

Inositol: Inositol or cyclohexane-1,2,3,4,5,6-hexol, Inositol hexaniacinate (inositol hexanicotinate), Myo-Inositol, or cis-1,2,3,5-trans-4,6-cyclohexanehexol

Common Indications:

- Metabolic syndrome; insulin resistance
- Diabetic neuropathy
- Obsessive-compulsive disorder
- Panic attacks
- Depression
- Polycystic ovary syndrome (PCOS)
- Psoriasis associated with lithium therapy

General Comments:

Inositol is part of the water-soluble B vitamin complex and esily found in many plants. Inositol is also synthesized in the intestinal tract by the beneficial flora. Inositol occurs as a component of phospholipids in animal tissue.

The popular "no flush" niacin that we read about is formed from mixing inositol with 6 niacin (Vitamin B3) molecules to form inositol hexaniacinate or inositol hexanicotinate. This form of no flush niacin does NOT have the same impact as true niacin and does nothing to alter or reduce adverse lipid panels.

Benefits & Mechanism of Action:

Metabolic syndrome; insulin resistance

As part of phospholipids in cellular membranes, phosphatidylinositol helps to mediate cellular responses to external stimuli. Myo-inositol may be helpful in improving insulin resistance and decreasing symptoms of metabolic syndrome.³ A study of postmenopausal women found that myo-inositol supplementation improved triglyceride levels, HDL levels, and diastolic blood pressure.^{3,4} When the observations were collected at one year, it was found that the benefits were greater the longer that the women were treated.⁴

Increasing urinary concentrations are directly correlated to reduced insulin receptor

activity and can been used as a biomarker for insulin resistance. 5,6

Diabetic neuropathy

Inositol Inositol is used in the development of peripheral nerves and has shown to be effective in the treatment of diabetic neuropathy.^{3,7,8}

Inositol was found to be more effective than placebo when used in patients with diabetic neuropathy associated erectile dysfunction.⁹

Obsessive-compulsive disorder

A double blind study of myo-inositol, at 18 grams per day, showed a significant reduction in symptoms of OCD, non-inferior to a selective serotonin reuptake inhibitor (SSRI), with less side effects. 10,11

Panic attacks

One study showed 18g of inositol daily was found to be as effective as fluvoxamine, an SSRI, in reducing anxiety symptoms associated with panic. 12,13,14 Inositol does not appear to be effective for panic disorders associated with PTSD. 15

Depression

A human study assessed patients with depression and found that 12g of inositol daily for 4 weeks showed an improvement of depression symptoms. 16,17 It appeared to be more helpful in females. 16,17

Polycystic ovary syndrome (PCOS)

Myo-inositol is reported to help restoring metabolic, hormonal and spontaneous ovarian activity in those with PCOS. Studies have shown that doses greater than 200mg daily improve oocyte and embryo quality prior to in vitro fertilization, improve androgen associated symptoms, improve glucose metabolism, restore menstruation in those who had lost cycles due to PCOS, and increase pregnancy rates. 19,20,21,22,23,24,25 It has also been shown to help with hirsutism associated with PCOS. 26,27

Psoriasis associated with lithium therapy

Inositol supplementation has been shown to improve peripheral side effects of lithium therapy, including flares in psoriasis, without interfering with lithium's mechanism of action in the brain. ^{28,29}

Dose: in different applications:

General

- 1,000 mg 18 grams daily in divided doses
- Inositol hexaniacinate 540 mg daily

Obsessive-Compulsive disorder/Panic Attacks

• 18g dailv^{10,11,12,13,14}

Depression

• 12g daily^{16,17}

PCOS

200-4,000mg once daily before breakfast^{19,20,21,22,23,24,25}

Symptoms of Deficiency:

No inositol deficiency has been identified in humans. A deficiency is not likely because of its widespread occurrence in foods and the body's ability to synthesize inositol from glucose-6-phosphate.

The following symptoms have been associated with reduced inositol intake

- Anxiety
- Depression
- Difficulty falling asleep
- Uterine fibroids
- PMS

Food and supplements that may decrease inositol include:

Caffeine

Cautions & Side Effects:

Medication interactions

Medications with increased effects while taking inositol include:

- Oral contraceptives in the treatment of PCOS
- Melatonin melatonin increases the effects of inositol supplementation

Medications that can decrease the amount of inositol include:

- Antibiotics
- Lithium

Food Sources:

Inositol occurs in foods in three different forms: as free myo-inositol, phytic acid, and inositol-containing phospholipids. The richest plant sources of myo-inositol are cantaloupe and oranges, followed by seeds such as beans, grains, and nuts. The richest animal sources are organ meats. Free myo-inositol predominates in brain and kidney, whereas phospholipid-inositol is concentrated in skeletal muscle, heart, liver, and pancreas.

