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Is vegan diet advisable for children?

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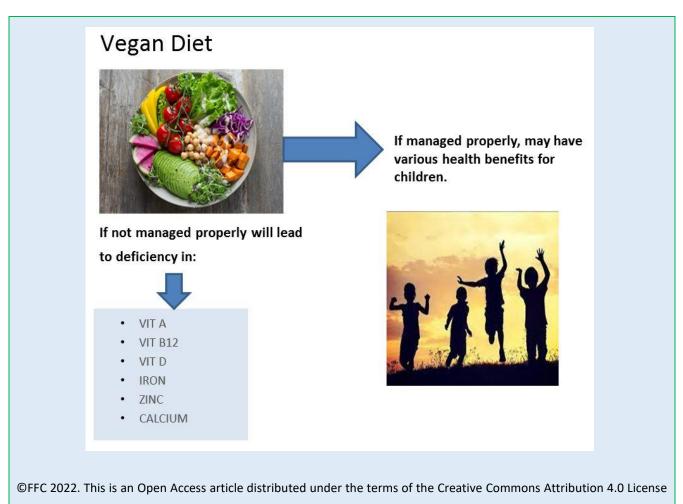
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ABSTRACT

Vegan diet (VD) is a diet that consists of only plant-based foods. This diet completely excludes all animal products; meat, fish, poultry, and food gotten from milk, oil, and honey. Vegan diet differs from vegetarian diet, although they are similar; some vegetarians include egg (ovo-vegetarians), milk from animals (lacto-vegetarians) or fish (pescovegetarian) in their diet. VD is believed to be healthier than diets containing animal produce and generally have a higher diet quality than non-vegan diets. VD has many health benefits and may also reduce the risk of certain conditions such as hypertension, diabetes, and cancer. However, these benefits seen in adults may not be the same case for toddlers and young children. Their diet directly affects their height, weight, and psychomotor and neurocognitive development. VD may not supply all the nutrients necessary for development and may lead to nutrient deficiency. Vegan children are at a risk of insufficient supply and deficiency of some critical nutrients such as protein, long chain fatty acids, cholesterol, rion, zinc, iodine, calcium, and vitamin A, B₁₂ and D. Deficiency of these nutrients could lead to various developmental and sometimes irreversible disorders. Apart from nutrition, VD also seems to alter the metabolomics and gut microbiota constitution of a vegan. Overall VD may have health benefits for children if it is properly fortified and supplemented. VD in children will also have to be carefully monitored to ensure the diet is healthy, nutritious and promote healthy growth and development.

Keywords: children, vegan diet, nutrition, metabolomics, inborn errors of metabolism



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INTRODUCTION

In a vegan diet, all food products obtained from animals including meat, fish, milk, egg, honey, and seafood are completely excluded [1]. The practice of this kind of diet is known as veganism and those who practice it are known as vegans [2]. Reasons for being a vegan maybe ethical, religious, or ecological [3] or to improve health, prevent non-communicable disease or could be as a result of general concern for welfare of animals [4-5]. It must be put into account that although vegan diet and vegetarian diet are basically almost the same, they have some differences. While vegans exclude all animal products from their diet, some vegetarians may choose to eat egg (ovo-vegetarians) or milk (lacto-vegetarians) from animals like goat and cow or both and egg from animals (lacto-ovo vegetarians). Some vegetarians may also have fish contained in their diet (pescovegetarian) [6]. Vegan diet is made up of plant sources such as fruits and vegetables, legumes such as peas, beans, and lentils, nuts and seeds, bread, cereals, olive, meat substitutes, potatoes, vegetable oils, breads, brown rice, pasta, and also dairy alternatives such as soymilk, coconut milk, and almond milk [7-9]. About 1%-5% of the populations in the western countries follow a vegan diet and this diet is common among people between 15-34 years [10-11].

