

Chapter 9

Vitamins and Minerals: Types, Sources and their Functions



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9.1 Introduction

Nutrients are grouped into two major classes: macronutrients and micronutrients. Macronutrients are nutrients that are needed by the body in large amounts while micronutrients are those needed by the body in minute amounts. Macronutrients such as carbohydrates, proteins, and lipids provide molecules for the structural and metabolic activities of the human body, while micronutrients (vitamins and

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minerals) are essential for the body's proper functioning. The need for micronutrients depends on the metabolic activities as well as on the life cycle of an individual. Even in intrauterine life, the need for micronutrients is essential for the normal development of the fetus. In particular, vitamin D, iodine, iron, and folic acid deficiencies could lead to congenital disorders or even death. The daily requirements of these micronutrients are not fixed, although many scientific papers have mentioned the daily-required allowance of various vitamins and minerals. Factors such as physical exercise, pregnancy, childhood, adolescence, old age or specific diets (e.g. Vegan) influence the need for micronutrients. Therefore, the evaluation of the micronutrients' requirements and the consequences of micronutrients' deficiencies are critical to explain their role in health and disease (Derbyshire 2018; Hans and Jana 2018).

Fortification of food is one of the most effective and safe strategies used to enhance nourishment. For example, mother milk feeding can be considered a type of fortification, which is essential for the healthy growth of babies up to 2 years of age (Allen and Dror 2018). Micronutrients play an important role in reduction in the risk of disease and the maintenance of good health. Micronutrients and vitamins are vital for the proper functioning and proliferation of all dividing cells in the body. Therefore, micronutrients are essential for growth and metabolism. Balanced diet aids in maintaining good health by increasing the body's resistance to infection and, in the case of disease, can be a useful component of the therapy (Piccardi and Manissier 2009). Nutrients provide basic molecules which humans are unable to synthesize. Vitamins and minerals are required in small quantities (<100 mg/day) and are known as micronutrients (Ormsbee et al. 2014). The two important classes of micronutrients are shown in Fig. 9.1.

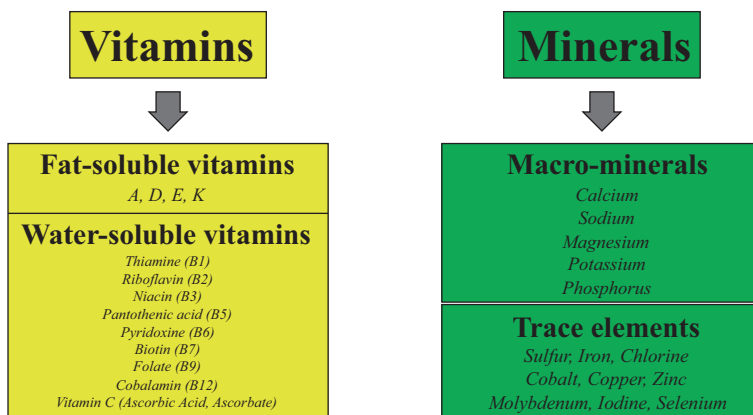


Fig. 9.1 Classification of micronutrients

9.2 Micronutrients: Health Benefits, Deficiencies and Risks

The human body needs micronutrients only in small quantities (in milligrams or micrograms) although they are as important as the macronutrients (Welch and Graham 2004). They are essential for the body's healthy development. Some vitamins for example work as cofactors or coenzymes in many metabolic processes. Trace elements like Zinc can help to improve immune function and minerals like Iron can prevent anemia among others (www.cdc.gov). As the body cannot synthesize micronutrients, therefore, they need to be supplied in the diet in adequate amounts. If their supply is not enough deficiency is possible and this can result to a myriad of diseases. Vitamin A deficiency for example can lead to blindness. A folic acid deficient pregnant woman can give birth to infants with neural tube defects. Iodine deficiency can lead poor brain fetal development. Micronutrient deficiency can increase the risk of infections and can lead to more micronutrient deficiencies (www.unicef.org). These are just the tip of the iceberg. Micronutrient deficiency may lie hidden and usually, the signs and symptoms develop late in the disease and are most often irreversible. The details of the impact of these micronutrients on health and diseases are discussed in the succeeding parts of the chapter.

9.3 Vitamins

Vitamins are essential substances for the normal functioning and development of the body. There are two classes of vitamins namely: Fat-soluble and Water-soluble vitamins (Table 9.1). The known vitamins include A, C, D, E, and K, and the B vitamins: thiamin (B₁), riboflavin (B₂), niacin (B₃), pantothenic acid (B₅), pyridoxine (B₆), cyanocobalamin (B₁₂), biotin, and folate/folic acid.

9.3.1 Fat-Soluble Vitamins

As earlier mentioned, vitamins can be classified as either a water-soluble or fat-soluble vitamins. The fat-soluble vitamins are very vital for the smooth functioning of the body. Their deficiencies have been implicated in several health disorders. The recommended daily allowance (RDA) of the fat soluble vitamins A, D, E, and K is 8000–1000 µg/day, 8000–5000 µg/day, 8–10 µg/day, and 70–140 µg/day respectively (Kamangar and Emadi 2012). The RDA for water soluble vitamins including thiamin, riboflavin, pyridoxine, niacin, biotin, ascorbic acid, and pantothenic acid are 1 mg/day, 1.2 mg/day, 2–2.2 mg/day, 13 mili-equivalents, 100–200 µg/day, 60 µg/day, and 4–7 mg/day respectively (Kamangar and Emadi 2012).

Table 9.1 Types, functions and sources of vitamins

Vitamins	Types/Subclasses	Functions	Sources	References
Fat Soluble Vitamins	A	Vision, growth and development, Immune function Reproduction, Red blood cell formation, Skin and bone formation.		
	Preformed Vitamin A		Animal products, such as human milk, glandular meats, liver and fish liver oils (especially), egg yolk, whole milk, and other dairy products.	Rodriguez-Amaya (2004)
	Pro Vitamin A		Red palm oil, green leafy vegetables (e.g. spinach, amaranth, broccoli), yellow vegetables (e.g. pumpkins, squash, and carrots), and yellow and orange non-citrus fruits (e.g. mangoes, apricots, and papayas), red pepper, sweet potatoes. Also found in Brazilian palm fruit known as <i>Buriti</i> and the Vietnam fruit known as <i>Gac</i> .	Booth et al. (1992) and Vuong (1997)
	D (active form: 1,25-dihydroxyvitamin D)	Maintains bone health, muscle and nerve contraction and and general cellular function in all cells of the body.	Made in the skin from a cholesterol like precursor (7-dehydrocholesterol) by exposure to sunlight or can be provided pre-formed in the diet. Diets such as Fortified milk, cheese, and cereals; egg yolks; salmon.	WHO/FAO (2004) and Feldman et al. (1997)
	E (α -tocopherol)	Antioxidant, formation of blood vessels and boosting of immune function	Good dietary sources of vitamin E include nuts, such as almonds, peanuts and hazelnuts, and vegetable oils, such as sunflower, wheat germ, safflower, corn and soybean oils. Also sunflower seeds and green, leafy vegetables such as spinach and broccoli contain vitamin E.	Bellizzi et al. (1994) and WHO/FAO (2004)
	K	Helps in blood clotting, bone metabolism and regulation of blood calcium levels.		Shearer et al. (1996), Booth et al. (1994), Booth et al. (1993), Conly and Stein (1992) and WHO/FAO (2004)

