

Supplemental Table 1. Association between long-term and recent weight change and HR-pQCT indices at the tibia and radius, adjusted for age, sex, height, current smoking, diabetes and initial weight (i.e. initial weight = weight at enrollment for long-term weight change; and initial weight = weight at preceding clinic visit for recent weight change). Beta coefficients are presented for weight change treated as a continuous variable, and least square means are presented for weight change group.

	Weight Change ^a											
	Long-term weight change						Recent weight change					
	Continuous		Categorical Measure				Continuous		Categorical Measure			
	Beta	P-value ^b	≥5% loss (N=166)	<5% change (N=275)	≥5% gain (N=920)	P-value ^b	Beta	P-value ^b	≥5% loss (N=301)	<5% change (N=834)	≥5% gain (N=226)	P-value ^b
HR-pQCT indices^c												
Tibia (N=1328)^d			Mean ^e					Mean ^e				
Ct.BMD	0.7246	<.0001	832.2	838.6	858.2	<.0001	1.2199	<.0001	836.8	852.8	858.5	0.0005
Ct.Po	-0.0115	0.0356	10.312	10.629	10.086	0.4490	-0.0365	0.0046	10.666	10.251	9.936	0.0143
Ct.Th	0.0033	<.0001	1.100	1.158	1.228	<.0001	0.0042	<.0001	1.159	1.201	1.232	0.0009
Tb.BMD	0.2920	<.0001	162.0	171.1	174.9	<.0001	0.2151	0.1297	170.4	173.6	175.0	0.1562
Tb.N	0.0060	<.0001	1.880	1.980	2.088	<.0001	0.0074	<.0001	1.978	2.054	2.119	<.0001
Tb.Th	-0.0001	0.0009	0.072	0.072	0.070	0.1036	-0.0001	0.0039	0.072	0.071	0.069	0.0121
Tt.Ar	0.1562	0.3160	776.5	770.1	769.6	0.4229	-0.0487	0.8965	775.8	777.2	768.3	0.3833
Tt.BMD	0.7033	<.0001	259.8	275.3	288.2	<.0001	0.8319	<.0001	273.3	283.1	289.5	0.0005
FL	12.5653	<.0001	5980	6274	6496	<.0001	14.6088	0.0001	6267	6467	6478	0.0159
Radius (N=1263)^d												
Ct.BMD	0.2150	0.0506	943.4	944.9	946.0	0.6808	0.4581	0.0798	941.3	946.1	946.0	0.4410
Ct.Po	-0.0004	0.8773	3.894	4.078	4.001	0.5066	-0.0009	0.8891	4.011	4.050	3.983	0.8556
Ct.Th	0.0019	<.0001	0.819	0.844	0.882	0.0005	0.0031	<.0001	0.835	0.873	0.891	0.0016
Tb.BMD	0.3292	<.0001	154.8	159.0	166.3	0.0015	0.4309	0.0040	159.9	163.2	168.7	0.0113
Tb.N	0.0041	<.0001	1.948	1.991	2.109	<.0001	0.0046	0.0007	2.011	2.080	2.095	0.0075
Tb.Th	0.0000	0.7852	0.066	0.066	0.066	0.6826	0.0000	0.9939	0.066	0.065	0.066	0.7935
Tt.Ar	0.0214	0.7826	307.4	312.1	312.2	0.2757	0.0317	0.8647	314.0	313.0	310.3	0.3903
Tt.BMD	0.6626	<.0001	294.3	301.9	315.1	0.0005	0.9162	0.0002	300.8	310.3	319.0	0.0016
FL	3.5057	<.0001	2407	2512	2572	0.0002	5.0189	0.0065	2492	2560	2584	0.0298

^a Long-term (40 year) weight change was determined between the index examination (2011-2014) and enrollment (1971-1974) and expressed as a percentage of the weight at the

^b Bold indicates p-values <0.05

^c Ct.BMD=cortical volumetric bone mineral density, Ct.Po=cortical porosity, Ct.Th=cortical thickness, Tb.BMD=trabecular volumetric bone mineral density, Tb.N=trabecular

^d The sample includes 1,361 individuals with HR-pQCT measures at the tibia (N=1,328) and/or radius (N=1,263).

