

Selenium

Common Indications:

- Antioxidant
- Cancer Prevention
- Thyroid Autoimmune Modulatory
- Immunomodulatory
- Cardiovascular Health
- Metal detoxification
- Male fertility
- Mental health
- Gestational diabetes

General Comments:

Until the late 1950s, selenium was thought to be toxic. Although it can be toxic at high doses, it is now recognized as an important nutritional trace mineral. Selenium plays important roles in detoxification and antioxidant defense mechanisms in the body. Selenium functions as a redox "gatekeeper", improving antioxidant defense mechanisms. Although selenium is best absorbed in selenomethionine and selecysteine as organic forms, selenium is prepared commonly as sodium selenite and sodium selenate in variety of products.

Benefits & Mechanism of action:

Antioxidant:

- Selenium helps to reduce lipid peroxidation and neutralize the destructive hydrogen peroxide radicals.
- Selenium potentiates the antioxidant activity of vitamin E.
- Selenocysteine insertion complex suggests Selenium's activity in cancer prevention¹.

Immunomodulatory:

- Selenium is a co-factor for glutathione peroxidase, which is an important antioxidant enzyme in the immune system.
- Selenium has reported anti-viral activity, may increase T- lymphocytes, and enhances
 natural killer cell activity¹⁸. A 2013 meta-analysis reported that use of high-dose selenium
 (IV and PO) in patients with sepsis helped reduce mortality²⁰.

PHONE: (513) 366-2100 FAX: (513) 351-3800 WEB: www.hippevo.com

 A study in 2007 has found that selenium helped to suppress HIV progression and improve CD4 counts¹⁹. Selenium also helps to improve immune function in HIV patients and other morbidity associated with the disease²¹.

Cancer prevention:

- Epidemiological studies have correlated low dietary selenium intakes with higher rates of cancer. Studies have shown that selenium reduced the risk and inversely intervened the progression of cancers in patients who have low selenium level or are smokers. Human meta-analysis report selenium has a protective effect on various cancers^{1,10}:
 - Colorectal¹¹
 - o Prostate⁹
 - o Lung⁷
 - o Bladder²
 - Breast⁴
- Selenium may also help to reduce the side effects commonly found in cancer chemotherapy treatments including cisplatin-induce nephrotoxicity⁷.

Cardiovascular Health:

• Selenium's anti-oxidant activities are reported to enable it to protect against heart attacks and strokes. A 2006 meta-analysis reported that increased levels of selenium correlate with a decreased risk of cardiovascular disease¹³. However, a 2013 systematic review reported no benefit in preventing CV disease when using selenium¹⁴. Some studies suggest that the cardiovascular benefits from selenium are dependent on the baseline selenium status. It is more beneficial for the patients who have serum selenium levels below 120 ng/mL for selenium supplements being impactful in reducing cardiovascular and coronary heart disease mortality; however, high serum selenium level would cause elevated blood pressure or hypertension^{16,17}.

Metal detoxification:

• Helps detoxify heavy metal toxins such as mercury and cadmium^{27,28}.

Thyroid Autoimmune Modulatory:

• Recently discovered that the deiodinase enzyme that converts thyroid hormone (T4) to triiodothyronine (T3, the active form) is a selenium- dependent enzyme²³. A systematic review of the literature in 2010 reported that selenium supplementation reduced thyroid peroxidase antibodies and may be beneficial for those with autoimmune thyroiditis, including Hashimoto's thyroiditis²².

Fertility:

Several literatures suggest that low selenium level linked to low fertility in man.
 Testosterone synthesis and spermatogenesis require selenium as an essential source of energy. Providing selenium level in low selenium male individuals would increase sperm motility and fertility rate^{29,30}. Nonetheless, an animal study showed that either excessive or insufficient amount of selenium would detrimentally affect fertility³¹.

Mental Health

 Improved symptoms of depression, elevated mood, decreased anxiety and tiredness shown in 2 studies^{25,26}.

Diabetes

 Some studies suggest selenium intake helps improve serum glucose concentration^{32,33} and other studies have found the opposite evidence in diabetes patients^{15,16}. However, many studies support selenium intake in improving serum glucose concentration in gestational diabetes^{35,36,37}.

Dose:

- DRI* 50mcg daily
- ODA** 50 200mcg daily
- * The Dietary Reference Intakes (DRI) are the most recent set of dietary recommendations established by the Food and Nutrition Board of the Institute of Medicine, 1997-2001. They replace previous RDAs, and may be the basis for eventually updating the RDIs.
- **The Optimum Daily Allowance (ODA) represents a reference level beyond the RDI, and is often many times higher than the RDI to prevent diseases such as aging or cancer. These numbers are based on clinical use.