There are different guidelines regarding vegan diets for children, some believe that a vegan diet lacks essential diets needed for growth and development in children while some suggest that a well-balanced vegan diet with adequate amount of vital nutrients like calcium, zinc, iron, supplemented with Vitamins B₁₂, D and polyunsaturated fatty acid docosahexaenoic acid (DHA) is appropriate for children [12-13]. The American Dietetic Association approves vegan diets for all age groups including children provided they are properly planned while the German Nutrition Society does not support children taking vegan diet [14]. According to Weder et al. [1], there was no significant difference in the rate of growth of children who consume vegetarian, vegan, and omnivore diets. Vegan diets are believed to be healthier than diets containing animal products and generally have a higher diet quality than non-vegan diets. Vegan diet has been found to have many health benefits and may also reduce the risk of certain conditions such as hypertension, type-2 diabetes, obesity, non-alcoholic fatty liver disease and cancer [15-16]. According to some data from the Seventh-Day Adventist Health Study, it was reported that vegan diet had positive effects against the development and progression of obesity, hypertension, diabetes, and cardiovascular related diseases [17]. Veganism has been widely recommended and accepted as a model of healthy diet associated with gut microbiota [18-19], cardiovascular diseases [20], diabetes, obesity, cancer, chronic kidney disease, and metabolic syndrome [21-23]. The higher fiber, low energy density, lower saturated fat, antioxidants, potassium, and sodium contents in vegan diet can contribute to a lower body mass index (due to the reduced protein intake). This is because the vegan diet is high in fiber, omega-6 polyunsaturated fatty acids but low in cholesterol, total fat and saturated fatty acids as compared with omnivorous diet [22]. With a lower cholesterol level, this further helps to regulate the blood pressure.

Diets rich in fruits, vegetables, and whole grains, which lead to an increase in blood antioxidant capacity, could reduce blood pressure in hypertensive patients. Due to the consumption of more fruits and vegetables, children who consume vegan diet have lower cholesterol levels, lower rates of obesity and overweight and high levels of antioxidants in their blood. Atherosclerosis begins in the early stage of life and gradually progresses into risk factors of cardiovascular disease in the adult life. Since vegan diet has been shown to reduce the risk factor of cardiovascular disease (CVD), this could potentially improve the cardiometabolic health from childhood to adulthood [24].

In a trial involving obese and hypercholesterolemic children, vegan diet in children was discovered to be more effective than the American Heart Associationrecommended diet in reducing the adult risk factors of CVD [24-25]. Vegan diet reduces high blood pressure via several mechanisms, such as improving blood viscosity, vasodilation and insulin sensitivity; by altering the baroreceptors, renin-angiotensin and sympathetic nervous system; by its anti-oxidant and antiinflammatory properties and by changing the colony and strain of gut microflora. Individuals who consume vegan diets have lower systolic blood pressure, diastolic blood pressure and reduced risk of hypertension compared to those who consume omnivores diet [26]. Vegan diet has also been discovered to be associated with increased lifespan and cancer protection and this is due to the low level of methionine found in this diet [27]. When compared with the omnivores diet, vegan diet has been discovered to be associated with lower bone mineral density which is linked with higher risk of bone fracture [28-29] and this could be because of the deficiency in nutrients and minerals like Vitamins D, B₁₂, A, iodine, zinc, folate, selenium, and calcium which are important for proper bone health [30-31]. Restricted intake of whole food groups in vegan children could however be of great concern due to the fact that their nutrient and energy needs are higher, and the growth might be impaired due to nutrient deficiency at some sensitive points of development [32].



Fig 1. Vegan diet [33]

It is also important to note that most of the studies supporting these benefits are observational. How the diet directly performs its actions and the mechanistic basis for these observations are unclear. The direct action of metabolism, in addition to systemic effects such as regulation of hormonal actions and antioxidation, serve as a link between diet and health [14].

Owing to differences in protein composition in these diets, amino acid intake and plasma levels may account for one of the main differences between vegan and vegetarian diets as some vegetarians may consume egg, milk from animal, fish and even meat occasionally.

Vegan Diet and The Gut Microbiota: The human gut microbiota is made up of numerous microorganisms like viruses, bacteria, fungi and protozoa [34]. The bacterial composition and diversity can be altered continuously by diets, lifestyle habits, environmental factors, stress, infant transitions, use of probiotics, prebiotics and antibiotics, intestinal and metabolic diseases [19, 34-35]. High-fat, low-fiber and high-protein diets have been reported to increase intestinal inflammation by modifying the translocation of bacterial populations and metabolites involved in modulating inflammatory response [36]. An imbalance in the activity and composition of the microorganisms in the gut is referred to as 'gut microbiota dysbiosis' [37] and it is associated with disorders like chronic kidney, hepatic and gastrointestinal diseases (Ulcerative colitis, Crohn's disease), colorectal cancer, allergy, autoimmune disease, obesity, type 2 diabetes and CVD [34,38-39].

Long term nutritional patterns can change both function and diversity of the gut microbiota and fibers, fats and proteins are commonly involved in the metabolic pathways in the gut microbiota [40-41]. Vegan diets are sources of nutrients for microorganisms while an omnivorous diet greatly alters the human gut microbiota which is made up of bile-tolerant potentially harmful microorganisms due to the fact that omnivores diet has increased levels of fecal bile acids [35]. These bile acids alter the composition of the gut microbiota through metabolic and inflammatory pathways [42-43].

