

Zinc Deficiency in First Year Female Students of Tehran University of Medical Sciences

Fedyeh Haghollahi, M.Sc.;¹ Fatemeh Ramezanzadeh, M.D.;¹ Mansoureh Norouzi, M.Sc.;¹ Mamak Shariat, M.D.;² Atossa Mahdavi, M.D.;¹ Abbas Rahimi Foroshani, Ph.D.;³ Abdolfattah Sarafnejad, Ph.D.;⁴ Ahmad Reza Sadeghi, M.D.;⁵ Fatemeh Shahsavari, M.Sc.;⁶ Maryam Chamari, M.Sc.⁶

1 Vali-e-Asr Reproductive Health Research Center, Tehran University of Medical Sciences, Tehran, Iran

2 Maternal-Fetal-Neonatal Health Research Center, Tehran University of Medical Sciences, Tehran, Iran

3 Department of Biostatistics & Epidemiology, School of Public Health, Tehran University of Medical Sciences, Tehran, Iran

4 Department of Immunology, School of Public Health, Tehran University of Medical Sciences, Tehran, Iran

5 Student office, Tehran University of Medical Sciences, Tehran, Iran

6 Department of Biochemistry & nutrition, Tehran University of Medical Sciences, Tehran, Iran

Received February 2008; Revised and accepted March 2008

Abstract

Objective: Zinc deficiency is an important health problem in developing countries. The aim of this study was diagnosis of zinc deficiency among young female students of Tehran University of Medical Sciences.

Materials and methods: This cross sectional study evaluated 420 medical and paramedical students of Tehran University in 2005. All of them were female and in the first year of their education. Serum concentrations of zinc were measured by enzymatic method students in whom had sufficient criteria to enroll in the study. Zinc concentration more than 85 μ /dl was considered as normal value. Concentration less than 50 μ /dl was defined as severe zinc deficiency. Mild and moderate deficiencies were described as zinc levels between 50 to 85 μ /dl. Analysis was performed by t-test, chi-square and ANOVA using SPSS software.

Results: Zinc deficiency was found in 7.1% of the participants. In women with body weights between 50.1-60 kg, zinc deficiency was less than patients with body weight of \leq 50 Kg ($p=0.04$). In zinc deficient group acne was observed more frequently ($p=0.01$). Also anemic participants ($Hb<12$) had lower zinc level than non anemic women ($Hb\geq 12$) ($p=0.001$).

Conclusion: This study found severe and mild to moderate zinc deficiency in 1.4% and 5.7% of female students, respectively. Zinc deficiency is common in anemic students. Zinc supplementation is recommended in anemic patients and in cases of confirmed zinc deficiency.

Key Words: Zinc deficiency, Nutrition, Acne, BMI, Zinc

Correspondence:

Fedyeh Haghollahi, Vali-e-Asr Reproductive Health Research Center, Imam Khomeini Hospital, Keshavarz Blvd., Tehran,

14194, Iran.

Tel: 98-21-66939320 Fax: 98-21-66581658

E-mail : Vrhrc@tums.ac.ir, Fedyeh_hagh@yahoo.com

Introduction

Zinc is a trace mineral with widespread roles to sustain human health. It is a component of many enzymes and is involved in cellular division (DNA and protein synthesis), insulin activation, metabolism of ovarian and testicular hormones, liver function, behavioral development, learning, wound healing, heavy metal (e.g. lead) poisoning protection, eye accommodation and regulation of taste sensation. Also it is important in energy production, protein, carbohydrate and fat metabolism. Zinc is found in almost all body fluids and tissues. Approximately every human being contains 2 grams of this element, distributing 60% in skeletal muscles and 30% (100-200 mg) in the bone mass. High concentrations are seen in retina (274 mg/gr) and prostatic fluid (300-500mg/l). Zinc is mainly excreted in the stool (0.5-3 mg/ day, depending on its intake) and in a lesser degree in the urine, skin, seminal fluid, menstrual blood, hair and nail (1, 2). There is not a specific organ for zinc reserve, so its daily intake is needed. Dietary fibers and phytates have inhibitory effect on zinc absorption (only 20% of the dietary zinc is absorbed), so its deficiency is more common in regions with high consumption of rice and unleavened bread (1, 2).

High levels of cadmium, phosphates and ethylene-diamine-tetra-acetic acid (EDTA) in fast foods could decrease zinc concentration in plasma (2). Dietary phytates, iron supplements, gastrointestinal disorders, cigarette smoking and chronic stress could decrease serum zinc concentration (2). Daily intake of 350 µg folate decreases zinc absorption (2, 3).