Cautions & Side Effects:

Selenium is a trace mineral that could be toxic if excessive amounts were ingested on a regular basis. Symptoms of selenium toxicity include loss of hair and nails, skin lesions, nervous system abnormalities, digestive dysfunction, and a garlicky breath odor. Although deaths from selenium toxicity have been reported in livestock, no deaths have occurred in humans.

References:

Antioxidant

1. Rayman MP. Selenium in cancer prevention: a review of the evidence and mechanism of action. Proc Nutr Soc. 2005;64:527–542.

Cancer prevention

- 2. Amarai AF, Cantor KP, Silverman DT, et al. Selenium and bladder cancer risk: a meta-analysis. Cancer Epidemiol Biomarkers Prev. 2010;19(9):2407-15.
- 3. Bardia A, Tleyjeh IM, Cerhan JR, et al. Efficacy of antioxidant supplementation in reducing primary cancer incidence and mortality: systematic review and meta-analysis. Mayo Clin Proc. 2008 Jan;83(1):23-34.
- 4. Chen YC, Prabhu KS, Das A, et al. Dietary selenium supplementation modifies breast tumor growth and metastasis. Int J Cancer. 2013.

PHONE: (513) 366-2100 FAX: (513) 351-3800 WEB: www.hippevo.com

- 5. Duffield-Lillico AJ, Slate EH, Reid ME, et al. Selenium supplementation and secondary prevention of nonmelanoma skin cancer in a randomized trial. J Natl Cancer Inst 2003;95(19):1477-1481.
- 6. Etminan M, FitzGerald JM, Gleave M, et al. Intake of selenium in the prevention of prostate cancer: a systematic review and meta-analysis. Cancer Causes Control 2005 Nov;16(9):1125-31.
- 7. Fritz H, Kennedy D, Fergusson D, et al. Selenium and lung cancer: A systematic review and meta-analysis. PLoS One. 2011;6(11):e26259.
- 8. Hoque A, Albanes D, Lippman SM, et al. Molecular epidemiologic studies within the Selenium and Vitamin E Cancer Prevention Trial (SELECT) Cancer Causes Control 2001 Sep;12(7):627-33.
- 9. Hurst R, Hooper L, Norat T, et al. Selenium and prostate cancer: systematic review and meta-analysis. Am J Clin Nutr. 2012;96(1):111-22.
- 10. Lee EH, Myung SK, Jeon YJ, et al. Effects of selenium supplements on cancer prevention: a meta-analysis of randomized controlled trials. Nutr Cancer. 2011;64(8):1185-95.
- 11. Ou Y, Jiang B, Wang X, et al. Selenium and colorectal adenomas risk: a meta-analysis. Nutr Cancer. 2012;64(8):1153-9.
- 12. Reid ME, Duffield-Lillico AJ, Sunga A, et al. Selenium supplementation and colorectal adenomas: an analysis of the nutritional prevention of cancer trial. Int J Cancer 2006 Apr 1;118(7):1777-81.

Cardiovascular Health

- 13. Flores-Mateo G, Navas-Acien A, Pastor-Barriuso R, et al. Selenium and coronary heart disease: a meta- analysis. Am J Clin Nutr. 2006;84(4):762-73.
- 14. Rees K, Hartley L, Day C, et al. Selenium supplementation for the primary prevention of cardiovascular disease. Cochrane Database Syst Rev. 2013;1:CD009671.
- 15. Stranges S, Marshall JR, Trevisan M, et al. Effects of selenium supplementation on cardiovascular disease incidence and mortality: secondary analyses in a randomized clinical trial. Am J Epidemiol 2006 Apr 15;163(8):694-9.
- 16. Bleys J, Navas-Acien A, Guallar E. Serum selenium levels and all-cause, cancer, and cardiovascular mortality among US adults. Arch Intern Med. 2008 Feb 25;168(4):404-10.
- 17. Nawrot T, Staessen J, Roels H, et al. Blood pressure and blood selenium: a cross-sectional and longitudinal population study. Eur Heart J. 2007 Mar;28(5):628-33. Epub 2007 Jan 22.

