



Research Article

Links between Leukemia, Obesity and Vitamin-D Deficiency in a Cross Sectional Study in the North of Saudi Arabia

Alenezy A¹, Alenzi FQB², Alenzi SM³, Alenzi FQA⁴, Alenzi SMM⁵, Abo El-Fetoh NM⁶, Alhowaish JA¹, Alhowaish MA¹, Al-bukhari TA⁷ and Arrowaili A⁸

¹Department of Family and Community Medicine, college of Medicine Northern Border University, Arar, Saudi Arabia

²College of Applied Medical Sciences, Prince Sattam bin Abdulaziz University, Al-Kharj, Saudi Arabia

³Department of Imaging, KFMC, Riyadh, Saudi Arabia

⁴Riyadh College of Dentistry and Pharmacy, Riyadh, Saudi Arabia

⁵Dept. of Pharmacy, KAUH, KSU, Riyadh, Saudi Arabia

⁶Department of Community medicine, college of medicine, Sohage University, Egypt

⁷Department of Immunology, College of Medicine, UQU, Makkah, Saudi Arabia

⁸Department of Surgery, College of Medicine, Imam Mohamed bin Saud University, Riyadh, Saudi Arabia

*Corresponding author: Faris Q Alenzi, Professor of Immunology, College of Applied Medical Sciences
Prince Sattam bin Abdulaziz University (PSAU), Saudi Arabia, E-mail: fqalenzi@ksu.edu.sa

Rec date: June 11, 2016 Acc date: June 14, 2016 Pub date: June 23, 2016

Abstract

Introduction: Vitamin D deficiency has recently been recognized as a worldwide concern and still linked to obesity. Despite its high prevalence, no previous community based studies have been conducted in Arar; northern border of Kingdom of Saudi Arabia (KSA), addressing this issue could be traced. This study was carried out to show the magnitude of the problem of vitamin D deficiency and how such a deficiency can eventually has associated with overweight and obesity in female Arar population, KSA.

Methods: A cross-sectional study was carried out during the period from 1, November 2015 to 29, February 2016. A total of 185 female subjects aged 18–35 years, attending five randomly selected primary healthcare centers were selected using a systemic random sampling procedure. Data were collected by personal interview using a predesigned questionnaire including the relevant questions for obtaining the needed data. Anthropometric examination included height and weight measurements were obtained. Body Mass Index (BMI) was calculated. Serum levels of 25-OH vitamin D were measured by ELISA.

Results: Majority (58.9%) of the participants had deficient - lower than 12 ng/ml Vitamin Level, 18.9% had insufficient level (12-20 ng/ml) and about fifth (19.5%) only had normal level (20-50 ng/ml). More than one third (34.9) of cases of severe Vitamin D deficiency were severely obese, more than third 31.2% were obese, 14.7% were overweight and less than fifth (19.3%) were normal weight. Most (42.9%) of cases with

Vitamin D level 12-20 were overweight, most (38.9) of normal Vitamin D level participants had normal body weight and only 16.7% of them were obese.

Conclusion: Vitamin D deficiency is a common problem in females in Arar, KSA and vitamin D deficiency is strongly associated with obesity. Nutritional education programs are needed.

Keywords: Vitamin D
Kingdom of Saudi Arabia

Overweight; Obesity; Females;

Introduction

Apparently, obesity is associated with a low-grade of white adipose tissue (WAT) resulting from chronic activation of the innate immune system. Excess fat is thought to stimulate white blood cells (WBCs) that produce molecules as a part of the normal immune response upon injury or infection. Obese adipose tissue mainly releases cytokines including: TNF- α , IL-6 and leptin. At biology level, it can be by a variety of complement components. As the common factor linking adipose tissue to the metabolic context of obesity, insulin resistance and atherosclerosis are associated with a low-grade chronic status, to which the complement system is an important contributor (Figure 1A and Figure 1B).

