Association Of Vitamin D Deficiency With Risk Factors In Postmenopausal Women

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ABSTRACT

In conditions of age-related decrease in sex hormones and a number of pathological conditions and diseases in postmenopausal women, there is a deficiency of D hormone. In our country, the geographic location of which is below northern latitude and sufficient ultraviolet radiation, an insufficient content of vitamin D is seen in postmenopausal women. There are a lot of risk factors leading to vitamin D deficiency - the presence of smog and dust in cities, insufficient consumption of vitamin-fortified foods, the presence of problems with the gastrointestinal tract and excretory system and a number of others. One of the important factors that reduce vitamin storage is overweight and obesity, especially in combination with old age, when all absorption processes are reduced. The aim of our research was to study risk factors in women with vitamin D deficiency with subsequent correction of the deficiency state. We examined the level of total 25 (OH) D in the blood serum in 46 postmenopausal women, and identified risk factors. Vitamin D deficiency was detected in 86.96% of women, and its deficiency was registered in 10.87%. At the same time, a pronounced vitamin deficiency was not registered in any patient. Overweight was registered in 32.6%, obesity of varying degrees in 26.1%. Given the indicators, recommendations were given for correcting vitamin D deficiency. All postmenopausal women, especially those with risk factors for deficiency, are recommended to determine the basic level of vitamin D. In case of deficiency, drug correction is recommended to reduce the risk of cardiovascular and oncological diseases.

KEYWORDS

Vitamin D, postmenopausal women, risk factors, obesity, hypovitaminosis, correction
INTRODUCTION

Vitamin D was recently known as a regulator of calcium-phosphorus metabolism. Recently, however, a lot of information has appeared on the effect of vitamin D on other physiological parameters of the body, both women and men [9, 14]. Vitamin D is chemically similar to progesterone. Today it is considered a steroid hormone with progesterone-like activity and its effect on the body is great [3]. There is enough information in the literature on the effect on reproductive function - maintaining the luteal phase, ensuring endometrial receptivity, anti-inflammatory effect, as well as the effect on pregnancy - reducing the frequency of premature birth and late spontaneous miscarriages, low birth weight [1,2,4,12]. Given the association of vitamin D deficiency with a decrease in fertility, as well as with a worsening of maternal and perinatal pregnancy outcomes, it is recommended to normalize its indicators in pregravid preparation [4,10]. There is more evidence that vitamin D hypovitaminosis leads to an increase in the incidence of cancer and autoimmune diseases, diabetes mellitus, there is a connection with obesity, the development of these complications is most typical for older age. In recent years, it is increasingly recognized that vitamin D has a significant effect on the functioning of the organs of the cardiovascular system. Epidemiological researches have shown that there is an inverse correlation between vitamin levels and diseases such as hypertension, atherosclerosis, coronary heart disease, myocardial infarction, heart failure, stroke and hyperlipidemia [6,11]. Prevention of the above complications reduces the incidence of postmenopausal problems. Vitamin D deficiency is indeed a global problem.

THE MAIN RESULTS AND FINDINGS

Vitamin D in 80-90% enters the body through the skin, a small part of it comes with food. There are varieties of vitamin D - D3 (cholecalciferol) and D2 (ergocalciferol). D3 (cholecalciferol) is called "solar" because it is formed in the skin through exposure to ultraviolet light. To do this, children and adults should be in the sun for about 15-30 minutes several times a week. However, the intense production of vitamin D occurs during unhealthy hours - from 11:00 to 14:00, when the skin needs protection. Ultraviolet radiation from the sun is composed of UVA and UVB rays. Vitamin D synthesis is more actively triggered in the presence of UVB radiation (although it is considered more aggressive, does not allow glass to pass through). D2 (ergocalciferol) enters the body along with the usual food; unfortunately, there are not many animal foods enriched with vitamin D [13].

Despite the fact that the geographical location of Uzbekistan is at 41 latitude, it is below 35 parallel north latitude; in a zone of moderate insolation, a sunny country, where there is plenty of ultraviolet radiation, but vitamin D deficiency is common. What is the reason for this? There are many reasons for the deficit in Uzbekistan. First of all, residents of large cities in the atmosphere have a large layer of smog and dust ("dust cushion") - this prevents UV rays from breaking through. Secondly, insufficient consumption of foods with polyunsaturated fatty acids (fatty fish). Thirdly, insufficient sun exposure, dark skin color, closed clothing are also risk factors for vitamin D deficiency. Also, impaired absorption of vitamin D due to gastrointestinal diseases
(malabsorption syndrome, Crohn's disease, pancreatitis, cholestasis, etc.), long-term use of glucocorticoids, liver and kidney disease, when the processes of vitamin D hydroxylation are disrupted [7,8]. Obesity is also a factor in which vitamin D is deposited in adipose tissue. These risk factors are most often found in the older age group, respectively, the risk group is women over the age of 50. Old age is also a risk factor for hypovitaminosis, this fact is associated with a decrease in the production of vitamin D in the skin, absorption of vitamin D in the intestine, the number of VDR receptors in muscle tissue and functional activity of the kidneys, which in turn aggravates the deficiency state.