Nutrient Interactions:

Calcium

Phytic acid or inositol hexaphosphate, which is the form of inositol that occurs in plants, binds with divalent cations decreasing the absorption of these minerals from the gastrointestinal tract.

Zinc

Phytic acid or inositol hexaphosphate, which is the form of inositol that occurs in plants, binds with divalent cations decreasing the absorption of these minerals from the gastrointestinal tract.

Iron

Phytic acid or inositol hexaphosphate, which is the form of inositol that occurs in plants, binds with divalent cations decreasing the absorption of these minerals from the gastrointestinal tract.

References:

General Comments

- 1. Smith, P. What You Must Know About Vitamins, Minerals, Herbs & More. New York: Square One Publishing, 2008.
- 2. Coombs, G. The Vitamins: Fundamental Aspects in Nutrition and Health, 3rd Ed. Burlington, MA: Elsevier Academic Press, 2008, 265-381.

Metabolic syndrome; insulin resistance

- 3. Santamaria A, Giordano D, Corrado F, et al. One-year effects of myo-inositol supplementation in postmenopausal women with metabolic syndrome. Climacteric. 2012;15(5):490-5.
- 4. Giordano D, Corrado F, Santamaria A, Quattrone S, Pintaudi B, DiBenedetto A, D'Anna R (2011). "Effects of myo-inositol supplementation in postmenopausal women with metabolic syndrome: a perspective, randomized, placebo-controlled study". Menopause: the Journal of the North American Menopause Society 18 (1): 102–104.
- 5. Stull AJ, et al. Relationships between urinary inositol excretions and whole-body glucose tolerance and skeletal muscle insulin receptor phosphorylation . Metabolism. (2008)

6. Larner J, Craig JW. Urinary myo-inositol-to-chiro-inositol ratios and insulin resistance . Diabetes Care. (1996)

Diabetic Nephropathy

- 3. Santamaria A, Giordano D, Corrado F, et al. One-year effects of myo-inositol supplementation in postmenopausal women with metabolic syndrome. Climacteric. 2012;15(5):490-5.
- 7. Chau JF, et al Sodium/myo-inositol cotransporter-1 is essential for the development and function of the peripheral nerves . FASEB J. (2005)
- 8. Salway JG, et al. Effect of Myo-inositol on Peripheral-nerve Function in diabetes. Lancet. Dec1978;2(8103):1282-84.
- 9. Agostini R, Rossi F, Pajalich R. Myoinositol/folic acid combination for the treatment of erectile dysfunction in type 2 diabetes men: a double-blind, randomized, placebo-controlled study . Eur Rev Med Pharmacol Sci. (2006)

Obsessive-Compulsive Disorder

- 10. Fux M, et al. Inositol Treatment of Obsessive-compulsive Disorder. Am J Psychiatry. Sep1996;153(9):1219-21.
- 11. Carey PD, Warwick J, Harvey BH, Stein DJ, Seedat S. Single photon emission computed tomography (SPECT) in obsessive-compulsive disorder before and after treatment with inositol. Metab Brain Dis. Jun2004;19(1-2):125-34.

Panic Attacks

- 12. Benjamin J, et al. Inositol Treatment in Psychiatry. Psychopharmacol Bull. 1995;31(1):167-75.
- 13. Benjamin J, et al. Double-blind, Placebo-controlled, Crossover Trial of Inositol Treatment for Panic Disorder. Am J Psychiatry. Jul1995;152(7):1084-86.
- 14. Palatnik A, Frolov K, Fux M, Benjamin J. Double-Blind, controlled, crossover trial of inositol versus fluvoxamine for the treatment of panic disorder. J Clin Psychopharmacol. Jun2001;21(3):335-9.
- 15. Kaplan Z, et al Inositol treatment of post-traumatic stress disorder . Anxiety. (1996)

Depression

- Levine J, et al Double-blind, controlled trial of inositol treatment of depression. Am J Psychiatry. (1995)
- 17. Levine J, et al Follow-up and relapse analysis of an inositol study of depression . Isr J Psychiatry Relat Sci. (1995)

Polycystic ovary syndrome (PCOS)