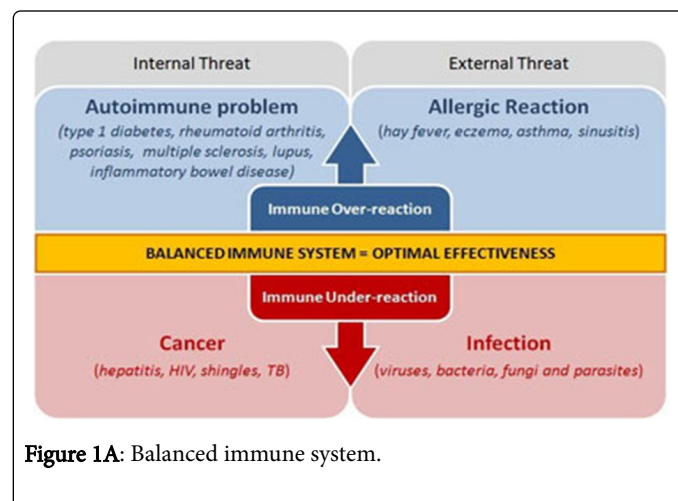
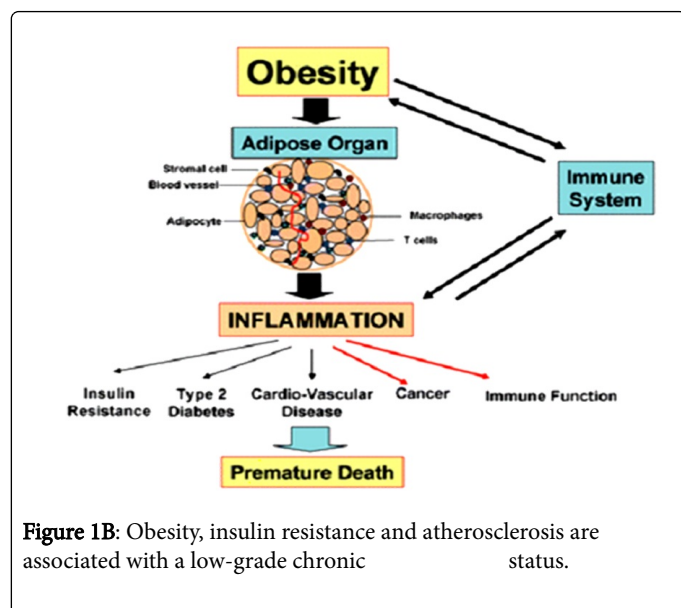


Figure 1A: Balanced immune system.

increasing prevalence of obesity [1] is turning type 2 diabetes into one of the most frequent causes of death [2]. Similarly, vitamin D

has recently been recognized as a worldwide concern [3], still linked to obesity. Approximately 1 billion people worldwide from vitamin D [4], which may result from limited exposure to sunlight, long-term wearing of covering clothes, use of sunscreen, age as well as low consumption of food containing ergocalciferol, and mal-absorption syndrome [5].

Several studies have found an association between vitamin D and a cluster of metabolic abnormalities called the 'metabolic syndrome', including abdominal obesity, insulin resistance, dyslipidemia, and hypertension, with the consequent risk of developing cardiovascular diseases and/or type 2 diabetes [6].



concomitant association of vitamin D with insulin resistance, impaired insulin secretion, and their important metabolic consequences has generated the hypothesis of a possible role of vitamin D in the pathogenesis of type 2 diabetes [7].

prevalence of vitamin D was 35% higher in obese subjects compared to the eutrophic group and 24% higher than in the overweight group. results indicate that the prevalence of vitamin D was more elevated in obese subjects [8].

is a link between 25(OH)D levels and insulin responsiveness of tissues as well as between glucose levels and glycosylated hemoglobin in people without diabetes mellitus type 2 [9].

However, data from other authors controvert the relation of vitamin D and metabolic syndrome factors [10,11]. Given these contradicting data, we sought to determine whether serum 25(OH)D concentration in late reproductive age healthy women is associated with body composition and glucose metabolism.