Clinical signs of zinc deficiency include reduced immune system activity, skin disorders, hair loss, diarrhea, fatigue, delayed wound healing, and decreased brain development in neonates, learning difficulties and infertility. Regular daily need for men and women is at least 15 mg and 12 mg, respectively. Severe zinc deficiency adversely affects the pregnancy outcome, and results in prolonged labor, intra uterine growth retardation and fetal demise (4-7). Severe zinc deficiency could result in growth retardation, delayed puberty and behavioral disturbances (8, 9). Serum zinc is a good determinant for predicting the zinc level of the body; it ranges from 10.7 µmole/l (70 µg/dl) to 15.3 µmole/l (150 µg/dl) in adults (10).

In some studies mild to moderate deficiencies are defined as 50-85 µg/dl (10-12). Due to high phytate diets in Iran zinc deficiency is expected to be high.

According to a study conducted in Zahedan, zinc deficiency was 42.8% in teen girls and 49% in pregnant women (13). In a study in Tehran, its prevalence was 65% in high school students (14).

The present study aimed to evaluate zinc deficiency in female medical students as the future mothers in order to diagnose and treat the disorder as soon as possible.

Materials and methods

In this cross sectional (descriptive analytic) study, which was performed in 2005, all newly entered female medical students of Tehran University of Medical Sciences (Tehran, Iran) were evaluated. All of them were in the first year of the education. Exclusion criteria were defined according to students' positive answers to the following conditions: gastrointestinal and hepatic disorders, acrodermatitis entropatica, renal diseases, smoking, addiction, oral contraceptives (within three months ago), post operative period and severe burns.

From each student a 3 mL blood sample was collected and hemoglobin (Hb), hematocrit, and serum zinc concentration was measured. Serum zinc concentration was measured by enzymatic method. This method has the same sensitivity and precision in comparison with atomic absorption (15).

Zinc concentrations more than 85 µ/dl were considered as normal values. Concentrations less than 50 µ/dl were defined as severe zinc deficiency. Mild and moderate deficiencies were described as zinc levels between 50 to 85 µ/dl (16-18). When zinc deficiency was detected, the patients were treated under supervision of chancellor of student affairs.

This project was approved by the ethical committee of chancellor for research of the university and was conducted after receiving the written consents of the students.

Analysis was performed by t-test, Chi-square, Pearson correlation coefficient. Spss 11 (Spss Inc. Chicago IL.) software was used for statistical analysis and p-value less than 0.05 was considered as statistical significance.

Results

After interviewing and completion of the related questionnaires, 420 participants entered the survey. The participants had mean and standard deviation of age 18.87 ± 1.31 years, height 160.40 ± 5.41 cm, weight 56.71 ± 8.09 kg, body mass index (BMI) 21.93 ± 3.37 kg/m², hemoglobin of 13.39 ± 5.10 mg/dl, hematocrit

Table 1: Comparison of anthropometric variables between Zinc deficient and normal groups

	Zinc deficiency n (%)	Normal Zinc level n (%)	P-Value
Height (cm)			
≤156	10 (9.9)	91 (90.1)	
156.1-163	19 (10.1)	169 (89.9)	0.43
≥163.1	6 (5.8)	97 (94.2)	
Weight (kg)			
≤50	14 (12.6)	97 (87.4)	
50.1-60	8 (4.8)	158 (95.2)	0.04
≥60.1	13 (11.3)	102 (88.7)	
BMI (kg/m ²)			
≤20 (thin)	12 (10.3)	104 (89.7)	
20-25 (Normal)	17 (8.4)	186 (91.6)	0.94
25-30 (Over weight)	5 (9.3)	49 (90.7)	
≥30 (Obese)	1 (11.1)	8 (88.9)	

of 40.32±3.24%.

Mean and standard deviation of serum zinc level in participants were 104.69±18.07 µg/dl. In 24 cases (5.7%) zinc level was between 50-85 µg/dl and in 6 students (1.4%) was less than 50 µg/dl.

Hematocrit was not significantly different between the zinc deficient and normal groups (39.7±3.2 vs 40.2±3.4, respectively) (p=0.5). Also hemoglobin concentration was not significantly different between the zinc deficient and normal groups (13.3±2.5 vs 13.4±5.3, respectively) (p=0.9). Significant correlation was found between zinc level and hematocrit (p=0.024, r=0.11). Mean zinc concentration was 98.53±19.23 µg/dl in anemic students (Hb <12 mg/dl) and 105.24±18.4 µg/dl in students with normal Hb (Hb>12) (p=0.016). The mean weight (p=0.59), height (p=0.38) and BMI (p=0.31) were not significantly different between zinc deficient and normal groups.