Immunomodulatory

DUONE, (512) 200 2100 - 5AV, (512) 251 2000 - WED. history com

- 18. Gazdik F, Horvathova M, Gazdikova K, et al. The influence of selenium supplementation on the immunity of corticoid-dependent asthmatics. Bratisl Lek Listy. 2002;103(1):17-21.
- 19. Hurwitz BE, Klaus JR, Llabre MM, et al. Suppression of human immunodeficiency virus type 1 viral load with selenium supplementation: a randomized controlled trial. Arch Intern Med. 2007 Jan 22;167(2):148-54.
- 20. Alhazzani W, Jacobi J, Sindi A, et al. The effect of selenium therapy on mortality in patients with sepsis syndrome: A systematic review and meta-analysis of randomized controlled trials. Crit Care Med. 2013;41(6):1555-1564.
- 21. Baum M, Campa A, Lai S, et al. Effect of micronutrient supplementation on disease progression in asymptomatic, antiretroviral-naive, HIV-infected adults in Botswana: a randomized clinical trial. JAMA. 2013 Nov 27;310(20):2154-63.

Thyroid autoimmune modulatory

- 22. Toulis KA, Anastasilakis AD, Tzellos TG, et al. Selenium supplementation in the treatment of Hashimoto's thyroiditis: a systematic review and meta-analysis. Thyroid. 2010;29(1):1163-73.
- 23. Turker O, Kumanlioglu K, Karapolat I, et al. Selenium treatment in autoimmune thyroiditis: 9-month follow-up with variable doses. J Endocrinol 2006 Jul;190(1):151-6.

Mental health

- 24. Rayman M, Thompson A, Warren-Perry M, et al. Impact of selenium on mood and quality of life: a randomized, controlled trial. Biol Psychiatry 2006 Jan 15;59(2):147-54.
- 25. Mokhber N, Namjoo M, Tara F, et al. Effect of supplementation with selenium on postpartum depression: a randomized double-blind placebo-controlled trial. J Matern Fetal Neonatal Med. 2011 Jan;24(1):104-8.
- 26. Benton D, Cook R. Selenium supplementation improves mood in a double-blind crossover trial. Psychopharmacology (Berl). 1990;102(4):549-50.

Metal detoxification

- 27. Zwolak I, Zaporowska H. Selenium interactions and toxicity: a review. Selenium interactions and toxicity. Cell Biol Toxicoll 2012;28(1):31-46.
- 28. Bolkent S, Sacan O, Yanardag R, et al. Effects of vitamin E, vitamin C, and selenium on gastric fundus in cadmium toxicity in male rats. Int J Toxicol. 2008 Mar-Apr;27(2):217-22.

Male fertility

29. Turk S, Mandar R, Mahlapuu R, et al. Male infertility: decreased levels of selenium, zinc and antioxidants. J Trace Elem Med Biol. 2014 Apr;28(2):179-85.

PHONE: (513) 366-2100 FAX: (513) 351-3800 WEB: www.hippevo.com

- 30. Yao DF, Mills J. Male infertility: lifestyle factors and holistic, complementary, and alternative therapies. Asian J Androl. 2016 May-Jun;18(3):410-8.
- 31. Shalini S, Bansal M. Dietary selenium deficiency as well as excess supplementation induces multiple defects in mouse epididymal spermatozoa: understanding the role of selenium in male fertility. Int J Androl. 2008 Aug;31(4):438-49.

Diabetes

- 32. Campbell S, Aldibbiat A, Marriott C, et al. Selenium stimulates pancreatic beta-cell gene expression and enhances islet function. FEBS Lett. 2008 Jun 25;582(15):2333-7.
- 33. Alizadeh M, Safaeiyan A, Ostadrahimi A, et al. Effect of L-arginine and selenium added to a hypocaloric diet enriched with legumes on cardiovascular disease risk factors in women with central obesity: a randomized, double-blind, placebo-controlled trial. Ann Nutr Metab. 2012;60(2):157-68.
- 34. Stranges S, Marshall J, Natarajan R, et al. Effects of long-term selenium supplementation on the incidence of type 2 diabetes: a randomized trial. Ann Intern Med. 2007 Aug 21;147(4):217-23.
- 35. Kilinc M, Guven M, Ezer M, et al. Evaluation of serum selenium levels in Turkish women with gestational diabetes mellitus, glucose intolerants, and normal controls. Biol Trace Elem Res. 2008 Summer;123(1-3):35-40.
- 36. Bo S, Lezo A, Menato G, et al. Gestational hyperglycemia, zinc, selenium, and antioxidant vitamins. Nutrition. 2005 Feb;21(2):186-91.
- 37. Molnar J, Gagamvolgyi Z, Herold M, et al. Serum selenium concentrations correlate significantly with inflammatory biomarker high-sensitive CRP levels in Hungarian gestational diabetic and healthy pregnant women at mid-pregnancy. Biol Trace Elem Res. 2008 Jan;121(1):16-22.