In a study conducted in St. Petersburg (North-West region of Russia), to show if the glucose metabolism and body fat content depend on serum levels of 25-hydroxyvitamin D [25(OH)D], it was considered that, 25(OH)D levels were from 19.4 to 134.0 nMol/L (mean 52.9 ± 22.7). Vitamin D (lower than 50 nMol/L) and (50-75 nMol/L) was revealed in 59.1% and 27.8% of women, respectively. study showed also that low 25(OH)D levels were associated with obesity [12]. study was carried out to show the magnitude of the problem of vitamin D and how such a can eventually has associated with overweight and obesity in female Arar population, KSA.

Methods

Study sitting

present study was conducted in Arar. Arar is the regional headquarter of the Northern Border Province of Saudi Arabia.

Study type

A cross-sectional study was carried out during the period from 1, November 2015 to 29, February 2016.

Sampling

Total of 185 female subjects aged 18–35 years attending randomly selected primary healthcare centers in Arar city. were selected using a systemic random sampling procedure. Each woman was interviewed separately, and was assured. Health centers provide healthy and sick women with healthcare services in an acceptable atmosphere of both privacy and

Data collection methods

Data were collected by means of personal interview with the sampled women using a predesigned questionnaire covering the following items:

Socio-demographic characteristics of the woman, including age, educational and marital status.

Data related to Vitamin D level and body weight as physical activity, exposure to sunlight, and consumption of milk, dairy products and egg, performing muscular exercise, history of consumption of fatty meals, consumption of fruits and vegetables and family history of obesity was obtained.

exclusion criteria were, Menopause, Diabetes mellitus, liver or kidneys disease and mal-absorption syndrome. Anthropometric examination included height and weight measurements with the use of a calibrated balance beam scale and a wall-mounted stadiometer and calculation of body mass index (BMI). Normal weight was as $BMI < 25 \text{ kg/m}^2$, overweight as $25 \leq BMI < 30 \text{ kg/m}^2$ and obesity as $BMI \geq 30 \text{ kg/m}^2$ [13].

Serum levels of 25-OH vitamin D were measured by ELISA. Status of vitamin D was as: normal - 25(OH)D levels higher than 50(ng/ml); - 20 to <50(ng/ml); and - low than 20(ng/ml) and severely if <12(ng/ml)[5].

Ethical considerations

study was reviewed and approved by the Research Ethics Committee of Faculty of Medicine, Northern Border University. Participants were informed that participation is completely voluntary, and written consent was obtained from each participant before being subjected to the questionnaire and discussing the objective with the participants. No names were recorded on the questionnaires. Adequate training of data collectors took place to ensure protection of and all questionnaires were kept safe.

Statistical analysis

Collected data were coded and analyzed using statistical package for the social sciences (SPSS, version 15). w2-test was used as a test of and were considered at P value 0.05 or less.

Results

Table 1 illustrates some socio-demographic characteristics, Vitamin D level and BMI of the studied women. majority of women participating in the study were between 18 and 28 years of age, and

only 35% were between 28 and 35 years of age. As regards the educational status of the participants, most of participants (82.1%) had university graduates and about one quarter (17.8%) had lower educational levels. As marital status, about two-thirds (66.4%) of the were single, nearly one-third (29.2%) were married. As regards Vitamin D level, unfortunately, the majority (58.9%) of the participants had - lower than 12 ng/ml Vitamin D level, 18.9% had level (12-20 ng/ml); about (19.5%) only had normal level (20-50 ng/ml) and 2.7% had level >50 ng/ml. As regards the BMI, only 26.5% of the participants had normal body weight, more than (23.2) were overweight, about quarter (23.8%) were obese, nearly quarter (24.9%) were very obese with BMI>35 and only 1.6% of participants were underweight.

