The Impact of Body Weight Change on Bone Mineral Density of the Lumbar Spine in Perimenopausal Women: A Retrospective, One-year Follow-up Study

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Background: Low body weight was one of the risk factors of osteoporosis. Little is known about the correlation between body weight change and bone mineral density (BMD) in Korean women. Therefore, this study was designed to reveal the impact of body weight change on BMD of the lumbar spine in perimenopausal women.

Methods: 105 healthy perimenopausal women aged between 44 and 50 years old were enrolled from August 2002 to March 2009. BMD was measured by dual energy X-ray absorptiometry. Partial correlation coefficients between body weight change and BMD change were calculated after the adjustments for several variables. BMD changes among groups based on BMI and the percentage change in body weight during 1-year follow-up period were compared.

Results: At both baseline and year 1, BMD of lumbar spine tended to be associated more with body weight. There was a significant association between body weight change and BMD change in lumbar spine during 1-year follow-up period. The weight gain group relatively showed an increase in BMD of lumbar spines than weight loss group. There was no BMD change in BMI less than 23 kg/m² group, but in case of BMI more than 23 kg/m² group, BMD in weight gain group increased more than the weight maintaining group.

Conclusion: This study demonstrated that body weight change is associated with change in BMD of lumbar spine in perimenopausal women especially if they are overweight.

Keywords: Perimenopause; Body Weight; Bone Density; Women

INTRODUCTION

Menopausal transition is a dynamical hormone changing period that occurs because of a decrease in the function of the ovaries. Increased level of follicular stimulating hormone and irregular patterns of menstruation are common and remarkable characteristics of menopausal transition. On average, women in the middle of menopausal transition gained almost 0.5 kg per year. This could affect not only high level of cholesterol but also body metabolic system, like basal metabolic rate, so they are recommended to maintain their weight for normal-weight
perimenopausal women and to lose their weight to decrease the risk of cardiovascular diseases for overweight or obese perimenopausal women.\textsuperscript{3,4)}

Losing body weight might help control blood lipid profile but low body weight could affect bone mineral density because low body weight itself is one of known risk factors of osteoporosis.\textsuperscript{5,7)} A large sectional study has shown that there was a positive correlation between body weight and bone mineral density (BMD) and especially weight barring part of body, lumbar spine and femur neck tended to be associated more with body weight than no weight barring part like radius.\textsuperscript{8)} According to some of domestic studies, all three groups - male adults (average age of 49.6), women over 50 years old and young females (average age of 34) - had a correlation between body weight and lumbar spine \textsuperscript{(r = 0.295, r = 0.24–0.29, and r = 0.222).\textsuperscript{9,11}} A prospective study followed up women in menopausal transition for 2 years and inferred its relation between body weight change rate and BMD change rate on a graph. The diagram proposed that about 3% of weight gain per 2 years was need in order not to lose bone mineral density, and even though body weight was maintained, it also caused 1% loss of bone mineral density every 2 year.\textsuperscript{12)}

Premenopausal women with osteopenia tend to have faster process of losing bone mineral density and diagnosed with osteoporosis more after than premenopausal women with normal BMD.\textsuperscript{13)} If premenopausal women with osteopenia are treated early enough, that may help not to proceed its process to reach osteoporosis.\textsuperscript{14)} Therefore, when life style modification like losing body weight is planned for a premenopausal woman to reduce cardiovascular risk, it's correlation with BMD must be considered first. However, little is known about the correlation between body weight change and BMD for Korean women. Therefore, this study was designed to reveal the impact of body weight change on BMD of the lumbar spine in perimenopausal women.

**METHODS**

1. Study Subjects

Healthy premenopausal women aged between 44 and 50 years old who visited University Hospital in Busan and had annual health check-ups including BMD at least twice were enrolled from August 2002 to March 2009. Total 105 participants were eligible to participate if they met all of the following criteria: 1) did not suffer from osteoporosis and diseases which could cause secondary osteoporosis, like anorexia nervosa, chronic malabsorption, hyperparathyroidism, hyperthyroidism, hypogonadism, prolactinoma, Cushing syndrome, osteogenesis imperfecta, rheumatoid arthritis, chronic obstructive pulmonary disease, chronic neuropathy, chronic renal failure, diabetes mellitus, acromegaly, alchollic, chronic liver disease, collagenous disease and metabolic disease; 2) had no history of transplanted organs, hysterectomy with or without ovariotomy, fracture or taking medicine affecting BMD such as hormonal therapy, oral contraceptive pill, steroid, anti conversant drugs, thyroid hormone or anti coagulant drugs and 3) had no period for at least 2 months and were diagnosed with menopausal transition.\textsuperscript{15)}

