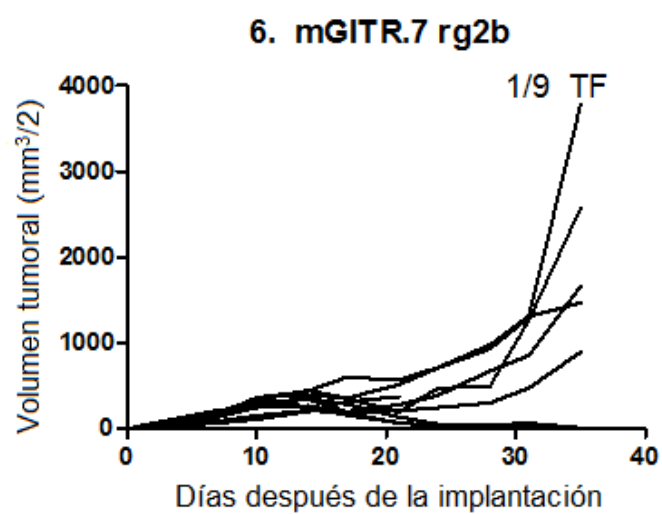


Figura 47F

**Figura 48A****Figura 48B**

**Figura 48C****Figura 48D**

**Figura 48E****Figura 48F**

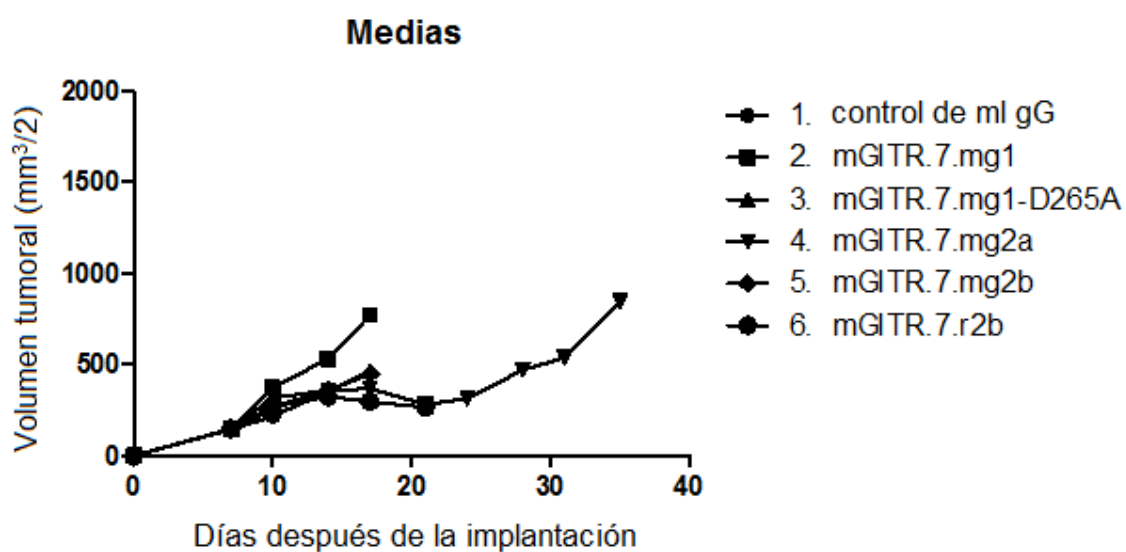


Figura 49A

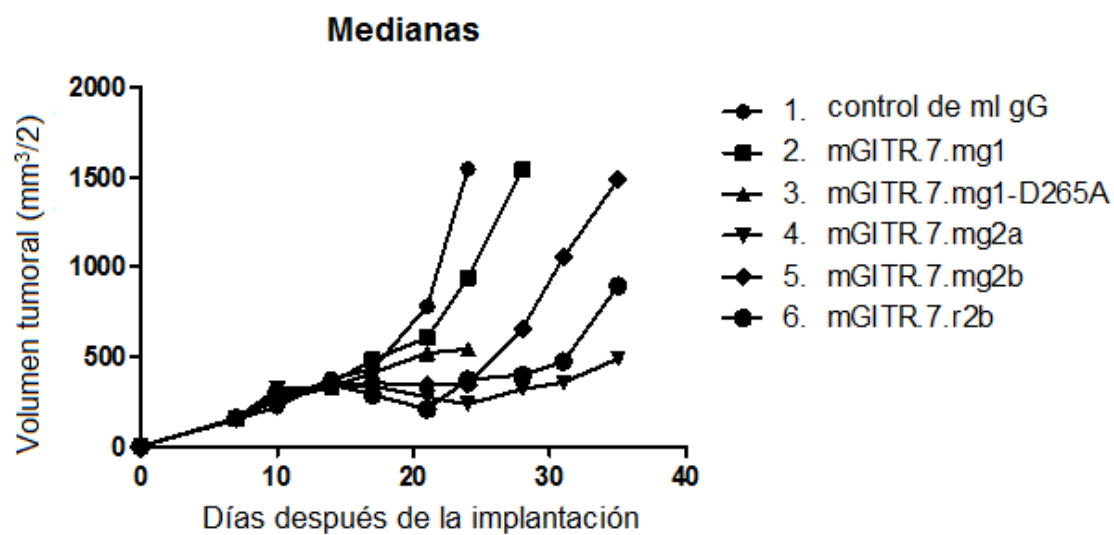


Figura 49B

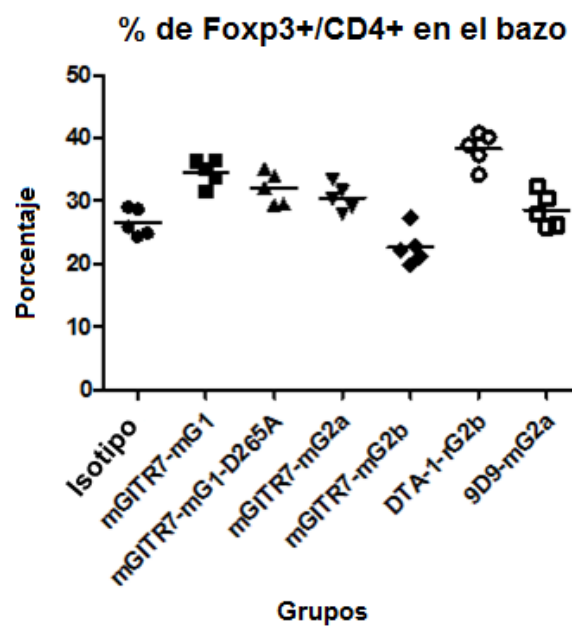


Figura 50A

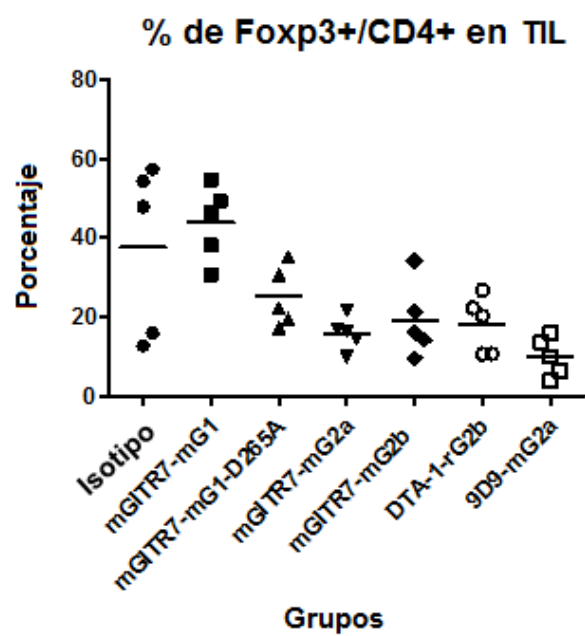