| Parameter | No. (%) |
|----------------------|------------|
| Age group | |
| 18- | 120 (64.8) |
| 28-35 | 65 (35.0) |
| Mean±SD | 27±4.3 |
| Educational level | |
| University education | 152 (82.1) |
| Others | 33 (17.8) |
| Marital status | |
| Single | 123 (66.4) |
| Married | 54 (29.2) |
| Divorced | 5 (2.7) |
| Widowed | 3 (1.6) |

| Vitamin D level (ng/ml) | |
|---|---------------|
| <12 | 109 (58.9) |
| (12-20) | 35 (18.9) |
| (20-50) | 36 (19.5) |
| >50 | 5 (2.7) |
| Mean ± SD | 19.55 ± 11.28 |
| Body mass index (BMI) kg/m ² | |
| <18.5 | 3 (1.6) |
| 18.5-24.9 | 49 (26.5) |
| 25-29.9 | 43 (23.2) |
| 30-34.9 | 44 (23.8) |
| >35 | 46 (24.9) |
| Mean ± SD | 27.8 ± 10.24 |

Table 1: Some socio-demographic characteristics, Vitamin D level and BMI of the studied women, Arar, KSA (n=185).

Table 2 illustrates the relationship between Vitamin D level and age group and educational level of the studied population. study revealed that severe Vitamin D was most common in the age group 18-28 years (58.7%) compared with age group 28-35 years, with a statistically (Po 0.05). majority of the studied population (75.2%) with sever Vitamin D (<12) completed their university education, and so most of the participants (97.1%) with Vitamin D level 12-20 with a statistically

| Parameter | Vitamin D level (ng/ml) | | | | Total(n=185) | P value | |
|----------------------|-------------------------|-------------------|-------------------|--------------|--------------|---------|--|
| | < 12 (n=109) | (12-20) (n=35) | (20-50) (n=36) | >50 (n=5) | | | |
| | No. (%) | No. (%) | No. (%) | No. (%) | No. (%) | | |
| Age group | | | | | | | |
| 18- | 64 (58.7) | 29 (82.9) | 23 (63.9) | 4 (80.0) | 120 (64.8) | 0.049 | |
| 28-35 | 45 (41.3) | 6 (17.1) | 13 (36.1) | 1 (20.0) | 65 (35.0) | | |
| Educational level | | | | | | | |
| University education | 82 (75.2) | 34 (97.1) | 32 (88.9) | 4 (80.0) | 152-82.1 | 0.156 | |
| Others | 27 (24.7) | 1 (2.9) | 4 (11.1) | 1 (20.0) | 33-17.8 | | |

Table 2: Relationship between Vitamin D level and age group and educational level of the studied women, Arar, KSA.

Table 3 illustrates the relationship between Vitamin D level and BMI, Exposure to sunlight, Consumption of milk, dairy products and egg, Performing muscular exercise, Consumption of fruits and vegetables, Preferring fatty meals in the studied women. More than one third (34.9) of cases of sever Vitamin D were severely obese, more than third 31.2% were obese, 14.7% were overweight and less than (19.3%) were normal weight. Most (42.9%) of cases with

Vitamin D level 12-20 were overweight, most (38.9) of normal Vitamin D level participants had normal body weight and only 16.7% of them were obese, most (57.8%) of sever Vitamin D and 71.4 of 12-20 serum level, don't exposed to sunlight with a statistically (Po0.05).More than half (55.6%) of participants with normal Vitamin D level consumed milk and dairy products and egg daily compared with 43.1% of sever Vitamin D with a

statistically More than half (52.8%) of participants with normal Vitamin D level were performing muscular exercise compared with 40% of cases of Vitamin D level 12-20. 49.6% of sever Vitamin D cases rarely consume fruits and vegetables compared with 41.7% of participants with normal Vitamin D level with a statistically Most of (61.1%) of participants with normal Vitamin D level don't like fatty meals compared with 50.4% of case of sever Vitamin D and 40% of cases of Vitamin D level 12-20 with a statistically