	K ₁ (Phylloquinone)		Green leafy vegetables like spinach, broccoli. Vegetable oils from soyabean, rapeseed and olive. Also found in peanut, corn, sunflower and safflower	
	K ₂ (Menaquinone)		Animal liver, fermented foods such as cheese, fermented soyabean (<i>Japanese natto</i>), intestinal microflora	
Water-Soluble Vitamins	B ₁ (Thiamine)	Cofactor for several enzymes involved in energy metabolism, plays central role in cerebral metabolism.	Beef, liver, dried milk, nuts, oats, oranges, pork, eggs, seeds, legumes and yeast.	Aviva (2010) and Makarchikov (2009)
	B ₂ (Riboflavin)	Converts carbohydrates into glucose for energy production, neutralizes free radicals hence acts as anti-oxidant	Plant foods and animal sources, namely poultry, meat, fish and dairy products such as eggs, milk and cheese. Green vegetables such as collard greens, turnips as well high quality protein-rich foods.	Pinto and Rivlin (2013)
	B ₃ (Niacin)	Helps lower LDL cholesterol, lowers risk of cardiovascular diseases, eases arthritis.	Animal foods such as lean red meat, poultry and liver. Pea butter is an excellent source of niacin. Other useful sources include whole grain cereals, bread tea, coffee, maize (sweet corn).	Lule et al. (2016)
	B ₅ (Pantothenic acid)	Pantothenic acid is a key component of CoA, a cofactor that carries acyl groups for many enzymatic processes, and of phosphopantetheine within acyl carrier proteins, a component of the fatty acid synthase complex.	Peanut butter, liver, kidney, peanuts, almonds, wheat bran, cheese and lobster. Vast majority of is already incorporated into Coenzyme A (CoA) and as phosphopantetheine	Gregory (2011)
	B ₆ (Pyridoxine)	Acts as a critical co-factor for a diverse range of biochemical reactions that regulate basic cellular metabolism.	Richest sources of vitamin B6 include fish, beef liver and other organ meats, potatoes, starchy vegetables and fruits.	Marcelina et al. (2018) and Juan et al. (2018)

Table 9.1 (continued)

Vitamins	Types/Subclasses	Functions	Sources	References
	B ₇ (Biotin)	<p>Biotin is a water-soluble vitamin and serves as a coenzyme for five carboxylases in humans.</p> <p>It helps to convert food into glucose, which is used for energy production. It also helps to produce fatty acids and amino acids (the building blocks of protein). Biotin activates protein/amino acid metabolism in the hair roots and fingernail cells.</p>	Biotin is found in egg yolk, almonds, sweet potatoes, mushroom, cauliflower, cheese, spinach.	Janos et al. (2009); www.nutri-facts.org/Vitamin_B7
	B ₉ (Folate)	Helps in DNA replication, metabolism of vitamins and amino acids, proper cell division. Folic acid helps to reduce risk of spina bifida (neural tube defects) in neonates when taken by pregnant mothers.	Occurs naturally in dark green leafy vegetables, spinach. Also found in liver, avocado pear, paw-paw fruits, orange, beans and nuts	Kunisawa et al. (2012)
	B ₁₂ (Cyanocobalamin)	Plays a significant role in cellular metabolism, especially in DNA synthesis, methylation and mitochondrial metabolism. It helps in brain function and synthesis of red blood cells.	Vitamin B12 is naturally found in foods including meat (especially liver and shellfish), eggs, and milk products.	Kelly et al. (2006)
	C (Ascorbic acid)	<p>One of the important properties of vitamin C is its antioxidant activity.</p> <p>Vitamin C functions in enzyme activation, oxidative stress reduction, and immune function</p> <p>Antioxidant activity of vitamin C helps to prevent certain diseases such as cancer, cardiovascular diseases, common cold, age-related muscular degeneration and cataract.</p>	Vitamin c is abundantly available in many natural sources, including fresh fruits and vegetables. The richest sources of ascorbic acid including Indian gooseberry, citrus fruits such as limes, oranges and lemons, tomatoes, potatoes, papaya, green and red peppers, kiwifruit, strawberries and cantaloupes, green leafy vegetables such as broccoli, fortified cereals and its juices are also rich sources of vitamin C. Another source of vitamin C is animals. They usually synthesize their own vitamin C and are highly concentrated in the liver part	Sudha and Raveendran (2017) and Schlueter and Johnston (2011)

9.3.1.1 Vitamin A

Vitamin A was the first fat-soluble vitamin identified in 1913. Beta-carotene is converted to vitamin A in the liver. It protects the eyes in case of infections and contributes to the vision in dim light. Vitamin A combines with the protein opsin to form rhodopsin in the retinal rod cells. When vitamin A levels are inadequate, the lack of rhodopsin makes it difficult to see in dim light (Gräslund et al. 2008). Vitamin A is also involved in the physiological function of the epithelia and glands, and in normal cell differentiation. Vitamin A supports bone growth and the immune functions. Being oil-soluble, it is prescribed as retinol palmate every 6 months to pre-school children. The prevalence of vitamin A deficiency is 33.3% in pre-school children worldwide, while in Africa it is estimated at 44.4%. In Ethiopia, 80,000 deaths occur due to vitamin A deficiency. In this country, vitamin A deficiency prevalence is listed at 61% in preschool children (Abrha et al. 2016). Vitamin A deficiency is linked to the following diseases:

Vitamin A insufficiency leads to visual impairment, xerophthalmia, Bitot's spots (triangular, frothy, harsh and raised patches seen on the bulbar conjunctiva), keratomalacia (softening of the thickness of the cornea), follicular hyperkeratosis, anorexia, growth retardation, respiratory and intestinal infections, and the degeneration of the myelin sheaths. The excess of vitamin A may cause nausea, vomiting, and anorexia. The functions of vitamin A in the body include: development and upkeep of the epithelial tissue, support of the skin, upkeep of visual sharpness (in diminishing light), bone development and immunity. Sources of vitamin A include: green leafy vegetables, germination cereals and pulses, meat, fish, green mango, papaya, pumpkin, yellow fruits and vegetables. Other sources are eggs, spinach, cabbage, carrots, amaranth, angle liver oils, cod liver oil, liver or halibut liver oil (Abrha et al. 2016; Gräslund et al. 2008).

9.3.1.2 Vitamin D

The best-known utility of vitamin D is to control the concentration of phosphorus and calcium in the blood. This anti-rachitic vitamin occurs in a number of forms. Four crystalline D vitamins are isolated and at least 10 pro-vitamins D are known. For humans, most of the vitamin D supply comes from animal sources or synthesized following sun exposure. Vitamin D₂ is formed by exposing ergosterol to ultraviolet light. Vitamin D₃ is the natural vitamin D found in fish oil and formed in the skin of man and animals following exposure to sunlight (Nair and Maseeh 2012). It may be formed by the irradiation of 7-dehydrocholesterol. Vitamin D₂ and D₃ are known as vitamers. Vitamin D enhances the absorption of phosphorus and calcium from the intestine and their deposition in bones. Vitamin D sources include exposure to sunlight, liver, eggs, butter, cheese, fish liver oil, fortified foods, milk, and margarine. Vitamin D also stimulates the normal mineralization of bones and increases the tubular reabsorption of phosphate. It also possesses antioxidant

properties (Christakos et al. 2015; Gaman et al. 2019). Ageing decreases the capacity of the skin to produce vitamin D₃. After the age of 70, vitamin D levels decrease to about 25% of the normal value in adults (Kennel et al. 2010). In the temperate zone, the cutaneous production of vitamin D₃ is limited by the reduced ultraviolet radiations availability. In the absence of solar exposure, about 400–600 IU of vitamin D seems to be required daily to maintain the normal bone metabolism. Deficiency of vitamin D can affect bone development. In children, vitamin D deficiency causes rickets, a disorder in which bones deteriorate and bend under pressure. In adults, vitamin D deficiency results in osteomalacia (soft bones) which enhances the risk of bones fractures (Kennel et al. 2010; Nair and Maseeh 2012; Christakos et al. 2015).

9.3.1.3 Vitamin E

Vitamin E is an anti-sterility factor and a natural antioxidant. Structurally related names of vitamin E include tocopherols or tocotrienols (Gheorghe et al. 2019). It is involved in the healing of wounds and in immunity. Sources of vitamin E include wheat germ oil, nuts, cereals, meat, eggs, milk, green leafy vegetables, and other vegetables. The deficiency of vitamin E is associated in humans with cystic fibrosis, ataxia and abetalipoproteinemia (disorder that interferes with the normal absorption of fat and fat-soluble vitamins from food) and with habitual abortions and testicular degeneration in laboratory animals. Also, vitamin E deficiency can lead to increased hemolysis of the red blood cells and macrocytic anaemia in premature infants. The effects of excessive vitamin E doses lead to interference with the utilization of vitamins A and K, prolonged prothrombin time, intestinal irritability, headache, fatigue, and dizziness (Schmölz et al. 2018; Birringer et al. 2019; Gomez-Pomar et al. 2018; Moisa et al. 2018; Kennic and Coleman 2019; Reddy and Jialal 2018).