2. Measures

Each participant also underwent a complete physical examination including anthropometric measurements. Height and body weight were measured using a digital scale, with the examinee wearing a light gown. Body mass index (BMI) was calculated as follows: BMI = weight (kg)/height squared (m$^2$). Resting blood pressure was measured using an automatic sphygmomanometer (BP203RV-II; Nippon Colin, Komaki, Japan) after a minimum of 10 minutes at rest in sitting position at 8 am to 10 am. Study subjects refrained from smoking or ingesting caffeine for 30 minutes prior to measurement. Blood samples were collected from the antecubital vein to determine serum concentrations of alkaline phosphatase (ALP) after overnight fasting. The serum biochemical test was conducted by Hitachi autoanalyzer model 7600-110 (Hitachi Co., Tokyo, Japan) and commercially available kits. Total body fat percentage was determined using a bioelectrical impedance analyzer (Inbody 3.0; Biospace Co., Seoul, Korea). The examinees were asked not to take any food and any water from 12 am to 2 pm, respectively, before the measurements. BMD of L1–4th lumbar spine was measured by using dual energy X-ray absorptiometry (DEXA; Lunar Prodigy, GE Medical System, Waukesha, WI, USA).

Before the health check-up began, all information like past histories, exercise state, current drinking state and smoking state were collected by self-reports. In case of alcohol state, it was divided into 3 groups, no alcohol intake group, less than 15 g a
RESULTS

1. Basal Characteristics of the Study Subject
Mean age of 105 people aged between 44 and 50 was 45.8 ± 1.7 years old. Average BMI was 23.4 ± 2.6 kg/m². Mean systolic and diastolic blood pressure was 117.7 ± 18.0 mm Hg and 72.0 ± 10.9 mm Hg. Mean daily calcium consumption was 917.2 ± 243.6 mg (range, 431.5 to 1,911.3 mg) which were more than the standard daily calcium consumption recommendation by the Korean Nutrition Society, 700 mg, in 2005. Only 8.6% of them smoked and 23.8% and 3.8% of them were modest and heavy drinkers, relatively (Table 1).

3. Statistical Analyses
Statistical analyses were performed using the SPSS ver. 10.0 (SPSS Inc., Chicago, IL, USA). Changes in BMI, total body fat percent, fat free mass, waist circumference, ALP and BMD of the lumbar spine during 1 year follow-up period were determined by paired t-test. Partial correlation coefficients between body weight change and BMD change were calculated after adjustment for age and changes in calcium intake, exercise, alcohol drinking and smoking status. One-way analysis of variance (ANOVA) and the Scheffe’s post hoc test for pairwise comparisons were used to compare BMD changes among groups based on the percentage change in body weight during 1-year follow-up period; weight maintaining group (within 3% change of initial weight), weight gain group (the increase of initial weight by 3% or more) and weight lost group (the reduction of initial weight by 3% or more).16) Then, we compared two group divided on the basis of initial BMI ≥ 23 kg/m² vs. BMI < 23 kg/m².
2. Association between Anthropometric Data and Lumbar BMD

There was no significant difference between weight, BMI and waist circumference, while significant differences between percent body fat, ALP, fat free mass and 2nd lumbar spine BMD during 1-year follow-up period (Table 2). After adjustment for age, calcium intake, exercise status, smoking status, and alcohol drinking, partial correlation coefficients of between 2nd lumbar spine and average BMD change and weight change were 0.32 (P < 0.01) and 0.28 (P < 0.01) relatively, but there was no significant correlation between weight change and ALP change (Table 3).

3. Differences in the Mean BMD Change between Groups

One-way ANOVA showed that there was at least one significant difference in the mean BMD change between groups: weight maintaining, weight gain, and weight lost group. According to post-hoc test, the weight gain group relatively showed an increase in BMD of lumbar spine than weight lost group (Table 4). There was no BMD change in BMI less than 23 kg/m² group, but in case of BMI more than 23 kg/m² group, BMD in weight gain group was increased more than the weight maintaining group.

