

FIGURA 1

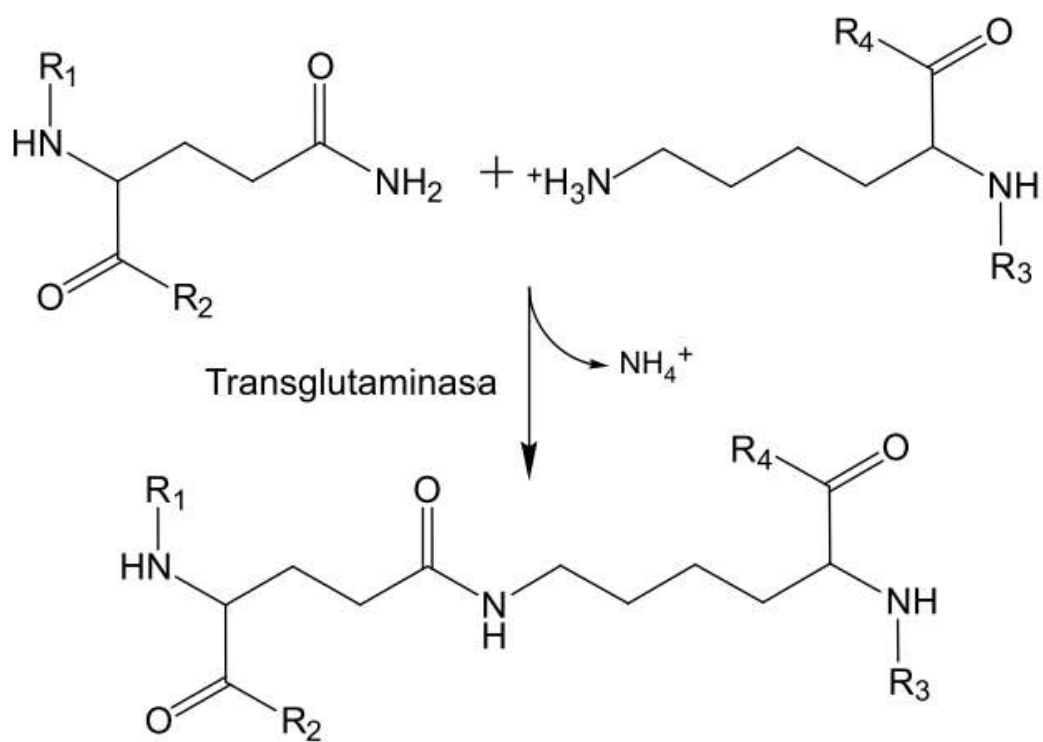


FIGURA 2

Z-Gln-Gly	
Z-Gln-Gly-CAD-Biotina	
Z-Gln-Gly-N ₃	
Z-Gln-Gly-PEG ₂ -BCN	
Z-Gln-Gly-PEG ₂ -AuristatinaF	