Dietary fibers serve as substrates for bacterial metabolism in the intestine and the end products of the metabolism such as short chain fatty acids (SCFAs), which play a major role in immunoregulation [34,44], providing anti-inflammatory activities in the intestine [45], improving glucose tolerance, blood lipid profiles and insulin sensitivity, absorption of water and sodium, inhibiting cancer cell proliferation [37-38, 46], reduction of body weight [47] and also protection from inflammatory bowel disease, type 2 diabetes and immune diseases [19].

Vegan diet which is low in fats contains monounsaturated and polyunsaturated fats thereby altering the intestinal microbial composition [44] while animal saturated fats promote inflammation leading to metabolic disorder [19] and this has been discovered to be the driving force in CVD as a result of high levels of total serum cholesterol and LDL [45].

Dietary proteins in vegan diet cause an increase in the level of SCFAs in the intestine while animal proteins are involved in inflammatory bowel disease and are also associated with CVD [45].

Vitamins like vitamin K, biotin, cobalamin, riboflavin, thiamin are involved in bacterial metabolism and can be synthesized by the gut microbiota [48].

Metabolic Profile of Vegan Diet In Children: Adequate knowledge of the metabolic concerns of a strict vegan diet is still limited [49]. Children who consume vegan diet have been discovered to have a unique metabolic profile characterized by modification in biosynthesis of bile acids including increased levels of unconjugated primary bile acids and reduced taurine to glycine conjugation ratio of bile acid. It is also unknown whether the differences in the bile acid biosynthesis have any impact on endocrine functions, digestion, absorption and gut microbiome [14]. Low levels of polyunsaturated fatty acid docosahexaenoic acid (DHA), HDL-C and LDL-C, total cholesterol, variations in the level of circulating amino acids, low levels of vitamins A and D, low level of circulating leucine/isoleucine, phenylalanine, valine/betaine, and aspartate and higher levels of alanine, arginine, and glycine and lower protein intake were also observed [50].

A study was conducted in Finland to assess the difference in the metabolomics of omnivores, vegetarian and vegan children and to check if vegan diet is sufficient to support normal growth and development of vegan children. The aim of this exercise was also to assess the effect of vegan diet in dietary management of childhood disorders. At the end of the study it was found that vegan children had a distinct metabolic profile from that of omnivorous children. This distinction was characterized by lower levels of circulating fatty acids, lower levels of cholesterol and low density and high density lipoproteins, change in circulating amino acids, almost devoid of DHA and EPA and also lower levels of vitamin A and D. In the study, the low sufficiency of vitamin A and even lesser DHA could be a cause of concern in the visual health of vegan children. Vegan children also had higher folate intake and serum concentration than omnivorous children while some vegan children had higher folate levels than the recommended range. Although there have not been much adverse effects of high folate and low vitamin B12 levels as seen in some vegans, this might nevertheless pose a problem in neurocognitive health of the child [50].

The most striking alteration of pathway however was that of the bile acid. In vegan children, there were higher levels of unconjugated primary bile acids and significantly lower levels of conjugated primary bile acids as compared to omnivores. Although this is so, it is

uncertain how exactly this alteration affect the usefulness of bile acid in digestion and absorption [50]. Vegan diet has also been shown to influence the gut microbiota composition in vegans as compared to omnivores. Plant based diet like the vegan diet seems to be beneficial by enhancing the development of a more diverse and stable microbial system [19].

Nutrients of Interest: Children need proper nutrients to grow up and stay healthy and strong. The dietary requirements for children are quite different from adults, they require more nutrients and energy to ensure normal

growth and development of the brain, immune and endocrine systems [51]. Nutrition for children can also help establish a foundation for healthy eating habits and nutritional knowledge that children can apply throughout life. There are various nutrients required for health and growth of children. All the classes of food, which include; carbohydrates, protein, fat and oil, minerals and vitamins, have important functions in growth, health protection and energy metabolism and must be present during the growth of a child and if not present, will result in malnutrition or even death in worst cases. Below are some nutrients and their functions in the body:

Table 1. Nutrients of interest in vegan diet, their functions, and sources.