DISCUSSION

The aim of this study was to evaluate the impact of body weight change on BMD of the lumbar spine in apparently healthy perimenopausal women during the 1-year follow-up period.
Their BMD change by their weight change was retrospectively monitored to give general and clinical information for adequate weight control among perimenopausal women to prevent BMD loss. Body weight is well-known as a very important determinant of BMD.\(^{17}\) Weight baring exercises place continuously more stress on the skeleton and then protect the skeleton from external forces or impacts. In addition, some estrogens from fat tissue prevent bone atrophy from bone.\(^{18}\) Low body weight is associated with an increased risk of low bone mass, while obesity is considered as a protective factor for osteoporosis in postmenopausal women.\(^{19-22}\)

Most previous studies about the relationship between body weight and BMD for Korean women have not used DEXA measurement (g/cm\(^2\)) in their statistical analyses and have not showed correlation coefficients in their studies. This study accumulated DEXA measurement (g/cm\(^2\)) for 1 year and used the measures to analyze its correlation between anthropometric parameters and BMD. These results revealed a significant correlation between the 2nd lumbar spine BMD change, the mean lumbar spine BMD change and body weight change during the 1-year follow-up period.\(^{9-11}\) According to body weight change rate, lumbar BMD change was assessed. As a result, only 3% body weight change in perimenopausal women over 1 year impacted on lumbar spine BMD. Had it followed up the weight and BMD of participants for more than 1 year, the results may have shown more significant correlation between them.

Both body fat mass and fat free mass influence on bone metabolism and act mechanical forces on the skeleton, although which body compartment of them relates more strongly to BMD is still controversial.\(^{23}\) Previous studies presumed that increase in both body fat mass and skeletal muscular mass causes increase in BMD,\(^{7,24,25}\) while recent studies have shown that BMD was decreased by increase in body fat mass after the adjustments for body weight.\(^{26,27}\) In this study, however, both body fat mass and fat free mass had no significant correlations with BMD. The reason of this result is still not clear but this maybe explained in part by bioelectrical impedance analysis based on the principle that the resistance to an applied electric current. This result may be affected by patient’s hydration or time spent on measuring. These errors could not reflect enough to correct body composition change due to body weight change. Although more accurate measurement such as DEXA, computed tomography or magnetic resonance imaging can be considered to solve this problem, there still exist so many limitations in clinical field. In Rancho Bernardo Study, the correlation coefficient between body fat mass and skeletal muscular was 0.5\(^{4}\) and it is too strong to separate them individually for analyzing them statistically. Because of these limitations, it was hard to implicate its correlation.

This study still has some limitations, although most potential confounders were carefully controlled to lower the possibility of residual confounding effects. First of all, the subjects of this study were all examinees who visited University Hospital and so it has yet to be seen whether our results can be generalized to entire population. Second, one-year follow-up period might not be enough to demonstrate the correlation between body weight change and BMD change. Third, while many kinds of nutrition such as calcium, protein, vitamin D, vitamin K, and minerals are needed to build up and maintain bone mass, this study obtained only calcium intake information from the subjects. Forth, only current eating habit was invested so that previous calcium accumulations were not considered. Fifth, low body weight group was not included in this study, even though same amount of body change would affect more on BMD in people with lower body weight than others.\(^{29}\) Many previous studies have shown that if one experienced `weight cycling,’ even though one gained weight again, it did not recover to its initial BMD.\(^{28,30}\) There is a possibility that this study might fail to identify the weight fluctuation during the follow-up period. Maybe this influences BMD change.

In conclusion, we found that only 3% body weight change during a 1 year period impacts on lumbar spine BMD in perimenopausal women and that in case of especially overweight women, the weight gain relatively shows an increase in BMD of lumbar spines than the weight loss in a retrospective cohort study. Our findings suggest that weight change in perimenopausal overweight women may contribute to developing BMD loss and much more longitudinal studies and a more comprehensive approach are needed.

**CONFLICT OF INTEREST**

No potential conflict of interest relevant to this article was reported.
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REFERENCES