According to height and BMI classification in zinc deficient and students with normal zinc level there was not significant difference between the two groups (p=0.43 and p=0.94, respectively). Although zinc deficiency was more common in BMI ≤ 20 kg/m² and >25 kg/m² (10.3% and 20.4%, respectively) (p=0.4). There was a significant relation between body weight and zinc deficiency (p=0.04) (Table 1).

Despite the significant relationship between acne and zinc deficiency (27.8% in zinc deficient and 12.7% in healthy students, p=0.01), there was not significant relation with irregular menses, menorrhagia (heavy bleeding) and anorexia (Table 2).

Discussion

This study showed severe and mild to moderate

zinc deficiency in 1.4% and 5.7% of female students, respectively. Al-timimi reported the prevalence of this deficiency 2.7% for severe cases and 55.7% for Mild to moderate cases (12). Haralambie observed decreased plasma zinc was 23.3% in male athletes and 43% in female athletes (11). In Hettiarachchim's research in serilanca 51.5% of boys and 58.3% of girls (aged 12-16 years old) had severe zinc deficiency (19). Zinc concentration less than 70 µ/dl was seen in 41.5% of nonpregnant females in Pathak's study (20). Prevalence of zinc deficiency in 15-80 years old population was 0.53 percent in Kuwait, which implies sufficient zinc consumption in that region (21). In a review by Kaji, one of the important health problems in the world is various levels of zinc deficiency in children and female adults. He suggested that pregnant women have inadequate intake in 80 percent of cases (and 100% in developing countries) (22). Andriollo-Sanchez showed this deficiency was less than 5 percent in Europe (23).

Despite the high prevalence of zinc deficiency in the aforementioned studies, our study on newly entered medical students showed that 7.1% of them were zinc deficient. Although the students were from different regions of the country (and probably with better nutritional, socioeconomic, cultural situations to leave their homes in order to educate in the largest university of Iran), but it seems that they are not representative of the whole population.

Although mean hematocrit and hemoglobin concentrations were not significantly different between the zinc deficient and normal groups, but significant correlation was found between zinc level and hematocrit. Mean zinc concentration was lower in anemic students. Other studies confirm these results (24-27).

Table 2: Comparison of Clinical findings between Zinc deficient and normal groups

	Zinc deficiency n (%)	Normal Zinc level n (%)	P-Value
Acne			
No	26 (72.2)	317 (87.3)	0.01
Yes	10 (27.8)	46 (12.7)	
Irregular Menstrual Cycle			
No	28 (77.8)	290 (79.9)	0.45
Yes	8 (22.2)	73 (20.1)	
Menorrhagia			
No	30 (83.3)	331 (91.4)	0.16
Yes	6 (16.7)	31 (8.6)	

Regular Menstrual Cycle: Menstruation with 21-35 days interval

Menorrhagia: Menstrual bleeding more than eight days and bleeding with clot

Ai-Guoma described the relation of lowered plasma zinc and anemia (24), So he recommended that in iron deficiency anemia, zinc deficiency must be considered.

In this research, there was a significant relation between body weight and zinc deficiency (especially in ≤ 50 kg and ≥ 60 kg), which implies the possible role of malnutrition such as anorexia or bulimia. Gokbel showed lowered leptin levels in zinc deficient people (28). The study of Ghayour-Mobarhan showed lower serum zinc level in people with obesity (due to zinc role in catabolism). He described that in zinc deficiency, leptin is lowered and catabolism decreases (29, 30). Smidt described that the effect of zinc on fat metabolism, although subsequent regulation of leptin secretion is not well known (31). In some studies its role on decreasing appetite (1, 3, 31, 32) and in others its role on regulating the appetite is validated (1,3,21,27,33,34). In our research 13.9 % of zinc deficient students had anorexia. Some researchers believe that in menstruation periods decrease in hematocrit and hemoglobin concentrations results in lower zinc levels (3, 25, 26 and 35). Although in this study 16.7 % of zinc deficient students had menorrhagia. Significant relationship between acne and zinc deficiency (27.8% in zinc deficient and 12.7% in healthy students, $p=0.01$), was observed. Zinc deficiency is related to skin disorders (3, 4). Nigman described the relationship of this deficiency and psoriasis (10).