^e Adjusted for age, sex, height, current smoking, and diabetes and initial weight

Supplemental Table 2. Association between long-term and recent weight change and HR-pQCT indices at the tibia and radius, adjusted for age, sex, height, current smoking, diabetes. Similar to the analysis shown in Table 3 but **here the covariates information were based on initial/baseline examination** (i.e. baseline = time at enrollment for long-term weight change; and baseline = time at preceding clinic visit for recent weight change). Beta coefficients are presented for weight change treated as a continuous variable, and least square means are presented for weight change group.

	<u>Weight Change^a</u>											
	<u>Long-term weight change</u>						<u>Recent weight change</u>					
	<u>Continuous</u>		<u>Catogorical Measure</u>				<u>Continuous</u>		<u>Catogorical Measure</u>			
HR--pQCT indices ^c	Beta	P-value ^b	≥5% loss (N=166)	<5% change (N=275)	≥5% gain (N=920)	P-value ^b	Beta	P-value ^b	≥5% loss (N=301)	<5% change (N=834)	≥5% gain (N=226)	P-value ^b
<u>Tibia</u> (N=1328) ^d			Mean ^e						Mean ^e			
Ct.BMD	0.7986	<.0001	780.6	786.7	807.5	<.0001	1.2011	<.0001	834.3	849.2	854.8	0.0012
Ct.Po	-0.0146	0.0111	12.790	13.096	12.495	0.3373	-0.0398	0.0023	10.855	10.446	10.102	0.0124
Ct.Th	0.0035	<.0001	1.088	1.138	1.210	<.0001	0.0039	<.0001	1.152	1.189	1.219	0.0025
Tb.BMD	0.3441	<.0001	136.9	147.0	151.8	<.0001	0.2146	0.1356	171.4	174.4	176.4	0.1285
Tb.N	0.0063	<.0001	1.680	1.780	1.896	<.0001	0.0073	<.0001	2.006	2.081	2.147	<.0001
Tb.Th	-0.0001	0.0070	0.069	0.069	0.067	0.2535	-0.0001	0.0040	0.072	0.070	0.069	0.0157
Tt.Ar	0.0361	0.8258	753.9	751.1	752.4	0.8647	0.2316	0.5354	773.0	778.2	771.3	0.8416
Tt.BMD	0.7859	<.0001	230.3	245.6	259.6	<.0001	0.7855	0.0001	273.4	282.2	288.8	0.0009
FL	13.2809	<.0001	5092	5406	5668	<.0001	15.1860	<.0001	6244	6441	6473	0.0098
<u>Radius</u> (N=1263) ^d												
Ct.BMD	0.2925	0.0111	918.3	920.4	922.3	0.5291	0.4759	0.0720	937.8	942.3	942.5	0.4401
Ct.Po	-0.0024	0.4210	4.350	4.505	4.408	0.7244	-0.0021	0.7642	4.080	4.122	4.038	0.7895
Ct.Th	0.0020	<.0001	0.721	0.743	0.779	0.0021	0.0031	<.0001	0.833	0.870	0.887	0.0023
Tb.BMD	0.3299	<.0001	135.3	138.8	146.1	0.0035	0.4019	0.0081	161.0	163.9	169.4	0.0163
Tb.N	0.0044	<.0001	1.649	1.689	1.810	<.0001	0.0043	0.0021	2.031	2.099	2.110	0.0138
Tb.Th	0.0000	0.4812	0.069	0.069	0.069	0.5455	0.0000	0.9591	0.066	0.065	0.066	0.7703
Tt.Ar	-0.0047	0.9537	332.3	338.7	340.3	0.0783	0.0854	0.6496	311.7	311.0	309.2	0.5544
Tt.BMD	0.6858	<.0001	257.1	262.8	275.5	0.0026	0.8693	0.0006	301.6	310.8	318.9	0.0030
FL	3.3483	<.0001	2112	2217	2289	<.0001	5.1057	0.0063	2491	2556	2585	0.0279

^a Long-term (40 year) weight change was determined between the index examination (2011-2014) and enrollment (1971-1974) and expressed as a percentage of the weight at the

^b Bold indicates p-values <0.05

^c Ct.BMD=cortical volumetric bone mineral density, Ct.Po=cortical porosity, Ct.Th=cortical thickness, Tb.BMD=trabecular volumetric bone mineral density, Tb.N=trabecular

^d The sample includes 1,361 individuals with HR-pQCT measures at the tibia (N=1,328) and/or radius (N=1,263).

