

**Zinc as an essential nutritional component of human body (a systemic review)**Mehrangiz Ghaffari¹, Mostafa Arabyaghoubi²¹ Assistant Professor of Pathology, Department of Pathology, Zabol University of Medical Sciences, Zabol, Iran² Assistant Professor of Anesthesiology, Department of Anesthesiology, Zabol University of Medical Sciences, Zabol, Iran**ABSTRACT****Introduction:**

Second to iron, zinc, which is widely distributed in the human body, is the most abundant element in human body. The human body has about 2 to 3 grams of zinc, with the highest concentrations in the liver, pancreas, kidneys, bones and muscles. Other tissues with a higher concentration of zinc include parts of the eye, prostate gland, spermatozoa, skin, hair, fingernail and toenails.

Methods:

Searches were conducted by two independent researchers in international (PubMed, Web of science, Scopus and Google scholar) and national (SID, Magiran) databases for related studies from the inception of the databases to September 2017 (without time limitation) in English and Persian languages. To ensure literature saturation, the reference lists of included studies or relevant reviews identified through the search were scanned.

Discussion: Absorption and excretion of zinc are carried out through hemostatic mechanisms that are not quite well known yet. The absorption mechanism consists of two paths. Albumin is the most important zinc plasma carrier. The amount transported in blood, in addition to zinc, depends on the availability of albumin. Zinc is a single intracellular ion with structural, catalytic and regulatory roles. Zinc plays important structural roles as part of a multi-protein structure.

Key words: Zinc , essential nutritional ,component , human body

INTRODUCTION:

Second to iron, zinc, which is widely distributed in the human body, is the most abundant element in human body. The human body has about 2 to 3 grams of zinc, with the highest concentrations in the liver, pancreas, kidneys, bones and muscles. Other tissues with a higher concentration of zinc include parts of the eye, prostate gland, spermatozoa, skin, hair, fingernail and toenails (1). Zinc is a mainly intracellular ion associated with more than 300 different enzymes (in different enzyme groups and categories). Although zinc is abundant in cytosol, it is almost attached to proteins; however, the attached portion is in balance with a small ionic component (2).

Methods:**1.1. Search strategy**

Searches were conducted by two independent researchers in international (PubMed, Web of science, Scopus and Google scholar) and national (SID, Magiran) databases for related studies from the inception of the databases to September 2017 (without time limitation) in English and Persian languages. To ensure literature saturation, the reference lists of included studies or relevant reviews identified through the search were scanned. The specific search strategies were created by a Health Sciences Librarian with expertise in systematic review search using the MESH terms and free terms according to the PRESS standard . After the MEDLINE strategy was finalized, it was adapted to search in other databases.

Accordingly, PROSPERO was searched for ongoing or recently related completed systematic reviews. The key words used in the search strategy were “Zinc , essential nutritional ,component , human body” which were combined with Boolean operators including AND, OR, and NOT.

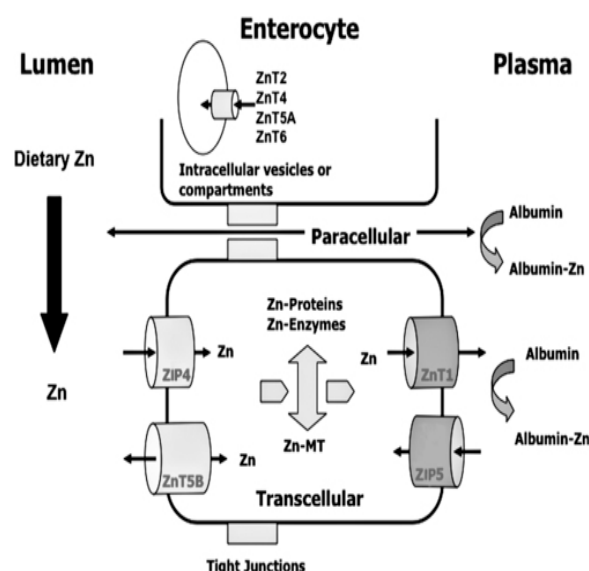
1.2 .Study selection

Results of the Literature review were exported to Endnote. Prior to the formal screening process, a calibration exercise was undertaken to pilot and refine the screening. Formal screening process of titles and abstracts were conducted by two researchers according to the eligibility criteria, and consensus method was used for solving controversies among the two researchers. The full text was obtained for all titles that met the inclusion criteria. Additional information was retrieved from the study authors in order to resolve queries regarding the eligibility criteria. The reasons for the exclusion criteria were recorded. Neither of the review authors was blinded to the journal titles, the study authors or institutions.