NUTRIENT	FUNCTION	SOURCES
PROTEIN	Protein which is made up of amino acids is necessary for growth. These amino acids are either essential or non-essential. Essential amino acid can't be made or stored in the body; therefore these amino acids have to be present in the diet. The nine essential amino acids are: histidine, isoleucine, leucine, lysine, methionine, phenylalanine, threonine, tryptophan, and valine. Essential and nonessential amino acids both produce energy and build proteins, and some form neurotransmitters and hormones. Amino acids such as tryptophan, tyrosine, histidine, and arginine are used by the brain for the synthesis of various neurotransmitters and neuromodulators [53].	Essential amino acids are found in animal sources in sufficient quantities while food such as vegetables, grains, and nuts, are lower in essential amino acids. But they can be combined to create complementary proteins that do provide enough essential amino acids [52].
CHOLESTEROL	Cholesterol can be biosynthesized in the body and ingested through diet. It is synthesized mainly by the liver and also other cells of the body as it has very essential biological functions. Some uses of cholesterol are: building of cell membrane which is highly essential in growing children, cell transporters and signaling molecules, production of hormones, assist in metabolism and production of vitamin D, production of bile acids necessary for digestion of fat and absorption of essential nutrients. It also serves as precursor for several biochemical pathways [54]. Cholesterol is carried around the body by low density lipoprotein (LDL) and high density lipoprotein (HDL). Although cholesterol has several biological functions, excess cholesterol may increase the risk of cardiovascular diseases.	Cholesterol in diet is mainly gotten from animal sources like meat, milk and egg [55].
CALCIUM	Calcium, the most abundant mineral in the body is the basic component of bone and needed for healthy bone and teeth in adult and particularly children. Calcium must be in optimum supply because children are still in the growth phase. Calcium plays a role in muscle movement, cardiovascular health (as it plays a key role in clot formation in blood) and is necessary for maintaining communication between the brain and other body parts [56].	Major sources of calcium include cheese, yogurt, milk and other diary food, fish. It can also be obtained from leafy greens and soybean [57].

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VITAMIN D	Vitamin D is a fat-soluble vitamin produced by the body after being exposed to sunlight. Vitamin D can also be taken through food but the body might not get sufficient amounts of vitamin D through diet alone. Vitamin D is necessary for maintaining healthy bones and teeth. It promotes the absorption of calcium and phosphorus by the body and it is important to maintain a proper immune system [58].	Vitamin D is necessary for maintaining healthy bones and teeth. It promotes the absorption of calcium and phosphorus by the body.
VITAMIN A	Vitamin A is another one of the four fat soluble vitamins. Vitamin A helps in the growth and maintenance of healthy teeth, skeletal and soft muscles, skin and also maintains the integrity of the mucus. Vitamin A promotes good eyesight especially when there is insufficient light [59].	There are two types of vitamin A present in food; Preformed vitamin A which is found in animal products like meat, beef liver, fish, poultry, egg, and dairy food. The most abundant natural source of vitamin A is cod liver oil which is pressed from the liver of cod fish. Provitamin A is found in plant sources such as fruit like carrots and minerals. Provitamin is also usually called beta carotene which is a powerful antioxidant and can alleviate the risk of cancers [60].
IODINE	lodine is a trace element necessary for the production of thyroid hormones T3 and T4. lodine is needed for proper growth and also for proper development of the brain. It also aids the regulation of body temperature, metabolism and heart rate [61].	The most abundant sources of iodine are egg, seaweed, fish such as cod and tuna, shrimp, and basically mostly seafood. Iodine can also be found in iodized salt, dairy products such as milk, yogurt and cheese [62].
ZINC	Zinc is a trace mineral associated with very numerous cellular functions in the body. It is required in the catalytic activity of many enzymes and plays a role in immune function, protein synthesis, wound healing, DNA synthesis and cell division [63]. Zinc must be provided through the daily diet since the body does not have a specialized zinc storage system.	The most abundant source of zinc in diet is oyster which contains high amount of zinc. Other food sources high in zinc are meat, poultry, crab, crab, clams, lobster, pork, beans, nuts, whole grains, fortified cereals and dairy products [64].
VITAMIN B ₁₂	Vitamin B_{12} is a water-soluble vitamin that plays a key role in the functioning of brain and nervous system, as well as formation of red blood cells. It is involved in the metabolism of every cell in the body. It helps to create and regulate DNA [65]. It also affects fatty acid and amino acid metabolism.	Sources of Vitamin B_{12} are beef, pork, ham, poultry, lamb, fish, dairy products, eggs and fortified cereals. Vitamin B_{12} is found in plants but an unrealistic amount needs to be consumed in order to meet the body requirement [66].
FOLATE	Folate is essential for thymidine synthesis and thus for DNA formation and any cell division hence its role in the formation of red blood cells and healthy cell growth and function [67]. It is also an important nutrient involved in bone protection [69-71].	Main sources of folate include dark green leafy vegetables, beans, peas and nuts. Fruit like oranges, lemons, banana, melons and strawberries are also rich in folate [68].
IRON	Iron is a trace mineral needed for proper formation and functioning of hemoglobin [72].	The best source of iron which is bioavailable are beef, liver, canned clams, ground beef, chicken, mussel, fortified cereal, white beans, dark chocolate, oyster, tofu, chickpeas [73].