Figura 50B

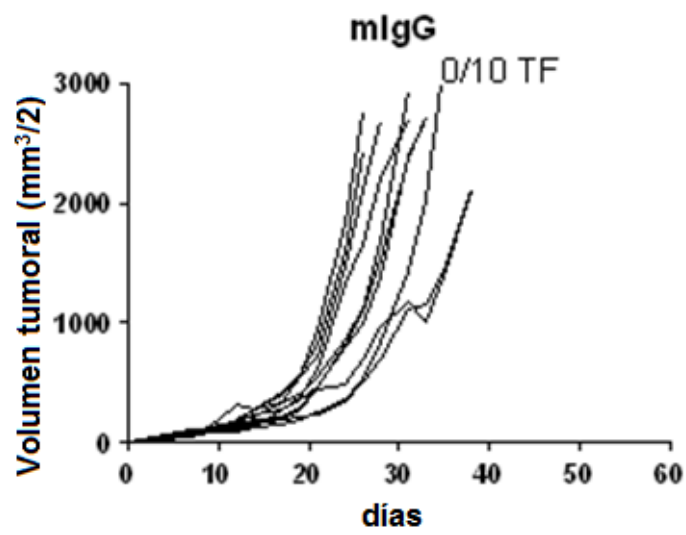


Figura 51A

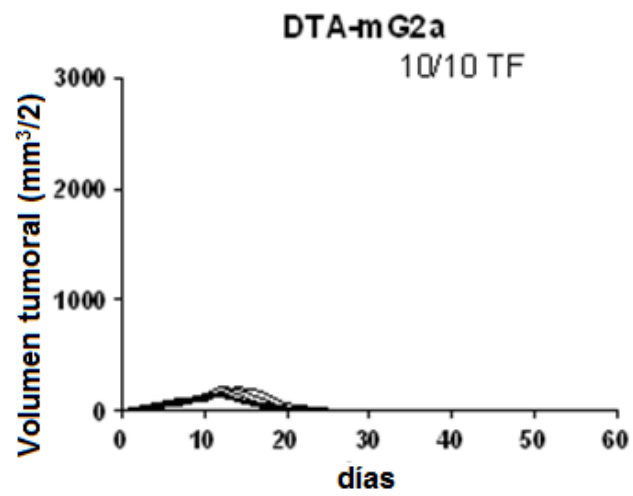


Figura 51B

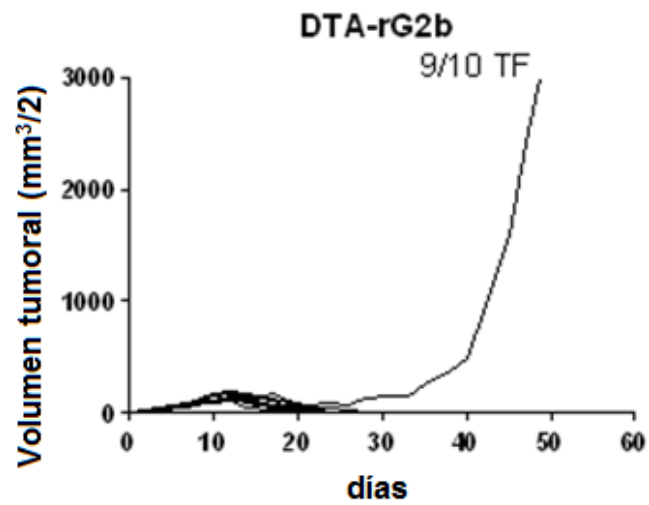


Figura 51C

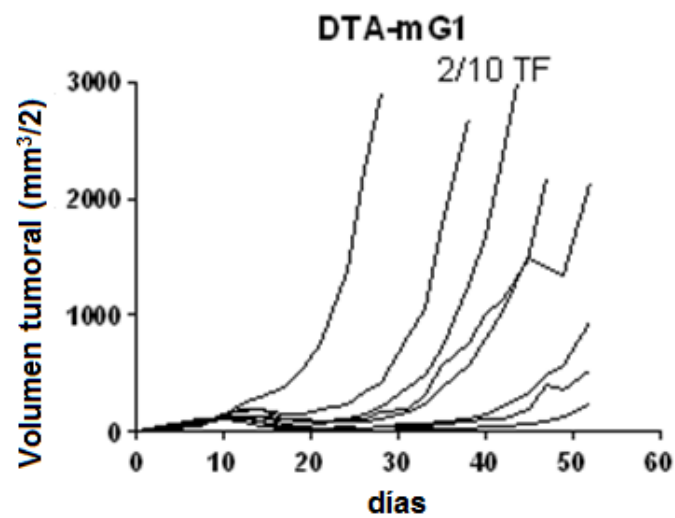


Figura 51D

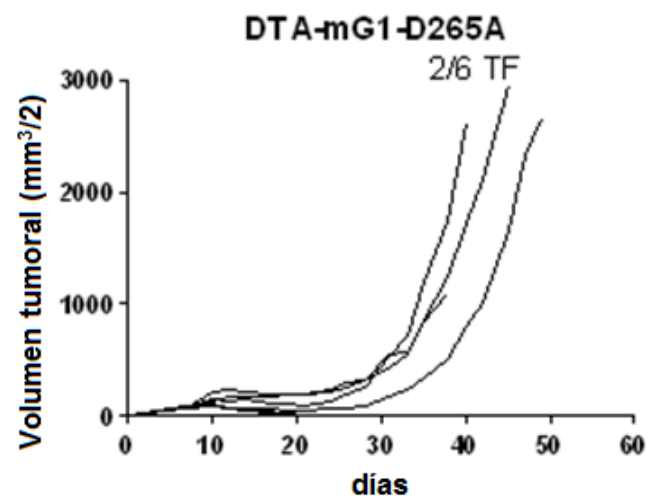


Figura 51E

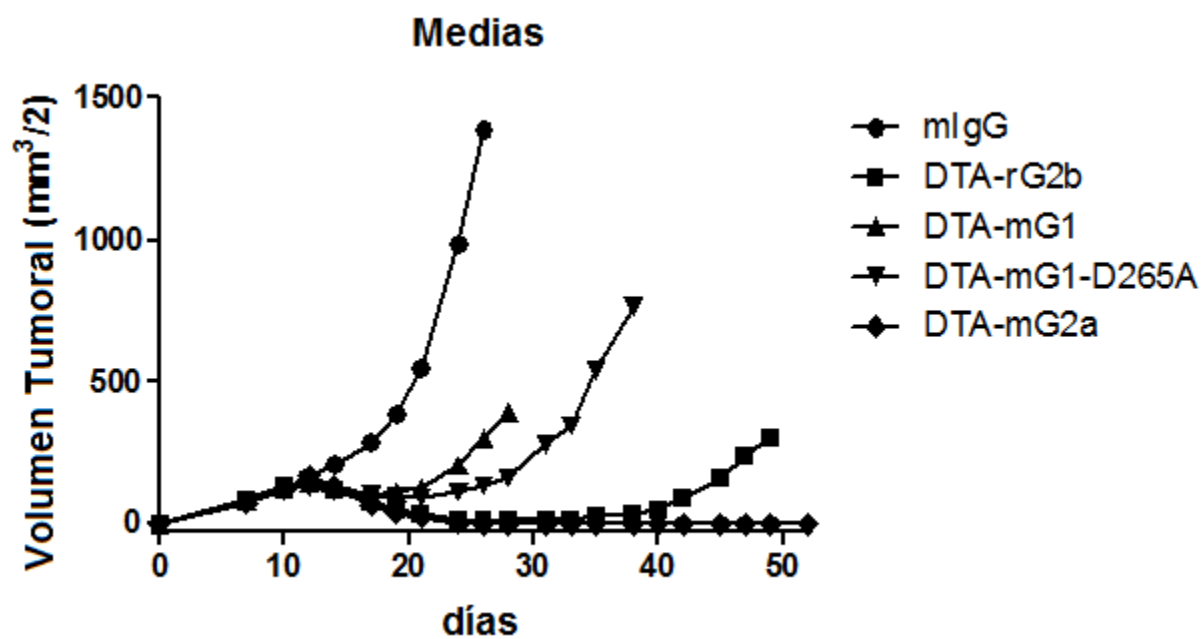


