

# USANA Technical Bulletin

**Disclaimer:** The information provided in this Technical Bulletin is strictly educational. It may not be used to promote USANA products, nor is it intended as medical advice. For diagnosis and treatment of medical disorders, consult your health care professional. When there are references to third party websites, addresses, and/or phone numbers, USANA, Inc. makes no claim, actual or implied, regarding the content or validity of the information obtained from such sources. This Technical Bulletin may be copied and freely distributed only if all text remains intact and unchanged.

## Vitamin D<sub>3</sub> (Cholecalciferol)

### Technical Background

- Vitamin D is a fat-soluble nutrient that plays a central role in bone growth and health.<sup>1</sup> It can be acquired either through the diet or through exposure to sunlight, when ultraviolet light reacts with a form of cholesterol and converts it to vitamin D. Cholecalciferol is the form of vitamin D normally found in humans and the form typically found in nutritional supplements.
- A unique property of vitamin D is that it functions very much like a hormone. Its target tissues in the human body include the kidneys, intestines, and bones, where it acts to regulate calcium and phosphorus homeostasis.<sup>1</sup>
- In the intestines, vitamin D plays an important role in the absorption of calcium. In bone tissue, vitamin D plays a role in regulating calcium deposition (bone mineralization) and mobilization.<sup>1</sup>
- Given the above functions, vitamin D is essential for normal bone development, particularly in children. Without it, bones do not calcify properly, leading to the condition known as “rickets”. Vitamin D also plays an important role in tooth development. It is necessary for proper tooth eruption, growth, and ultimate strength.
- Research is currently being done to examine the role of vitamin D and its analogues in inhibiting prostate cancer<sup>2</sup> and breast cancer.<sup>3</sup> Recently, a research team reviewed 63 studies on the relationship between vitamin D and certain types of cancer. In addition to prostate and breast cancer, the majority of studies found a relationship between vitamin D status and lower risk of colon and ovarian cancers as well.<sup>4</sup> In another recent study, women taking a calcium and vitamin D supplement had a 60 percent lower incidence of all cancers than women not taking the supplement.<sup>5</sup>
- Vitamin D intake is especially important for women. In addition to its possible role in breast cancer and osteoporosis prevention,<sup>6</sup> recent studies have found that vitamin D and calcium supplementation in pre-menopausal women may also decrease in the incidence of pre-menstrual syndrome (PMS).<sup>7</sup>
- A recent study found that over half of the women in North America receiving treatment for osteoporosis had an inadequate intake of vitamin D.<sup>8</sup> New research indicates that vitamin D deficiencies are widespread among pregnant women and infants despite prenatal vitamin usage.<sup>9</sup>
- Vitamin D is also being researched for its role as an immune system regulator and modulator,<sup>10,11</sup> for its role in reducing insulin resistance and type-2 diabetes,<sup>12,13</sup> and for its role in healthy heart and lung function.<sup>14,15</sup>

## Sources and Recommended Intake

- The Recommended Dietary Allowance (RDA) for vitamin D is 5 micrograms (200 IU) per day for children, and 5-15 micrograms (200-600 IU) per day for adults.<sup>16</sup> These amounts can be obtained with 15-30 minutes of exposure to sunlight on the hands, arms, and face. The use of sunscreens interferes with vitamin D synthesis. Dark skin (i.e. having a high melanin content) requires longer exposure than lighter skin to achieve the same degree of vitamin D synthesis<sup>17</sup>. Furthermore, the capacity of skin to synthesize vitamin D decreases with age.<sup>18</sup>
- The only significant dietary source of vitamin D is fortified milk. Other sources include fish and fish liver oils.<sup>19</sup>
- Although the upper limit established by the Food and Nutrition Board is 2,000 IU (or 50 mcg), many prominent researchers view this as being too restrictive. Human clinical trial data published after the establishment of the UL support a significantly higher UL. Absence of toxicity in trials conducted in healthy adults support the level of 10,000 IU as a more reasonable UL.<sup>20</sup>
- Importantly, the benefits of vitamin D supplementation are achieved only if adequate calcium and phosphorus are provided in the diet.

## Abstracts

**Augier P and Gandini S. Vitamin D Supplementation and Total Mortality: A Meta-analysis of Randomized Controlled Trials. Arch Intern Med 2007 Sep 10;167(16):1730-7.** Ecological and observational studies suggest that low vitamin D status could be associated with higher mortality from life-threatening conditions including cancer, cardiovascular disease, and diabetes mellitus that account for 60% to 70% of total mortality in high-income countries. We examined the risk of dying from any cause in subjects who participated in randomized trials testing the impact of vitamin D supplementation (ergocalciferol [vitamin D(2)] or cholecalciferol [vitamin D(3)]) on any health condition. METHODS: The literature up to November 2006 was searched without language restriction using the following databases: PubMed, ISI Web of Science (Science Citation Index Expanded), EMBASE, and the Cochrane Library. RESULTS: We identified 18 independent randomized controlled trials, including 57 311 participants. A total of 4777 deaths from any cause occurred during a trial size-adjusted mean of 5.7 years. Daily doses of vitamin D supplements varied from 300 to 2000 IU. The trial size-adjusted mean daily vitamin D dose was 528 IU. In 9 trials, there was a 1.4- to 5.2-fold difference in serum 25-hydroxyvitamin D between the intervention and control groups. The summary relative risk for mortality from any cause was 0.93 (95% confidence interval, 0.87-0.99). There was neither indication for heterogeneity nor indication for publication biases. The summary relative risk did not change according to the addition of calcium supplements in the intervention. CONCLUSIONS: Intake of ordinary doses of vitamin D supplements seems to be associated with decreases in total mortality rates. The relationship between baseline vitamin D status, dose of vitamin D supplements, and total mortality rates remains to be investigated. Population-based, placebo-controlled randomized trials with total mortality as the main end point should be organized for confirming these findings.

