

Table 14.2.10.1
Summary of Pharmacodynamic Parameters - Bone Turnover
Safety Population

Parameter: Type I Collagen C-Telopeptides (pg/mL)

Visit	Statistic	Treatments				Total (N=48)
		Dose Group 1 0.25 mg/kg/day (N=12)	Dose Group 2 0.75 mg/kg/day (N=12)	Dose Group 3 2.0 mg/kg/day (N=12)	Dose Group 4 6.0 mg/kg/day (N=12)	
Baseline	n	11	10	12	12	45
	Mean	871.0	935.8	936.8	889.3	907.8
	SD	160.85	286.50	256.25	186.68	220.54
	Median	891.0	857.0	988.5	834.0	891.0
	Minimum	571	611	641	581	571
	Maximum	1143	1383	1477	1223	1477
	Interquartile Range	743.0, 941.0	711.0, 1228.0	703.0, 1055.5	775.5, 1017.5	743.0, 1016.0
Day 1	n	12	12	12	11	47
	Mean	974.8	940.8	838.3	786.8	887.3
	SD	252.77	227.60	233.30	331.68	265.30
	Median	930.0	915.5	853.0	746.0	833.0
	Minimum	647	642	479	365	365
	Maximum	1356	1390	1346	1533	1533
	Interquartile Range	777.5, 1239.5	785.5, 1030.0	695.0, 979.5	536.0, 812.0	703.0, 985.0

(1) Percentage change is calculated as $100 \times (\text{change from baseline} / \text{baseline})$.
Baseline is defined as the last response during Screening prior to Study Day -1. P-values are from paired t-tests testing statistical significance of the change from baseline.

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Day 1 Change from Baseline	n	11	10	12	11	44
	Mean	72.4	-12.9	-98.5	-115.7	-40.6
	SD	241.87	233.44	237.69	421.04	294.13
	Median	76.0	19.0	-120.0	-198.0	-14.5
	Minimum	-360	-557	-492	-644	-644
	Maximum	450	237	346	693	693
	Interquartile Range	-133.0, 235.0	-13.0, 168.0	-230.0, 21.5	-367.0, -16.0	-221.5, 108.5
	P-Value	0.3445	0.8651	0.1789	0.3834	0.3645
Day 1 Percent change from Baseline (1)	n	11	10	12	11	44
	Mean	9.9	2.7	-8.0	-6.7	-0.8
	SD	25.38	23.76	23.93	53.72	33.78
	Median	11.6	2.8	-14.4	-28.1	-1.6
	Minimum	-31	-45	-36	-64	-64
	Maximum	50	36	35	112	112
	Interquartile Range	-12.6, 25.3	-1.4, 22.9	-25.2, 3.4	-33.6, -1.9	-26.2, 15.4

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Week 2	n	12	12	12	12	48
	Mean	963.7	903.3	710.4	625.7	800.8
	SD	157.68	251.01	180.03	203.19	239.13
	Median	975.5	862.5	686.5	615.5	834.0
	Minimum	647	623	387	331	331
	Maximum	1225	1626	1023	933	1626
	Interquartile Range	878.0, 1070.0	777.5, 950.5	595.5, 811.0	456.5, 796.0	639.0, 955.5
Week 2 Change from Baseline	n	11	10	12	12	45
	Mean	85.0	-19.9	-226.4	-263.7	-114.3
	SD	185.90	266.99	185.99	229.69	256.40
	Median	147.0	76.0	-209.5	-299.0	-120.0
	Minimum	-300	-470	-650	-561	-650
	Maximum	334	250	18	166	334
	Interquartile Range	-72.0, 228.0	-245.0, 191.0	-338.5, -79.5	-442.0, -85.0	-333.0, 140.0
	P-Value	0.1604	0.8189	0.0014	0.0022	0.0045

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Week 2 Percent change from Baseline (1)	n	11	10	12	12	45
	Mean	11.9	2.2	-22.5	-27.7	-10.0
	SD	20.52	26.44	15.27	26.50	27.47
	Median	15.6	9.8	-19.1	-32.2	-13.7
	Minimum	-26	-36	-48	-60	-60
	Maximum	37	35	2	29	37
	Interquartile Range	-6.8, 30.7	-25.7, 23.8	-32.4, -12.2	-49.0, -10.3	-30.1, 15.6
Week 4	n	12	12	12	12	48
	Mean	915.9	983.5	949.8	989.2	959.6
	SD	263.13	298.47	303.76	216.29	265.48
	Median	900.0	944.5	881.5	928.0	909.5
	Minimum	564	512	667	737	512
	Maximum	1433	1450	1817	1351	1817
	Interquartile Range	688.5, 1090.5	791.0, 1256.5	771.5, 1007.0	808.5, 1174.0	784.5, 1102.5

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Week 4 Change from Baseline	n	11	10	12	12	45
	Mean	17.4	34.8	12.9	99.8	42.0
	SD	267.45	271.50	181.45	211.90	228.25
	Median	-7.0	-1.5	22.0	90.0	45.0
	Minimum	-453	-262	-271	-181	-453
	Maximum	527	691	340	644	691
	Interquartile Range	-186.0, 165.0	-198.0, 114.0	-121.0, 124.0	-12.5, 176.5	-120.0, 149.0
	P-Value	0.8338	0.6947	0.8098	0.1309	0.2231
Week 4 Percent change from Baseline (1)	n	11	10	12	12	45
	Mean	3.6	6.8	2.7	14.5	7.0
	SD	28.96	34.06	19.03	32.98	28.52
	Median	-1.2	-0.2	3.4	9.3	6.0
	Minimum	-43	-28	-27	-18	-43
	Maximum	58	91	37	111	111
	Interquartile Range	-20.0, 17.5	-19.6, 11.9	-11.7, 16.8	-1.3, 18.5	-11.1, 17.3

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