| Parameters | Vitamin D level (ng/ml) | | | | Total (n=185) | P value |
|--|-------------------------|-------------------|-------------------|--------------|---------------|---------|
| | <12 (n=109) | (12-20) (n=35) | (20-50) (n=36) | >50 (n=5) | No. (%) | |
| | No. (%) | No. (%) | No. (%) | No. (%) | | |
| BMI | | | | | | |
| <18.5 | 0(0.0) | 1(2.9) | 2(5.6) | 0(0.0) | 3(1.6) | *NA |
| 18.5-24.9 | 21(19.3) | 12(34.3) | 14(38.9) | 2(40.0) | 49(26.5) | |
| 25-29.9 | 16(14.7) | 15(42.9) | 10(27.8) | 2(40.0) | 43(23.2) | |
| 30-34.9 | 34(31.2) | 4(11.4) | 6(16.7) | 0(0.0) | 44(23.8) | |
| >35 | 38(34.9) | 3(8.6) | 4(11.1) | 1(20.0) | 46(24.9) | |
| Exposure to sunlight | | | | | | |
| No | 63(57.8) | 25(71.4) | 18(50.0) | 3(60.0) | 109(58.9) | 0.000 |
| Yes | 46(42.2) | 10(28.6) | 18(50.0) | 2(40.0) | 76(41.1) | |
| Consumption of milk, dairy products and egg | | | | | | |
| Daily | 47(43.1) | 22(62.9) | 20(55.6) | 1(20.0) | 90(48.6) | 0.769 |
| Weekly | 32(29.4) | 7(20.0) | 7(19.4) | 2(40.0) | 48(25.9) | |
| Rare | 30(25.5) | 6(17.1) | 9(25.0) | 2(40.0) | 47(25.1) | |
| Performing muscular exercise for keeping Vitamin D level | | | | | | |
| Yes | 40(63.3) | 14(40.0) | 19(52.8) | 3(60.0) | 76(41.1) | 0.607 |
| No | 69(36.7) | 21(60.0) | 17(47.2) | 2(40.0) | 109(58.9) | |
| Consumption of fruits and vegetables | | | | | | |
| Yes daily | 22(20.2) | 8(22.9) | 10(27.8) | 1(20.0) | 40(21.6) | 0.253 |
| Weekly | 33(30.3) | 17(48.6) | 11(30.6) | 2(40.0) | 64(34.6) | |
| Rarely | 54(49.6) | 10(28.6) | 15(41.7) | 2(40.0) | 81(43.8) | |
| Preferring fatty meals | | | | | | |
| Yes | 54(49.6) | 16(45.7) | 14(38.9) | 1(20.0) | 85(45.9) | 0.857 |
| No | 55(50.4) | 17(48.6) | 22(61.1) | 4(80.0) | 80(43.3) | |

Table 3: Relationship between Vitamin D level and BMI, exposure to sunlight, consumption of milk, dairy products and egg, performing muscular exercise, consumption of fruits and vegetables and preferring fatty meals in the studied women, Arar, KSA

Discussion

Accumulating evidence suggests vitamin D plays a role in the development of obesity. People usually do not used to supplementing vitamin D, since vitamin D can be produced when their skin is exposed to the sunlight. Nevertheless, even in highly sunny regions, vitamin D exists; suggesting vitamin D is a global problem. cross sectional study was conducted in Arar city, the

capital of the Northern Province of Saudi Arabia, we aimed to determine the association between vitamin D and overweight and obesity. A total of 185 female subjects with mean age (\pm SD) 27 years (\pm 4.3), were included in the study. 25-hydroxy vitamin D (25(OH)D), was measured in blood samples. Body mass index (BMI) was calculated. majority (58.9%) of the participants had (lower than 12 ng/ml) Vitamin D level, 18.9% had level (12-20 ng/ml) and 19.5% only had normal level (20-50 ng/ml),

with Mean level (\pm SD) were 19.55 ± 11.28 ng/dL. are in accordance with of Russian study which reported that, vitamin D (lower than 50 nMol/L) and (50-75 nMol/L) was revealed in 59.1% and 27.8% of women, respectively. study showed also that low 25(OH)D levels were associated with obesity [12]. Our are also in accordance with Kavadar et al. [13] who found that, decreased physical activity (PA) and obesity have been associated with the low vitamin D levels. Mean \pm SD Vitamin D level in the non-diabetic population in his study were 17.62 ± 10.47 ng/dL [13].

