

Table 14.2.10.1
Summary of Pharmacodynamic Parameters - Bone Turnover
Safety Population

Parameter: Osteocalcin (ng/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	12	12	12	12	48
	Mean	37.94	35.66	41.17	44.36	39.78
	SD	11.622	6.800	5.617	5.979	8.320
	Median	32.15	34.80	41.95	42.80	39.55
	Minimum	25.4	25.6	33.5	37.0	25.4
	Maximum	59.1	47.6	53.5	55.2	59.1
	Interquartile Range	29.50, 49.30	30.55, 41.10	36.75, 43.65	39.30, 48.70	32.90, 45.15
Day 1	n	12	12	12	11	47
	Mean	39.37	35.89	34.93	33.52	35.98
	SD	14.189	7.526	6.881	9.135	9.792
	Median	34.40	36.75	36.45	32.50	35.60
	Minimum	21.8	22.9	25.5	23.4	21.8
	Maximum	60.0	53.0	46.6	45.6	60.0
	Interquartile Range	27.70, 54.95	31.30, 38.55	27.35, 39.75	24.30, 44.60	27.60, 40.40

(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	12	12	12	11	47
	Mean	1.43	0.23	-6.23	-11.37	-3.83
	SD	8.197	5.304	10.930	9.137	9.805
	Median	1.00	2.75	-2.65	-14.60	-2.20
	Minimum	-10.5	-7.5	-25.9	-22.7	-25.9
	Maximum	20.0	6.0	13.1	4.2	20.0
	Interquartile Range	-2.50, 5.90	-5.45, 4.70	-15.85, -0.75	-16.20, -2.20	-11.20, 3.40
	P-Value	0.5592	0.8816	0.0738	0.0020	0.0102
Day 1 Percent change from Baseline (1)	n	12	12	12	11	47
	Mean	3.95	1.41	-12.63	-25.05	-7.72
	SD	22.015	15.449	25.020	20.228	23.388
	Median	2.38	7.86	-6.20	-34.39	-5.28
	Minimum	-32.5	-23.9	-48.4	-45.6	-48.4
	Maximum	50.8	19.4	39.1	10.1	50.8
	Interquartile Range	-8.11, 14.87	-13.41, 14.21	-37.43, -2.07	-40.81, -4.61	-30.32, 9.94

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Week 2	n	12	12	12	12	48
	Mean	38.53	35.10	37.51	29.04	35.04
	SD	12.025	8.238	8.624	4.853	9.300
	Median	34.70	33.55	39.30	28.95	33.10
	Minimum	24.1	24.7	19.8	21.9	19.8
	Maximum	61.6	50.4	48.9	36.7	61.6
	Interquartile Range	29.45, 49.40	29.30, 40.10	30.50, 43.80	24.90, 33.10	28.00, 40.85
Week 2 Change from Baseline	n	12	12	12	12	48
	Mean	0.58	-0.56	-3.66	-15.32	-4.74
	SD	5.986	7.934	9.818	6.450	9.797
	Median	0.70	-0.50	-2.40	-17.55	-3.50
	Minimum	-8.5	-16.7	-24.9	-23.7	-24.9
	Maximum	13.0	15.8	13.6	-4.2	15.8
	Interquartile Range	-2.95, 2.30	-4.15, 3.85	-8.00, 1.70	-19.65, -10.30	-10.45, 1.00
	P-Value	0.7421	0.8119	0.2232	<.0001	0.0016

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Week 2 Percent change from Baseline (1)	n	12	12	12	12	48
	Mean	2.48	-0.07	-7.81	-33.87	-9.82
	SD	16.756	22.408	23.644	12.548	23.717
	Median	2.02	-1.25	-5.69	-39.67	-8.08
	Minimum	-20.0	-37.3	-46.5	-45.1	-46.5
	Maximum	34.1	49.8	40.6	-10.9	49.8
	Interquartile Range	-6.93, 5.37	-13.10, 10.84	-20.63, 4.44	-43.32, -25.31	-25.51, 3.26
Week 4	n	12	12	12	12	48
	Mean	39.20	42.24	47.91	42.81	43.04
	SD	14.136	8.426	6.648	10.851	10.549
	Median	33.80	41.45	48.30	38.60	41.80
	Minimum	25.0	25.1	38.7	26.4	25.0
	Maximum	66.7	60.1	58.5	58.5	66.7
	Interquartile Range	27.70, 49.90	38.90, 44.40	42.20, 52.55	34.60, 54.20	36.25, 51.90

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Week 4 Change from Baseline	n	12	12	12	12	48
	Mean	1.26	6.58	6.74	-1.55	3.26
	SD	5.650	9.341	9.747	11.176	9.602
	Median	0.30	4.40	8.10	-1.80	2.05
	Minimum	-4.9	-4.5	-13.7	-19.2	-19.2
	Maximum	14.1	28.4	25.0	17.2	28.4
	Interquartile Range	-3.70, 5.45	0.20, 10.90	0.75, 10.90	-9.00, 4.35	-2.50, 8.60
	P-Value	0.4567	0.0327	0.0355	0.6403	0.0230
Week 4 Percent change from Baseline (1)	n	12	12	12	12	48
	Mean	2.72	20.91	18.74	-2.43	9.98
	SD	14.063	28.644	25.095	25.932	25.416
	Median	1.04	12.32	19.09	-4.59	4.83
	Minimum	-15.3	-9.5	-25.6	-42.1	-42.1
	Maximum	26.8	89.6	74.6	41.6	89.6
	Interquartile Range	-10.76, 14.41	0.03, 35.74	1.61, 27.24	-19.70, 9.68	-6.10, 24.53

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