Menopause is characterized by ovarian depletion, a decrease in the production of steroid hormones. About 1/3 of a woman's life falls on the period of menopause, on average it is at least 15-20 years. In turn, the menopausal age is characterized by certain problems: from early psychovisceral disorders, urogenital problems and changes in the skin to late cardiovascular disorders, osteoporosis and Alzheimer's disease [11]. For this age, the most common cause of death is coronary heart disease. Thus, the combination of the problems of menopausal age and vitamin D deficiency dictates the need to search for risk factors and timely correction in this age group.

Considering the above, we examined 46 postmenopausal women. The aim of our study was to study risk factors in women with vitamin D deficiency with subsequent adequate correction of hypovitaminosis. The examination of women was carried out in the polyclinic of the City Perinatal Center.

**RESEARCH METHODS**

Assessment of the level of total 25 (OH) D in blood serum. Determination of the quantitative 25-OH vitamin D in blood serum was carried out by chemiluminescence immunoassay in vitro using an automatic chemiluminescence immunological analyzer of the MAGLUMI series. To interpret the results, we used the most frequently recommended standards for the concentration of 25 (OH) vitamin D: an indicator up to 10 ng / ml (25 nmol / l) is interpreted as a pronounced deficiency, 11-20 ng / ml (50 nmol / l) indicates a deficiency, 21-30 ng / ml (51-75 nmol / l) - insufficiency, with values over 30 ng / ml (75 nmol / l), the concentration of vitamin D has an adequate level. For prevention and treatment, it is recommended to adhere to the target level in the range of 30-60 ng / ml (75-150 nmol / L).

**RESULTS OF THE RESEARCH**

The age of the examined women ranged from 49 to 70 years. The median age was 56.1 years. There were 8 elderly women (over 60 years old) in the study group (17.4%), they are a risk group for severe vitamin D deficiency. The menopause age ranged from 1 to 20 years. From the group of women examined, early menopause (40-45 years old) was diagnosed in 9 (19.6%) women, in 1 (2.2%) women at 56 years old, in the remaining 36 (78.3) women - timely.
Determination of the level of total 25 (OH) D in blood serum in 46 postmenopausal women revealed the following values (Table 1). An adequate level of vitamin D of 67.77 ng/ml was registered only in a 53-year-old patient who had been taking drugs in a prophylactic dosage for the last 4 months. The remaining 45 women were diagnosed with decreased vitamin D levels. The mean serum 25 (OH) D levels in the deficient group were 16.39 ± 3.87, and in the deficient group - 23.96 ± 2.4.

We found that 97.8% of the women undergoing examination had reduced levels of vitamin D in the blood serum (Table 2). Of these, vitamin D deficiency was registered in 10.87% of women in the surveyed group. At the same time, a pronounced vitamin deficiency was not registered in any patient from the group, which indicates the intake of vitamin into the body. In 86.96% of women, a deficiency of vitamin D in the blood serum was found, which corresponds to the idea of the frequent occurrence of a reduced vitamin D-status.

### Table 1. Vitamin D-status of the studied women.

| Serum 25 (OH) D level          | Number of people | % of all investigated |
|-------------------------------|------------------|-----------------------|
| Severe vitamin D deficiency   | 0                | 0%                    |
| Vitamin D deficiency          | 40               | 86.96%                |
| Insufficiency of vitamin D    | 5                | 10.87%                |
| Adequate vitamin D levels     | 1                | 2.17%                 |

### Table 2. Vitamin D-status by identified groups
The next stage of the examination was to identify risk factors that lead to vitamin D deficiency. Given possible adjustments of risk factors, recommendations were given for correcting hypovitaminosis in order to further reduce complications in this category of women.

The risk group for vitamin D deficiency includes older age; people with a dark skin tone; other factors in which insufficient exposure to ultraviolet rays occurs. That is, these are factors in which, even with adequate availability or intake of "solar" vitamin, vitamin deficiency develops. It was said above that in the surveyed group, 17.4% of the deficiency state was associated with one of the factors, such as old age. 11 (23.9%) women from the study group had a dark skin tone. The main source of vitamin D is its formation under the influence of UV rays, which is aggravated by insufficient exposure to them associated with closed clothing. This fact was revealed in every second patient, in 24 (52.2%) postmenopausal women.