9.3.1.4 Vitamin K

The major function of vitamin K is the formation of prothrombin in the liver along with other vitamin K dependent clotting factors namely: VII, IX, X, protein C and S which are essential for normal blood coagulation or blood clotting. It is found in fresh green leafy vegetables, lettuce, cabbage, egg yolk, soybean oil, and liver. Our body can also produce its own vitamin K courtesy of the normal bacterial flora of our intestines. Premature babies receive 0.5–1 mg vitamin K intramuscularly or 1–2 mg orally at birth to prevent Vitamin K deficiency. Vitamin K deficiency can result to generalized bleeding, the development of hemorrhagic disease of the newborn, and prolonged clotting time in adults. Excessive doses of vitamin K can lead to hyperbilirubinemia in infants and vomiting in adults (Reddy and Jialal 2018).

9.3.2 Water Soluble Vitamins

9.3.2.1 Thiamine (Vitamin B1)

Thiamine or vitamin B1 is water-soluble and acts as coenzyme known thiamine pyrophosphate. Thiamine pyrophosphate is involved in carbohydrate metabolism. Thiamine pyrophosphate is also involved in the hexose monophosphate shunt. It is a neuro-protective agent (Ikeda et al. 2016). Thiamine is found in nuts, potatoes, meat, beans, and cereals. Its deficiency leads to a disease known as beriberi. It is often observed in polished rice-eating communities because, in polished rice, the seed coat, which contains this vitamin, is removed. There are three types of beriberi: dry, wet and infantile (Romagnoli et al. 2012). Thiamine is a co-catalyst in sugar digestion and is necessary to the function of the heart, nerves, and muscles. Vitamin B1 sources include unmilled oat, wheat germ, beets, nuts, meat, lentils, potatoes, pork, eggs, poultry, dried beans, green peas, beans, green verdant vegetables. Thiamine insufficiency can cause beriberi, polyneuritis, mental disarray, ataxia or Wernicke-Korsakoff syndrome in alcoholics. Excess can lead to tachycardia, migraines or peevishness, a sleeping disorder (Wiley and Gupta 2019).

9.3.2.2 Riboflavin (Vitamin B2)

Riboflavin (or lactoflavin) is a yellow crystalline substance. Riboflavin is found in grain, milk, eggs, liver, oats, and green verdant vegetables. It is involved in tissue respiration. Its derivatives are FAD (flavin adenine dinucleotide in its oxidized state) and FADH₂ (FAD in its reduced form). FADH₂ gives two ATP in the electron transport chain. FAD and FADH₂ are involved in oxidation-reduction reactions. One FADH₂ is obtained in the TCA cycle (Barile et al. 2016). These act as coenzymes in the alpha-keto glutarate dehydrogenase and succinate dehydrogenase complexes. Its deficiency results in a condition known as ariboflavinosis. Ariboflavinosis is characterized by cheilosis (textured desquamation of the skin around the mouth), glossitis (sparkly red and sore tongue), soreness of the lips, eye disturbances and photophobia (light sensibility), oily skin the nose, scrotal dermatitis (Henriques et al. 2010).

9.3.2.3 Niacin (Vitamin B3)

It is a water-soluble vitamin essential to the human diet, but can be synthesized in the body from tryptophan. Its deficiency leads to a condition known as pellagra. It is found in some cereals, yeast extracts, and meat. In the body, niacin is converted to the NAD (nicotinamide adenine dinucleotide) coenzyme. These co-enzymes are involved in oxidation-reduction reactions. They act as coenzymes of the isocitrate

dehydrogenase, alpha-ketoglutarate dehydrogenase and malate dehydrogenase complexes (Ronsein et al. 2016). Niacin has lipid-lowering effects and can be used in the treatment of diabetes mellitus (Elam et al. 2000).

9.3.2.4 Pantothenic Acid (B5)

It is a member of the vitamin B complex group. As part of the fatty acid synthase and of coenzyme A, it is involved in hormonogenesis and energy production. Patients with pantothenic acid deficit may develop adrenal insufficiency, enteritis or dermatological conditions (alopecia, dermatitis etc) (Lykstad and Sharma 2019).

9.3.2.5 Biotin (Vitamin B7)

Biotin is also considered as a member of the B-complex group. It helps in the synthesis of fatty acids, utilization of glucose, metabolism of proteins, and utilization of vitamin B₁₂ and folic acid. The effects of biotin deficiency or excess have remained unknown. It is obtained from green beans, egg yolk, dark green vegetables, kidneys and liver (Hsu et al. 2016).

9.3.2.6 Folic Acid (Vitamin B9)

Folic acid is the synthesized form of folate, a water-soluble vitamin, found in green leafy vegetables, fruits and liver. Following conversion, vitamin B9 or folate becomes tetrahydrofolate, its active form. It is an essential molecule in the synthesis of nucleic acids (DNA and RNA). Folate deficiency can lead to neural tube defects, thus pregnant women should receive folate supplementations as a preventive method. Folate deficiency can also cause megaloblastic anemia, a type of macrocytic anemia, and prompts a differential diagnosis with vitamin B₁₂ deficiency, which also causes megaloblastic anemia (Lykstad and Sharma 2019; Ankar and Kumar 2019).

Methylenetetrahydrofolate is involved in the synthesis of thymidine, an important component of the DNA. Neural tube defects are common in children whose mothers were deficient in folic acid during pregnancy (Hodgetts et al. 2015). Neural tube defects can be prevented in children whose mothers receive folic acid supplements during pregnancy. Methyl vitamin B₁₂ mediates the reaction in which the amino acid methionine, which is required for the synthesis of myelin, is generated from homocysteine. During this process, methyl tetrahydrofolate is also converted to tetrahydrofolate. The normal generation of methyl tetrahydrofolate depends upon an adequate supply of both folic acid and vitamin B₁₂. Deficiency of either of them can produce a defect in all the tissues with rapid rate of cellular proliferation, e.g. the bone marrow (resulting in megaloblastic anaemia) and the gastrointestinal tract.

The administration of large doses of folic acid in a patient with vitamin B₁₂ deficiency may alleviate the anaemia but it cannot cure or may even aggravate the neurological deficit, by increasing the tissue demand of vitamin B₁₂ (Reynolds 2014).

9.3.2.7 Cobalamin (Vitamin B12)

Vitamin B₁₂ (cyanocobalamin) is a water-soluble vitamin found in meat, dairy products, fish and eggs. (Hariz and Bhattacharya 2019). It is essential for the normal production of red blood cells by the bone marrow and for the growth of nervous cells. Vitamin B₁₂ deficiency causes megaloblastic anemia and, *via* myelin damage, neurological deficits (ataxia, neuropathy) or even neuropsychiatric symptoms such as dementia. Vegan diets contain insufficient amounts of vitamin B₁₂. Folate deficiency can also cause megaloblastic anemia and should be taken into account when looking for the potential cause of the anemic syndrome (Lykstad and Sharma 2019; Ankar and Kumar 2019).

9.3.2.8 Vitamin C

Vitamin C (ascorbic acid) is a crystalline solid which is soluble in water (Halliwell 2001). Most animals and plants synthesize ascorbic acid; however, primates and humans lack gluconolactone oxidase, a key enzyme in the final step of ascorbate synthesis. Thus, in these species, the daily requirements of vitamin C should come from the diet. Oranges, lemons, grapefruit, leafy green vegetables and beef liver are the best sources of vitamin C (Ferraro et al. 2016). Vitamin C is needed to form collagen that gives strength to the connective tissues and required for wound healing and a normal immune function. Vitamin C deficiency causes changes in the connective, leading to the development of scurvy, a disease in which the synthesized collagen is unstable. The symptoms of scurvy include muscle pain, joint swelling, and bleeding. Vitamin C acts as antioxidant and free oxygen radical scavenger and can be used topically in skin disorders, including those caused by photo-aging (Sorice et al. 2014; Luis Gomez et al. 2018). The hyperpigmentation of the skin can be treated with vitamin C, since it inhibits the activity of melanocytes, i.e. the cells involved in the synthesis of melanin (Telang 2013).