Zinc deficiency prevention is not only considered as an important issue for preventing maternal mortality, malaria and other diseases, but also nutritional state of developing countries is one of the third thousand targets (36,37). So evaluation of plasma zinc or its daily intake is indicated in order to schedule future interventional measures (38).

Future studies are recommended to explain the low prevalence in these students compared to general population. According to the effects of zinc deficiency on public health (especially maternal health), well designed studies are warranted to evaluate the true prevalence of this health problem in our community. It will also be interesting to repeat this survey, on the same sample of students just before completion of their education to compare the findings and explain any changes that might occur during the period of their education.

Acknowledgement

The authors wish to thank the deputy of research of Tehran University of medical sciences who provided the financial support for performing this study, and Vali-Asr health research center and staff of Noor laboratory (especially Mr. K. Majidi) for their cooperation in this survey.

There exists no conflict of interest to declare.

References

1. Tuormaa TE. Adverse effects of zinc deficiency: A review from the literature. *The Journal of Orthomolecular Medicine* 1995; 10: 150-64.
2. Brown KH, Wuehler SE, Peerson JM. The importance of Zinc in human nutrition and estimation of the global prevalence of Zinc deficiency. *Food Nutr Bull* 2001; 22: 113-25.
3. Tamura T, Goldenberg RL, Johnston KE, DuBard M. Maternal plasma zinc concentrations and pregnancy outcome. *Am J Clin Nutr* 2000; 71: 109-13.
4. King JC. Determinants of maternal zinc status during pregnancy. *Am J Clin Nutr* 2000; 71 Suppl 5: S1334-43.
5. Rounsipragarn R, Borirug S, Herabutya Y. Plasma zinc level and intrauterine growth retardation: a study in pregnant women in Ramathibodi Hospital. *J Med*

