OMEGAPRO DHA

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Clinical Applications

- Supports Early Brain Development*
- Promotes Brain, Eye, Heart, and Cell Membrane Function Throughout the Lifespan*
- Supports Immune Health and Eicosanoid Metabolism*

OMEGAPRO DHA features patented MaxSimil® DHA monoglyceride fish oil. This readily absorbed omega-3 fatty acid formula provides concentrated docosahexaenoic acid (DHA) plus eicosapentaenoic acid (EPA). DHA is often recommended during pregnancy to support healthy fetal brain development. Throughout the life span, both DHA and EPA support healthy brain, heart, immune, and eye function as well as healthy eicosanoid metabolism.*

All Inspired Health LLC. Formulas Meet or Exceed cGMP Quality Standards

Discussion

The two most well-researched omega-3 polyunsaturated fatty acids (PUFAs) are docosahexaenoic acid (DHA) and eicosapentaenoic acid (EPA). DHA is an essential structural component of the central nervous system, and EPA plays a role in supporting healthy cardiac and circulatory systems. OMEGAPRO DHA features MaxSimil DHA. This particular form of the highly absorbable, patented MaxSimil monoglyceride fish oil contains 1,350 mg of concentrated DHA and 236 mg of EPA.*

Absorption

Studies conducted by the manufacturer of MaxSimil provide promising results suggesting better absorption than other fish oils. An unpublished, double-blind, crossover, pharmacokinetic study was performed in healthy overnight-fasted male and female subjects (N = 20) ages 19 to 60. Each subject was administered a single dose of six softgels (containing ~2000 mg EPA and ~1500 mg DHA) of either ethyl ester (EE) fish oil or MaxSimil. Compared to the EE form, MaxSimil EPA and DHA reached a peak concentration more than three times higher than that reached by EE fish oil. Moreover, MaxSimil not only reached maximum concentration faster but also maintained plasma levels longer. Additional peer-reviewed research related to bioavailability is warranted.*[1-3]

Early Brain and Eye Development

Polyunsaturated fatty acids (PUFAs) play a critical role in the normal development of the eye, brain, and central nervous system, and DHA is the most abundant structural fatty acid in these tissues. DHA is transferred directly from mother to fetus during pregnancy, especially during the last trimester, and is available to infants through breast milk. This fatty acid plays a vital role in both prenatal and postnatal brain development. Based on recommendations made by various organizations, pregnant and nursing women should consume 100-300 mg of DHA daily to meet basic fetal and infant needs.*[4,5]

Research indicates that DHA-supplemented and breastfed infants score significantly better on mental and psychomotor development tests and that PUFAs may support normal activity levels and learning capacity during preschool years. [4,6] A study of infants (N = 229) enrolled in three randomized controlled trials suggested that a dose of 0.36% of total fatty acids as DHA (a concentration representative of human breast milk) contributed to favorable problem-solving performance, a parameter found to correlate with later IQ and vocabulary development. [7] However, the outcome of a systematic review and meta-analysis showed that although omega-3 supplementation does appear to improve childhood psychomotor and visual development, more studies are needed to confirm these conclusions and to explore the significance of IQ later in childhood.*[8]

DHA has also been researched for its role in developing and maintaining eye health and function during early life. A randomized trial of healthy formula-fed infants (N = 244) suggested that infants who were not given DHA had poorer visual acuity than those who were supplemented with DHA amounting to 0.32% of total fatty acids. (9) Optimal retinal and visual cortex maturation was understood to depend upon dietary DHA during development, and visual acuity and mental development were "seemingly improved by extra DHA."*(6)

Nervous System, Brain, Eye, and Cell Membrane Function

Functioning via cell membranes and anchored by phospholipid molecules, DHA helps to optimize signal transmission in the tissues of the brain, eyes, and the overall nervous system.^[10] Levels of this fatty acid decline with aging. DHA supplementation may play a role in maintaining myelin and neuronal health, supporting healthy eicosanoid metabolism in the brain, and exerting pleiotropic effects to support healthy metabolism and aging throughout the lifespan.*^[6,11,12]