Figura 52A

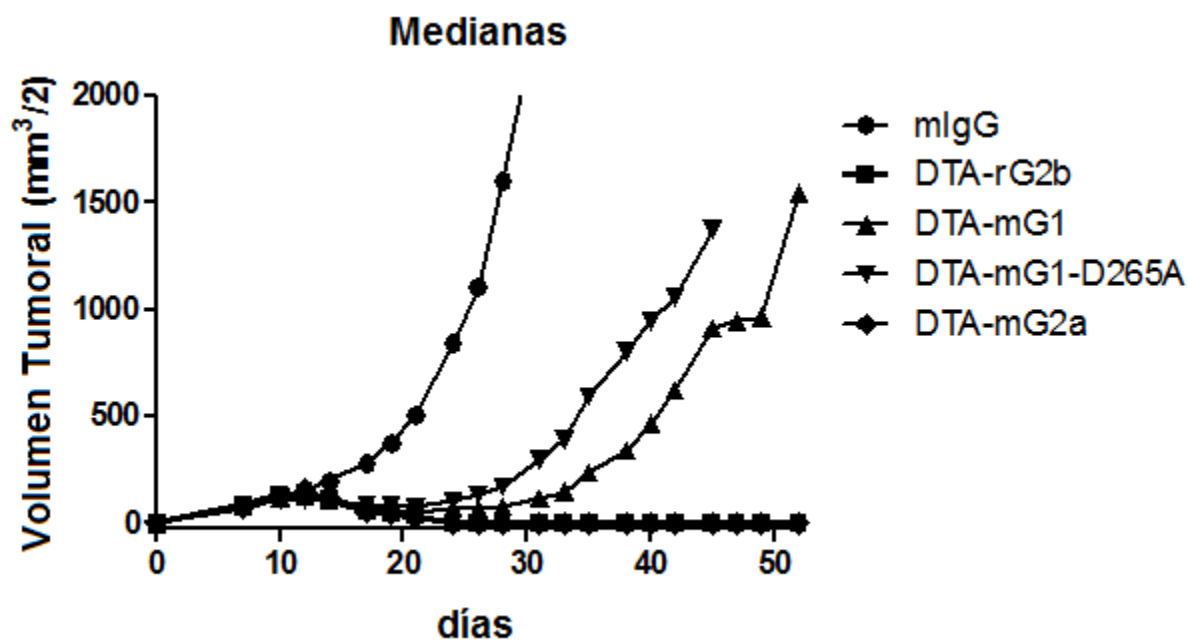


Figura 52B

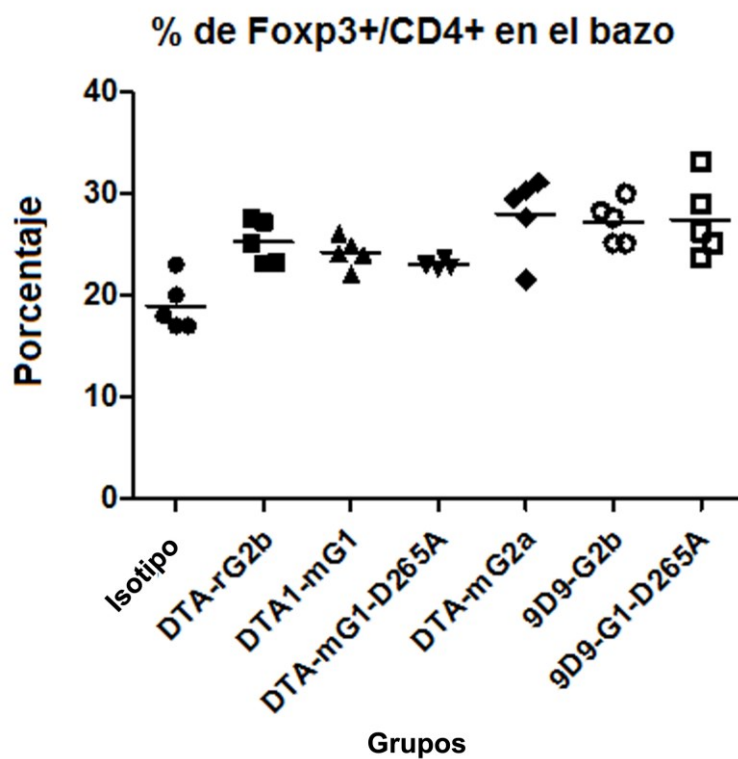


Figura 53A

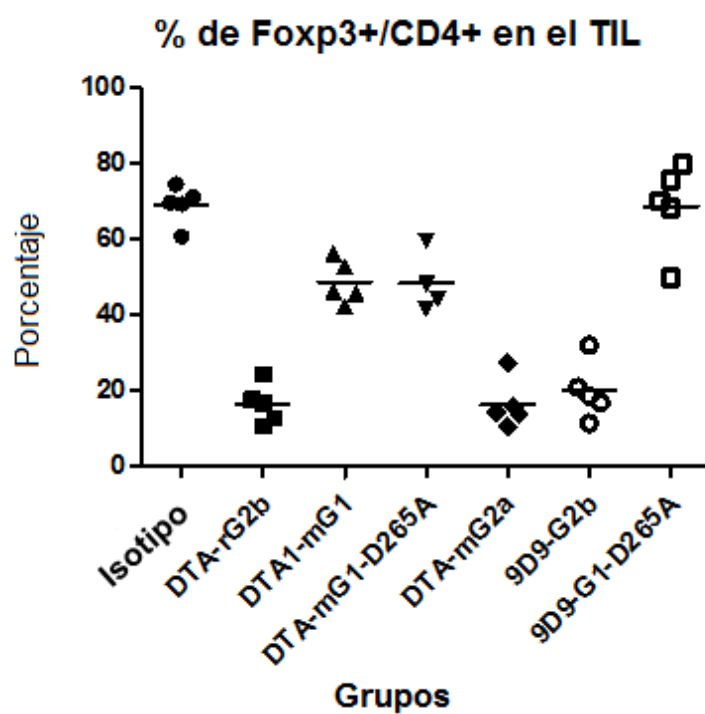
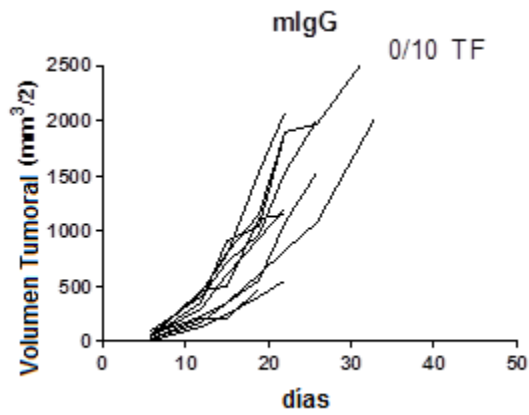
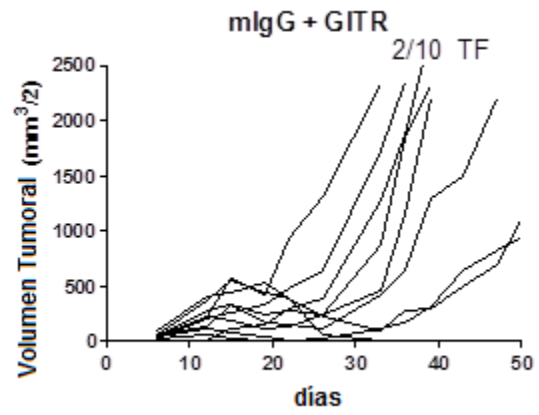
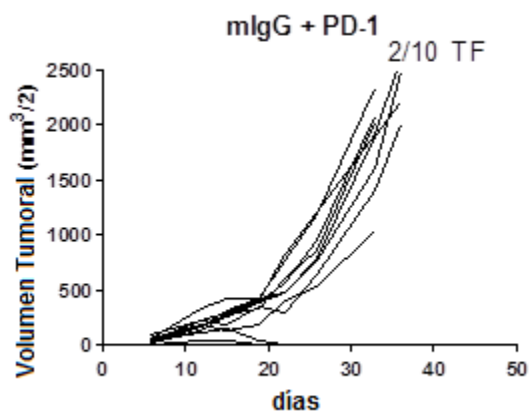
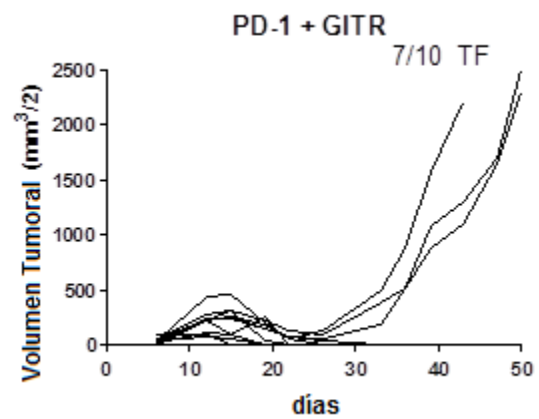