As regards the BMI, only 26.5% of the participants had normal body weight, 23.2% were overweight, 23.8% were obese, nearly quarter (24.9%) were very obese with BMI >35 and only 1.6% of participants were underweight with Mean (\pm SD) of BMI was 27.8 ± 10.24 .

is in accordance with Kavadar et al. [14] who found Mean (\pm SD) of BMI in non-diabetic population were 28.2 ± 3.16 kg/m².

In the current study, (34.9) of cases of sever Vitamin D were severely obese, 31.2% were obese, 14.7% were overweight and (19.3%) were normal weight. 42.9% of cases with Vitamin D level 12-20 were overweight, 38.9 of normal Vitamin D level participants had normal body weight and only 16.7% of them were obese.

are supported with of systematic review of Santos M et al. [8] who reported that, the prevalence of vitamin D was 35% higher in obese subjects compared to the eutrophic group and 24% higher than in the overweight group. are also supported by other epidemiological studies which demonstrated that vitamin D is closely related to obesity [15].

are again supported by of Karlsson et al. [14] free 25(OH)D was 13.3 ± 5.5 (obese) and 23.7 ± 10.7 (normal-weight). obese women had a 20.1 nmol/L lower mean 25(OH)D concentration compared to normal-weight women. 56 % of obese women and 12 % of normal-weight women had 25(OH)D concentrations ≤ 50 nmol/L [16]. Moreover, Meta-analysis results showed that the prevalence of vitamin D was between obesity group and control group, and the prevalence of vitamin D was associated with obesity in Asians and European-Americans [17].

Several cross-sectional studies revealed a strong correlation between the low vitamin D levels and obesity, which is also commonly associated with insulin resistance [18-20]. It was found that the combination of vitamin D and obesity had an impact on the risk of insulin resistance [21]. In adolescents with mean age (\pm SD) was 14.9 ± 1.4 Ys; the mean (\pm SD) body mass index (in kg/m²) was 36 ± 5 . Seventeen of the adolescents were vitamin D [22]. In the PCOS patients, 97.2% were obese and vitamin D was noted among 62.2% them. mean serum 25(OH)D level was 18.4 and 21.6 ng/mL in PCOS and control groups respectively. in mean 25(OH)D levels between the 2 groups was not statistically [23].

In conclusion, there are and immune system responses in overweight and obese individuals unique opportunities for intervention strategies to help ameliorate the risk of obesity-associated disease such as hypertension and DM.

Acknowledgements

project was supported by a research grant from the Deanship of research at Prince Sattam bin Abdulaziz University (PSAU) (REF NO 2014-01-2758).