A study of healthy community volunteers (N = 280), ages 35-54, showed that higher DHA levels were significantly associated with performance improvement in the areas of nonverbal reasoning, mental flexibility, working memory, and vocabulary. (13) Other research has focused on neuroprotectin D1 (NPD1), an important mediator derived from DHA through the action of 15-lipoxygenase-1. This mediator appears to have a positive effect on neurotrophic cell signaling, normal cell-life cycles, beta-amyloid neurobiology, and prostaglandin formation. A systematic review and meta-analysis suggested that omega-3 supplements play a role in improving cognitive development in infants. Although the research summarized in those studies indicated a functional role of DHA in brain health and an effect on attention domain, additional research is needed to further define DHA's role in cognitive function for groups other than infants.*

DHA is also required for the functional integrity of retinal pigment epithelium (RPE) cells and may play an ongoing role in eye health and function throughout life. [10,16] Studies on human RPE cells suggest that NPD1 orchestrates cell-protective mechanisms (including inhibition of caspase-3 activation and COX-2 expression) and thus promotes a healthy response after cellular insult.*(16,17)

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Supplement Facts

Serving Size: 2 Softgels Servings Per Container: 15

	Amount Per Serving	%Daily Value
Calories	25	
Total Fat	2.5 g	3%⁺
Cholesterol	15 mg	5% [†]
MaxSimil® DHA Fish Oil Concent		**
Total Omega-3 Fatty Acids	1.58 g	**
DHA (docosahexaenoic ac		**
EPA (eicosapentaenoic aci	d) 236 mg	**

† Percent Daily Values are based on a 2,000 calorie diet. ** Daily Value not established.

Other Ingredients: Softgel (fish gelatin, vegetable glycerin, and purified water), GRAS enteric coating (ethylcellulose, sodium alginate, purified water, medium-chain triglycerides, oleic acid, vegetable stearic acid, and ammonium hydroxide), and natural mixed tocopherols.

Contains: Fish (anchovy and/or sardine and/or mackerel [sources of fish oil] and tilapia [source of fish gelatin]).

Manufactured using MaxSimil® fish oil. MaxSimil® is a registered trademark of Ingenutra Inc. Protected under US patents 8,119,690 and 8,198,324; Canadian patents 2672513 and 2677670.

Directions

Take one to two softgels daily, or as directed by your healthcare

Consult your healthcare practitioner prior to use. Individuals taking medication should discuss potential interactions with their healthcare practitioner. Do not use if tamper seal is damaged.

Eicosanoid Metabolism, Heart Health, and Immune Health

In addition to the functional effects discussed above, research has established that omega-3 fatty acids antagonize arachidonic acid-induced eicosanoid formation. They also help generate resolvins and protectins known as specialized pro-resolving mediators (SPMs)-EPA and DHA metabolites that are naturally produced in vivo through enzymatic conversion of EPA and DHA-to promote a healthy response to the arachidonic acid cascade. Omega-3s have also been widely researched for their promotion of cardiovascular health, including maintaining healthy lipid levels already within the normal range and supporting normal resistance to oxidative stress.[10] Although not excluding similar benefits of EPA and docosapentaenoic acid (DPA), DHA may have a direct or indirect benefit of keeping the heart rhythm healthy and consistent. In multivariable-adjusted cross-sectional analyses among older adults (N =~3000), plasma phospholipid DHA—but not EPA or DPA—was inversely associated with resting heart rate and a trend toward lower blood pressure (BP).[18] In one short trial, DHA reduced 24-hour ambulatory BP and heart rate[19]; in another small trial (n = 59), the effect was not seen. Researchers speculated that statistical power may have been limited by the size of the latter study.[20] This may have been the case since in another, larger study (N = 224) in healthy men receiving 4 g/ day each of EPA and DHA, lower heart rate and blood pressure was achieved. In this study, plasma phospholipid analysis revealed that only DHA was responsible for lowering heart rate. [21] Additionally, due to the preponderance of findings that EPA and DHA can help moderate blood pressure, the FDA has approved a qualified health claim with regard to this function. [22] Specifically for Omega MonoPure DHA EC: Research shows that consuming EPA and DHA combined may be beneficial for moderating blood pressure, a risk factor for CHD (coronary heart disease). However, the FDA has concluded that the evidence is inconsistent and inconclusive. One serving (2 softgels) of Omega MonoPure DHA EC provides 1.58 grams of EPA and DHA.

