Omega-3 as an Anti-Inflammatory Modality: Literature Review

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Abstract

Omega-3 fatty acids are polyunsaturated fatty acids that have multiple double bonds, the first double bond is located on the third carbon atom of the omega-methyl group, the next double bond is located at the third carbon atom from the previous double bond. The methyl omega group is the last group in the fatty acid chain. Omega-3 fatty acids are nutrients that play a vital role in the growth and development process of brain neuron cells for the intelligence of the born baby.

1. Introduction

These omega-3 fatty acids are derived from their precursors, namely linoleic and linolenic essential fatty acids. Essential fatty acids cannot be formed in the body and must be supplied directly from food.¹⁴ Then the precursors enter the elongate and desaturate processes which produce three forms of omega-3 fatty acids: LNA (alpha-linolenic acid (C 18: 3, n -3)), EPA (eicosapentaenoate (C20: 5, n- 3)), and DHA (docosahexaenoate (C22: 6, n-3)).⁵⁻⁸

The parent of omega-3 fatty acids is alpha linolenic acid (ALA). ALA with the help of the delta-6-desaturase enzyme can be converted into stearidonic acid, then by the delta-5-desaturase enzyme the body is converted to eicosapentaenoic acid (EPA) and by the delta4-desaturase enzyme it is converted to docosahexaenoic acid (DHA).⁹⁻¹⁰ The process of making DHA and AA facilitated by the enzymes desaturase and elongase. The activity of these two enzymes is still lacking in premature infants, even in term infants up to the age of 4-6 months. Therefore, the addition of DHA and AA in premature infants is highly recommended at a dose that refers to the fatty acid content in breast milk.¹¹

The activity of both desaturase and elongase enzymes is influenced by the fatty acids found in food. Fish oil that contains a lot of DHA will inhibit the activity of this enzyme so that it can inhibit the formation of AA.¹² In contrast, corn oil or safflower stimulates the activity of the desaturase enzyme, thereby increasing the formation of AA.¹¹ The physical and chemical properties of the metabolism, digestion, absorption and secretion of omega-3s are the same as those of fat.¹³⁻¹⁵

The physical properties of triglycerides are determined by the proportion and chemical structure of the fatty acids that make up them. The more short-chain fatty acids and unsaturated bonds there are, the softer and more fluid the fats are. Preferably, the more long-chain saturated fatty acids, such as plammic acid (C1 6: 0) and stearic acid (18: 0) that are found in animal
fat, the denser the fat. Triglyceride properties are also determined by the position (omega) and the position of fatty acids on the glycerol molecule.\textsuperscript{16,17}

**Classification, source of omega-3 and nomenclature**

Fatty acids are distinguished according to the number of carbon they contain, namely short chain fatty acids (6 carbon atoms or less), medium chain (8 to 12 carbon), long chain (14 - 18 carbon), and very long chain (20 carbon atoms or more).\textsuperscript{18} Essential fatty acids actually consist of linoleic acid (AL) / "linoleic acid" (LA), linolenic acid (ALN) / "- linolenic acid" (ALA) and arachidonic acid / "arachidonic acid" (AA), fatty acids it cannot be made by the body either from other fatty acids or from carbohydrates or amino acids. Arachidonic acid can be made from linolenic acid (n-6 series), therefore only linolenic fatty acids and linoleic fatty acids are considered as essential fatty acids.\textsuperscript{19-21}

These two essential fatty acids cannot change from one another and differ both in metabolism and function, even physiologically they both have opposite functions, contain one double bond, whereas polyunsaturated fatty acids contain two or more double bonds. The classification of fatty acids according to the length of the carbon chain and the level of saturation in fat which is abundant in nature can be seen in table 2.3.\textsuperscript{22}