Figura 53B

**Figura 54A****Figura 54B****Figura 54C****Figura 54D**

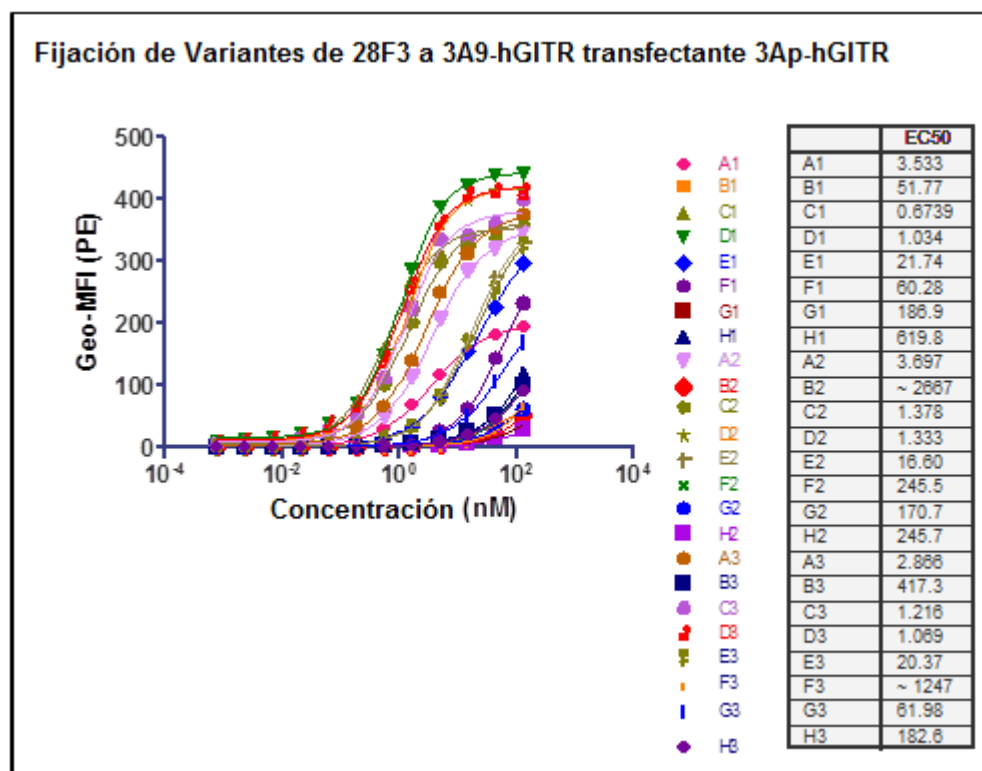


Figura 55A

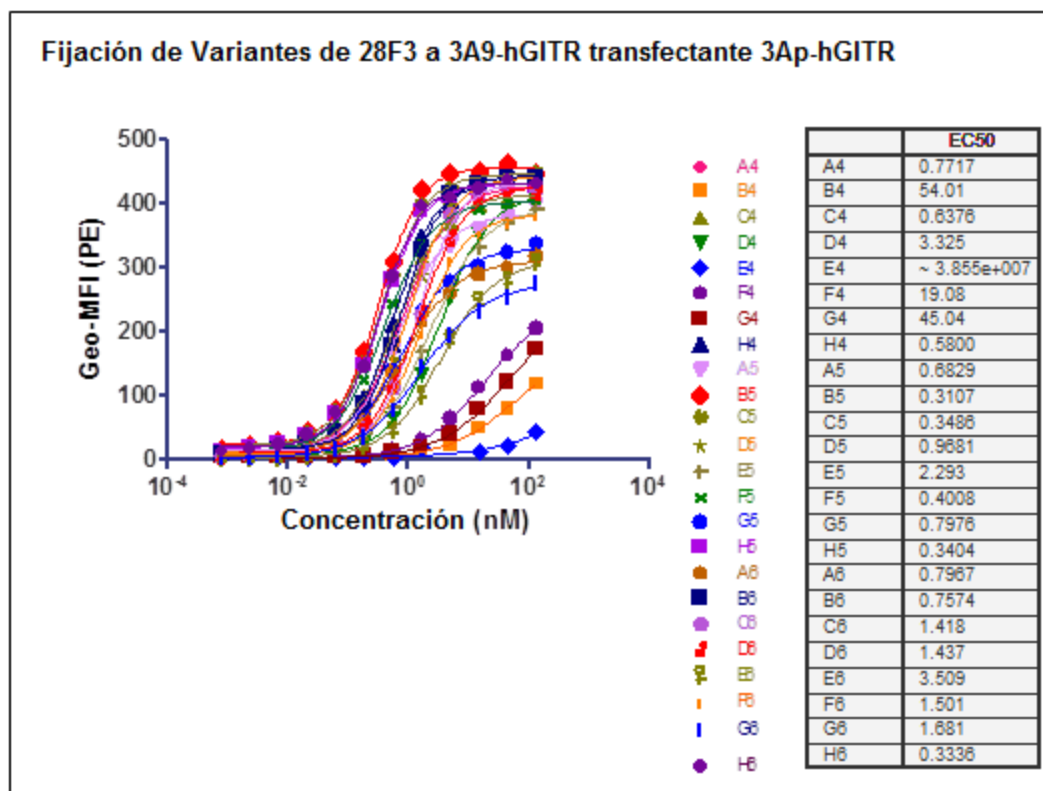


Figura 55B

**Variantes de 28F3 en secreción de IL-2 de células 3A9-hGITR
en presencia de anti-CD3 unido a placas**

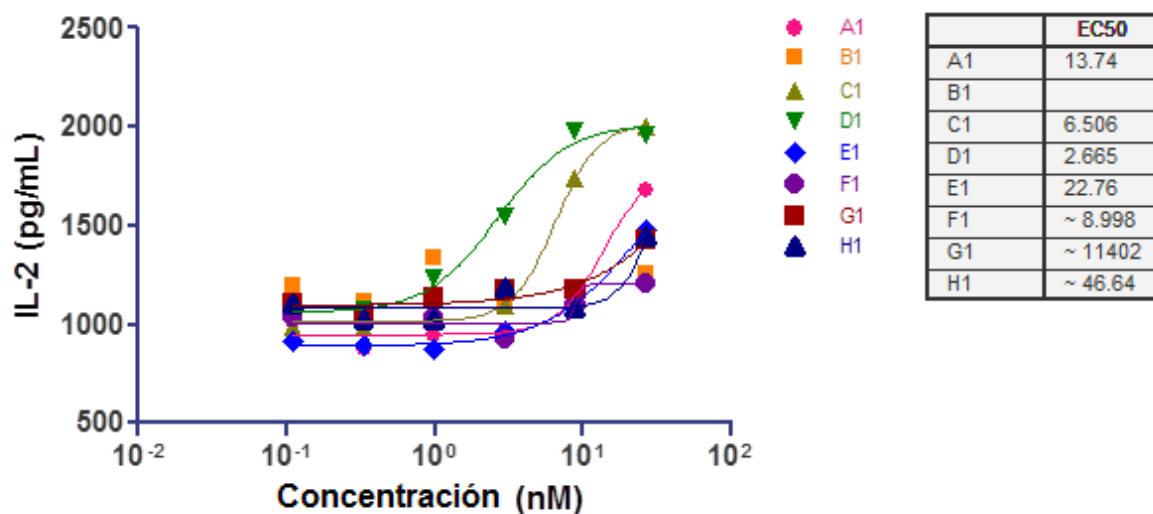


Figura 56A

**Variantes de 28F3 en secreción de IL-2 de células 3A9-hGITR
en presencia de anti-CD3 unido a placas**

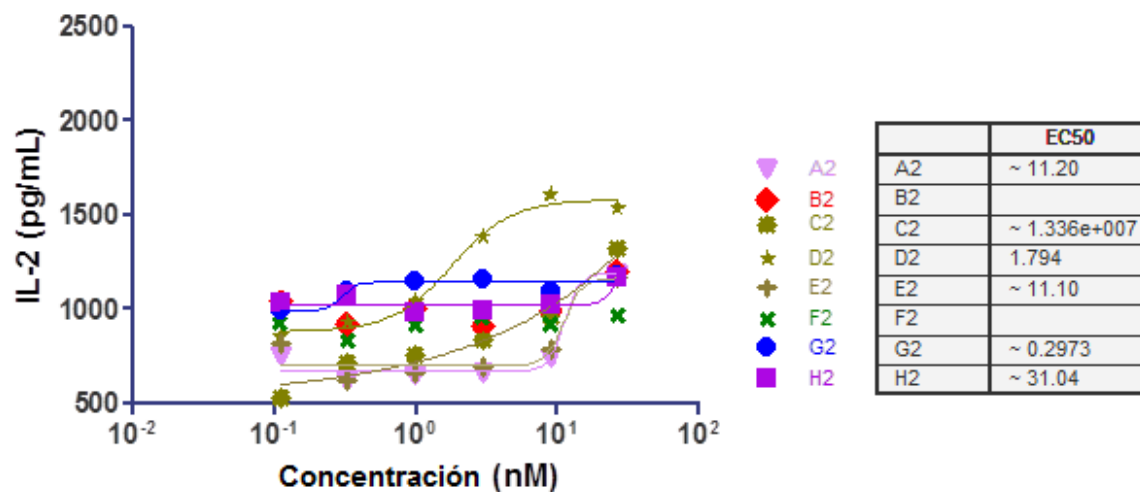


Figura 56B

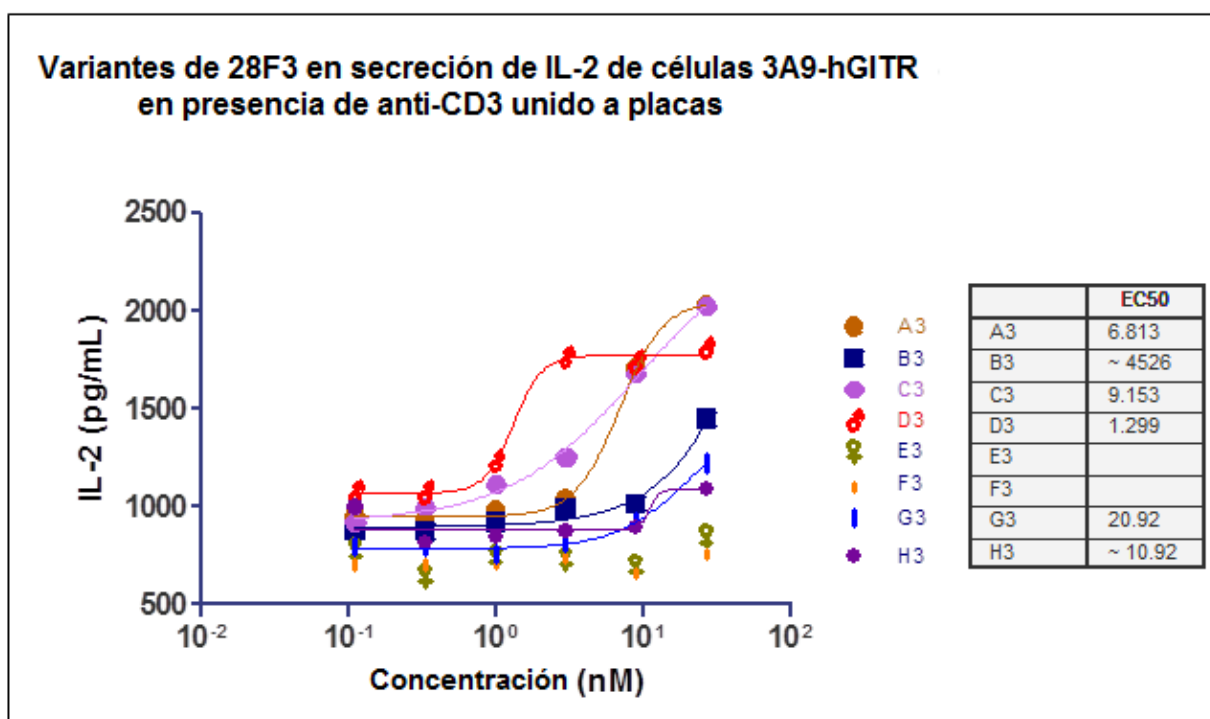


Figura 56C