9.4 Minerals

Minerals are inorganic elements that cannot be synthesized in the body but obtained from the diet. They are naturally present in soil and water. Some are essential to living organisms while some are very toxic. Plants absorb significant amount of minerals from the environment and usually passed them along the food chain to animals. The deficiency of such nutritionally important minerals usually proves fatal.

Minerals are key elements of the body. They are needed in the buildup and function of important biomolecules in the human body. Although, minerals are not a source of energy in the body but they are necessary for the maintenance of normal biochemical processes in the body (Zhao et al. 2016). Based on the body needs, these essential minerals can be classified as either a macro or micro (trace) minerals.

9.4.1 Macro-Minerals

Macro-minerals are nutritionally important minerals such as sodium, calcium, phosphorus, magnesium and potassium. They are classified as macro because the average adult daily requirement is greater than 100 mg/day (Prashanth et al. 2015).

9.4.1.1 Calcium

It is obtained from hard water, shellfish, fish, dark green leafy vegetables and milk (Jeurnink and De Kruif 1995). Calcium is a mineral which is essential for an adequate growth and bone development (Matkovic and Ilich 1993). It is a common mineral found in blood (McCarron and Reusser 1999). Cells require an adequate amount of calcium to perform various functions (Miller et al. 2001). Teeth and bones are rich in calcium (Vallet-Regí and González-Calbet 2004). It is also involved in blood clotting (Hall et al. 1991). Most of the calcium is found in bones (Reid et al. 2015). The total extracellular fluid space contains about 900 mg of calcium which is in dynamic equilibrium with the skeleton (Bronner and Stein 1995). About 1% of the skeletal calcium (10 g) is readily exchangeable with the calcium in extracellular fluids and constitutes a large calcium reservoir. The remaining 99% of the bone calcium is only slowly exchangeable. Nearly 500 mg of calcium are deposited in and mobilized from the bones daily in a continuous process of remodelling. Calcium is secreted into the intestine through bile, pancreatic juice, and intestinal secretion but is completely reabsorbed. As the blood level of calcium decreases, bones start to release calcium to increase the calcemia. When the calcium level increases in the blood, it is deposited in bones or excreted in urine. Calcium is involved in the nerve function, contraction of muscles and clotting of blood. Its level in blood is maintained by the parathyroid hormone and calcitonin. The daily-recommended allowance for calcium is 1 g per day. Vitamin D is involved in the uptake of calcium by the body. Its deficiency results in rickets, osteoporosis, and osteomalacia. Tetany may occur due to the deficiency of calcium in the blood. Causes of hypocalcaemia include chronic renal failure (Naveh-Many et al. 1995), phosphate therapy etc. (Välimäki et al. 1999). The excess amount of calcium in the blood can result in the development of kidney stones (Pettifor 2008; Sahay and Sahay 2012).

9.4.1.2 Sodium

Sodium (natrium) is present in most foods and its dietary deficiency is rare. Sodium is involved in the control of blood. Sodium chloride is the most common form of sodium which is marketed as table salt (Cogswell et al. 2016). Kidneys are the main regulators of body sodium and normally 98% of the body loss of sodium occurs in urine. If more sodium is ingested, its excretion in the urine increases. If less sodium is ingested or if plasma sodium falls due to any reason, sodium may totally disappear from the urine. This is usually through the adrenocortical hormone aldosterone, which increases the tubular reabsorption of sodium in the renal tubules. An increased level of sodium in the blood defines hyponatremia, and is characterized by seizures, oedema, neuromuscular excitability, irritability, weakness and lethargy (Kalogeropoulos et al. 2015; Xi et al. 2015).

9.4.1.3 Magnesium

Magnesium is obtained from hard water, spices, apricots, bananas, soybeans, nuts, green leafy vegetables, and whole grains. It aids in the maintenance of bone growth and integrity and is involved in the regulation of the cardiac cycle and the functioning of muscles and nerves. Deficiency diseases are hypomagnesaemia and neuromuscular irritability. Toxicity symptoms are hypotension, respiratory failure, and cardiac disturbances (Allen and Sharma 2019; Gragossian and Friede 2019).

9.4.1.4 Potassium

It is obtained from whole and skimmed milk, meat, bananas, raisins and prunes. A proper plasma potassium level is essential for the normal heart functioning. Potassium ions also take part in the normal functioning of skeletal muscle fibers. Potassium is needed for many enzyme reactions (Weaver 2013). Glycogenesis requires the presence of potassium. Insulin administration causes a fall in plasma potassium level because the deposition of glycogen brought about by insulin is also accompanied by the deposition of potassium. Moreover, insulin also increases protein synthesis within the cells, which by binding potassium ions can lead to a low plasma potassium level. Potassium deficiency leads to hypokalemia, paralysis, and cardiac disturbances. Excessive potassium levels lead to hyperkalemia, paralysis and cardiac disturbances (He and MacGregor 2008).

9.4.1.5 Phosphorus

It is obtained from legumes, nuts, cereals, fish, meat, cheese and poultry (Takeda et al. 2012). The bioavailability of minerals such as iron and zinc may be low in a total vegetarian diet because of the presence of substances such as phytic acid. Besides, large amounts of dietary fiber may interfere with its proper absorption.

Because humans are omnivorous, trace element deficiencies are unlikely to develop. Studies have shown that mineral deficiencies are not more frequent among vegetarians than among non-vegetarians. In fact, man's need for trace elements has not yet been precisely determined. Trace elements should not be used as dietary supplements, since excessive amounts can have injurious effects. Phosphorus is involved in the formation of bone and teeth, ATP, GTP and UTP. It is a component of DNA and RNA, it is present in phospholipids and it forms part of cell membranes. Hyperphosphatemia can occur in renal failure (Arai and Sakuma 2015).

9.4.2 Trace Elements

As the name implies, trace elements are essential group of minerals, which are needed, in small quantity for the day-to-day metabolic processes in man. They are regarded, as trace elements because their daily requirement should be below 100 mg, above which can be toxic to health. However, the deficiency of any of these trace elements can lead to serious health challenges (Prashanth et al. 2015). Trace minerals include iron, copper, zinc, iodine, manganese etc. (Table 9.2).

Table 9.2 Trace minerals, functions and sources

S/N	Minerals	Functions	Sources	References
1	Iron (Fe)	Helps in the formation of heme proteins needed for the transport of oxygen to the red blood cells, flavoproteins and other enzymes.	Fe is widely distributed in organ meats, red meats (30–70% is haem iron), egg yolks; legumes; dried fruits; dark, leafy greens; iron-enriched breads and cereals; and fortified cereals; fish; poultry; shellfish	Fairweather-Tait and Hurrell (1996)
2	Copper (Cu)	Part of many enzymes including metalloenzymes; needed for red blood cell formation, connective tissues	Foods high in copper include liver, kidney, shellfish, wholegrain cereals and nuts. Soft or acidic water passing through copper pipes can also contribute copper to the diet.	Fairweather-Tait and Hurrell (1996)
3	Zinc (Zn)	It plays main roles in the cell-mediated immunity, bone formation, tissue growth, brain function, growth of the fetus and child. It also has roles in pathogenesis of some dermatological disorders.	Meats, fish, poultry, oysters, leavened whole grains, vegetables	Bagherani and Smoller (2016)

(continued)

Table 9.2 (continued)