Impact of Vegan Diet in Children Nutrients and Metabolomics: Plant-based diet is becoming more prevalent in recent times especially in European and western countries. Families or individuals who choose to adhere to vegan diet may do so for various reasons. Choosing vegan or vegetarian diet may be due to ecological, ethical health related or religious reasons [74]. This diet has been shown to have many positive effects on health in adults, and even children could benefit from this diet, however, veganism in children must be paid close attention, as there are many factors to consider [15]. Adults require different set of nutritional values compared to children because children require more energy and nutrients per body weight [75]. The more restrictive the diet and the younger the child is, the greater is the risk of nutritional deficiency [76].

Toddlers, children and teens who are still growing and developing require nutrient in sufficient qualities to ensure proper growth and development, in not only height and weight but also neurocognitive and psychomotor development. The influence of diet on the gut microbiota can also impact the emotional, epigenetic development and cognitive aspects of an individual [77-78]. Proper development of endocrine, neural and immunological systems need to be considered [79]. Vegan diet may not provide all the necessary nutrients needed for growth and development of a child which may lead to deficiencies if not properly checkmated.

Nutritional Composition of a Vegan Diet Cholesterol: Cholesterol, saturated fatty acids, LDL and HDL altogether are low in vegan diet. Vegan diet has very low amount of cholesterol due to the fact that most cholesterol in diet is consumed from animal source. This results in various health benefits such as reduced body fat, reduced risk of heart disease, reduced risk of blood pressure to name a few. This is also true for children however; children are still growing and cholesterol has many roles in the growth of children. As mentioned above cholesterol is needed in formation of cell membrane and is also a precursor for several steroidal hormones. Progesterone, glucocorticoid, mineralocorticoid, androgens and estrogens are produced from cholesterol [80]. Although cholesterol is produced in the body and isn't necessarily needed from diet, cholesterol is needed in higher levels in children development therefore children may need cholesterol from diet. Therefore, although not as much as animal sources, plant sources of cholesterol such as nuts and seeds, olive oil and coconut oil must be included in the diet of vegan children [81].

Polyunsaturated fatty acid (DHA and EPA): EPA (Eicosapentaenoic acid) and DHA (Docosahexaenoic acid) are omega-3 fatty acids. They are needed in infants for proper development of brain, eyes, and nervous system. It also has been identified to ease depression and improve heart functioning. Vegan diet is very low in EPA and DHA due to the fact that they are majorly found in oil fishes like cod liver, herring, mackerel, salmon, menhaden and sardine. Though the body produces EPA from an essential fatty acid known as alpha-linolenic acid (ALA), the efficiency of conversion of ALA to EPA is much lower than obtaining EPA and DHA from the diet [82]. This is because EPA is a precursor of DHA. Therefore, maintaining the levels of EPA and DHA in the body will be more difficult because of the extra metabolic work needed for the body to produce EPA then use part of it to metabolize DHA. It is advisable that vegans consume food rich in ALA such as flaxseed, walnut, canola oil, soy products and hemp seed. EPA and DHA can also be derived from seaweed algae and omega-3 supplements [83].

Energy and essential amino acids: Vegan children have lower protein intake when compared with omnivorous children. The percentage of energy from protein is considerably lower which is usually portrayed in both vegan adults and children in form of lower body mass and lower BMI although it is still in normal range [84]. A study has found that vegan children tend to be smaller than non-vegan children, and have a deficit in calorie intake [1]. Another study shows that there is an overall reduced level of circulating essential amino acids in the blood pool. It is unclear whether lower levels of several essential amino acids could have adverse or beneficial effects as this has not yet been fully studied and there are no specific guidelines for healthy blood levels of individual amino acids especially for infants and children [50]. In some vegan children, growth velocity was discovered to be lower compared to the omnivorous children although not statistically significant. The vegan children also had similar levels of Insulin-growth factor-1 (IGF-I) and Insulin-like growth factor binding protein-3 (IGFBP-3) with the omnivorous children [85]. Vegans get protein in their diet from beans, soy beans, lentils, tofu, hempseed etc. Several of the vegetable products are characterized by low digestibility; plant cells and some substances like tannins, enzyme inhibitors, phytates can inhibit digestibility of protein [86]. Appropriate combination of vegan food can provide all the essential amino acids needed by the body [87]. In a study by Weder et al. [1], there was no difference in the total energy intake between children on vegan and omnivores diet, with the omnivorous children getting their caloric intake particularly from protein, added sugar and fats while the