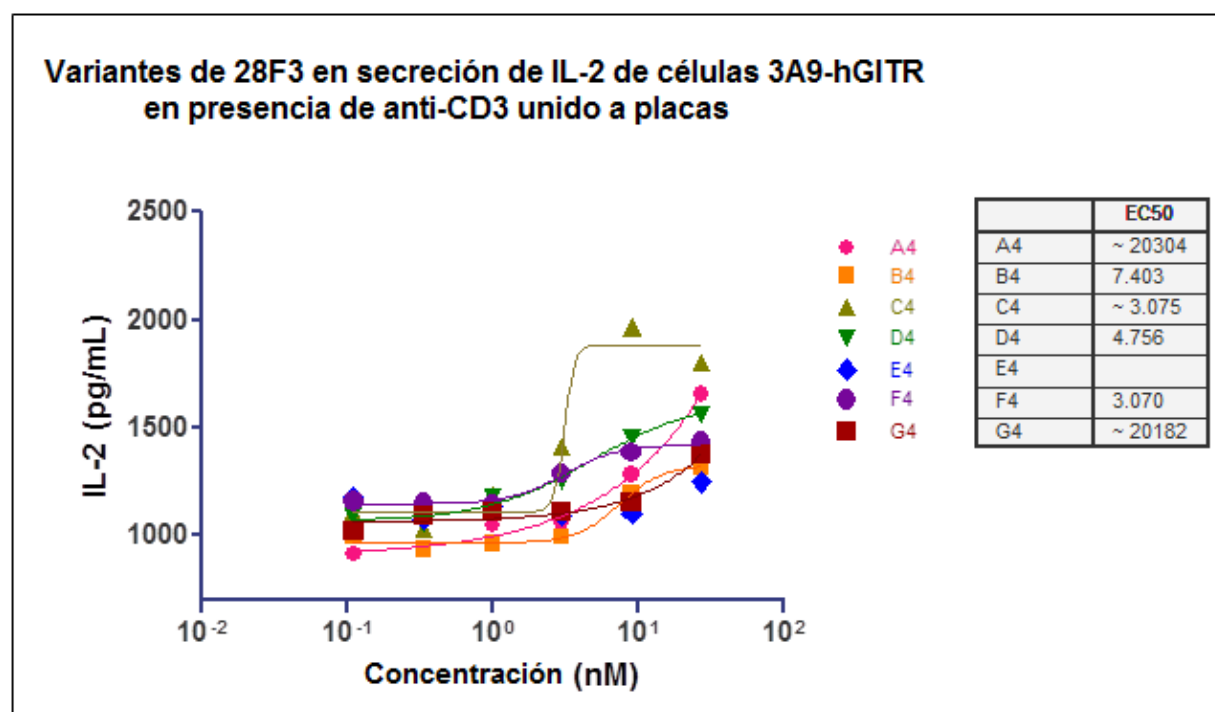


Figura 56D

Variantes de 28F3 en secreción de IL-2 de células 3A9-hGITR en presencia de anti-CD3 unido a placas

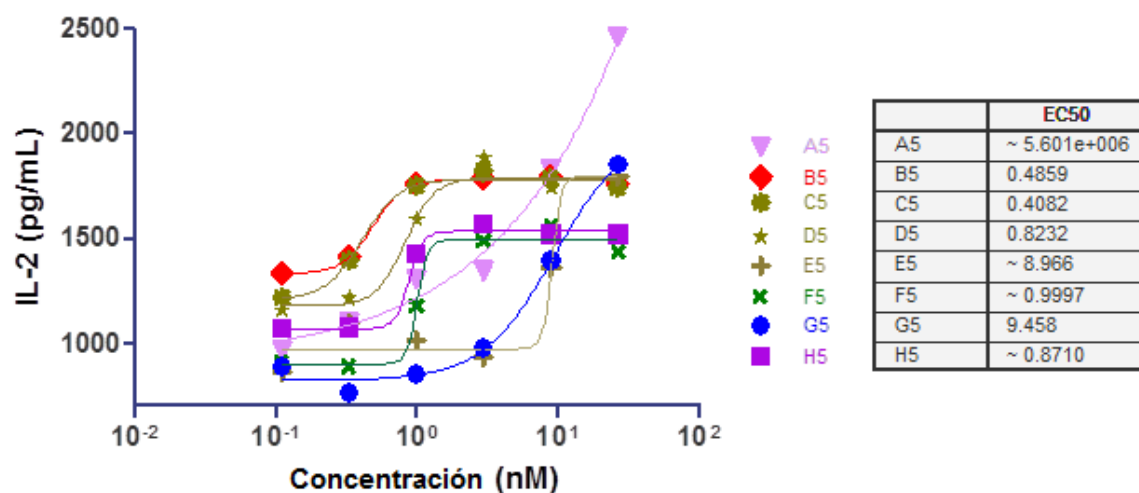


Figura 56E

Variantes de 28F3 en secreción de IL-2 de células 3A9-hGITR en presencia de anti-CD3 unido a placas

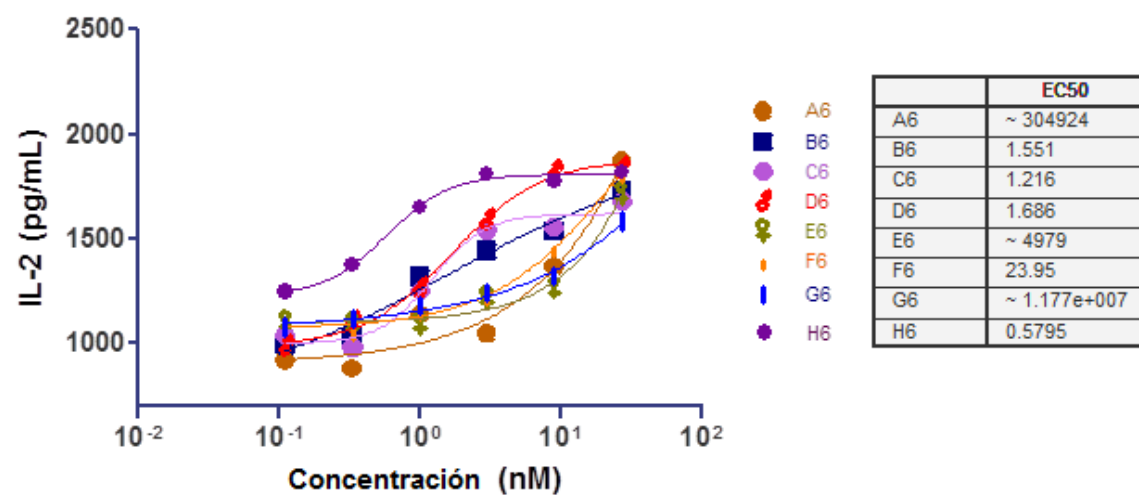
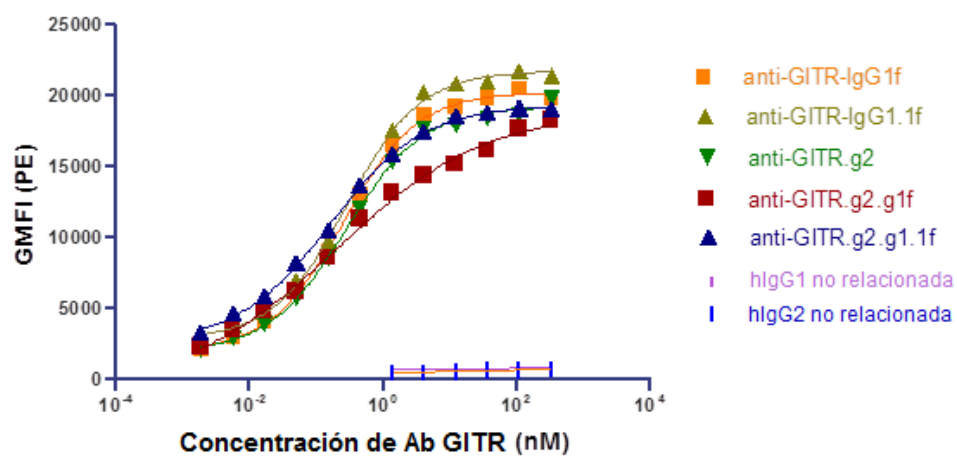


Figura 56F



	EC50
anti-GITR-IgG1f	0.2566
anti-GITR-IgG1.1f	0.2865
anti-GITR.g2	0.2773
anti-GITR.g2.g1f	0.2034
anti-GITR.g2.g1.1f	0.1628
hlgG1 no relacionada	131.8
hlgG2 no relacionada	6.725