1. Lobo RA. Treatment of the postmenopausal woman: basic and clinical aspects. 2nd ed. Philadelphia: Lippincott Williams & Wilkins; 1999.
2. Wing RR, Matthews KA, Kuller LH, Meilahn EN, Plantinga PL. Weight gain at the time of menopause. Arch Intern Med 1991;151:97-102.
3. Matthews KA, Meilahn E, Kuller LH, Kelsey SF, Caggiula AW, Wing RR. Menopause and risk factors for coronary heart disease. N Engl J Med 1989;321:641-6.
4. Santoro N, Chervenak JL. The menopause transition. Endocrinol Metab Clin North Am 2004;33:627-36.
5. De Laet C, Kanis JA, Oden A, Johanson H, Johnell O, Delmas P, et al. Body mass index as a predictor of fracture risk: a meta-analysis. Osteoporos Int 2005;16:1330-8.
6. Blumenthal JA, Sherwood A, Gullette EC, Babyak M, Waugh R, Georgiades A, et al. Exercise and weight loss reduce blood pressure in men and women with mild hypertension: effects on cardiovascular, metabolic, and hemodynamic functioning. Arch Intern Med 2000;160:1947-58.
7. Villareal DT, Shah K, Banks MR, Sinacore DR, Klein S. Effect of weight loss and exercise therapy on bone metabolism and mass in obese older adults: a one-year randomized controlled trial. J Clin Endocrinol Metab 2008;93:2181-7.
8. Edelstein SL, Barrett-Connor E. Relation between body size and bone mineral density in elderly men and women. Am J Epidemiol 1993;138:160-9.
9. Kim CJ, Rhee EJ, Kim HM, Kim HS, Lee EA, Kim YS, et al. Relationship between body composition and metabolic bone disease in Korean male adults. Korean J Bone Metab 2008;15:25-32.
10. Kim HK, Bae KM, Lim H, Oh CY. Bone mineral density of women in Namwon: comparison by age, pre and postmenopausal status and body weight. J Korean Acad Fam Med 2001;22:171-7.
11. Han JH, Cho KH. Correlation between body composition and spinal bone density in young women. J Korean Acad Fam Med 2002;23:215-23.
12. Brot C, Jensen LB, Sorensen OH. Bone mass and risk factors for bone loss in 4 perimenopausal Danish women. J Intern Med 1997;242:505-11.
13. Newton-John HF, Morgan DB. The loss of bone with age, osteoporosis, and fractures. Clin Orthop Relat Res 1970;71:229-52.
14. Chung YJ, Lee KM, Park JW, Chung SP. Factors related with BMD in premenopausal women. J Korean Acad Fam Med 2001;22:363-70.
15. Simkin-Silverman L, Wing RR, Hansen DH, Klem ML, Pasagian-Macaulay AP, Meilahn EN, et al. Prevention of cardiovascular risk factor elevations in healthy premenopausal women. Prev Med 1995;24:509-17.
16. Stevens J, Truesdale KP, McClain JE, Cai J. The definition of weight maintenance. Int J Obes (Lond) 2006;30:391-9.
17. Lee KS, J, Park JS, Ku ES, Bae CY, Shin DH. A study for osteoporosis of postmenopausal women. J Korean Acad Fam Med 2001;26:609-13.
18. Song YD, Lim SK, Chung YS, Park SW, Chung HC, Ahn KJ, et al. The effect of body fat on bone density in pre- and postmenopausal women. J Korean Soc Endocrinol 1993;8:251-8.
19. Daniell HW. Osteoporosis of the slender smoker. Vertebral compression fractures and loss of metacarpal cortex in relation to postmenopausal cigarette smoking and lack of obesity. Arch Intern Med 1976;136:298-304.
20. Aloia JF, Cohn SH, Vaswani A, Yeh JK, Yuen K, Ellis K. Risk factors for postmenopausal osteoporosis. Am J Med 1985;78:95-100.
21. Moon JH, Lee JK, Lee OH, Lee SY, Kim YJ. The association between indicators of central obesity and bone mineral density in women. J Korean Acad Fam Med 2001;22:192-9.
22. Zhao LJ, Jiang H, Papasian CJ, Maulik D, Drees B, Hamilton J, et al. Correlation of obesity and osteoporosis: effect of fat
23. Mass on the determination of osteoporosis. J Bone Miner Res 2008;23:17-29.

24. Reid IR, Plank LD, Evans MC. Fat mass is an important determinant of whole body bone density in premenopausal women but not in men. J Clin Endocrinol Metab 1992;75:779-82.

25. Chen Z, Lohman TG, Stini WA, Ritenbaugh C, Aickin M. Fat or lean tissue mass: which one is the major determinant of bone mineral mass in healthy postmenopausal women? J Bone Miner Res 1997;12:144-51.

26. Zhao LJ, Liu YJ, Liu PY, Hamilton J, Recker RR, Deng HW. Relationship of obesity with osteoporosis. J Clin Endocrinol Metab 2007;92:1640-6.

27. Hsu YH, Venners SA, Terwedow HA, Feng Y, Niu T, Li Z, et al. Relation of body composition, fat mass, and serum lipids to osteoporotic fractures and bone mineral density in Chinese men and women. Am J Clin Nutr 2006;83:146-54.

28. Nguyen TV, Sambrook PN, Eisman JA. Bone loss, physical activity, and weight change in elderly women: the Dubbo Osteoporosis Epidemiology Study. J Bone Miner Res 1998;13:1458-67.

29. Fogelholm GM, Sievanen HT, Kukkonen-Harjula TK, Pasanen ME. Bone mineral density during reduction, maintenance and regain of body weight in premenopausal, obese women. Osteoporos Int 2001;12:199-206.

30. Fogelholm M, Sievanen H, Heinonen A, Virtanen M, Uusi-Rasi K, Pasanen M, et al. Association between weight cycling history and bone mineral density in premenopausal women. Osteoporos Int 1997;7:354-8.