Bone health: The vegan diet may cause an intake of calcium below recommended levels. In a study by Weikert et al. [88], the rate of calcium excretion and

vegan children got theirs mainly from fibers.

serum level of parathyroid hormones (PTH) in vegans were measured. A low rate of calcium excretion and increased PTH level were observed which is the physiological result of low calcium intake. Vegan toddlers fed on breast milk may not experience the problem of not getting their calcium requirements because the calcium from the mother's bones will enrich the breast milk [89]. Lower calcium levels were observed in vegan infants not exclusively breastfed when compared with omnivores infants. Therefore, infants who are not breastfed should not be administered with inadequate plant-based food because the majority of them will lack sufficient calcium supply [90]. As they grow older it will be imperative to monitor their calcium intake and make sure they get adequate calcium supply because it is necessary for the healthy growth of their bones and is needed at higher levels during development [91]. Vegan adults and vegan children alike have higher risks of bone fractures due to lower intake of calcium from their diet. The calcium in their diet also faces the problem of bioavailability. Some sources of bioavailable calcium for vegans are Choy, Chinese cabbage, kale and collard [92]. Vegan diet has been discovered to prevent the children from attaining ideal height or bone mineral status [32]. Consuming nondairy drinks enriched with calcium is recommended for vegan children in order to derive a good level of calcium in their body [93].

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Fiber: Vegans usually tend to consume plenty amounts of fiber. Consumption of fiber might be beneficial because it aids bowel movement and can make up the bulk of food and pass through the system with little to no absorption by the body. However excess consumption of fiber especially in vegans will cause loss of nutrients because the fiber might hinder the absorption of nutrients such as fats, iron, and calcium from food [94]. Therefore, it is imperative that less fiber is consumed in the diet for

vegan children. Due to the high fiber content in vegan food, there is premature satisfaction and fullness which can lead to an insufficiency in the energy uptake especially in children [95].

Iron: Sources of iron in vegan diet is generally not as bioavailable as non-vegan sources and this can lead to a lower iron status and this was observed in a study by Desmond et al. [32] involving children who consume vegan diet. Bioavailability of non-heme iron is also reduced due to the presence of phytates and dietary fiber in vegan diet. If the iron is not sufficiently absorbed this may lead to an iron deficiency. Non-heme iron found in plant food [96]. This however might not pose a problem because vegans often consume large amounts of vitamin C rich foods and vitamin C improves the absorption of non-heme iron. In addition to consuming vitamin C rich foods, vegans can also consume calcium fortified food and supplements [89].

Zinc: Vegans are often considered to be at risk for zinc deficiency. Zinc aids the immune system and is a cofactor necessary for the proper functioning of numerous enzymes such as carbonic anhydrase, phosphatases, nucleases, peptidases and many others in the body. Zinc is an essential trace element and needs to be provided in the diet, however, phytic acid found in beans, legumes, seeds, nuts and grains bind the present zinc and inhibit the absorption. Therefore, zinc is not easily absorbed from plants sources while animal sources supply about half of the zinc intake in omnivores [89]. For vegan children this might pose a serious problem on their wellbeing as their immunity might not be as strong as that of adults. Therefore, extra attention needs to be paid to zinc serum levels in vegan children. However, despite the low zinc availability to the body, vegans do not necessarily

show reduced immunological competence. This suggests that the body may have a compensatory mechanism for zinc absorption, but this is yet to be explored. Low plasma zinc levels could however lead to iron deficiency due to its involvement as a catalyst in iron metabolism [97].

Vitamin B2: Large quantities of Vitamin B2 are found in foods from animal origin, while the supply of this vitamin from plant-based food is low, as observed in some studies conducted in Finland, France and Germany (Berlin) where it was discovered that the amount of vitamin B2 absorbed in vegans is lower compared to omnivores because the absorption of vitamin B2 from plant-based food is low [88, 98-99].