Figura 57

Ab GTR en secreción de IFN-g mediante linfocitos T CD4 estimulados con células CHO que expresan OKT3 - Donante #2

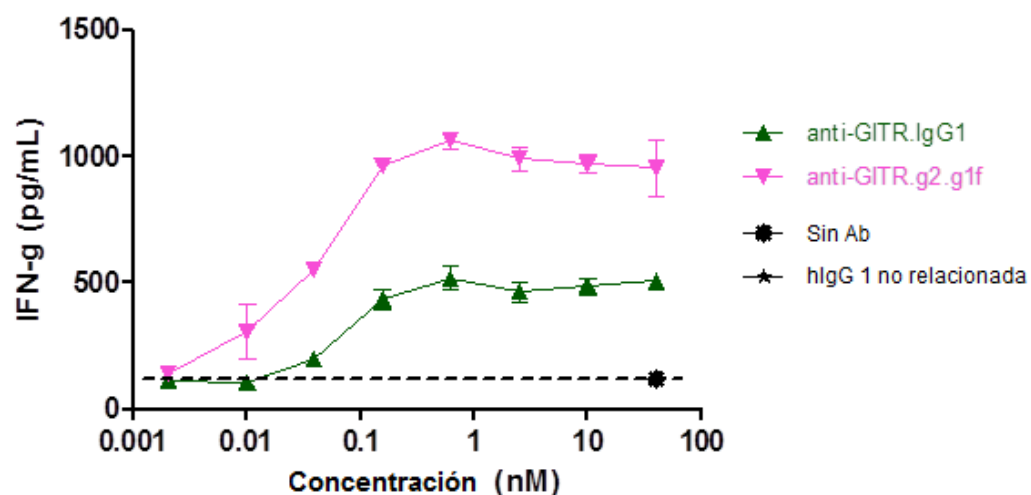


Figura 58A

Ab GTR en secreción de IL-2 mediante linfocitos T CD4 estimulados con células CHO que expresan OKT3 - Donante #3

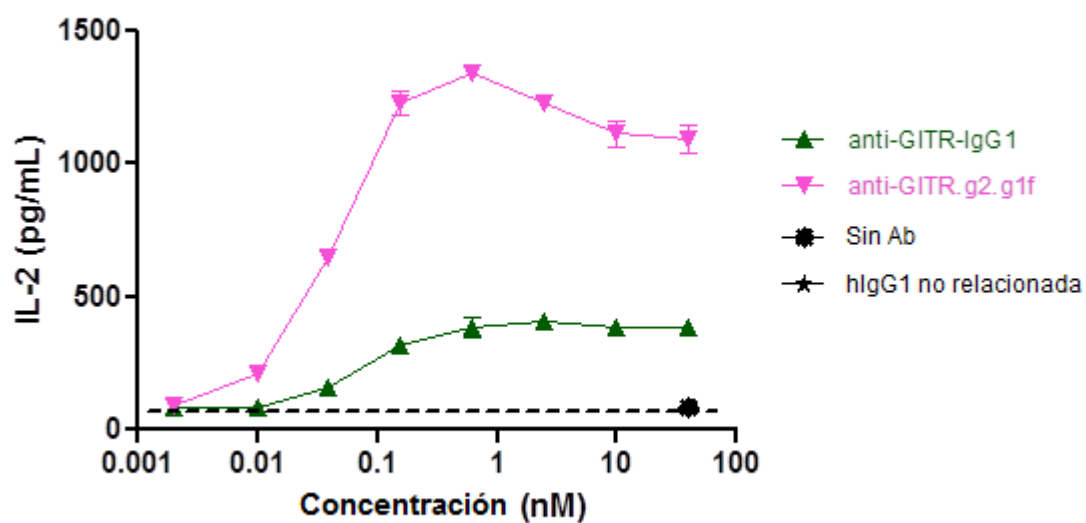


Figura 58B

**Ab GITR en secreción de IL-2 mediante linfocitos T CD4 estimulados
con células CHO que expresan OKT3 - Donante #3**

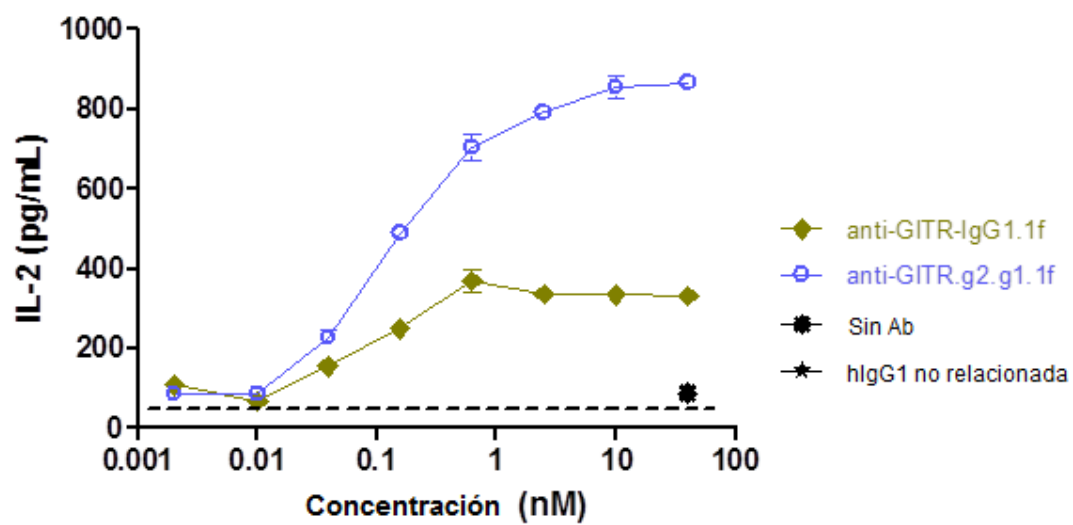


Figura 58C

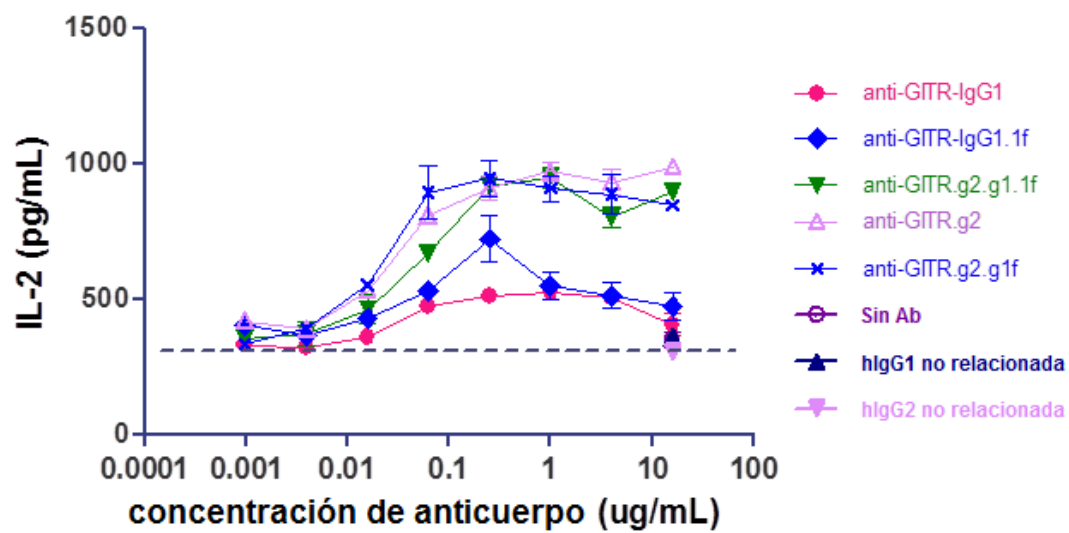
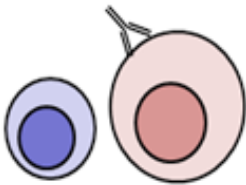
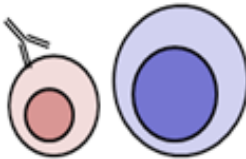