^e Adjusted for age, sex, height, current smoking, and diabetes, **assessed at initial/baseline examination**

Supplemental Table 3. Association between long-term weight change and HR-pQCT indices at the tibia and radius, adjusted for age, sex, height, current smoking, diabetes and the standard deviation of weight measurements over time. Beta coefficients are presented for weight change treated as a continuous variable, and least square means are presented for weight change group.

HR--pQCT indices ^c	Long-term weight change ^a					
	<u>Continuous</u>		<u>Catogorical Measure</u>			
	Beta	P-value ^b	≥5% loss (N=166)	<5% change (N=275)	≥5% gain (N=920)	P-value ^b
<u>Tibia</u> (N=1328)^d			Mean^e			
Ct.BMD	0.9493	<.0001	833.3	839.1	858.2	<.0001
Ct.Po	-0.0277	<.0001	10.577	10.820	10.046	0.0681
Ct.Th	0.0033	<.0001	1.126	1.176	1.225	<.0001
Tb.BMD	0.1136	0.1289	168.2	175.3	174.0	0.0750
Tb.N	0.0019	0.0079	1.979	2.066	2.065	0.0056
Tb.Th	0.0000	0.5915	0.071	0.071	0.071	0.4780
Tt.Ar	-1.2403	<.0001	801.3	789.3	764.7	<.0001
Tt.BMD	0.6415	<.0001	266.6	280.0	287.2	<.0001
FL	4.5794	0.0264	6260	6464	6460	0.0260
<u>Radius</u> (N=1263)^d						
Ct.BMD	0.5197	0.0001	940.8	942.4	946.8	0.3321
Ct.Po	-0.0065	0.0709	4.018	4.167	3.979	0.8095
Ct.Th	0.0017	<.0001	0.836	0.859	0.878	0.0202
Tb.BMD	0.2126	0.0068	158.4	162.3	165.3	0.0503
Tb.N	0.0027	0.0002	1.986	2.029	2.097	0.0007
Tb.Th	0.0000	0.6751	0.066	0.066	0.066	0.5447
Tt.Ar	-0.2103	0.0306	313.7	315.9	311.6	0.6344
Tt.BMD	0.5664	<.0001	299.0	306.7	313.5	0.0133
FL	1.5427	0.1224	2481	2564	2559	0.0770

^a Long-term (40 year) weight change was determined between the index examination (2011-2014) and enrollment (1971-1974) and expressed as a percentage of the weight at the index examination. Recent (6 year) weight change was determined between the index examination and the preceding clinic visit (2005-2008) and expressed as a percentage of the weight at the preceding clinic visit.

^b Bold indicates p-values <0.05

^c Ct.BMD=cortical volumetric bone mineral density, Ct.Po=cortical porosity, Ct.Th=cortical thickness, Tb.BMD=trabecular volumetric bone mineral density, Tb.N=trabecular number, Tt.BMD= total (integral) volumetric bone mineral density, Tt.CSA=total cross-sectional area (Tt.CSA, mm²), FEA=Micro finite element analysis, and FL=failure load

^d The sample includes 1,361 individuals with HR-pQCT measures at the tibia (N=1,328) and/or radius (N=1,263).