Vitamin D: Vitamin D is essential in regulating calcium and phosphate metabolism. Vegans are at higher risk of Vitamin D deficiency. Vitamin D occurs in two forms: vitamin D2 (ergocalciferol) and vitamin D₃ (cholecalciferol). Vitamin D₂ found in plant sources is less bioavailable than vitamin D_3 . Vitamin D_2 can also be produced by ultraviolet radiation in plants [87]. Vitamin D₃ is obtained from animal or plant sources [100]. The body also produces vitamin D₃ from cholesterol. Cholecalciferol is synthesized by the body through exposure to sunlight which stimulates the conversion of 7-hydrocholesterol to vitamin D_3 in the skin [101]. Although the body can synthesize its own vitamin D₃, it is important that it is present in the diet as the amount produced by the body may not be sufficient. It is essential for vegan children to have sufficient vitamin D supply because it is necessary for bone maintenance. Lower bone mineral content was observed in vegan children when compared with omnivore children and this could result from their low calcium, protein, vitamin D and vitamin B₁₂ intake [32]. Vitamin D can also be obtained from food supplements although, not all vitamin D₃ supplements are acceptable by vegans as most of them are made from cholecalciferol derived from lanolin which is extracted from sheep's wool. In a study by Hovinen et al, [50], low vitamin D status was reported in vegan Finnish children when compared with children consuming omnivores diet despite the daily intake of Vitamin D supplements.

Vitamin B₁₂ (cyanocobalamin): Vitamin B₁₂ deficiency is one of the greatest problems occurring in veganism. This is because sufficient vitamin B12 can only be obtained from animal sources and is produced solely by microorganisms in the herbivorous animals [30]. Plants do not have reliable sources of Vitamin B₁₂ [102]. Vitamin B₁₂ is bound to animal protein sources like meat. Upon reaching the stomach the stomach acid, HCL (Hydrochloric acid) separates vitamin B12 from the protein which it is bound to. After the separation, a protein secreted by the parietal cells of the stomach known as intrinsic factor, a glycoprotein, binds to the vitamin B₁₂ and activates it by removing the cyano group attached to the cobalt atom then it can be absorbed into the body. Children who practice veganism are at a very high risk of being affected by vitamin B_{12} deficiency because they need B₁₂ for brain development and also for proper formation of red blood cells [103]. The lack of vitamin B₁₂ leading to deficiency could bring about several problems such as growth, reflex and memory problems, tremors, numbness, dementia, depression and megaloblastic anemia characterized by improper formation of red blood cells. It could also cause permanent nerve and brain damage [104]. In some vegans among the Indian and Hong Kong population who rarely include supplements in their diets, high prevalence of vitamin B₁₂ deficiency have been reported [20]. In order to receive an adequate supply of Vitamin B₁₂,

supplements of Vitamin B₁₂ should be taken along with the vegan diet because inadequate vitamin B₁₂ status was observed in some children who consume vegan diet without supplementation [32, 105].

The problem of Vitamin B₁₂ deficiency is not only restricted to vegan toddlers and children. Even Veganism in pregnant mothers can affect the fetus if it is not getting enough Vitamin B₁₂ from the mother. The growth and development of the fetus might be impaired. Therefore, it is compulsory for vegan toddlers, children, pregnant women and adults to rely on fortified food like soy milk, cereals and also supplements [106]. Infants breastfed by vegan mothers have also been reported to have vitamin B₁₂ deficiency which could be accredited to the depletion of vitamin B₁₂ in their mother's body. Symptoms such as reduced growth rate, reduced bone mineral density, anorexia, weak muscles, delayed speech development, involuntary movement, enlarged liver and spleen, megaloblastic anemia are usually observed in those infants [107-108]. Impaired DNA synthesis and cell function is also associated with vitamin B₁₂ deficiency [109].

Vitamin B₁₂ is involved in the metabolism of methionine, fatty acids and amino acids, regulation of homocysteine concentration and formation of blood cells. Low serum level of vitamin B₁₂ is associated with high plasma level of homocysteine, and this is considered to be a risk factor of cardiovascular disease [110]. According to a study by Pawlak [111], the individuals on vegan diet had very low level of serum B₁₂, low status of Vitamin B₁₂ biomarkers like methylmalonic acid (MMA) or holotranscobalamin II (holoTC) and very high level of homocysteine (Hcy). Due to the deficiency of vitamin B₁₂ in vegan diet [110], vegans are on an increased risk of suffering from hyperhomocysteinemia.

Vegan Diet and Children's Immune System: Vegan diet has often been related to lowered levels of both white blood cells and red blood cells. The decline in protein and calories adversely affects the immune system. In growing children, nutrition directly affects the development of the immune system. Malnutrition, especially in vegan children will lead to an impaired immune system. However, nutrients that have critical roles in the maintenance of the immune function are best obtained from fruit, vegetables, nuts, whole grains and seeds which are the bulk of vegan diet. Also, the vegan diet is rich in phytochemicals which improve the immune function [112].