FIGURA 3

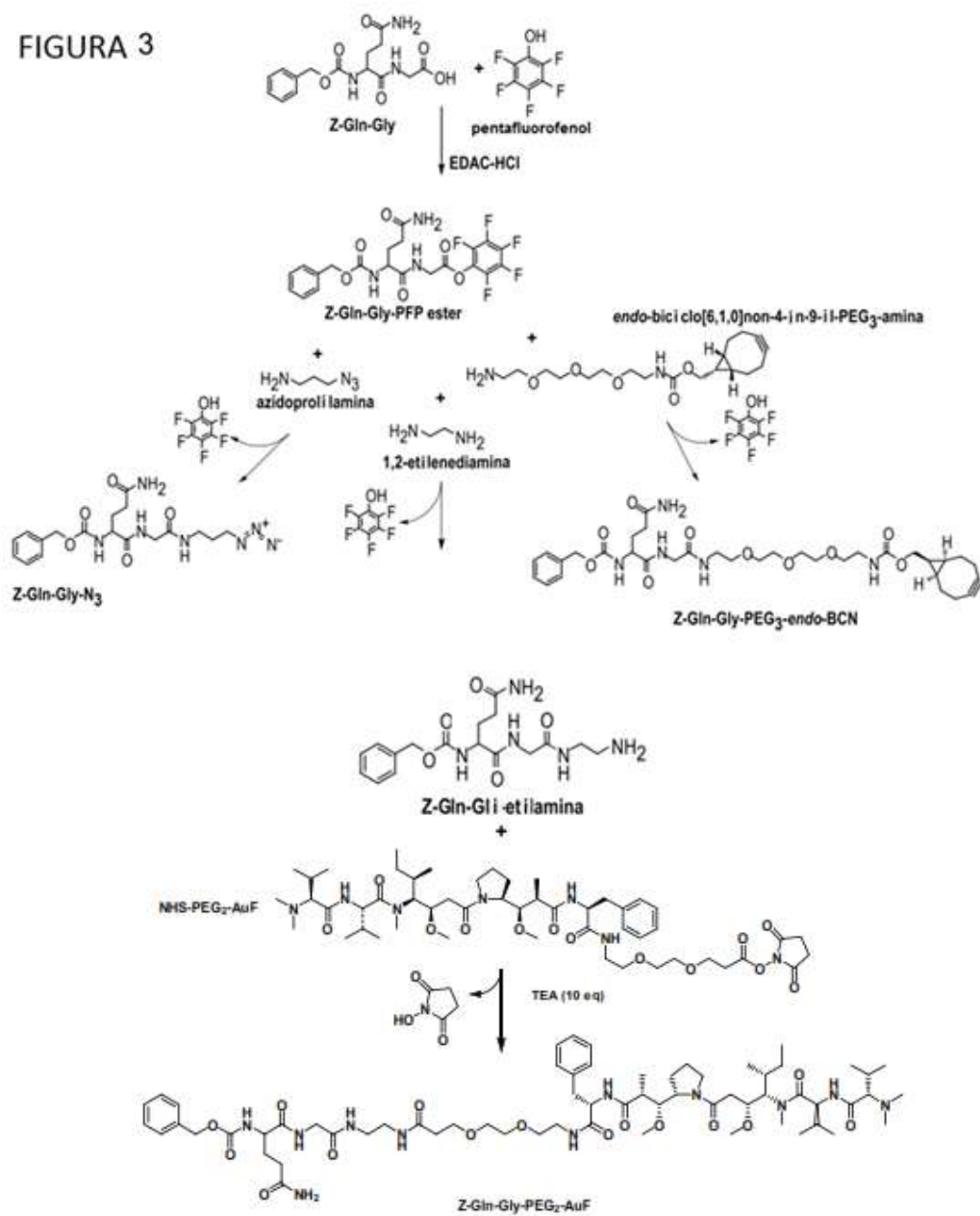


FIG. 4A. Fab-kappa



FIG. 4B. Fab-lambda

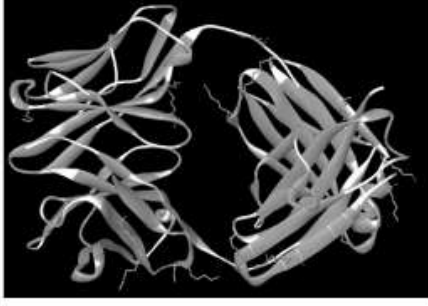


FIG. 4C. Fc



FIG. 5A. Anticuerpo 02

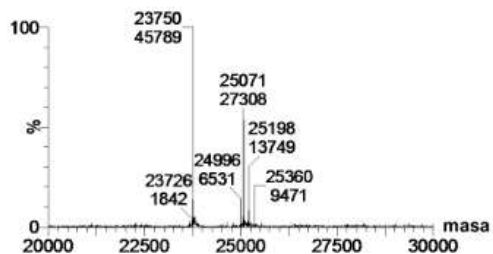


FIG. 5B. Anticuerpo 03

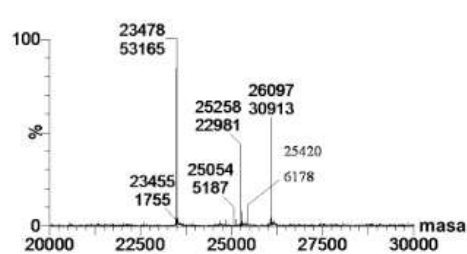


FIG. 5C. Anticuerpo 01

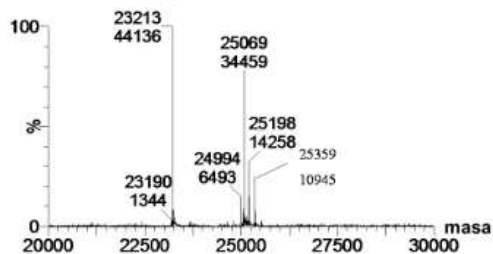


FIG. 5D. Anticuerpo 04

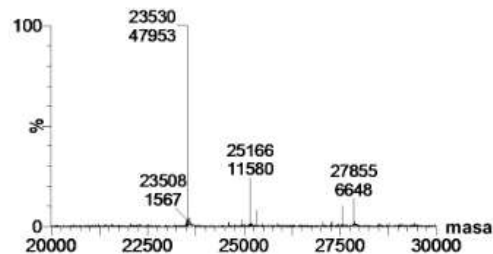


FIG. 5E. Anticuerpo 05

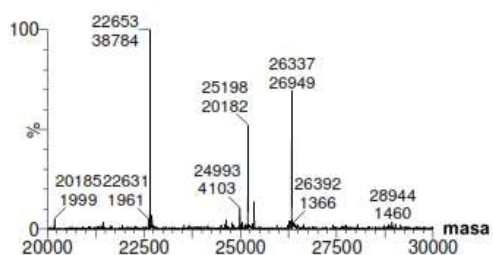


FIG. 5F. Anticuerpo 06

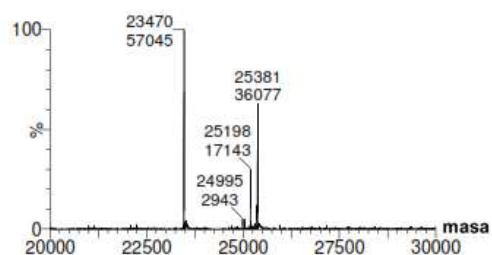


FIGURA 6

hu Cy
118
AST**K**GPSVFP LAPSS**K**STSG GTAALGCLVK DYFPEPVTVS WNSGALTSGV HIFPAVLQSS GLYSLSSVVT VPSSSLGTQT YICNVNH**K**PS NT**K**VD**K**KVEP