Figura 59



Treg	Teff
WT	huGITR



Treg	Teff
huGITR	WT

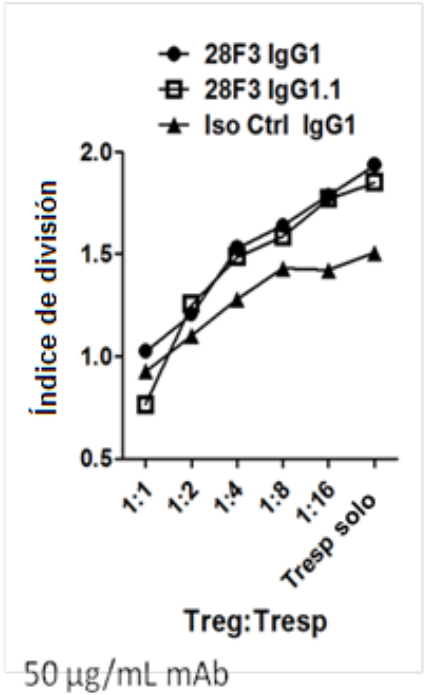


Figura 60A

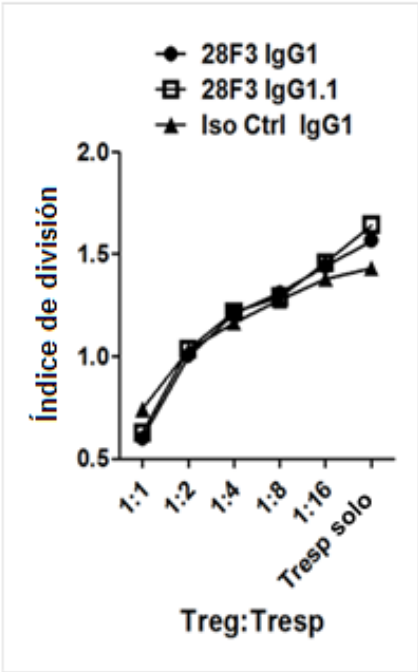


Figura 60B

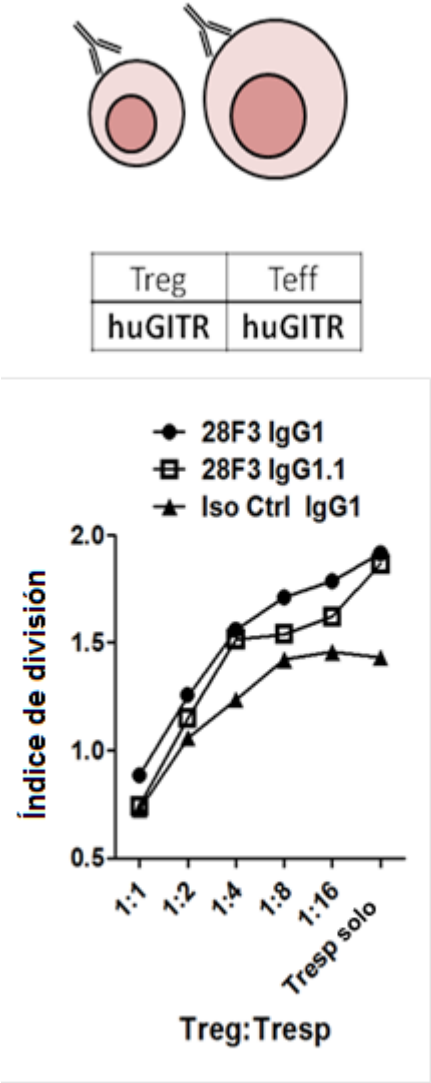


Figura 60C

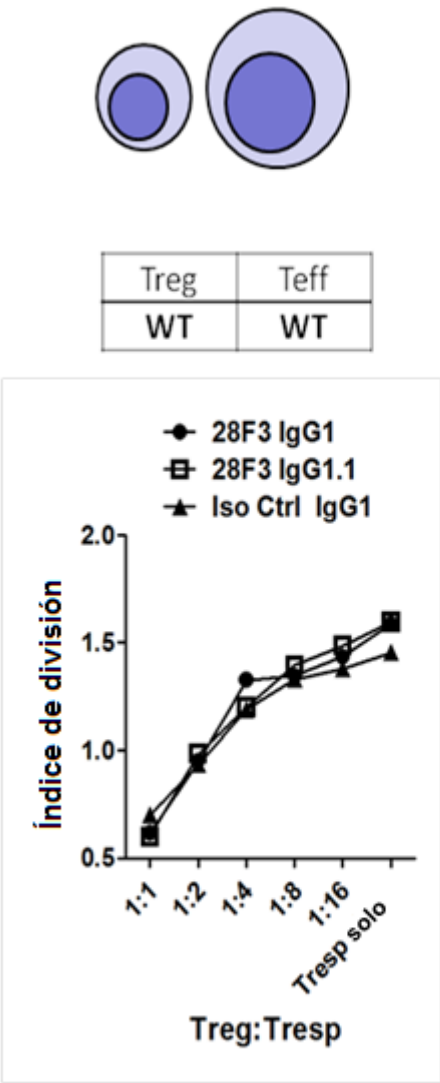


Figura 60D

Donante 1

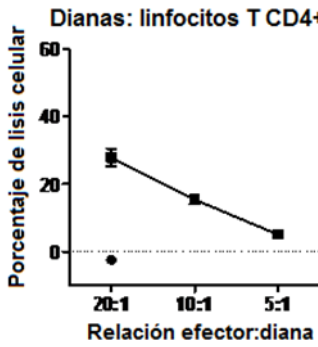


Figura 61A

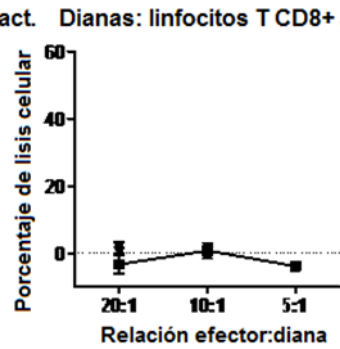


Figura 61B

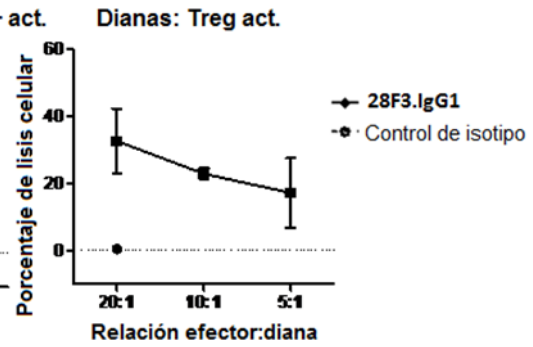


Figura 61C

Donante 2

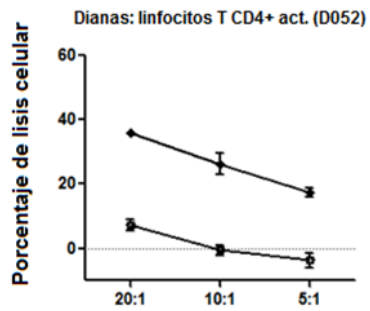


Figura 61D

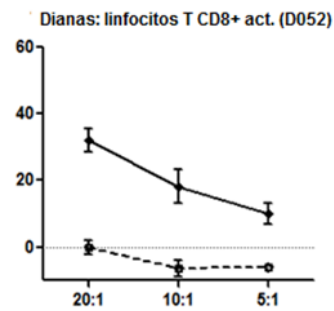


Figura 61E

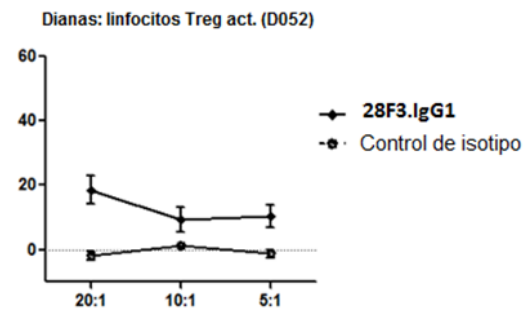


Figura 61F

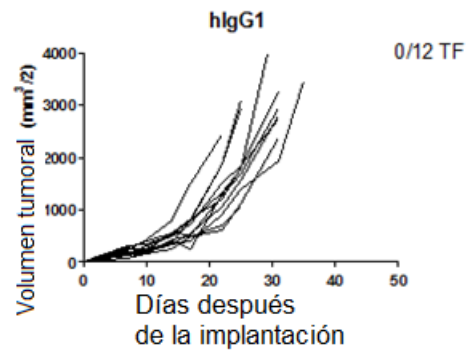


Figura 62A

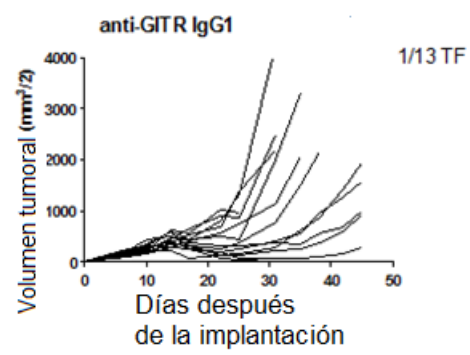


Figura 62B

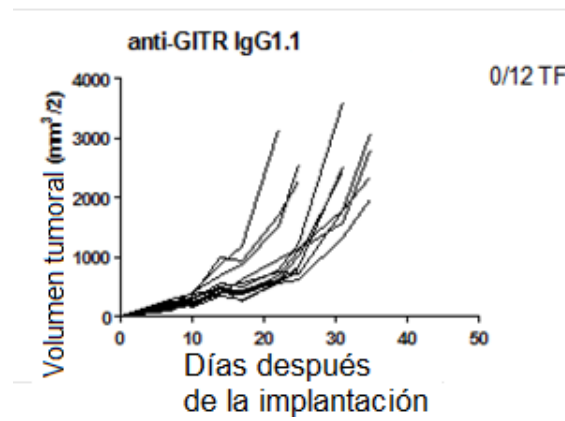
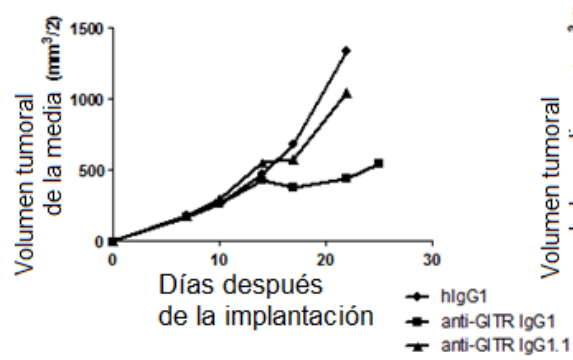
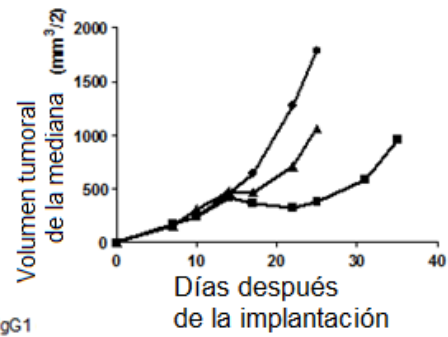