- 18. Unfer, V et al. "Effects of myo-inositol in women with PCOS: a systematic review of randomized controlled trials." Gynecological Endocrinology 28.7 (2012): 509-515.
- 19. Genazzani AD, et al Differential insulin response to myo-inositol administration in obese polycystic ovary syndrome patients . Gynecol Endocrinol. (2012)
- 20. Donà G, et al Inositol administration reduces oxidative stress in erythrocytes of patients with polycystic ovary syndrome . Eur J Endocrinol. (2012)
- 21. Carmina E. PCOS: metabolic impact and long-term management. Minerva Ginecol. (2012)
- 22. Artini PG, et al Endocrine and clinical effects of myo-inositol administration in polycystic ovary syndrome. A randomized study . Gynecol Endocrinol. (2013)
- 23. Costantino D, et al. Metabolic and hormonal effects of myo-inositol in women with polycystic ovary syndrome: a double-blind trial . Eur Rev Med Pharmacol Sci. (2009)
- 24. Genazzani AD, et al. Myo-inositol administration positively affects hyperinsulinemia and hormonal parameters in overweight patients with polycystic ovary syndrome. Gynecol Endocrinol. (2008)
- 25. Papaleo E, et al. Myo-inositol in patients with polycystic ovary syndrome: a novel method for ovulation induction . Gynecol Endocrinol. (2007)
- 26. Cupisti S, et al Influence of body mass index on measured and calculated androgen parameters in adult women with Hirsutism and PCOS. Exp Clin Endocrinol Diabetes. (2007)
- 27. Mueller A, et al. Endocrinological markers for assessment of hyperandrogenemia in hirsute women . Horm Res. (2007)

Psoriasis associated with lithium therapy

- 28. Bersudsky, Y., Vinnitsky, I., Grisaru, N., Yaroslavsky, U., Gheorghiu, S., Ivgi, D., Kofman, O. and Belmaker, R. H. (1992), The effect of inositol on lithium-induced polyuria—polydipsia in rats and humans. Hum. Psychopharmacol. Clin. Exp., 7: 403–407. doi: 10.1002/hup.470070606
- 29. Allan SJ, et al The effect of inositol supplements on the psoriasis of patients taking lithium: a randomized, placebo-controlled trial . Br J Dermatol. (2004)

Dosage - Obsessive-Compulsive disorder/Panic Attacks

- 10. Fux M, et al. Inositol Treatment of Obsessive-compulsive Disorder. Am J Psychiatry. Sep1996;153(9):1219-21.
- 11. Carey PD, Warwick J, Harvey BH, Stein DJ, Seedat S. Single photon emission computed tomography (SPECT) in obsessive-compulsive disorder before and after treatment with inositol. Metab Brain Dis. Jun2004;19(1-2):125-34.
- 12. Benjamin J, et al. Inositol Treatment in Psychiatry. Psychopharmacol Bull. 1995;31(1):167-75.
- 13. Benjamin J, et al. Double-blind, Placebo-controlled, Crossover Trial of Inositol

- Treatment for Panic Disorder. Am J Psychiatry. Jul 1995;152(7):1084-86.
- 14. Palatnik A, Frolov K, Fux M, Benjamin J. Double-Blind, controlled, crossover trial of inositol versus fluvoxamine for the treatment of panic disorder. J Clin Psychopharmacol. Jun2001;21(3):335-9.
- Levine J, et al Double-blind, controlled trial of inositol treatment of depression . Am J Psychiatry. (1995)
- 17. Levine J, et al. Follow-up and relapse analysis of an inositol study of depression . Isr J Psychiatry Relat Sci. (1995)
- 19. Genazzani AD, et al. Differential insulin response to myo-inositol administration in obese polycystic ovary syndrome patients . Gynecol Endocrinol. (2012)
- 20. Donà G, et al Inositol administration reduces oxidative stress in erythrocytes of patients with polycystic ovary syndrome . Eur J Endocrinol. (2012)
- 21. Carmina E. PCOS: metabolic impact and long-term management . Minerva Ginecol. (2012)
- 22. Artini PG, et al Endocrine and clinical effects of myo-inositol administration in polycystic ovary syndrome. A randomized study . Gynecol Endocrinol. (2013)
- 23. Costantino D, et al. Metabolic and hormonal effects of myo-inositol in women with polycystic ovary syndrome: a double-blind trial. Eur Rev Med Pharmacol Sci. (2009)
- 24. Genazzani AD, et al. Myo-inositol administration positively affects hyperinsulinemia and hormonal parameters in overweight patients with polycystic ovary syndrome . Gynecol Endocrinol. (2008)
- 25. Papaleo E, et al. Myo-inositol in patients with polycystic ovary syndrome: a novel method for ovulation induction . Gynecol Endocrinol. (2007)