Sources of omega-3 fatty acids and the amount of omega 3 content in food, namely: Mackerel (2.5 gr); Herring (1.7 gr); Salmon (1.2 gr) Crustacean / lobster (0.2 gr); Squid (0.6 gr); Salmon oil (19.9 gr); Cod liver oil (18.5 gr); Herring oil (11.4 gr). The need for omega-3 fats will produce fatty acids and cholesterol which are actually needed to form membrane cells in all organs. Important organs such as the retina and central nervous system are mainly composed of fat. Fatty acids that are needed by body tissues, especially essential fatty acids. Essential fatty acids are fatty acids that cannot be made in the body so they must be obtained from food, consisting of linoleic, linolenic and arachidonic acids.\textsuperscript{23-26}

Fatty acids consisting of carbon chains that hold all of the hydrogens that can be bound are called saturated fatty acids. Fatty acids that contain one or more double bonds to which a hydrogen atom can be added are called unsaturated fatty acids. WHO (1990) recommends that fat consumption as much as 15-30% of total energy needs is considered good for health. This amount meets the need for essential fatty acids and to aid in the absorption of fat-soluble vitamins.\textsuperscript{27-30}

Among the fats consumed in a day, it is recommended that at most 10% of total energy needs come from saturated fat, and 3-7% from polyunsaturated fat. The recommended cholesterol intake is <300 mg daily. The recommendations given by the FAO / WHO expert group regarding the consumption of saturated fatty acids, unsaturated fatty acids and cholesterol are: (1) the consumption of saturated fatty acids should not exceed 10% of the total energy (2) It is recommended that the consumption of linoleic fat contributes between 4-10 % of total energy. Consumption higher than this range is recommended if the consumption of saturated fat and high cholesterol, and (3) the consumption of cholesterol from food is recommended to be less than 300 mg / day.\textsuperscript{31-32}

WHO has set recommendations regarding the intake of omega 3 for each person, namely 0.3 -0.5 g / day (EPA + DHA). Which includes unsaturated fatty acids are: omega-3 (EPA & DHA), omega-6, AA, omega-9. Essential fatty acids are especially important for normal growth and development of fetuses and babies, as well as for brain development and vision.\textsuperscript{33}

**Benefits of Omega-3**

The advantage of omega-3 is very important for health, even most important among other fatty acids because it has anti-inflammatory and anti-clotting effects, is also good for the central nervous system and brain and can prevent CVD. The most abundant omega-3 fatty acids in fish are EPA and DHA. Consuming fish regularly can prevent CVD. Omega-3 unsaturated fatty acids play an important role in the morphological, biochemical, and molecular development of the brain and other organs.\textsuperscript{34}

Lack of omega-3 fatty acids, caused by insufficient intake or due to diseases that reduce absorption, can inhibit brain development, physical health and environmental interactions, which have a strong effect on the formation of cognitive development. Prolonged omega-3 deficiency can be fatal. Lack of omega-3 fatty acids causes nerve and vision problems and can
interfere with the development of the nervous system. As a result, there may be disturbances in the immune system, memory, mental and vision.35-36

Excessive provision of fat can lead to obesity and heart disease and can even lead to malignancy, can increase cholesterol levels, LDL which can spur the occurrence of atherosclerosis and coronary heart disease. This really depends on the amount of energy that comes from fat, the composition of the fatty acids, the composition from lipoproteins, dietary fiber consumed, antioxidants, activity, and degree of health. Saturated fatty acids such as lauric, myristic, and palmitic acid can increase cholesterol levels and LDL levels, while giving polyunsaturated fatty acids can reduce cholesterol and LDL levels. Monounsaturated oleic acids do not increase LDL levels but can increase HDL lipoproteins.37,38

Omega-6 and omega-3 fatty acids act as precursors or raw materials for eicosanoid compounds, which are highly reactive compounds. The eicosanoid compounds produced by omega-6 and omega-3 fats are often different, if not opposite. Thus, because omega-6 and omega-3 fatty acids compete as eicosanoid precursors and also have different biological roles, the balance between these two fatty acids in the daily diet is very important.39