^e Adjusted for age, sex, height, current smoking, and diabetes and **the standard deviation of weight measurements over time**

Supplemental Table 4. Association between long-term and recent weight change and HR-pQCT indices at the tibia and radius in men, adjusted for age, sex, height, current smoking, and diabetes. Beta coefficients are presented for weight change treated as a continuous variable, and least square means are presented for weight change group

Weight Change^a												
HR-pQCT indices ^c	Long-term weight change						Recent weight change					
	Continuous		Categorical Measure				Continuous		Categorical Measure			
	Beta	P-value ^b	≥5% loss (N=89)	<5% change (N=149)	≥5% gain (N=356)	P-value ^b	Beta	P-value ^b	≥5% loss (N=122)	<5% change (N=389)	≥5% gain (N=83)	P-value ^b
<u>Tibia</u> (N=580) ^d			Mean ^e						Mean ^e			
Ct.BMD	0.3765	0.0729	854.6	865.4	875.4	0.014	0.8285	0.0670	857.9	873.7	870.8	0.186
Ct.Po	-0.0100	0.3145	10.491	10.693	9.990	0.212	-0.0202	0.3478	10.450	10.127	10.264	0.692
Ct.Th	0.0019	0.0177	1.262	1.356	1.358	0.003	0.0037	0.0324	1.308	1.344	1.367	0.115
Tb.BMD	-0.1497	0.1815	186.9	189.9	186.5	0.916	-0.3206	0.1841	190.1	186.2	186.9	0.535
Tb.N	0.0022	0.0306	2.176	2.235	2.240	0.118	0.0013	0.5390	2.226	2.210	2.283	0.231
Tb.Th	-0.0001	0.0008	0.072	0.071	0.070	0.092	-0.0002	0.0363	0.072	0.071	0.068	0.052
Tt.Ar	-0.9320	0.0071	929.5	908.1	887.2	0.002	-1.4386	0.0544	912.5	902.8	872.9	0.014
Tt.BMD	0.1915	0.2369	291.5	305.1	306.1	0.025	0.2073	0.5525	300.6	302.8	307.3	0.375
FL	-6.6814	0.0678	7779	7914	7774	0.971	-5.3583	0.4961	7810	7849	7654	0.364
<u>Radius</u> (N=546) ^d												
Ct.BMD	0.0457	0.8030	942.5	938.6	946.1	0.635	-0.0861	0.8253	941.3	948.6	934.0	0.395
Ct.Po	-0.0041	0.4430	4.592	4.684	4.321	0.227	0.0062	0.5842	4.484	4.316	4.737	0.315
Ct.Th	0.0007	0.2988	0.946	0.974	0.982	0.183	0.0031	0.0204	0.939	0.975	1.008	0.021
Tb.BMD	-0.0166	0.8843	182.6	184.0	184.5	0.691	0.0253	0.9168	183.9	182.3	189.1	0.330
Tb.N	0.0003	0.6829	2.186	2.223	2.240	0.125	-0.0010	0.5648	2.246	2.223	2.213	0.403
Tb.Th	0.0000	0.3862	0.070	0.069	0.069	0.357	0.0000	0.9245	0.069	0.069	0.071	0.149
Tt.Ar	-0.2383	0.1681	373.8	380.8	374.6	0.913	-0.4417	0.2301	385.9	375.0	366.6	0.019
Tt.BMD	0.1332	0.4874	326.0	330.2	334.2	0.306	0.5440	0.1825	323.2	332.0	341.2	0.047
FL	-1.8066	0.3094	3145	3251	3202	0.443	0.9008	0.8111	3166	3210	3221	0.511

^a Long-term (40 year) weight change was determined between the index examination (2011-2014) and enrollment (1971-1974) and expressed as a percentage of the weight at the index examination. Recent (6 year) weight change was determined between the index examination and the preceding clinic visit (2005-2008) and expressed as a percentage of the weight at the preceding clinic visit.

^b Bold indicates p-values <0.05

^c Ct.BMD=cortical volumetric bone mineral density, Ct.Po=cortical porosity, Ct.Th=cortical thickness, Tb.BMD=trabecular volumetric bone mineral density, Tb.N=trabecular number, Tt.BMD= total (integral) volumetric bone mineral density, Tt.CSA=total cross-sectional area (Tt.CSA, mm²), FEA=Micro finite element analysis, and FL=failure load

^d The male sample includes individuals with HR-pQCT measures at the tibia (N=580) and/or radius (N=546).