Omega-3 fatty acids may also have immunomodulatory effects.[10] It has been suggested in several human studies that supplementing with a DHA-rich oil appears to decrease the activation of T lymphocytes and mononuclear cells and decrease levels of inflammatory mediators. [23,24] The immune-altering effects may be due to the composition of the cell membrane as well as to the activity of the DHA-derived resolvins.*[25]

References

- 1. Unpublished, internal data. Ingenutra.
- 2. Fortin S, inventor; Centre de Recherche sur les Biotechnologies Marines, assignee. Compositions comprising polyunsaturated fatty acid monoglycerides or derivatives thereof and uses thereof. US patent 8,198,324. June 12, 2012.

 3. Brunet S, Chamoun R, Fortin S, et al. MaxSimil®: A novel, patented natural platform for enhanced absorption of omega-3s. Single dose, double-blind, 2-way crossover pilot pharmacokinetic study on healthy subjects
- under normal diet. Sherbrooke (Québec), Canada: Ingenutra; 2018. [Unpublished]
 4. Singh M. Essential fatty acids, DHA and human brain. Indian J Pediatr. 2005 Mar;72(3):239-42. [PMID: 15812120]
- 5. Guesnet P, Alessandri JM. Docosahexaenoic acid (DHA) and the developing central nervous system (CNS) Implications for dietary recommendations. *Biochimie*. 2011 Jan;93(1):7-12. [PMID: 20478353] 6. Uauy R, Dangour AD. Nutrition in brain development and aging: role of essential fatty acids. *Nutr Rev*. 2006 May;64(5 Pt 2):S24-33; discussion S72-91. [PMID: 16770950]
- 7. Drover J, Hoffman DR, Castañeda YS, et al. Three randomized controlled trials of early long-chain polyunsaturated fatty acid supplementation on means-end problem solving in 9-month-olds. Child Dev. 2009 Sep-0ct;80(5):1376-84. [PMID: 19765006]
 8. Shulkin M, Pimpin L, Bellinger D, et al. n-3 fatty acid supplementation in mothers, preterm infants, and term infants and childhood psychomotor and visual development: a systematic review and meta-analysis. J Nutr.
- 2018 Mar 1;148(3):409-418. [PMID: 29546296]
 9. Birch EE, Carlson SE, Hoffman DR, et al. The DIAMOND (DHA intake and measurement of neural development) study: a double-masked, randomized controlled clinical trial of the maturation of infant visual acuity as a
- 9. Biol E2, Carlson SE, Indiminal on, et al. The Dishrotory Orbit make and ineasurement or neural developments study, a double-masked, randomized controlled clinical trial of the mediatation of limital visual acting function of the dietary level of docosahexaenoic acid. Am J Clin Nutr. 2010 Apr;91(4):848–59; [PMID: 20130095]
 10. Ghasemi Fard S, Wang F, Sinciair AJ, et al. How does high DHA fish oil affect health? A systematic review of evidence. Crit Rev Food Sci Nutr. 2018 Mar 1:1-44. [PMID: 29494205]
 11. Bazan NG, Rodriguez de Turco EB, Gordon WC. Pathways for the uptake and conservation of docosahexaenoic acid in photoreceptors and synapses: biochemical and autoradiographic studies. Can J Physiol Pharmacol. 1993 Sep;71(9):690-8. [PMID: 8313233]
- 12. Cole GM, Ma QL, Frautschy SA. Dietary fatty acids and the aging brain. Nutr Rev. 2010 Dec;68 Suppl 2:S102-11. [PMID: 21091943]