S/N	Minerals	Functions	Sources	References
4	Iodine (I)	Growth and development, Metabolism, Reproduction, Thyroid hormone production	Seafood, foods grown in iodine-rich soil, iodized salt, bread, dairy products	Fairweather-Tait and Hurrell (1996)
5	Selenium	Anti-oxidant; Selenium is needed for the proper functioning of the immune system, and appears to be a key nutrient in counteracting the development of virulence and inhibiting HIV progression to AIDS. It is required for sperm motility and may reduce the risk of miscarriage.	Meats, seafood, grains. Cereals, seafood and meat products are the richest sources of Se and are the main contributors to the daily Se intake, whereas vegetables, fruits and beverages are generally in Se	Combs (1988), Fairweather-Tait and Hurrell (1996) and Margaret (2000)
6	Chromium (Cr)	Chromium acts as an antioxidant. It also helps to decrease insulin resistance in diabetic patients	Dietary sources of chromium include brewer's yeast, cheese, pork kidney, whole grain breads and cereals, molasses, spices and some bran cereals. Lean beef, oysters, eggs, and turkey are sources of chromium	Anderson et al. (1992), Anderson (2000) and Tulasi and Rao (2014)
7	Manganese (Mn)	It activates numerous enzymes — such as hydrolases, transferases, kinases, and decarboxylases — and is a constituent of some enzymes. Manganese also plays a role in blood clotting and hemostasis in conjunction with vitamin K.	Manganese is present in a wide variety of foods, including whole grains, clams, oysters, mussels, nuts, soybeans and other legumes, rice, leafy vegetables, coffee, tea, and many spices, such as black pepper.	Watts (1990), Aschner and Aschner (2005) and Buchman (2014)
8	Flouride (F)	It is involved in formation of bones and teeth; helps prevent dental decay and caries.	Drinking water (either fluoridated or naturally containing fluoride), fish, and most beverages, from oral tooth pastes as well.	O'Mullane et al. (2016)
9	Molybdenum (Mo)	Functions as a cofactor for at least four enzymes: sulfite oxidase, xanthine oxidase, aldehyde oxidase, and mitochondrial amidoxime reducing component.	Legumes; nuts, breads and grains; leafy greens; leafy, green vegetables; milk; liver	Novotny (2011)

Source: Modified from <https://www.uofmhealth.org/health-library/ta3912>

9.4.2.1 Iron

Iron is obtained from green leafy vegetables, dried nuts, beans, peas, egg yolk, red meat, kidney, and liver. Iron is one of the most abundant metals present in the body and is essential for life. As a component of haemoglobin and myoglobin, it is involved in the transfer of oxygen between blood and tissues. In most of the cells, iron is present as a component of enzymes involved in oxidation-reduction reactions. At birth of an infant, large amount of iron (246 mg) accumulates in the body. This iron storage depends on the iron intake of the mother during pregnancy. The greatest demand for iron is during the last 3 months of pregnancy. The developing fetus requires about 20–30 mg/day of iron in the pregnant woman. The iron in diet comes in the form of haem and non-haem. Haem has a higher bioavailability and can be found in meat, fish, poultry, and milk (Abbaspour et al. 2014). Non-haem is found in varying degrees in plant products. Iron deficiency leads to hypochromic microcytic anaemia. Factors that reduce iron absorption are the surgical removal of the upper small intestine, subtotal gastrectomy, chronic infections, achlorhydria and anti-acid therapy, excess of phosphates and oxalates, diarrhoea and malabsorption. In the case of deficient intake of iron, blood loss or increased iron demand, iron deposits in the body depletes, and anaemia occurs. Symptoms include pallor, weakness, irritability, fissures of the angles of the mouth, heart murmurs and indigestion (Lopez et al. 2016). Iron excess leads to the development of cirrhosis, skin pigmentation, and hemochromatosis. In the United States of America, 5% of men and 2% of women are suffering from anaemia (Johnson-Wimbley and Graham 2011).

9.4.2.2 Chloride

Sources of chloride include cornbread, potato chips, green olives, and animal products. Chloride is necessary to maintain the composition of blood and to the formation of hydrochloric acid. It regulates the acid-base balance and the osmotic pressure. Deficiency symptoms include alkalosis and failure to thrive in infants. Toxicity symptoms include an increase in the extracellular volume and hypertension (Shrimanker and Bhattarai 2019; Sur and Shah 2019).

9.4.2.3 Cobalt

Intestinal bacteria synthesize cobalt. It takes part in the formation of vitamin B₁₂. It is stored in the body in amounts of 1–2 mg. The liver stores enough cobalt for 3–4 years as hydroxyl cobalamine and methyl cobalamine. The daily requirements are 2–3 microgram (Angelova et al. 2014).

9.4.2.4 Copper

It is obtained from organ meats, nuts, dried legumes, whole grains, and cereals. In food, it is present as copper complexes and released in the stomach due to the acidic pH of the gastric juice. It is involved in bone formation and hematopoiesis. It is absorbed in the small intestine mostly by diffusion and in small amounts using carriers. In the circulation, copper combines with albumin, reaches the liver, and incorporated into ceruloplasmin, which is distributed to the tissues. Copper is excreted through bile in faeces and in urine, skin, hair, and nails. It is transported by albumin and is bound to ceruloplasmin. It is part of certain enzymes like ferro-oxidase, catalase, cytochrome oxidase and tyrosinase (Leone et al. 2006). It is required for red blood cell production. Copper-containing proteins like ceruloplasmin contribute to the absorption of iron in the gastrointestinal tract. Copper deficiency can cause hypochromic anaemia. Copper toxicity manifests as hepatolenticular degeneration and biliary cirrhosis (Patil et al. 2013).

9.4.2.5 Zinc

It is obtained from liver, muscle, oysters, cereals, and pulses. Infants need 5 mg/day, children need 10 mg/day and adults need 35 mg/day. It is a constituent of metallo-enzymes and enables cell growth and proliferation, sexual maturity, and fertility. It improves immunity, appetite, and taste. Iron and copper reduce its absorption. Zinc deficiency is rare and can be seen in patients with kidney diseases and in alcoholic patients. The growth of children with zinc deficiency is reduced. Zinc toxicity symptoms include the development of gastrointestinal diseases and a decreased immune function (Prasad 2008; Bredholt and Frederiksen 2016).

9.4.2.6 Silicon

Elevated levels of silicon are present in the connective tissue of the aorta, i.e. collagen and elastin components of the aortic wall. Silicon levels decline with ageing and with the development of atherosclerosis. Silicon seems to reduce the levels of fatty acids in the blood and the aorta. Fatty acids are associated with the formation of atherosclerotic plaques. Studies have shown silicon supplements also help in the prevention of osteoporosis (Jugdaohsingh 2007).

9.4.2.7 Manganese

It is obtained from nuts, cereals, and tea leaves. It is required for enzymes, such as succinate dehydrogenase, arginase, and glucosyltransferase. It is required for the synthesis of chondroitin sulphate which is necessary for cartilage formation (Tuschl et al. 2013). It stabilizes the DNA and the RNA. The total body content is 15 mg and

the daily requirement is of 4–10 mg. It is excreted in the bile and the pancreatic juice. Its deficiency causes a defective bone formation and leads to glucose intolerance, alopecia, reddening of hair and dermatitis. Toxicity symptoms include functional abnormalities of the central nervous system (Pallauf et al. 2012).

9.4.2.8 Fluoride

It is obtained from seafood, vegetables, grains, tea, coffee (Moody et al. 1980) and fluorinated water (Buzalaf et al. 2001). It is necessary for the mineralization of bones, the development of the dental polish and the aversion of dental caries. Dental caries can be prevented by the fluoridation of water and the addition of fluorides to toothpaste. Fluoride salts can be administered to children in the form of drops, tablets, and mouthwash. Toxicity leads to dental fluorosis (Iheozor-Ejiofor et al. 2013).

9.4.2.9 Iodine

Sources of iodine include seafood, iodized salts, eggs, dairy products and water. It is an essential element for the normal growth and development of the body. Iodine is necessary for the formation of thyroid hormones. Iodine deficiency occurs in nearly 2.6 billion individuals. Various degrees of iodine deficiency affect 50 million children worldwide (Ershow et al. 2016). Iodine cannot be stored in the body and requires a lifetime intake in tiny amounts. Common salt fortified with iodine can be used as an effective intervention for a long time. Iodine deficiency can lead to the development of goiter. The daily-recommended dietary allowance of iodine is 40 microgram/day. It is used also as antiseptic, as skin disinfectant prescribed in the form of povidone-iodine (Durani and Leaper 2008).