Absorption, transfer, storage, disposal

Absorption and excretion of zinc are carried out through hemostatic mechanisms that are not quite well known yet. The absorption mechanism consists of two paths. The saturated carrier mechanism operates in low-level receptacles (low luminosity concentration), and the inactive mechanism also operates at high loading times and high lumens concentration. The solubility of zinc in the digestive lumen is vital (3). Zinc ions are mainly attached to small amino acids or peptides in the lumen of the intestine and are released on the tight junctions and are absorbed by the carrier mechanism (hZIP1 family). The entry into the tight junctions stem cells is associated with the binding of zinc ions to metallothionein and other cytosolic proteins of absorption cells (4). Metallothionein transfers zinc to the lateral part of the body, and zinc is removed from the cell and transported to the bloodstream. Since the concentration of zinc in the blood is much higher than cytosol of the absorption cells, the

removal stage is carried out through the active transfer mechanism (5).



The absorption of zinc is affected by the amount of food and the presence of other interventional agents (especially phytates) in the diet. After consuming food, the concentration is increased then decreases through a dose-response pattern process. High-protein foods result in increased zinc absorption through the formation of Zn-amino acids (which make zinc more absorbable) (6). The absorption of zinc increases slightly during pregnancy and lactation. Zinc is absorbed first through the portal vein to the liver; then, it is distributed among different tissues. Absorption disorders are closely associated with several intestinal disorders such as Crohn's disease or inappropriate pancreatitis (7)

Transmission in the blood

Albumin is the most important zinc plasma carrier. The amount transported in blood, in addition to zinc, depends on the availability of albumin. A small portion of zinc is also transported by transferrin and alpha 2 macroglobulins (8). The major part of the zinc is in the blood, inside the erythrocytes and leukocytes. Plasma is metabolically active and its levels change in response to dietary intake and physiological factors such as injury and inflammation. Zinc levels decrease down to 50% in response to the acute phase of the injury, which is a possible cause of retention in the liver (9) .

Intestinal excretion

Zinc is excreted through feces in healthy individuals. When zinc is received intravenously, about 10% of the dose is received in the liver after 30 minutes. However, in cases of hunger, nephrosis, diabetes, alcoholism, liver cirrhosis, and porphyria, the excretion of zinc increases through urine. Plasma and urinary concentrations of histidine and cysteine zinc and other metabolites may be associated with increased zinc loss in these patients (10).

Function

Zinc is a single intracellular ion with structural, catalytic and regulatory roles. Zinc plays important structural roles as part of a multi-protein structure. It is also associated with 300 different enzymes and is involved in the synthesis or decomposition of carbohydrates, fats, proteins, and nucleic acids. It also functions as an intracellular signal in the brain cells and is stored in synaptic vesicles; it is essential for the normal functioning of the central nervous system. In addition, zinc stabilizes the structure of proteins and nucleic acids, as well as the integrity of subcellular organelles and transport processes (11).

Metallothionein

Metallothionein is the most abundant non-enzymatic protein containing zinc. This low molecular weight protein is rich in cysteine and contains exceptionally large amounts of metal that has zinc; it, also, carries less amounts of copper, iron, cadmium and mercury. The biological role of metallothionein is not clearly known, but it plays a functional role in the absorption of zinc. Metallothionein, as an intracellular reservoir, may act as a protease inhibitor or a controlling agent and it may decrease the oxidative stress (especially in high-stressed cells) (12). Hence, metallothionein may play a role in detoxifying metals as well as absorbing them.

High dosage of zinc in the nucleus leads to the stabilization of the DNA and RNA structure and is essential for the activity of RNA polymerase in cell division. Zinc is present in the chromatin

proteins involved in transcription and replication and is protective against degenerative-macular degenerative disease. Although zinc and nasal spray gluconate tablets are widely used to treat or prevent common colds, they don't seem to be quite effective (13).

Using diet reference

The intake of zinc DRI is 11 mg / day in male and female adolescents. Due to the lower weight of adolescent and adult women, their DRI is 8 to 9 mg per day. The essential rate is 8 mg per day before adolescence. Infants' DRI is 2 mg per day for the first 6 months of life, and 3 mg per day for the second 6 months (14).

Nutrition resources and intake

In most Americans, the vast majority of zinc comes from consuming meat, fish, poultry, ready-to-eat zinc-enriched breakfast cereals, milk and its products. The shellfish contain a high amount of zinc; other Mollusca, grains, dry beans and nuts are good sources of zinc. In general, zinc intake is associated with protein intake.

The content on the normal diet of adults in our western countries varies from 10 to 15 grams per day. Women need to receive less energy due to lower energy consumption. The density of zinc within the diet of an adult American is about 5.6 grams per 1000 kilocalories (15).