 13. Muldoon MF, Ryan CM, Sheu L, et al. Serum phospholipid docosahexaenoic acid is associated with cognitive functioning during middle adulthood. J Nutr. 2010 Apr;140(4):848-53. [PMID: 20181791] 14. Jiao J, Li Q, Chu J, et al. Effect of n-3 PUFA supplementation on cognitive function throughout the life span from infancy to old age: a systematic review and meta-analysis of randomized controlled trials. Am J Clin Nutr. 2014 Dec;100(6):1422-36. [PMID: 25411277]
- 15. Lukiw WJ, Bazan NG. Docosahexaenoic acid and the aging brain. J Nutr. 2008 Dec;138(12):2510-4. [PMID: 19022980]
 16. Bazan NG. Neuroprotectin D1 (NPD1): a DHA-derived mediator that protects brain and retina against cell injury-induced oxidative stress. Brain Pathol. 2005 Apr;15(2):159-66. [PMID: 15912889]
- 17. Bazan NG, Calandria JM, Serhan CN. Rescue and repair during photoreceptor cell renewal mediated by docosahexaenoic acid-derived neuroprotectin D1. J Lipid Res. 2010 Auc;51(8):2018-31. [PMID: 20382842] 18. Mozaffarian D, Lemaitre RN, King IB, et al. Circulating long-chain ω-3 fatty acids and incidence of congestive heart failure in older adults: the cardiovascular health study: a cohort study. *Ann Intern Med.* 2011 Aug;155(3):160-70. [PMID: 21810709]
- 19. Mori TA, Bao DQ, Burke V, et al. Docosahexaenoic acid but not eicosapentaenoic acid lowers ambulatory blood pressure and heart rate in humans. Hypertension. 1999 Aug;34(2):253-60. [PMID: 10454450] 20. Woodman RJ, Mori TA, Burke V, et al. Effects of purified eicosapentaenoic and docosahexaenoic acids on glycemic control, blood pressure, and serum lipids in type 2 diabetic patients with treated hypertension. Am J
- Clin Nutr. 2002 Nov;76(5):1007-15. [PMID: 12399272]
 21. Grimsqaard S, Bonaa KH, Hansen JB, et al. Effects of highly purified eicosapentaenoic acid and docosahexaenoic acid on hemodynamics in humans. Am J Clin Nutr. 1998 Jul;68(1):52-9. [PMID: 9665096]
 22. FDA Announces New Qualified Health Claims for EPA and DHA Omega-3 Consumption and the Risk of Hypertension and Coronary Heart Disease [update]. Silver Spring, MD: Center for Food Safety and Applied Nutrition Constituent Update, US Food and Drug Administration; June 19, 2019. https://www.fda.gov/food/cfsan-constituent-updates/fda-announces-new-qualified-health-claims-epa-and-dha-
- omega-3-consumption-and-risk-hypertension-and. Accessed July 18, 2019.
 23. Kew S, Mesa MD, Tricon S, et al. Effects of oils rich in eicosapentaenoic and docosahexaenoic acids on immune cell composition and function in healthy humans. Am J Clin Nutr. 2004 Apr;79(4):674-81. [PMID:
- 150516141 24. Vedin j, Cederholm T, Freund Levi Y, et al. Effects of docosahexaenoic acid-rich n-3 fatty acid supplementation on cytokine release from blood mononuclear leukocytes: the OmegAD study. Am J Clin Nutr. 2008 Jun;87(6):1616-22. [PMID: 18541548]
- 25. Calder PC. The relationship between the fatty acid composition of immune cells and their function. Prostaglandins Leukot Essent Fatty Acids. 2008 Sep-Nov;79(3-5):101-8. [PMID: 18951005]

Does Not Contain

Wheat, gluten, corn, yeast, soy protein, dairy products, shellfish, peanuts, tree nuts, egg, ingredients derived from genetically modified organisms (GMOs), artificial colors, artificial sweeteners, or artificial preservatives.

> *These statements have not been evaluated by the Food and Drug Administration. This product is not intended to diagnose, treat, cure, or prevent any disease.

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