9.4.2.10 Selenium

Selenium is a mineral with antineoplastic effects against colon cancer, breast cancer and possibly other malignancies (Hoffmann and Berry 2008). It is a natural antioxidant (Gaman et al. 2014). The amount of selenium present in food depends on how much selenium was found in the soil where the food had grown. It is obtained from animal and vegetal products. The richest food sources of selenium include muscle meat, cereals, grains, and dairy products like eggs. It partakes in the activity of metallo-enzymes and protein synthesis, prevents liver necrosis, stimulates pancreatic lipase secretion and is involved ATP generation. Selenium deficiency causes hemolytic anaemia, muscle necrosis, and cardiomyopathy. The excess of selenium causes toxicity-dermatitis, loss of hair and gives a garlic odour in breath (Rayman 2012).

9.5 Summary and Conclusion

Vitamins and minerals are essential substances for the normal functioning and development of the body. There are two classes of vitamins namely: Fat-soluble and Water-soluble vitamins. The known vitamins include A, C, D, E, and K, and the B vitamins: thiamin (B₁), riboflavin (B₂), niacin (B₃), pantothenic acid (B₅), pyridoxine (B₆), cyanocobalamin (B₁₂), biotin, and folate/folic acid. A number of minerals are essential for health: calcium, phosphorus, potassium, sodium, chloride, magnesium, iron, zinc, iodine, sulfur, cobalt, copper, fluoride, manganese, and selenium. The essential minerals of the body are divided into two viz-a-viz: macrominerals and trace minerals. The trace minerals are those that are required in small amounts by the body but do not imply they have little significance to the body. Micronutrients satisfy both the qualitative and quantitative requirements in the human diet and are consumed in moderate amounts.

References

- Abbaspour N, Hurrell R, Kelishadi R (2014) Review on iron and its importance for human health. *J Res Med Sci: Off J Isfahan Univ Med Sci* 19(2):164
- Abhra T, Girma Y, Haile K, Hailu M, Hailemariam M (2016) Prevalence and associated factors of clinical manifestations of vitamin a deficiency among preschool children in asgede-tsimbla rural district, North Ethiopia, a community based cross sectional study. *Arch Public Health* 74(1):4
- Advisory Committee on Technology Innovations (1975) Buriť palm. In: Underexploited tropical plants with promising economic value. Report of an Ad Hoc Panel of the Advisory Committee on Technology Innovations, Board on Science and Technology for International Development, Commission on International Relations. Washington, DC, National Academy of Sciences, pp 133–137
- Allen LH, Dror DK (2018) Introduction to current knowledge on micronutrients in human milk: adequacy, analysis, and need for research. *Adv Nutr* 9(suppl_1):275S–277S
- Allen MJ, Sharma S (2019) Magnesium. [Updated 2019 Apr 4]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK519036/>
- Anderson RA (2000) Chromium in the prevention and control of diabetes. *Diabetes Metab* 26(1):22–27
- Anderson RA, Bryden NA, Polansky MM (1992) Dietary chromium intake: freely chosen diets, institutional diets and individual foods. *Biol Trace Elem Res* 32:117–121
- Angelova MG, Petkova-Marinova TV, Pogorielov MV, Loboda AN, Nedkova-Kolarova VN, Bozhinova AN (2014) Trace element status (iron, zinc, copper, chromium, cobalt, and nickel) in iron-deficiency anaemia of children under 3 years. *Anemia* 2014:1–8
- Ankar A, Kumar A (2019) Vitamin B12 Deficiency (Cobalamin) [Updated 2019 Jan 11]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK441923/>
- Arai H, Sakuma M (2015) Bone and nutrition. Bone and phosphorus intake. *Clin Calcium* 25(7):967–972
- Aschner JL, Aschner M (2005) Nutritional aspects of manganese homeostasis. *Mol Asp Med* 26:353–362

- Aviva FV (2010) Thiamine (Vitamine B1). *J Evid Based Compliment Altern Med* 16(1):12–20
- Bagherani N, Smoller BR (2016) An overview of zinc and its importance in dermatology- Part I: importance and function of zinc in human beings. *Glob Dermatol* 3(5):330–336. <https://doi.org/10.15761/GOD.1000185>
- Barile M, Giancaspero TA, Leone P, Galluccio M, Indiveri C (2016) Riboflavin transport and metabolism in humans. *J Inherit Metab Dis* 39(4):545–557
- Bellizzi MC et al (1994) Vitamin E and coronary heart disease: the European paradox. *Eur J Clin Nutr* 48:822–831
- Birringer M, Blumberg JB, Eggersdorfer M, Frank J, Weber P (2019) History of Vitamin E research. *Vitam E Human Health*:7–18. https://doi.org/10.1007/978-3-030-05315-4_2
- Booth SL, Johns T, Kuhnlein HV (1992) Natural food sources of vitamin A and provitamin A. *UNU Food Nutr Bull* 14:6–19
- Booth SL et al (1993) Vitamin K1 (phyloquinone) content of foods: a provisional table. *J Food Compos Anal* 6:109–120
- Booth SL, Davidson KW, Sadowski JA (1994) Evaluation of an HPLC method for the determination of phyloquinone (vitamin K1) in various food matrices. *J Agric Food Chem* 42:295–300
- Bredholt M, Frederiksen JL (2016) Zinc in multiple sclerosis: a systematic review and meta-analysis. *ASN Neuro* 8(3):1759091416651511
- Bronner F, Stein WD (1995) Calcium homeostasis—an old problem revisited. *J Nutr* 125(suppl_7):1987S–1995S
- Buchman AR (2014) Manganese. In: BC ACR, Cousins RJ, Tucker KL, Ziegler TR (eds) *Modern nutrition in health and disease*, 11th edn. Lippincott Williams & Wilkins, Baltimore, pp 238–244
- Buzalaf M, Granjeiro JM, Damante CA (2001) Fluoride content of infant formulas prepared with deionized, bottled mineral and fluoridated drinking water. *ASDC J Dent Child* 68(1):37–41. 10
- Christakos S, Dhawan P, Verstuyf A, Verlinden L, Carmeliet G (2015) Vitamin D: metabolism, molecular mechanism of action, and pleiotropic effects. *Physiol Rev* 96(1):365–408
- Cogswell ME, Mugavero K, Bowman BA, Frieden TR (2016) Dietary sodium and cardiovascular disease risk—measurement matters. *N Engl J Med* 375(6):580
- Combs Jr, G. F. (1988). Selenium in foods. *Adv Food Res* 32:85–113
- Conly JM, Stein K (1992) Quantitative and qualitative measurements of K vitamins in human intestinal contents. *Am J Gastroenterol* 87:311–316
- Derbyshire E (2018) Micronutrient intakes of British Adults Across Mid-Life: A secondary analysis of the UK national diet and nutrition survey. *Frontiers in Nutrition* 5
- Durani P, Leaper D (2008) Povidone–iodine: use in hand disinfection, skin preparation and antiseptic irrigation. *Int Wound J* 5(3):376–387
- Elam MB, Hunninghake DB, Davis KB, Garg R, Johnson C, Egan D, Kostis JB, Sheps DS, Brinton EA, ADMIT Investigators (2000) Effect of niacin on lipid and lipoprotein levels and glycemic control in patients with diabetes and peripheral arterial disease: the ADMIT study: a randomized trial. *JAMA* 284(10):1263–1270
- Ershow AG, Goodman G, Coates PM, Swanson CA (2016) Research needs for assessing iodine intake, iodine status, and the effects of maternal iodine supplementation. *Am J Clin Nutr* 104(suppl_3):941S–949S
- Fairweather-Tait S, Hurrell RF (1996) Bioavailability of minerals and trace elements: Members of EC flair concerted action no. 10: Measurements of micronutrient absorption and status. *Nutr Res Rev* 9(1):295–324
- Feldman D, Glorieux FH, Pike JW (1997) *Vitamin D*. Academic, New York
- Ferraro PM, Curhan GC, Gambaro G, Taylor EN (2016) Total, dietary, and supplemental vitamin C intake and risk of incident kidney stones. *Am J Kidney Dis* 67(3):400–407
- Gaman AM, Buga AM, Gaman MA, Popa-Wagner A (2014) The role of oxidative stress and the effects of antioxidants on the incidence of infectious complications of chronic lymphocytic leukemia. *Oxid Med Cell Longev* 2014: Article ID 158135, 6 p. <https://doi.org/10.1155/2014/158135>