Despite the consistent reduced number of white blood cells found in vegans, the antioxidants provided by their food might be helpful in compensating the lack in leucocytes. Therefore, a very well planned and supplemented vegan diet may be beneficial to the immune system. Although there is a high antioxidant content in the vegan diet which may offer some immunity enhancing benefits, vegans don't necessarily exhibit a better immune system than omnivores [113].

Vegan Diet and Inborn Errors of Metabolism: Inborn errors of metabolism are rare disorders caused by inherited (genetic) factors or spontaneous mutations of genes necessary for the conversion of food into energy. Inborn errors of metabolism are heterogeneous and very diverse in nature [114]. Although specific inborn errors in metabolism are quite rare, they are more common when all different types are considered. Inborn errors in metabolism have been observed in 1 out of 2500 births [115]. It is imperative to consider the impact of vegan diet on the infants and children with these conditions.

Vegan diet in children can represent a new means to develop various strategies for dietary management of childhood disorders.

Phenylketonuria: Phenylketonuria is an autosomal recessive inborn metabolic disorder characterized by the

inability of the body to metabolize phenylalanine due to the absence of its metabolizing enzyme phenylalanine hydroxylase [116]. This leads to hyperphenylalaninemia which is the accumulation of phenylalanine in the blood, resulting in a series of health problems including retardation, poor skin pigmentation, growth impairment, convulsion and hypersensitivity [117]. In order to manage this condition, phenylalanine intake must be kept to the barest minimum. Phenylalanine is found mainly in high protein food such as meat, pork, chicken, egg and milk. Some other foods containing phenylalanine are beans, grains, nuts, fish, tofu, cheese and soybeans. Also, other food like soda containing aspartame, an artificial sweetener is made with phenylalanine. Consumption of low protein foods such as fruits, vegetable, juices, potatoes, peas, low protein bread and pasta will be more suitable for a child with phenylketonuria. Vegan diet may not be able to completely manage this condition but it serves as a good start.

Maple Syrup Urine Disease: Maple syrup urine disease is an inherited disorder characterized by the inability of the body to metabolize some amino acids properly. Isoleucine, leucine and valine cannot be metabolized due to the deficiency of the enzyme complex branched-chain alpha-keto acid dehydrogenase. This causes the accumulation of these amino acids in the body which leads to neurological problems, vomiting and dehydration. To avoid accumulation of these amino acids in the body, it is imperative to reduce the amount of them consumed in the diet. Foods that are high in protein should be avoided. Vegan diet is an advisable diet in managing this condition [118].

CONCLUSION

Vegan diet has a wide range of health benefits both in adults and children. It can be useful in managing some inborn errors of metabolism and other health conditions. However veganism in children must be closely monitored to prevent malnutrition. Depending on the severity of the malnutrition it could lead to impaired growth both bodily and cognitive, in worse cases it could even lead to death. In some cases of malnutrition owing to poorly planned vegan diet, the child was rushed to the hospital and diagnosed with vitamin D deficiency, anemia, failure to thrive and an overall poor health condition. There was also a case of a father and mother who lost custody of their child due to malnutrition because the child was fed with poor vegan diet.

This shows that for a child who is fed with vegan diet, the composition of the diet has to be well planned and accordingly supplemented with some vitamins, minerals and trace elements. Supervision of a nutritionist and pediatrician is advisable and also regular checkups to ensure that the vegan children are getting all their needs [119].

According to the Academy of Nutrition and Dietetics and also the American Dietetic Association, well-planned vegetarian and vegan diets supplemented appropriately are suitable for all life stages, but the German Nutrition Society include strong recommendations to parents that vegan diet should not be adopted by children without medical and dietetic supervision [120].

Also, if a child proposes to go on a vegan diet, the parent or guardian should find out why the child wants to go on a vegan diet. While some choose the vegan diet for a healthier diet and all its health benefits, some might not. Without knowing the reason for choosing a vegan diet, eating disorders like anorexia nervosa can easily be hidden by a vegan diet [121].

List of Abbreviations: DHA: Docosahexaenoic acid, VD: Vegan diet, CVD: Cardiovascular diseases, SCFAs: Short chain fatty acids, HDL-C: High-density lipoproteincholesterol, LDL: Low-density lipoprotein, ALA: Alphalinolenic acid, PTH: Parathyroid hormone, EPA: Eicosapentaenoic acid, MMA: Methylmalonic acid, holoTC: holotranscobalamin II, Hcy: Homocysteine, HCL: Hydrochloric acid.

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