- Assoc Thai 1999; 82: 178-81.
6. Martin-Lagos F, Navarro-Alarcon M, Terres-Martos C, Lopez-Garcia de la Serrana, Perez-Valero V, Lopez-Martinez MC. Zinc and copper concentrations in serum from Spanish women during pregnancy. *Biol Trace Elem Res* 1998; 61: 61-70.
 7. Kumru S, Aydin S, Simsek M, Sahin K, Yaman M, Ay G. Comparison of serum copper, zinc, calcium, and magnesium levels in preeclamptic and healthy pregnant women. *Biol Trace Elem Res* 2003; 94: 105-12.
 8. Gabrichidze G, Kintrajia P, Eristvi Z. Dynamic of zinc containing at physiological pregnancy. *Annals of Biomedical Research and education* 2001; 1: 248-50.
 9. Solomons NW. Trace elements. In: Rombeau JL, Caldwell MD, eds. *Clinical Nutrition: Parenteral Nutrition*. Philadelphia, Pa: WB Saunders, 1992; 150-83.
 10. Nigam PK. Serum zinc and copper levels and Cu: Zn ratio in psoriasis. *Indian J Dermatol Venereol Leprol* 2005; 71: 205-6.
 11. Haralambie G. Serum zinc in athletes in training. *Int J Sports Med* 1981; 2: 135-8.
 12. Al-Timimi DJ, Al-Sharbatti SS, Al-Najjar F. Zinc deficiency among a healthy population in Baghdad, Iraq. *Saudi Med J* 2005; 26: 1777-81.
 13. Salimi S Yaghmaei M, Joshaghani HR, Mansourian AR. Study of Zinc Deficiency in Pregnant Women. *Iranian J Publ Health* 2004; 33: 15-8.
 14. Mahmoodi MR. Epidemiologic assessment of zinc deficiency in high school students of Tehran city. Thesis of MSc in Nutrition, Tehran University of Medical Sciences, 1996.
 15. Erel O, Avci S. Semi-automated enzymatic measurement of serum zinc concentration. *Clin Biochem* 2002; 35: 41-7.
 16. Salgueiro MJ, Weill R, Zubillaga M, Lysionek A, Caro R, Goldman C, et al. Zinc deficiency and growth: current concepts in relationship to two important points: intellectual and sexual development. *Biol Trace Elem Res* 2004; 99: 49-69.
 17. FAO/WHO. Human vitamin and mineral requirements. In: *Human vitamin and mineral requirements*. Bangkok, Thailand: FAO/WHO, 2002.
 18. Pathak P, Kapil U, Kapoor SK, Dwivedi SN, Singh R. Magnitude of zinc deficiency among nulliparous nonpregnant women in a rural community of Haryana State, India. *Food Nutr Bull* 2003; 24: 368-71.
 19. Hettiarachchi M, Liyanage C, Wickremasinghe R, Hilmers DC, Abrahams SA. Prevalence and severity of micronutrient deficiency: a cross-sectional study among adolescents in Sri Lanka. *Asia Pac J Clin Nutr* 2006; 15: 56-63.
 20. Rwebembera AA, Munubhi EK, Manji KP, Mpembeni R, Philip J. Relationship between infant birth weight <2000 g and maternal zinc levels at Muhimbili National Hospital, Dar Es Salaam, Tanzania. *J Trop Pediatr* 2006; 52: 118-25.
 21. Abiaka C, Olusi S, Al-Awadhi A. Reference ranges of copper and zinc and the prevalence of their deficiencies in an Arab population aged 15-80 years. *Biol Trace Elem Res* 2003; 91: 33-43.
 22. Kaji M. Zinc in Endocrinology. *International Pediatrics* 2001; 16: 1-7.
 23. Ndrillo-Sanchez M, Hinger-Favier I, Meunier N, Toti E, Zaccaria M, Brandolini-Bunlon M, et al. Zinc intake and status in middle-aged and older European subjects: the ZENITH study. *Eur J Clin Nutr* 2005; 59 Suppl 2: S37-41.
 24. Ma AG, Chen XC, Xu RX, Zheng MC, Wang Y, Li JS. Comparison of serum levels of iron, zinc and copper in anaemic and non-anaemic pregnant women in China. *Asia Pac J Clin Nutr* 2004; 13: 348-52.
 25. Wright EA, Kapu MM, Isichei UP. Zinc depletion and menorrhagia in Nigerians using copper T-200 intra-uterine device. *Trace Elem Med* 1989; 6: 147-9.
 26. Aschkenazi S, Kaplan B, Shaklai M, Blickstein D, Bar J, Ben-Rafael Z. Red blood cell zinc protoporphyrin measurement for assessment of peripartum iron deficiency. *Clin Exp Obstet Gynecol* 1999; 26: 183-4.
 27. Ahamed M, Singh S, Behari JR, Kumar A, Siddiqui MK. Interaction of lead with some essential trace metals in the blood of anemic children from Lucknow, India. *Clin Chim Acta* 2007; 377: 92-7.
 28. Gokbel H, Baltaci AK, Uçok K, Okudan N, Mogulkoc R. Changes in serum leptin levels in strenuous exercise and its relation to zinc deficiency in rats. *Biol Trace Elem Res* 2005; 106: 247-52.
 29. Ghayour-Mobarhan M, Taylor A, New SA, Lamb DJ, Ferns GA. Determinants of serum copper, zinc and selenium in healthy subjects. *Ann Clin Biochem* 2005; 42: 364-75.
 30. Di MG, Matera MG, De MB, Vacca C, Di MS, Rossi F. Relationship between zinc and obesity. *J Med* 1993; 24: 177-83.
 31. Smidt K, Pedersen SB, Brock B, Schmitz O, Fisker S, Bendix J, et al. Zinc-transporter genes in human visceral and subcutaneous adipocytes: lean versus obese. *Mol Cell Endocrinol* 2007; 264: 68-73.
 32. Gibson RS. Zinc: the missing link in combating micronutrient malnutrition in developing countries. *Proc Nutr Soc* 2006; 65: 51-60.
 33. Marreiro DN, Geloneze B, Tambascia MA, Lerario AC, Halpern A, Cozzolino SM. Effect of zinc supplementation on serum leptin levels and insulin resistance of obese women. *Biol Trace Elem Res* 2006; 112: 109-18.
 34. Lee SL, Kwak EH, Kim YH, Choi JY, Kwon ST, Beattie JH, et al. Leptin gene expression and serum leptin levels in zinc deficiency: implications for appetite regulation in rats. *J Med Food* 2003; 6: 281-9.
 35. Songchitsomboon S, Komindr S. Serum zinc and copper in healthy adults living in Bangkok and

- surrounding districts. *J Med Assoc Thai* 1996; 79: 550-7.
36. Muller O, Krawinkel M. Malnutrition and health in developing countries. *CMAJ* 2005; 173: 279-86.
37. Faber M, Kvalsvig JD, Lombard CJ, Benade AJ. Effect of a fortified maize-meal porridge on anemia, micronutrient status, and motor development of infants. *Am J Clin Nutr* 2005; 82: 1032-9.