- Gaman AM, Egbuna C, Gaman MA (2019) Natural bioactive lead compounds effective against haematological malignancies. In: Egbuna C, Kumar S, Ifemeje JC, Ezzat SM, Kaliyaperumal S (eds) *Phytochemicals as Lead compounds for new drug discovery*. Elsevier, Amsterdam/Cambridge MA, pp 95–115. <https://doi.org/10.1016/B978-0-12-817890-4.00006-8>
- Gheorghe G, Stoian AP, Gaman MA, Socea B, Neagu TP, Stanescu AMA, Bratu OG, Mischianu DLD, Suceveanu AI, Diaconu CC (2019) The benefits and risks of antioxidant treatment in liver diseases. *Rev Chim* 70(2):651–655
- Gomez-Pomar E, Hatfield E, Garlitz K, Westgate PM, Bada HS (2018) Vitamin E in the preterm infant: a forgotten cause of Hemolytic Anemia. *Am J Perinatol* 35(03):305–310
- Gragossian A, Friede R (2019) Hypomagnesemia. [Updated 2019 Feb 23]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK500003/>
- Gräslund S, Nordlund P, Weigelt J, Hallberg BM, Bray J, Gileadi O et al (2008) Protein production and purification. *Nat Methods* 5(2):135
- Gregory SK (2011) Pantothenic acid. *Monogr Altern Med Rev* 16(3):263–274
- Hall D, Cromwell G, Stahly T (1991) Effects of dietary calcium, phosphorus, calcium: phosphorus ratio and vitamin K on performance, bone strength and blood clotting status of pigs. *J Anim Sci* 69(2):646–655
- Hans KB, Jana T (2018) Micronutrients in the life cycle: Requirements and sufficient supply. *NFS Journal*, 11:1–11
- Halliwell B (2001) Vitamin C and genomic stability. *Mutat Res/Fundam Mol Mech Mutagen* 475(1):29–35
- Hariz A, Bhattacharya PT (2019) Megaloblastic Anemia. [Updated 2019 Jan 23]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK537254/>
- He FJ, MacGregor GA (2008) Beneficial effects of potassium on human health. *Physiol Plant* 133(4):725–735
- Henriques BJ, Olsen RK, Bross P, Gomes CM (2010) Emerging roles for riboflavin in functional rescue of mitochondrial β -oxidation flavoenzymes. *Curr Med Chem* 17(32):3842–3854
- Hodgetts V, Morris R, Francis A, Gardosi J, Ismail K (2015) Effectiveness of folic acid supplementation in pregnancy on reducing the risk of small-for-gestational age neonates: a population study, systematic review and meta-analysis. *BJOG: Int J Obstet Gynaecol* 122(4):478–490
- Hoffmann PR, Berry MJ (2008) The influence of selenium on immune responses. *Mol Nutr Food Res* 52(11):1273–1280
- Hsu S-M, Raine L, Fanger H (2016) A comparative study of the peroxidase-antiperoxidase method and an avidin-biotin complex method for studying polypeptide hormones with radioimmunoassay antibodies. *Am J Clin Pathol* 75(5):734–738
- Iheozor-Ejiofor Z, O'Malley LA, Glenny AM, Macey R, Alam R, Tugwell P et al (2013) Water fluoridation for the prevention of dental caries. *Cochrane Database Syst Rev* 12
- Ikeda K, Liu X, Kida K, Marutani E, Hirai S, Sakaguchi M et al (2016) Thiamine as a neuroprotective agent after cardiac arrest. *Resuscitation* 105:138–144
- Janos Z, Wijeratne SS, Yousef IH (2009) Biotin. *Biofactors* 35(1):36–46
- Jeurnink TJ, De Kruijff KG (1995) Calcium concentration in milk in relation to heat stability and fouling. *Neth Milk Dairy J* 49:151–151
- Johnson-Wimbley TD, Graham DY (2011) Diagnosis and management of iron deficiency anemia in the 21st century. *Ther Adv Gastroenterol* 4(3):177–184
- Juan M, Raquel A, Josune O, Javier A, Agel G, Rose MO, Lluís SM, Gregorio VM, Marcela GG (2018) Dietary intake and food sources of niacin, riboflavin, Thiamin and Vitamin B6 in a representative sample of the Spanish population. The ANIBES study. *Nutrients* 10(7):846
- Jugdaohsingh R (2007) Silicon and bone health. *J Nutr Health Aging* 11(2):99
- Kalogeropoulos AP, Georgiopoulou VV, Murphy RA, Newman AB, Bauer DC, Harris TB et al (2015) Dietary sodium content, mortality, and risk for cardiovascular events in older adults: the health, aging, and body composition (Health ABC) study. *JAMA Intern Med* 175(3):410–419

- Kamangar F, Emadi A (2012) Vitamin and mineral supplements: do we really need them? *Int J Prev Med* 3(3):221
- Kelly RJ, Gruner TM, Furlong JM, Sykes AR (2006) Analysis of corrinoids in ovine tissues. *Biomed Chromatogr* 20:806–814
- Kemnic TR, Coleman M (2019) Vitamin E Deficiency. [Updated 2019 Nov 16]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK519051/>
- Kennel KA, Drake MT, Hurley DL (2010) Vitamin D deficiency in adults: when to test and how to treat. *Mayo Clin Proc* 85(8):752–758
- Kunisawa J, Hashimoto E, Ishikawa I, Kiyono H (2012) A pivotal role of Vitamin B9 in the maintenance of regulatory T cells In Vitro and In Vivo. *PLoS One* 7(2):e32094. <https://doi.org/10.1371/journal.pone.0032094>
- Leone N, Courbon D, Ducimetiere P, Zureik M (2006) Zinc, copper, and magnesium and risks for all-cause, cancer, and cardiovascular mortality. *Epidemiology* 17(3):308–314
- Lopez A, Cacoub P, Maccougall IC, Peyrin-Biroulet L (2016) Iron deficiency anaemia. *Lancet* 387(10021):907–916
- Luis Gomez A, Tchekalarova JD, Atanasova M, da Conceição Machado K, de Sousa Rios MA, Jardim MFP et al (2018) Anticonvulsant effect of anacardic acid in murine models: putative role of GABAergic and antioxidant mechanisms. *Biomed Pharmacother* 106:1686–1695. <https://doi.org/10.1016/j.biopha.2018.07.121>
- Lule VK, Garg S, Gosewade SC, Tomar SK (2016) Niacin. In: Caballero B, Fingelas P, Toldra F (eds) *The Encyclopedia of food and health*, vol 4. Academic, Oxford, pp 63–72
- Lykstad J, Sharma S (2019) Biochemistry, Water Soluble Vitamins. [Updated 2019 Feb 16]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK538510/>
- Makarchikov AF (2009) Vitamin B1: metabolism and functions. *Biochem (Moscow) Suppl Ser B Biomed Chem* 3(2):116–128
- Marcelina P, Seth S, Hanjo H (2018) Vitamin B₆ and its role in cell metabolism and physiology. *Cell* 7(84):1–28
- Margaret PR (2000) The importance of selenium to human health. *Lancet* 356(9225). [https://doi.org/10.1016/S0140-6736\(00\)02490-9](https://doi.org/10.1016/S0140-6736(00)02490-9)
- Matkovic V, Ilich JZ (1993) Calcium requirements for growth: are current recommendations adequate? *Nutr Rev* 51(6):171–180
- McCarron DA, Reusser ME (1999) Finding consensus in the dietary calcium-blood pressure debate. *J Am Coll Nutr* 18(sup5):398S–405S
- Miller GD, Jarvis JK, McBean LD (2001) The importance of meeting calcium needs with foods. *J Am Coll Nutr* 20(2):168S–185S
- Moisa C, Gaman MA, Pascu EG, Dragusin OC, Assani AD, Epingeac ME, Gaman AM (2018) The role of oxidative stress in essential thrombocythemia. *Arch Balk Med Union* 53(1):70–75
- Moody G, Ong B, Quinlan K, Riah A, Thomas J (1980) The determination of fluorine in coffee and tea using a microprocessor coupled with a fluoride ion-selective electrode. *Int J Food Sci Technol* 15(3):335–343
- Nair R, Maseeh A (2012) Vitamin D: the “sunshine” vitamin. *J Pharmacol Pharmacother* 3(2):118
- Naveh-Many T, Rahamimov R, Livni N, Silver J (1995) Parathyroid cell proliferation in normal and chronic renal failure rats. The effects of calcium, phosphate, and vitamin D. *J Clin Invest* 96(4):1786–1793
- Novotny JA (2011) Molybdenum Nutrition in Humans. *J Evid Based Complement Alter Med* 16(3):164–168
- Nutrient Data Laboratory. USDA National Nutrient Database for Standard Reference, Release 24 [Internet]. Available from <http://www.ars.usda.gov/ba/bhnrc/nd>
- O’Mullane DM, Baez RJ, Jones S, Lennon MA, Petersen PE, Rugg Gunn AJ, Whelton H, Whitford GM (2016) Fluoride and oral health. *Community Dent Health* 33:69–99