^e Adjusted for age, sex, height, current smoking, and diabetes

Supplemental Table 5. Association between long-term and recent weight change and HR-pQCT indices at the tibia and radius in women, adjusted for age, sex, height, current smoking, and diabetes. Beta coefficients are presented for weight change treated as a continuous variable, and least square means are presented for weight change group

Weight Change ^a												
HR-pQCT indices ^c	Long-term weight change						Recent weight change					
	Continuous		Categorical Measure				Continuous		Categorical Measure			
	Beta	P-value ^b	≥5% loss (N=77)	<5% change (N=126)	≥5% gain (N=564)	P-value ^b	Beta	P-value ^b	≥5% loss (N=179)	<5% change (N=445)	≥5% gain (N=143)	P-value ^b
Tibia (N=748)^d			Mean ^e					Mean ^e				
Ct.BMD	0.7929	<.0001	809.7	809.6	838.9	0.001	1.1128	0.0007	818.2	830.9	838.1	0.013
Ct.Po	-0.0193	0.0020	10.782	10.915	10.270	0.217	-0.0501	0.0014	10.996	10.472	9.830	0.002
Ct.Th	0.0030	<.0001	0.979	0.967	1.087	0.000	0.0023	0.0377	1.028	1.050	1.061	0.220
Tb.BMD	0.2914	<.0001	150.7	158.3	164.0	0.004	0.1777	0.3147	157.9	163.2	161.6	0.381
Tb.N	0.0046	<.0001	1.800	1.834	1.959	0.001	0.0050	0.0060	1.853	1.936	1.939	0.050
Tb.Th	0.0000	0.2713	0.070	0.072	0.070	0.962	-0.0001	0.2289	0.071	0.071	0.070	0.372
Tt.Ar	-0.1128	0.4812	692.8	674.2	671.6	0.046	-0.2521	0.5298	676.6	675.0	670.9	0.560
Tt.BMD	0.6909	<.0001	240.1	248.4	268.2	<.0001	0.6629	0.0071	252.8	263.1	264.9	0.043
FL	11.8677	<.0001	4856	4976	5307	<.0001	11.4711	0.0056	5056	5231	5235	0.074
Radius (N=717)^d												
Ct.BMD	0.3088	0.0201	937.2	947.2	945.0	0.404	0.7634	0.0246	940.2	943.9	951.8	0.155
Ct.Po	-0.0030	0.3527	3.480	3.697	3.747	0.238	-0.0094	0.2506	3.701	3.860	3.417	0.142
Ct.Th	0.0018	<.0001	0.727	0.733	0.784	0.021	0.0019	0.0369	0.748	0.775	0.770	0.314
Tb.BMD	0.3538	<.0001	134.3	137.6	147.9	0.009	0.4182	0.0288	140.4	144.9	147.1	0.142
Tb.N	0.0043	<.0001	1.795	1.813	1.990	0.000	0.0051	0.0097	1.849	1.961	1.958	0.021
Tb.Th	0.0000	0.9116	0.062	0.063	0.062	0.949	0.0000	0.8585	0.063	0.062	0.062	0.392
Tt.Ar	-0.0452	0.5474	262.1	256.7	257.3	0.361	0.0644	0.7372	255.4	259.2	257.5	0.652
Tt.BMD	0.6934	<.0001	269.9	276.1	293.8	0.005	0.7280	0.0196	281.5	288.1	292.0	0.162
FL	3.3349	<.0001	1863	1886	1974	0.029	3.5772	0.0533	1909	1956	1952	0.330

^a Long-term (40 year) weight change was determined between the index examination (2011-2014) and enrollment (1971-1974) and expressed as a percentage of the weight at the index examination. Recent (6 year) weight change was determined between the index examination and the preceding clinic visit (2005-2008) and expressed as a percentage of the weight at the preceding clinic visit.

^b Bold indicates p-values <0.05

^c Ct.BMD=cortical volumetric bone mineral density, Ct.Po=cortical porosity, Ct.Th=cortical thickness, Tb.BMD=trabecular volumetric bone mineral density, Tb.N=trabecular number, Tt.BMD= total (integral) volumetric bone mineral density, Tt.CSA=total cross-sectional area (Tt.CSA, mm²), FEA=Micro finite element analysis, and FL=failure load

^d The female sample includes individuals with HR-pQCT measures at the tibia (N=748) and/or radius (N=717).

^e Adjusted for age, sex, height, current smoking, and diabetes