218
KSCD**K**THTCP PCPAPELLGG PSVFLFP**K**P **K**DTILMISRTF EVTCVVVDVS HEDPEV**K**FNW YVDGVEVHNA **K**TKPREEQYN STYRVVSVLI VLHQDWLN**K**
318
EYK**K**VSN**K**A LPAPI**K**TIS KA**K**QPREPQ VYTLPPSRDE LT**K**NQVSLTC LVKGFYPSDI AVEWESNGQP ENNY**K**TTPPV LDSDGSFFLY SKLTVD**K**SRW
418 447
QQGNVFSCSV MHEALHNHYT **Q**KSLSLSP**K**

hu Ck
108
RTVAAPSVFI FPPSDEQL**K**S GTASVVCLLN NFYPRE**K**VQ **W**KVDNALQSG NSQESVTEQD **S****K**DSTYSLSS TLTLS**K**ADYE **K**H**K**VYACEVT HQGLSSP**V****T****K**
208
SFNRGEC

hu CA
110
GQP**K**AAAPSVI LFPSPSEELQ AN**K**ATLVCLI SDFYFGAVTV AWKADSP**V****K** AGVETITPSK QSN**N**KYAASS YLSLTP**E**Q**W****K** SHKSYSCQVT HEGSTVE**K**TV
210
APTECS