References

1. Flegal KM, Carroll MD, Ogden CL, Curtin LR (2010) Prevalence and trends in obesity among US adults, 1999-2008. *JAMA* 303: 235-241.
2. Mokdad AH, Ford ES, Bowman BA, Dietz WH, Vinicor F, et al. (2003) Prevalence of obesity, diabetes, and obesity-related health risk factors, 2001. *JAMA* 289: 76-79.
3. Binkley N, Ramamurthy R, Krueger D (2010) Low vitamin D status: prevalence, consequences, and correction. *Endocrinol Metab Clin North Am* 39: 287-301.
4. Adams JS, Hewison M (2010) Update in vitamin D. *J Clin Endocrinol Metab* 95: 471-478.
5. Bouillon R, Carmeliet G, Verlinden L, van Etten E, Verstuyf A, et al. (2008) Vitamin D and human health: lessons from vitamin D receptor null mice. *Endocr Rev* 29: 726-776.
6. National Cholesterol Education Program (200) Expert Panel on Detection, Evaluation, and Treatment of High Blood Cholesterol in Adults (Adult Treatment Panel III): Report of the National Cholesterol Education Program (NCEP) Expert Panel on Detection, Evaluation, and Treatment of High Blood Cholesterol in Adults (Adult Treatment Panel III) report. *Circulation* 106: 3143-3421.
7. Hyppönen E, Power C (2006) Vitamin D status and glucose homeostasis in the 1958 British birth cohort: the role of obesity. *Diabetes Care* 29: 2244-2246.
8. Pereira-Santos M, Costa PR, Assis AM, Santos CA, Santos DB (2015) Obesity and vitamin D a systematic review and meta-analysis. *Obes Rev* 16: 341-349.
9. Kositsawat J, Freeman VL, Gerber BS, Geraci S (2010) Association of A1C levels with vitamin D status in U.S. adults: data from the National Health and Nutrition Examination Survey. *Diabetes Care* 33: 1236-1238.
10. Robinson JG, Manson JE, Larson J, Liu S, Song Y, et al. (2011) Lack of association between 25(OH)D levels and incident type 2 diabetes in older women. *Diabetes Care* 34:628-634.
11. Gulseth HL, Gjelstad IMF, Tierney AC, Lovengrove JA, Defoort C, et al. (2010) Serum vitamin D concentration does not predict insulin action or secretion in European subjects with the metabolic syndrome. *Diabetes Care* 33: 923-925.
12. Grineva EN, Karonova T, Mischeva E, Belyaeva O, Nikitina IL (2013) Vitamin D is a risk factor for obesity and diabetes type 2 in women at late reproductive age. *Aging (Albany NY)* 5: 575-581.
13. WHO Expert Consultation (2004) Appropriate body-mass index for Asian populations and its implications for policy and intervention strategies. *Lancet* 363: 157-163.
14. Kavadar G, Demircio Lu DT, Emre TY (2015) relationship between vitamin D status, physical activity and insulin resistance in overweight and obese subjects. *Bosn J Basic Med Sci* 15: 62-66.
15. Li YX, Zhou L (2015) Vitamin D Obesity and Diabetes. *Cell Mol Biol (Noisy-le-grand)* 61: 35-38.
16. Karlsson, Amra Osmancevic, Nina Jansson, Lena Hulthén, Agneta Holmäng, et al. (2014) Increased vitamin D-binding protein and decreased free 25(OH)D in obese women of reproductive age. *European Journal of Nutrition* 53: 259-267.
17. Yao Y, Zhu L, He L, Duan Y, Liang W, et al. (2015) A meta-analysis of the relationship between vitamin D and obesity. *Int J Clin Exp Med* 8: 14977-14984.

18. Wortsman J, Matsuoka LY, Chen TC, Lu Z, Holick MF (2000) Decreased bioavailability of vitamin D in obesity. *Am J Clin Nutr* 72: 690-693.
19. Lu L, Yu Z, Pan A, Hu FB, Franco OH, et al. (2009) Plasma 25-hydroxyvitamin D concentration and metabolic syndrome among middle-aged and elderly Chinese individuals. *Diabetes Care* 32: 1278-1283.
20. Ou HY, Karnchanasorn R, Lee LZ, Chiu KC (2011) Interaction of BMI with vitamin D and insulin sensitivity. *Eur J Clin Invest* 41: 1195-1201.
21. Kabadi SM, Lee BK, Liu L (2012) Joint of obesity and vitamin D on insulin resistance and type 2 diabetes: results from the NHANES 2001-2006. *Diabetes Care* 35: 2048-2054.
22. Carine M Lenders, Henry A Feldman, Emily Von Scheven, Anne Merewood, Carol Sweeney, Darrell M Wilson, et al. (2009) Relation of body fat indexes to vitamin D status and among obese adolescents *Am J Clin* 90: 459-467.
23. Sadhir M, Kansra AR2, Menon S3 (2015) Vitamin D among Adolescent Females with Polycystic Ovary Syndrome. *J Pediatr Adolesc Gynecol* 28: 378-381.