Zinc deficiency

The clinical signs of zinc deficiency include shortness of height, hypogonadism, mild anemia, and low plasma zinc levels. Zinc deficiency leads to multiple immunological disorders. Severe zinc deficiency leads to thrombosis, lymphopenia, proliferative response of lymphocytes to mitochondria, selective reduction of T-helper cells, decreased activity of NK cells, insomnia and decreased thymus hormone activity; however, mild zinc deficiency can lead to reduced immune functions, such as interleukin 2 production impairment (16). Zinc supplements may improve immune function, but more evidence is required to do further studies. Mild zinc deficiency has been reported to be associated

with boredom, fatigue and decreased activity of NK cells; however, this form of deficiency is not related to thromboses and lymphopenia atrophy. The similarities between patients with sickle cell anemia and zinc deficiency indicate the possibility of a secondary zinc deficiency in anemia patients (17).

Poisoning

Poisoning due to oral intake of zinc (100 to 300 mg per day) is rare; however, in case of necessity, the maximum amount is 40 mg per day for adults. Excessive supplementation of zinc interferes with the absorption of copper. The major problem of zinc poisoning occurs in patients with renal failure who are undergoing dialysis due to the contamination of dialysis fluids or sticky plastics used in spiral dialysis or in galvanized tubes. The syndrome of poisoning in these patients is characterized by anemia, fever and dysfunction of the central nervous system. Taking more than 2 g or more zinc sulfate per day may cause gastric disorders and vomiting. Inhalation of zinc vapors during welding is toxic, but it can be prevented by taking precautionary measures (18).

References

1. Pereira PM, Vicente AF. Meat nutritional composition and nutritive role in the human diet. *Meat Science*. 2013 Mar 1;93(3):586-92.
2. Mann J, Truswell S, editors. *Essentials of human nutrition*. Oxford University Press; 2017.
3. Frossard E, Bucher M, Mächler F, Mozafar A, Hurrell R. Potential for increasing the content and bioavailability of Fe, Zn and Ca in plants for human nutrition. *Journal of the Science of Food and Agriculture*. 2000 May 15;80(7):861-79.
4. Bhowmik D, Chiranjib KP, Kumar S. A potential medicinal importance of zinc in human health and chronic. *Int J Pharm*. 2010;1(1):05-11.
5. Plum LM, Rink L, Haase H. The essential toxin: impact of zinc on human health. *International journal of environmental research and public health*. 2010 Mar 26;7(4):1342-65.
6. Indrayan AK, Sharma S, Durgapal D, Kumar N, Kumar M. Determination of nutritive value and analysis of mineral elements for some medicinally valued plants from Uttaranchal. *Current science*. 2005 Oct 10:1252-5.
7. Demirezen D, Aksoy A. Heavy metal levels in vegetables in Turkey are within safe limits for Cu, Zn, Ni and exceeded for Cd and Pb. *Journal of food quality*. 2006 Jun 1;29(3):252-65.
8. Lopez HW, Leenhardt F, Coudray C, Remesy C. Minerals and phytic acid interactions: is it a real problem for human nutrition?. *International journal of food science & technology*. 2002 Oct 1;37(7):727-39.
9. King JC. Zinc: an essential but elusive nutrient-. *The American journal of clinical nutrition*. 2011 Jun 29;94(2):679S-84S.
10. Mohamed AE, Rashed MN, Mofty A. Assessment of essential and toxic elements in some kinds of vegetables. *Ecotoxicology and environmental safety*. 2003 Jul 1;55(3):251-60.
11. Amiard JC, Amiard-Triquet C, Charbonnier L, Mesnil A, Rainbow PS, Wang WX. Bioaccessibility of essential and non-essential metals in commercial shellfish from Western Europe and Asia. *Food and Chemical Toxicology*. 2008 Jun 1;46(6):2010-22.
12. Hood MI, Skaar EP. Nutritional immunity: transition metals at the pathogen-host interface. *Nature Reviews Microbiology*. 2012 Aug;10(8):525.
13. Frassinetti S, Bronzetti GL, Caltavuturo L, Cini M, Della Croce C. The role of zinc in life: a review. *Journal of environmental pathology, toxicology and oncology*. 2006;25(3).
14. Vega-Gálvez A, Miranda M, Vergara J, Uribe E, Puente L, Martínez EA. Nutrition facts and functional potential of quinoa (*Chenopodium quinoa willd.*), an ancient Andean grain: a review. *Journal of the Science of Food and Agriculture*. 2010 Dec 1;90(15):2541-7.
15. Aberoumand A. Studies on nutritional values of some wild edible plants of Iran and India.

16. Maret W, Sandstead HH. Zinc requirements and the risks and benefits of zinc supplementation. *Journal of Trace Elements in Medicine and Biology*. 2006 May 10;20(1):3-18.
17. Oves M, Khan MS, Zaidi A, Ahmad E. Soil contamination, nutritive value, and human health risk assessment of heavy metals: an overview. In *Toxicity of heavy metals to legumes and bioremediation 2012* (pp. 1-27). Springer Vienna.
18. Lowe NM, Fraser WD, Jackson MJ. Is there a potential therapeutic value of copper and zinc for osteoporosis?. *Proceedings of the Nutrition Society*. 2002 May;61(2):181-5