FIGURA 7

CH1			conjugación			Bisagra superior			conjugación			CH2			conjugación			CH3			conjugación		
RFU												RFU						RFU					
A118K	1462.657		0.0%			E216K	1646.217		0.0%			G236K	2236.25					G341K	3075.741				
S119K	2972.125		0.0%			P217K	2532.893		0.0%			G237K	2304.169					Q342K	3572.799				
T120K	2162.525		43.2%			S219K	2986.836		0.0%			P247K	3386.941					P343K	5578.971	0.0%			
G122K	2389.853		0.0%			D221K	7438.612		87.6%			M252K	7153.469		75.1%			R344K	2447.62				
S131K	2489.956		0.0%			T223K	7519.935		90.8%			I253K	1947.679					E345K	1805.326				
S132K	2228.079		0.0%			H224K	8863.961		71.6%			S254K	2363.706					R355K	2694.082				
S134K	2310.796					T225K	4849.386		100.0%			R255K	1978.038					D356K	2725.487				
T135K	6392.472		80.4%									T256K	1834.634					L358K	2912.335				
S136K	10829.907		100.0%									D265K	3087.407					T359K	3624.477				
G137K	8619.62		38.9%									S267K	1705.879					N361K	2347.795				
G138K	1366.329		0.0%			C226K	2532.581		0.0%			H268K	175.398					Q362K	2385.153				
T139K	2209.782		23.4%			P227K	2354.523		0.0%			E269K	1357.406					S375K	2205.751				
E152K	1624.412		0.0%			P228K	2325.735		0.0%			D270K	1667.335					D376K	3507.073				
P153K	2304.666		0.0%			C229K	1763.406		0.0%			P271K	2086.99					E382K	1826.903				
S160K	2657.715		0.0%			P230K	2632.028		0.0%			E272K	2473.559					N384K	3079.738				
A162K	2984.247		0.0%									D280K	3097.36					G385K	6249.705	0.0%			
L163K	2882.938		0.0%									G281K	3099.032					Q386K	2659.827				
T164K	6282.873		0.0%									V282K	2597.079					P387K	2416.05				
S165K	1922.563		0.0%			A231K	2109.04		0.0%			E283K	4393.219		0.0%			N389K	3435.688				
G166K	1870.622		0.0%			P232K	1742.478		0.0%			V284K	2111.159					N390K	2623.177				
V167K	1557.957		0.0%			E233K	1873.353		0.0%			H285K	3070.38					L398K	1815.903				
S176K	2301.47		0.0%			L234K	1813.527		0.0%			N286K	2643.754					S400K	1725.105				
S177K	1807.244		0.0%			L235K	2517.083		0.0%			A287K	6908.539		0.0%			D401K	2674.659				
G178K	1964.895		0.0%									T289K	2614.413					G402K	2595.023				
L179K	1686.602		0.0%									Q295K	3647.973					D413K	2176.482				
P189K	1538.761		0.0%									Y296K	2717.136					S415K	2302.063				
S190K	1355.148		0.0%									N297K	8611.262					R416K	2435.814				
S191K			0.0%									S298K	2440.299					Q418K	2642.058				
S192K			35.6%									L309K	1830.155					Q419K	2225.251				
L193K			84.8%									H310K	1755.229					G420K	4998.647	0.0%			
G194K	1121.725		0.0%									Q311K	1682.229					N421K	3497.052				
T195K	2260.968											D312K	2683.16					V422K	3775.808				
Q196K	1529.04		0.0%									L314K	1121.615					A431K	2736.367				
T197K	1537.084		0.0%									N315K	1291.281					H433K	5569.945	0.0%			
P206K	1956.062		0.0%									G316K	1908.498					N434K	3378.801				
S207K	1436.57											E318K	2481.95					H435K	2867.337				
												A327K	2145.191					Y436K	3213.522				
												P329K	2561.945					S442K	3629.279				
												A330K	3577.203					L443K	4658.861	0.0%			
												P331K	2969.843					S444K	5481.729	0.0%			
												S337K	3239.634					P445K	10513.805	79.5%			
												A339K	3088.628					G446K	3399.864				

FIGURA 7 (continuación)

kappa			lambda			Controles		
		conjugación			conjugación		RFU	conjugación
R108K	1671.646		G110K	1357.393		M009	789.529	0.0%
T109K	1359.315		Q111K	1369.309		M009L	12003.892	100.0%
V110K	1330.375		P112K	1626.973				
A112K	1427.546		A115K	2027.041				
D122K	1271.206		S125K	1843.345				
E123K	1195.854		E126K	2251.417				
S127K	1468.245		L128K	2674.771				
G128K	1827.766		Q129K	2299.484				
T129K	1705.873		A130K	1514.892				
R142K	1705.734		N131K	1353.591				
E143K	1458.029		G145K	1975.998				
D151K	3575.421	0.0%	A146K	2082.148				
N152K	1380.271		V147K	3431.846	0.0%			
A153K	1840.456		S155K	2149.37				
L154K	1795.011		S156K	2284.603				
Q155K	1422.783		P157K	2165.887				
S156K	2061.606		A160K	1493.712				
G157K	2648.713		G161K	1482.482				
E165K	1591.482		S171K	2192.136				
D167K	1523.247		N172K	2115.781				
S168K	1385.903		N173K	2544.968				
D170K	1215.408		T184K	2422.712				
S182K	1377.623		E186K	2421.28				
A184K	1471.194		Q187K	3157.169	23.6%			
E187K	1400.015		S190K	1886.288				
H189K	1452.142		H191K	2053.506				
V191K	1788.621		E201K	2055.553				
Q199K	1345.66		G202K	1965.403				
G200K	2479.481		S203K	1813.785				
L201K	4245.729	84.6%	P211K	1984.017				
S202K	10192.973	100.0%	T212K	1923.191				
S203K	1941.088		E213K	3378.443	81.8%			
P204K	1579.603		S215K	2159.499				
N210K	1883.766							
R211K	1638.391							
G212K	1775.378							
E213K	3505.908	54.1%						