## References

- <sup>1</sup> Norman, AW. 1996. Vitamin D. Pp. 120-129. In EE Ziegler and LJ Filer (eds). Present Knowledge in Nutrition. ILSI Press, Washington, DC.
- <sup>2</sup> Woo TC, Choo R, Jamieson M, Chander S, Vieth R. Pilot study: potential role of vitamin D (Cholecalciferol) in patients with PSA relapse after definitive therapy. Nutr Cancer. 2005;51(1):32-6.
- <sup>3</sup> Mehta RR, Bratescu L, Graves JM, Green A, Mehta RG. Differentiation of human breast carcinoma cells by a novel vitamin D analog: 1alpha-hydroxyvitamin D5. Int J Oncol. 2000 Jan;16(1):65-73.
- <sup>4</sup> Garland CF et al. The Role of Vitamin D in Cancer Prevention. American Journal of Public Health, Vol 96, No. 2 252-261 February (2006).

- <sup>5</sup> Lappe JM et al. Vitamin D and calcium supplementation reduces cancer risk: results of a randomized trial. *The American Journal of Clinical Nutrition* 2007 June; 85(6):1586-91.
- <sup>6</sup> Papadimitropoulos E, Wells G, Shea B, Gillespie W, Weaver B, Zytaruk N, Cranney A, Adachi J, Tugwell P, Josse R, Greenwood C, Guyatt G. Meta-analyses of therapies for postmenopausal osteoporosis. VIII: Meta-analysis of the efficacy of vitamin D treatment in preventing osteoporosis in postmenopausal women. *Endocr Rev.* 2002 Aug;23(4):560-9.
- <sup>7</sup> Bertone-Johnson ER, Hankinson SE, Bendich A, Johnson SR, Willett WC, Manson JE. Calcium and vitamin D intake and risk of incident premenstrual syndrome. *Arch Intern Med.* 2005 Jun 13;165(11):1246-52.
- <sup>8</sup> Holick MF, Siris ES, Binkley N, Beard MK, Khan A, Katzer JT, Petruschke RA, Chen E, de Papp AE. Prevalence of vitamin D inadequacy among postmenopausal North American women receiving osteoporosis therapy. *J Clin Endocrinol Metab.* 2005 Jun;90(6):3215-24.
- <sup>9</sup> Bodnar LM et al. High prevalence of vitamin D insufficiency in black and white pregnant women residing in the northern United States and their neonates. *J. Nutr.* 137:447-452, February 2007
- <sup>10</sup> Cantorna MT, Zhu Y, Froicu M, Wittke A. Vitamin D status, 1,25-dihydroxyvitamin D<sub>3</sub>, and the immune system. *Am J Clin Nutr.* 2004 Dec;80(6 Suppl):1717S-20S.
- <sup>11</sup> Mathieu C, van Etten E, Decallonne B, Guilietti A, Gysemans C, Bouillon R, Overbergh L. Vitamin D and 1,25-dihydroxyvitamin D<sub>3</sub> as modulators in the immune system. *J Steroid Biochem Mol Biol.* 2004 May;89-90(1-5):449-52.
- <sup>12</sup> Chiu KC et al. Hypovitaminosis D is associated with insulin resistance and beta cell dysfunction. *Am J Clin Nutr* 2004 May;79(5):820-5.
- <sup>13</sup> Pittas AG et al. Vitamin D and calcium intake in relation to type 2 diabetes in women. *Diabetes Care* 29:650-656, 2006.
- <sup>14</sup> Major GC et al. Supplementation with calcium + vitamin D enhances the beneficial effect of weight loss on plasma lipid and lipoprotein concentrations. *Am J Clin Nutr*, Vol. 85, No. 1, 54-59, January 2007.
- <sup>15</sup> Black PN and Scragg R. Relationship between serum 25-hydroxyvitamin d and pulmonary function in the third national health and nutrition examination survey. *Chest* 2005 Dec;128(6):3792-8.
- <sup>16</sup> Institute of Medicine. *Dietary Reference Intakes for Calcium, Phosphorus, Magnesium, Vitamin D, and Fluoride.* Washington D.C.: National Academy Press. 1997
- <sup>17</sup> Clemens, TL and others. 1982. Increased skin pigment reduces capacity of skin to synthesize vitamin D<sub>3</sub>. *Lancet* 1: 74-76.
- <sup>18</sup> Webb, AR and others. 1988. Influence of season and latitude on the cutaneous synthesis of vitamin D<sub>3</sub>: exposure to winter sunlight in Boston and Edmonton will not promote vitamin D<sub>3</sub> synthesis in human skin. *J. Clin. Endocrinol. Metab.* 67: 373-378.
- <sup>19</sup> Cohen, RD and NS Braunstein. 1995. *Vitasearch.* Great Bay Nutrition Resources, New Market, NH. 49 pp.
- <sup>20</sup> Hathcock JN, Shao A, Vieth R, and Heaney R. Risk assessment for vitamin D. *Am J Clin Nutr* 2007 Jan;85(1):6-18