

SWAN Fact Sheet: Bone Health over the Menopause Transition*

Osteoporotic fractures usually occur in older postmenopausal women, however, bone loss leading to osteoporosis begins before menopause. The SWAN study followed a cohort of initially pre- and early perimenopausal women through the transition into postmenopause, and longitudinally tracked their bone health; we list below the major findings.

- A period of fast bone loss starts one year before the final menstrual period. This is generally (but not always) when there has been no menses for 3 or more months but some bleeding in the last year [1].Bone loss is fast for around 3 years, and continues in postmenopause, but more slowly [2].
- During the period of fast loss, bone density declines on average about 2% each year, with greater declines in the spine than in the hip. Over 10 years, cumulative bone density decline is around 10% [2]. This is accompanied by loss of bone quality and strength (the ability to resist breakage or fracture)[3,4].
- Women who lose more bone density during the menopause transition have more fractures in postmenopause [5].
- Women with earlier menopause have lower bone density in postmenopause and more fractures [6].
- Several SWAN findings demonstrate the importance of looking beyond bone density:
 - Women who are obese have greater bone density than women without obesity [7], but it does not necessarily translate to more bone strength [8]. Obese women had as many fractures as non-obese women in SWAN, but compared to non-obese women with similar bone density, obese women had significantly more fractures [8].
 - Bone density is higher in women with type 2 diabetes than in women without diabetes; yet, bone strength is lower in diabetic women [9] and they experience more fractures than women without diabetes [10]. This may be due to lower bone quality [11], earlier menopause, and faster bone lossin diabetic women [10].
 - A higher level of C-reactive protein, a marker of inflammation, that has been related to life stresses and adversity, is associated with greater bone density, but not with greater bone strength. SWAN women with higher levels of C-reactive protein had more fractures than women with lower levels of C-reactive protein [12]. In addition, increase in C-reactive protein level over time was associated with faster decline in bone density [13].

What can you do to maximize bone health?

- In premenopausal and early perimenopausal women, physical activity, be it doing housework or sports and regular exercise, is associated with greater bone density and bone strength [14,15]. Thus, women who are physically active enter the menopause transition (when bone is lost) with greater bone reserves.
- Vitamin D sufficiency (25-hydroxyvitamin D levels >20 ng/ml) increased over time in SWAN women [16]. SWAN women with adequate vitamin D levels had fewer fractures than women with low levels [17].
- Working with your health care provider to control body weight, blood glucose, and chronic inflammation is important for overall health. Whether doing so also lessens bone loss cannot be learned from SWAN; clinical trials of the effects of lowering glucose, inflammatory markers, and weight on bone loss are needed.



For more information, please check out:

- 1. Finkelstein JS, Brockwell SE, Mehta V, et al. Bone mineral density changes during the menopause transition in a multiethnic cohort of women. *J Clin Endocrinol Metab* 2008; 93: 861-8
- Greendale GA, Sowers MF, Han WJ, et al. Bone mineral density loss in relation to the final menstrual period in a multi-ethnic cohort: Results from the Study of Women's Health Across the Nation (SWAN). J Bone Miner Res 2012;27(1):111–8.
- 3. Greendale GA, Huang MH, Cauley JA, Han W, Harlow S, Finkelstein JS, Hans D, and Karlamangla AS. Trabecular bone score declines during the menopause transition: Results from the Study of Women's Health Across the Nation Trabecular Bone Score Study (SWAN-TBS). *J Clinical Endocrinology and Metabolism* 2020 April 4; 105(4): e1872-e1882
- Ishii S, Cauley JA, Greendale GA, Crandall CJ, Huang M-H, Danielson M, and Karlamangla AS. Trajectories of Femoral Neck Strength in Relation to the Final Menstrual Period in a Multi-Ethnic Cohort. *Osteop Intl.* 2013 Sep;24(9):2471-81.
- 5. Shieh A, Karlamangla AS, Huang MH, Han W, Greendale GA. Faster lumbar spine bone loss in midlife predicts subsequent fracture independent of starting bone mineral density. *The Journal of Clinical Endocrinology & Metabolism*. 2021 Jul;106(7):e2491-501.
- 6. Shieh A, Ruppert KM, Greendale GA, Lian Y, Cauley JA, Burnett-Bowie SA, Karvonen-Guttierez C, Karlamangla AS. Associations of age at menopause with postmenopausal bone mineral density and fracture risk in women. *The Journal of Clinical Endocrinology & Metabolism*. 2021 Sep 19.
- Finkelstein JS, Lee ML, Sowers M, et al. Ethnic variation in bone density in premenopausal and early perimenopausal women: effects of anthropometric and lifestyle factors. *J Clin Endocrinol Metab* 2002;87: 3057-67.
- 8. Ishii S, Cauley J, Greendale G, et al. Pleiotropic effects of obesity on fracture risk: The Study of Women's Health Across the Nation. *J Bone Miner Res* 2014; 29(12): 2561–70.
- 9. Ishii S, Cauley J, Crandall C, et al. Diabetes and femoral neck strength: Findings from The Hip Strength Across The Menopausal Transition Study. *J Clin Endocrinol Metab* 2012; 97(1): 190–7.
- Khalil N, Sutton-Tyrrell K, Strotmeyer ES, Greendale GA, Vuga M, Selzer F, Crandall CJ, and Cauley JA.Menopausal bone changes and incident fractures in diabetic women: a cohort study. *Osteoporosis international*. 2011 May; 22(5): 1367-76.
- 11. Yu EW, Putman MS, Derrico N, Abrishamanian-Garcia G, Finkelstein JS, Bouxsein ML. Defects in cortical microarchitecture among African-American women with type 2 diabetes. *Osteoporos Int* 2015; 26:673-9
- 12. Ishii S, Cauley JA, Greendale GA, et al. C-reactive protein, femoral neck strength, and 9-year fracture risk. Data from The Study of Women's Health Across the Nation. *J Bone Miner Res* 2013;28(7):1688–98.
- 13. Greendale GA, Jackson NJ, Han W, Huang M, Cauley JA, Karvonen-Gutierrez C, Karlamangla AS. Increase in C-Reactive Protein Predicts Increase in Rate of Bone Mineral Density Loss: The Study of Women's Health Across the Nation. *JBMR Plus*. 2021 Apr; 5(4): e10480.
- 14. Greendale GA, Huang MH, Wang Y, Finkelstein JS, Danielson ME, Sternfeld B. Sport and home physical activity are independently associated with bone density. *Medicine and Science in Sports and Exercise*2003 Mar 1;35(3):506-12.
- 15. Mori T, Ishii S, Greendale GA, et al. Physical activity as determinant of femoral neck strength in adult women. Findings from The Hip Strength Across The Menopausal Transition Study. *Osteoporos Int*2014; 25: 265–72.
- Mitchell DM, Ruppert K, Udupa N, et al. Temporal increases in 25-hydroxyvitamin D in midlife women: Longitudinal results from the Study of Women's Health Across the Nation. *Clin Endocrinol (Oxf)* 2019;91: 48-57
- 17. Cauley JA, Greendale GA, Ruppert K, et al. Serum 25 Hydroxyvitamin D, bone mineral density and fracture risk across the menopause. *J Clin Endocrinol Metab* 2015; 100 (5): 2046–54.

*SWAN recognizes that race is a social construct and that including race/ethnicity in describing our findings is complicated, with there being reasons for and against doing such. We and others are actively reviewing the best approach to ensure that we provide patients with the best information about their health.