FIGURA 8

Z-Gln-Gly-CAD-biotina: +631Da

	LC			Fd			Fc		
	Calculado	Observado	dMasa Porcentaje	Calculado	Observado	dMasa Porcentaje	Calculado	Observado	dMasa Porcentaje
T135K-CH1	23216	23216	0 100.0%	25099	25731	632 100.0%	25200	25202	2 70.4%
	23216		0.0%	25099	25731	632 100.0%	25363	25361	-2 29.6%
S136K-CH1	23216	23217	1 100.0%	25113	25113	0 16.2%	25200	25202	2 72.8%
	23216		0.0%	25113	25745	632 83.8%	25363	25362	-1 27.2%
D221K-hg	23216	23216	0 100.0%	25085	25086	1 16.1%	25200	25201	1 70.6%
	23216		0.0%	25085	25716	631 83.9%	25363	25361	-2 29.4%
T223K-hg	23216	23217	1 100.0%	25099	25100	1 57.4%	25200	25202	2 73.2%
	23216		0.0%	25099	25731	632 42.6%	25363	25361	-2 26.8%
H224K-hg	23216	23216	0 100.0%	25063	25064	1 43.6%	25200	25202	2 69.0%
	23216		0.0%	25063	25695	632 56.4%	25363	25362	-1 31.0%
T225K-hg	23216	23215	-1 100.0%	25099	25099	0 70.0%	25200	25200	0 72.2%
	23216		0.0%	25099	25729	630 30.0%	25363	25362	-1 27.8%
M252K-CH2	23216	23216	0 100.0%	25072	25071	-1 100.0%	23752	23753	1 1.7%
	23216		0.0%	25072		0.0%	23752	24384	632 98.3%
N297K-HC2	23216	23217	1 100.0%	25072	25071	-1 100.0%	23769		
	23216		0.0%	25072		0.0%	23769		
P445K-CH3	23216	23217	1 100.0%	25072	25071	-1 100.0%	23786	24418	632 100.0%
	23216		0.0%	25072		0.0%	23786	25201	1 64.2%
L201K-Ck	23231	23229	-2 18.0%	25072	25072	0 100.0%	25200	25201	1 35.8%
	23231	23864	633 82.0%	25072		0.0%	25363	25361	-2 72.2%
S202K-Ck	23257	23258	1 12.3%	25072	25071	-1 100.0%	25200	25201	1 72.2%
	23257	23890	633 87.7%	25072		0.0%	25363	25362	-1 27.8%

% de conjugación

T135K-CH1	100.0%
S136K-CH1	83.8%
D221K-hg	83.9%
T223K-hg	42.6%
H224K-hg	56.4%
T225K-hg	30.0%
M252K-CH2	98.3%
N297K-HC2	ND
P445K-CH3	100.0%
L201K-Ck	82.0%
S202K-Ck	87.7%

FIGURA 10

		X	X			
hu IgG1	GGPSVFLFPPKPKD	TLMSRTP	EVTCVVVDVS	HEDPEVQFNWYVDGVEVHNAKTKPREEQ	NSTYRVVSVLTVLHQDWLNGKEYCKVSNKALPADI	IEKT
hu IgG2	AGPSVFLFPPKPKD	TLMSRTP	EVTCVVVDVS	HEDPEVQFNWYVDGVEVHNAKTKPREEQ	NSTYRVVSVLTVLHQDWLNGKEYCKVSNKGLPADI	IEKT
hu IgG3	GGPSVFLFPPKPKD	TLMSRTP	EVTCVVVDVS	HEDPEVQFNWYVDGVEVHNAKTKPREEQ	NSTYRVVSVLTVLHQDWLNGKEYCKVSNKALPADI	IEKT
hu IgG4	GGPSVFLFPPKPKD	TLMSRTP	EVTCVVVDVS	QEDPEVQFNWYVDGVEVHNAKTKPREEQ	NSTYRVVSVLTVLHQDWLNGKEYCKVSNKGLPSSIEKT	
hu IgG1	ISKAKGQPREPQVY	TLPPSRDE	LITKNQVSLTCLVKGFYPSDIAVEWESN	GOPENNKKITPPVLDSDGSFFLYSKLTVDKSRWQQGNV	FSCSVMHEALHNH	
hu IgG2	ISKTKGQPREPQVY	TLPPSRDE	MTKNQVSLTCLVKGFYPSDIAVEWESN	GOPENNKKITPPMLDSDGSFFLYSKLTVDKSRWQQGNV	FSCSVMHEALHNH	
hu IgG3	ISKTKGQPREPQVY	TLPPSRDE	MTKNQVSLTCLVKGFYPSDIAVEWESS	GOPENNKKITPPMLDSDGSFFLYSKLTVDKSRWQQGNIF	FSCSVMHEALHNH	
hu IgG4	ISKAKGQPREPQVY	TLPPSQEEM	ITKNQVSLTCLVKGFYPSDIAVEWESN	GOPENNKKITPPVLDSDGSFFLYSKLTVDKSRWQQGNV	FSCSVMHEALHNH	
		X				
hu IgG1	YTOKSLSLSLSPGK					
hu IgG2	YTOKSLSLSLSPGK					
hu IgG3	FTOKSLSLSLSPGK					
hu IgG4	YTOKSLSLSLGLGK					