In premature babies who are underweight (2,500 g) and whose brain size is smaller than average. Because the number of neuron cells is also small, the baby can be deformed, have low quality and the process of brain cell growth and development is not normal or below optimal. Omega-3 fatty acids, EPA are also reported to play a role in preventing degenerative diseases since the fetus and during adulthood.40

When the fetus is in the womb, EPA is indispensable in the formation of blood vessel and heart cells. Meanwhile, as an adult, EPA functions to nourish blood, work mechanism of blood vessels and work of sacs that regulate blood circulation. Therefore, due to a deficiency of omega-3 EPA, you can be at risk of suffering from blood vessel and heart disease. The balance of the ratio of EPA, DHA, and AA in the blood of infants, adolescents, or adults can be used as an indicator to predict the risk of vascular system disorders and heart disease in the future.41

Preventive measures are highly recommended from an early age in order to avoid this degenerative disease. Food supply sources of omega-3, EPA, DHA, AA, and alpha-linolenic acid should be consumed in a balanced ratio. The ratio of consumption of omega 3: omega 6 is 1: 5 to 10:19. DHA is needed as a cup-forming element for the rhodopsin container, namely a vital compound for sensing and sending back signals received by the eye to the brain. Docosahexaenoic Acid (DHA) and Arachidonic Acid (AA) are nutritional elements which are also important in the growth and development of nerves in the brain and help form brain fat tissue (mylenisation) and maintain the interconnection of brain nerve cells, especially to influence brain development.42

DHA and AA are the largest components of long-chain polyunsaturated fatty acids (LC-PUFA), which are very important ingredients for the central nervous system organs. DHA is important for the formation of nerve tissue, while AA acts as a neurotransmitter and as an essential form of LC-PUFA fatty acids that must be added to food. Based on the results of research, supplementation of several fatty acids at an early age has shown improvement in the index of mental development and visual acuity but only at levels of 17 mg / 100 kkalDHA and 34 mg / 100 kkalAA.43

These levels are almost the same as the FAO / WHO recommendation for infant formula, which is based on average levels of breast milk throughout the world. The adequacy rate of DHA is 20 mg / kg BW / day. According to the POM, consuming excessive DHA and EPA can inhibit the formation of AA from linoleic acid, can suppress the activity of the cyclooxygenase enzyme that forms prostaglandins. Consuming excessive DHA can cause kidney damage as a result, the kidneys experience a decreased response to the inflammatory process so that the inflammation period is longer and there is a decrease in the production of enzymes that play a role in controlling kidney function.44

The consequences of consuming excess DHA have not been studied. However, these effects are believed by experts to persist and occur when children become adults. AA and DHA supply is needed, especially in the last trimester, post-birth and early childhood. Deficiency of these two types of essential fatty acids at birth is correlated with low body weight, small head circumference and low placental size as a result.
of which the development of the central nervous system and later cognitive abilities are affected. DHA deficiency is linked to attention deficit-hyperactivity disorder (ADD or ADHD) such as excessive physical activity, learning difficulties and lack of social skills.44

Table 1. Classification of Omega-3 and sources.

| General Nomenclature | Chemical term | Short Nomenclature | Source |
|-----------------------|---------------|--------------------|--------|
| Polyunsaturated Omega 3 Linoleic ** | Acid 9.12.15 - Octnearrieionic | 18:3(n-3/w-3) | Soybean oil, sprouts, wheat |
| Eicosapentaenoate / EPA | 5,8,11,14,17-Eikosapentaenoic acid | 20:5(N-3/w-3) | Certain fish oils (can be made from linoleic acid) |
| Docosahexanoate / DHA | 4,7,10,13,16,19 - 22: 6 (N-3 / w-3) Docosahexanoic acids | 22:6(n-3/w-3) | Breast milk, certain fish oil |

Figure 1. EPA and DHA as anti-inflammatory

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