Figura 62C

**Figura 63A****Figura 63B**

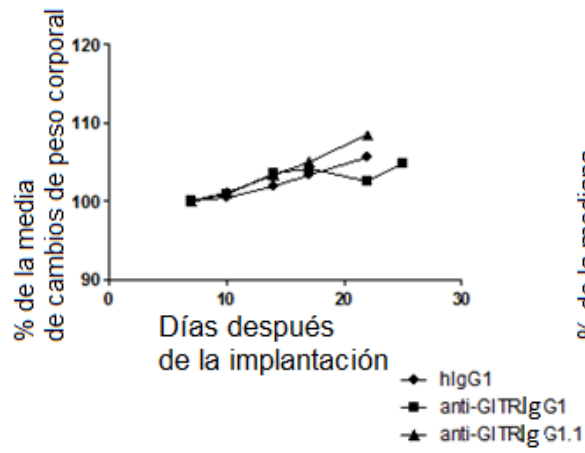


Figura 64A

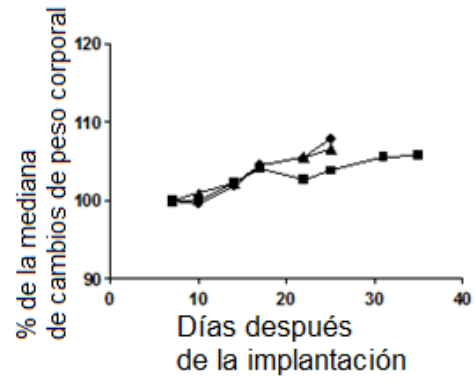
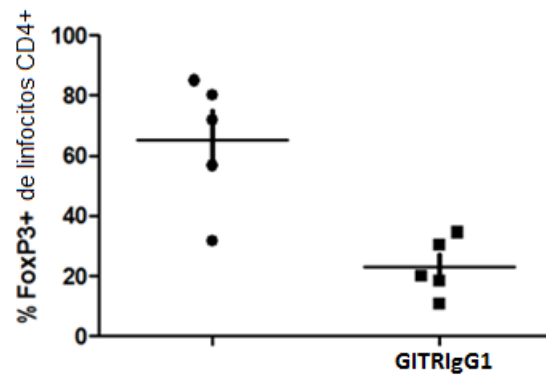


Figura 64B

**Figura 65**

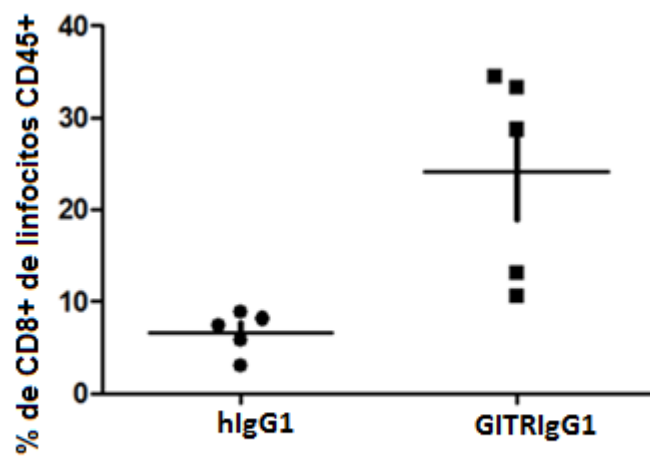


Figura 66

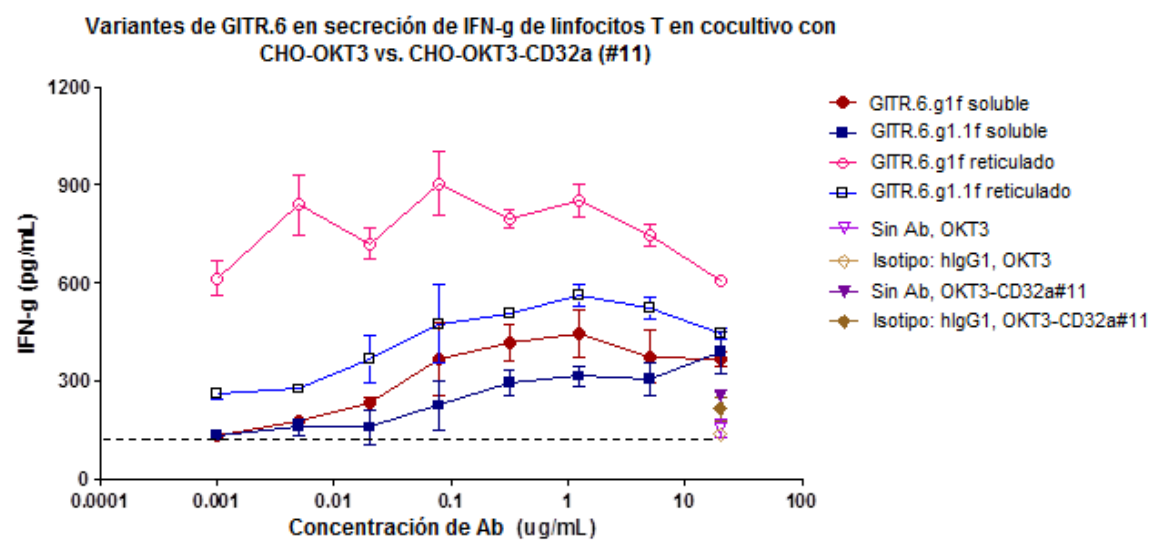


Figura 67

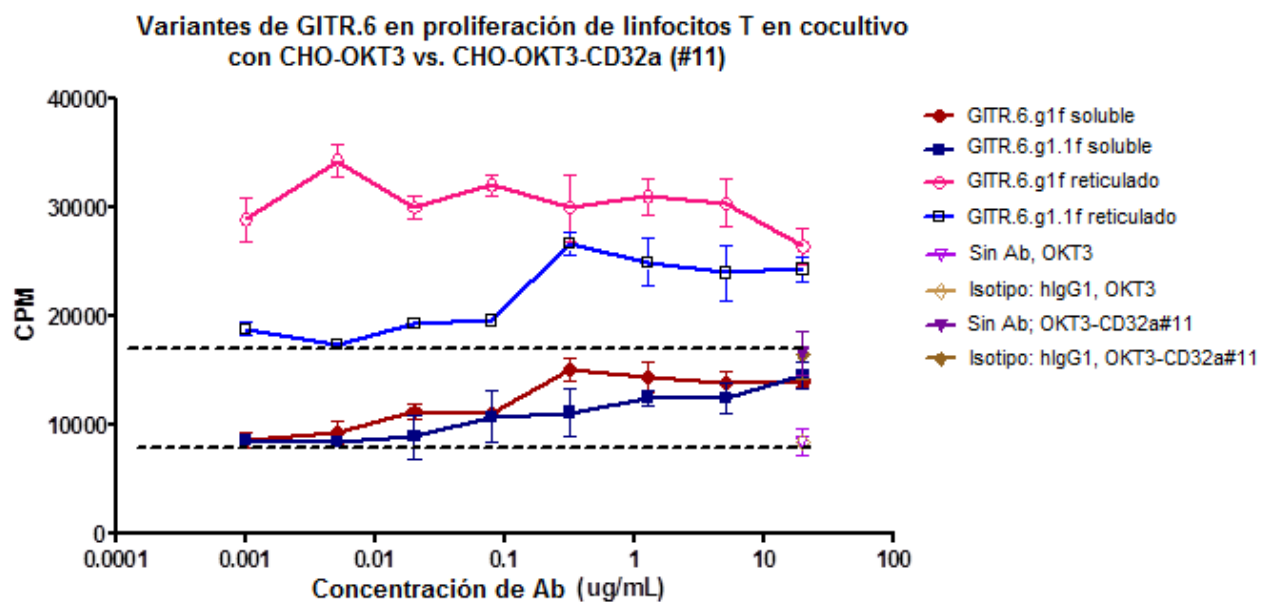


Figura 68