FIGURA 11

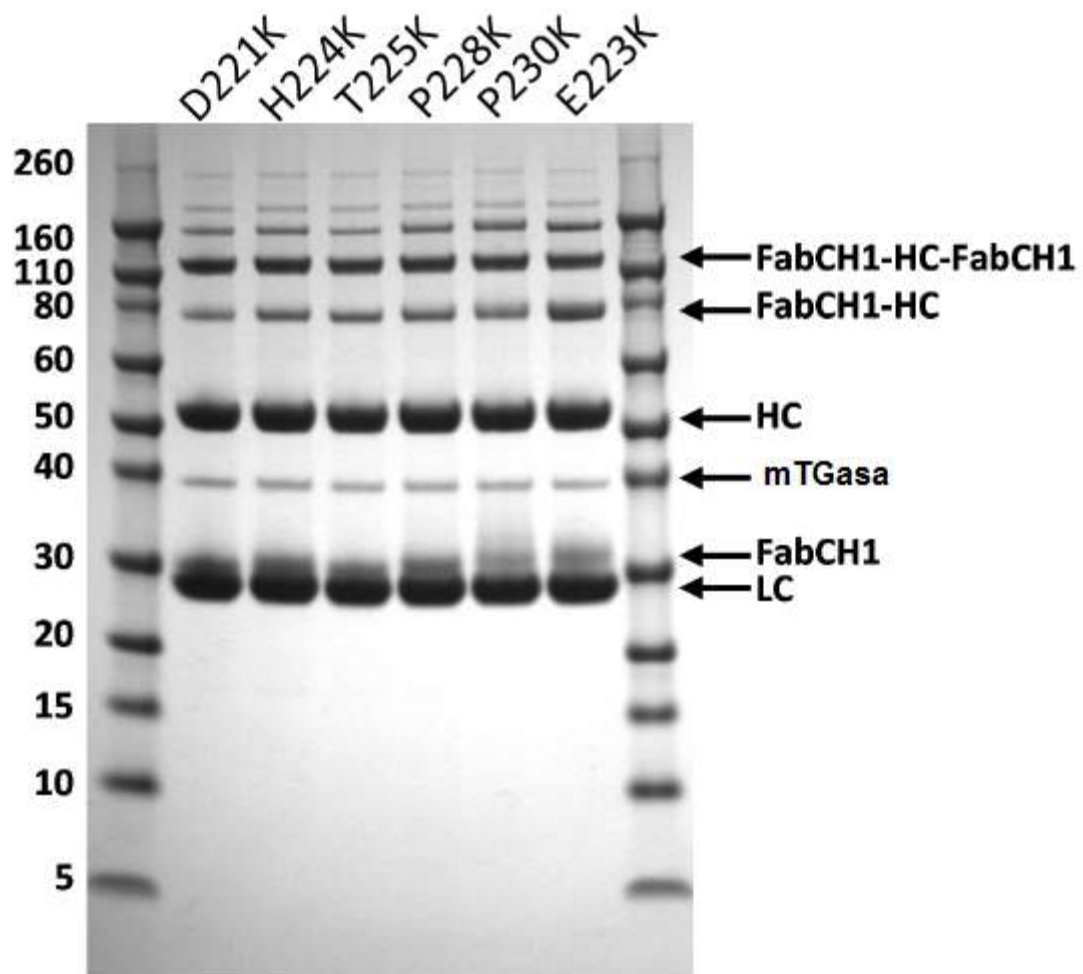


FIGURA 12