- Ormsbee MJ, Bach CW, Baur DA (2014) Pre-exercise nutrition: the role of macronutrients, modified starches and supplements on metabolism and endurance performance. *Nutrients* 6(5):1782–1808
- Pallauf J, Kauer C, Most E, Habicht S, Moch J (2012) Impact of dietary manganese concentration on status criteria to determine manganese requirement in piglets. *J Anim Physiol Anim Nutr* 96(6):993–1002
- Patil M, Sheth KA, Krishnamurthy AC, Devarbhavi H (2013) A review and current perspective on Wilson disease. *J Clin Exp Hepatol* 3(4):321–336
- Pettifor JM (2008) Vitamin D and/or calcium deficiency rickets in infants and children: a global perspective. *Indian J Med Res* 127(3):245
- Piccardi N, Manissier P (2009) Nutrition and nutritional supplementation: impact on skin health and beauty. *Dermato-endocrinology* 1(5):271–274
- Pinto JT, Rivlin R (2013) In Book: *Handbook of Vitamins*. 5th Ed. 191–265
- Prasad AS (2008) Zinc in human health: effect of zinc on immune cells. *Mol Med* 14(5–6):353
- Prashanth L, Kattapagari KK, Chitturi RT, Baddam VRR, Prasad LK (2015) A review on role of essential trace elements in health and disease. *J Dr NTR Univ Health Sci* 4(2):75
- Rayman MP (2012) Selenium and human health. *Lancet* 379(9822):1256–1268
- Reddy P, Jialal I (2018) Biochemistry, Vitamin, Fat Soluble. [Updated 2018 Nov 23]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK534869/>
- Reid I, Bristow S, Bolland M (2015) Calcium supplements: benefits and risks. *J Intern Med* 278(4):354–368
- Reynolds E (2014) The neurology of folic acid deficiency. In: *Handbook of clinical neurology*, vol 120. Elsevier, Amsterdam, pp 927–943
- Rodriguez-Amaya DB (2004) Carotenoids and food preparation: the retention of provitamin A carotenoids in prepared, processed, and stored foods. Arlington, John Snow and Opportunities for Micronutrient Interventions Project, 1997 <http://www.mostproject.org/carrots2.pdf>. Accessed 24 June 2004
- Romagnoli G, Luttk MA, Kötter P, Pronk JT, Daran J-M (2012) Substrate specificity of thiamine pyrophosphate-dependent 2-oxo-acid decarboxylases in *Saccharomyces cerevisiae*. *Appl Environ Microbiol* 78(21):7538–7548
- Ronsein GE, Hutchins PM, Isquith D, Vaisar T, Zhao X-Q, Heinecke JW (2016) Niacin therapy increases high-density lipoprotein particles and Total cholesterol efflux capacity but not ABCA1-specific cholesterol efflux in statin-treated SubjectsSignificance. *Arterioscler Thromb Vasc Biol* 36(2):404–411
- Sahay M, Sahay R (2012) Rickets–vitamin D deficiency and dependency. *Indian J Endocrinol Metab* 16(2):164
- Schlueter and Johnston (2011) Vitamin C: overview and update. *J Evid Based Complement Alter Med* 16(1):49–57
- Schmölz L, Schubert M, Kluge S, Birringer M, Wallert M, Lorkowski S (2018) The hepatic fate of Vitamin E, Vitamin E in health and disease, Jose Antonio Morales-Gonzalez, IntechOpen, <https://doi.org/10.5772/intechopen.79445>. Available from: <https://www.intechopen.com/books/vitamin-e-in-health-and-disease/the-hepatic-fate-of-vitamin-e>
- Shearer MJ, Bach A, Kohlmeier M (1996) Chemistry, nutritional sources, tissue distribution and metabolism of vitamin K with special reference to bone health. *J Nutr* 126(Suppl):S1181–S1186
- Shrimanker I, Bhattarai S (2019) Electrolytes. [Updated 2019 May 3]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK541123/>
- Sorice A, Guerriero E, Capone F, Colonna G, Castello G, Costantini S (2014) Ascorbic acid: its role in immune system and chronic inflammation diseases. *Mini Rev Med Chem* 14(5):444–452
- Sudha JD, Raveendran RL (2017) Vitamin C: Sources, functions, sensing and analysis. <https://doi.org/10.5772/intechopen.70162>

- Sur M, Shah AD (2019) Alkalosis. [Updated 2019 Aug 4]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK545269/>
- Takeda E, Yamamoto H, Yamanaka-Okumura H, Taketani Y (2012) Dietary phosphorus in bone health and quality of life. *Nutr Rev* 70(6):311–321
- Telang PS (2013) Vitamin C in dermatology. *Indian Dermatol Online J* 4(2):143
- Tulasi G, Rao KJ (2014) Essentiality of chromium for human health and dietary nutrition. *J Entomol Zoolgy Stud* 2(1):107–108
- Tuschl K, Mills PB, Clayton PT (2013) Manganese and the brain. *Int Rev Neurobiol* 110:277–312. Elsevier
- Välämäki M, Kinnunen K, Volin L, Tähtelä R, Löyttyniemi E, Laitinen K et al (1999) A prospective study of bone loss and turnover after allogeneic bone marrow transplantation: effect of calcium supplementation with or without calcitonin. *Bone Marrow Transplant* 23(4):355
- Vallet-Regí M, González-Calbet JM (2004) Calcium phosphates as substitution of bone tissues. *Prog Solid State Chem* 32(1–2):1–31
- Vuong LT (1997) An indigenous fruit of North Vietnam with an exceptionally high b-carotene content. *Sight Life Newsl* 2:16–18
- Watts DL (1990) The nutritional relationships of manganese. *J Orthomolecular Med* 5(4):219–222
- Weaver CM (2013) Potassium and health. *Adv Nutr* 4(3):368S–377S
- Welch RM, Graham RD (2004) Breeding for micronutrients in staple food crops from a human nutrition perspective. *J Exp Bot* 55(396):353–364
- WHO/FAO (2004) Handbook on vitamin and mineral requirements in human nutrition, 2nd ed.
- Wiley KD, Gupta M (2019) Vitamin B1 Thiamine Deficiency (Beriberi) [Updated 2019 Jan 2]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2019 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK537204/>
- Xi L, Hao Y-C, Liu J, Wang W, Wang M, Li G-Q et al (2015) Associations between serum potassium and sodium levels and risk of hypertension: a community-based cohort study. *J Geriatr Cardiol: JGC* 12(2):119
- Zhao A, Xue Y, Zhang Y, Li W, Yu K, Wang P (2016) Nutrition concerns of insufficient and excessive intake of dietary minerals in lactating women: a cross-sectional survey in three cities of China. *PLoS One* 11(1):e0146483

Website References

- Micronutrient Facts. <https://www.cdc.gov/nutrition/micronutrient-malnutrition/micronutrients/index.html>
- Micronutrients. https://www.unicef.org/nutrition/index_iodine.html