

TABLE 17-continued

Form	Final pH	Final Concentration (mg/mL)	Final Osmolality (mOsm/Kg)
B-4	6.05	40.35	286
B-5	6.17	40.02	293
B-6	6.20	39.84	274
B-7	6.21	40.13	300
B-8	6.18	39.58	306
B-9	6.45	39.39	290
B-10	6.48	39.08	291
B-11	6.98	39.51	288
B-12	7.02	39.49	289
B-13	6.21	40.08	289
B-14	6.27	39.73	285
B-15	6.19	40.15	295

Size Exclusion Chromatography

[0392] The stability of the Block B formulations was evaluated by measuring the amount of non-degraded aflibercept present in the formulation at time 0 (t₀) and after being subjected to storage conditions. Size Exclusion Chromatography was conducted on the formulations and the percentage of protein present in the main peak (non-degraded aflibercept) in each formulation is reported in Table 18-21 below.

TABLE 18

SEC results for Block B formulations at T = 0			
Form	Rel. Area (%), Before Main Peak	Rel. Area (%), Main Peak	Rel. Area (%), After Main Peak
B-1	0.98	99.02	0
B-2	0.99	99.01	0
B-3	1.17	98.83	0
B-4	1.14	98.86	0
B-5	1.20	98.80	0
B-6	1.17	98.83	0
B-7	1.25	98.75	0
B-8	1.20	98.80	0
B-9	1.30	98.70	0
B-10	1.29	98.71	0
B-11	1.44	98.56	0
B-12	1.51	98.49	0
B-13	1.28	98.72	0
B-14	1.27	98.73	0
B-15	1.29	98.71	0

TABLE 19

SEC results for Block B formulations after one week at 40° C.			
Form	Rel. Area (%), Before Main Peak	Rel. Area (%), Main Peak	Rel. Area (%), After Main Peak
B-1	2.44	97.56	0
B-2	1.44	98.56	0
B-3	2.53	97.47	0
B-4	2.03	97.97	0
B-5	2.73	97.27	0
B-6	4.26	95.74	0
B-7	2.37	97.63	0
B-8	2.99	97.01	0
B-9	2.90	97.10	0
B-10	3.05	96.95	0

TABLE 19-continued

SEC results for Block B formulations after one week at 40° C.			
Form	Rel. Area (%), Before Main Peak	Rel. Area (%), Main Peak	Rel. Area (%), After Main Peak
B-11	3.14	96.86	0
B-12	3.35	96.65	0
B-13	2.67	97.33	0
B-14	2.72	97.28	0
B-15	2.79	97.21	0

TABLE 20

SEC results for Block B formulations two weeks at 25° C.			
Form	Rel. Area (%), Before Main Peak	Rel. Area (%), Main Peak	Rel. Area (%), After Main Peak
B-1	1.28	98.72	0
B-2	1.24	98.76	0
B-3	1.61	98.39	0
B-4	1.57	98.43	0
B-5	1.70	98.30	0
B-6	1.60	98.40	0
B-7	1.76	98.24	0
B-8	1.64	98.36	0
B-9	1.84	98.16	0
B-10	1.90	98.10	0
B-11	2.18	97.82	0
B-12	2.39	97.61	0
B-13	1.69	98.31	0
B-14	1.75	98.25	0
B-15	1.74	98.26	0

TABLE 21

SEC results for Block B formulations after four weeks at 5° C.			
Form	Rel. Area (%), Before Main Peak	Rel. Area (%), Main Peak	Rel. Area (%), After Main Peak
B-1	1.40	98.60	0.00
B-2	1.33	98.62	0.10
B-3	1.65	98.35	0.00
B-4	1.60	98.40	0.00
B-5	1.70	98.30	0.00
B-6	1.58	98.42	0.00
B-7	1.70	98.30	0.00
B-8	1.61	98.39	0.00
B-9	1.81	98.19	0.00
B-10	1.81	98.19	0.00
B-11	2.07	97.93	0.00
B-12	2.17	97.83	0.00
B-13	1.61	98.39	0.00
B-14	1.65	98.35	0.00
B-15	1.66	98.34	0.00

[0393] The initial monomer contents were 98-99%. After storage for one week at 40° C., monomer contents decreased to about 97%, with small differences being apparent. The losses were entirely due to aggregation and not due to fragmentation, again emphasizing the nature of the primary degradation pathway.

[0394] After two weeks at 25° C., the monomer contents were mostly near 98%. The two nominal pH 7 formulations