Overweight and obesity also lead to an increased prevalence of vitamin D deficiency. Menopausal age is characterized by an increase in body weight due to reduced metabolism and insufficient physical activity. Since the termination of ovarian function in women, the number of women with metabolic syndrome has increased three times more. Obesity in menopause is accompanied by a decrease in fat-free mass, including muscle mass. Vitamin D deficiency is observed due to deposition in the fat mass, thus a vicious circle develops, which can only be broken with adequate correction of vitamin D deficiency and weight loss. In the group of surveyed women, body mass index was normal (BMI 19.8-24.9) in 19 women (41.3%), excess body weight (25-29.9) was found in 15 women (32.6%). In 12 (26.1%) women, obesity (30 or more) of varying degrees was registered. Moreover, class 1 obesity (BMI 30-34.9) was diagnosed in 8 (17.4%) women, class 2 obesity (BMI 35-39.9) - in 3 (6.5%) women, class 3 obesity (BMI 40-44.9) - in 1 (2.2%) woman. Noteworthy is the fact that in the group of obese women, vitamin D values ranged from 11.8 to 14.36 ng / ml, that is, all patients had vitamin D deficiency. It is well known that obesity leads to an increased risk of metabolic and cardiovascular diseases, cancer, and this leads to an increased risk of mortality in this group. Researches show that weight loss is problematic when vitamin D is deficient. According to the WHO, vitamin D deficiency increases the risk of cancers such as cancer of the endometrium, breast, ovaries, esophagus, stomach, liver, gallbladder, kidney, colon, Hodgkin's and non-Hodgkin's lymphomas [4,14]. In the surveyed group, 2 women were registered with breast cancer, 1 woman with non-Hodgkin's lymphoma, which amounted to 6.5%. The vitamin D concentration in these patients showed a deficiency of 14.36 to 17.76 ng / ml.

By the time of examination, in the study group of women, the consequences of both climacteric age and hypovitaminosis D were found: arterial hypertension - 14 (in 30.4% of cases), stroke and heart attack - 1 case (respectively, 2.2% in each case), diabetes mellitus -2 (4.3%). Diseases of the thyroid gland (hypothyroidism, nodular goiter) and the respiratory system have also been reported. The clinical parameters of hypovitaminosis were assessed. Clinical symptoms of vitamin D deficiency have no obvious manifestations and
are scarce. Most often, women are concerned about isolated or generalized pain, discomfort in bones and muscles. Vitamin D deficiency leads to myopathy, which can manifest itself as muscle weakness, especially in the proximal muscle groups, diffuse muscle pain, and difficulty walking. In the group of surveyed women, the above complaints of varying severity from mild to severe were observed in 37 patients, which amounted to 80.4%.

When studying the anamnesis in this category of women, it shows a number of problems in both obstetric-gynecological and somatic status (Table 3). Problems in reproductive health were observed in 65.2% of women (uterine myoma, fibroadenoma, fibrocystic breast disease, secondary infertility). A burdened obstetric history was observed in 54.3% of women, in the form of hypertensive conditions, postpartum hemorrhage, placental abruption, antenatal fetal death.

### Table 3. Assessment of patient anamnestic data

| Burdened gynecological anamnesis | % of women | Complicated obstetric anamnesis | % of women |
|----------------------------------|------------|---------------------------------|------------|
| Uterine fibroid                  | 45.6% (21) | Non-developing pregnancy        | 15.2% (7)  |
| Pelvic inflammatory diseases     | 13% (6)    | Premature birth                  | 17.4% (8)  |
| Sterility                        | 10.9% (5)  | Intrauterine fetal death         | 4.3% (2)   |
| Tumor diseases of the ovaries    | 10.8% (5)  | Preeclampsia                     | 47.8% (22) |
| Cervical disease                 | 6.5% (3)   | Cervical insufficiency           | 10.8% (5)  |
| Fibrocystic mastopathy           | 58.7% (27) | Placental abruption              | 19.6% (9)  |
| Fibroadenoma (operated on)       | 4.3% (2)   | Postpartum hemorrhage            | 23.9% (11) |

All patients with vitamin D deficiency were offered drug correction, the recommended drug was cholecalciferol (vitamin D₃). The proposed schemes for correcting vitamin D deficiency were selected individually, depending on the initial level and the achievement of target values. The recommended serum 25 (OH) D level is 40-60 ng / ml (100-150 nmol / L). Table 3 presents an algorithm that can be used to select the optimal daily dose of vitamin D preparations depending on its initial serum level and the desired one [8,10]. Prescribing the drugs was taking into account the contraindications to taking vitamin D preparation. The patient with normal vitamin levels was advised to continue prophylactic doses of the vitamin.
Table 3. Average change in blood concentration based on consumption per day (IU / day).

| Expected level, ng / ml | 20  | 30  | 40  | 50  | 60  |
|-------------------------|-----|-----|-----|-----|-----|
| The recommended daily amount of vitamin D₃, IU |
| Available level, ng/ ml |
| 10                      | 2000 | 4000 | 6000 | 10000 | 10000 |
| 15                      | 1000 | 3000 | 6000 | 9000  | 10000 |
| 20                      | 2000 | 5000 | 8000 | 10000 |
| 25                      | 1000 | 4000 | 7000 | 10000 |
| 30                      | 3000 | 6000 | 10000 |
| 35                      | 1000 | 5000 | 9000 |
| 40                      |     | 5000 | 9000 |

CONCLUSION

1. All postmenopausal women, especially those with risk factors for vitamin D deficiency, are recommended to determine total 25 (OH) D in serum.
2. All postmenopausal women with overweight and obesity are recommended to correct hypovitaminosis and weight loss.
3. All postmenopausal women with insufficient vitamin D content are recommended drug correction to reduce the risk of cardiovascular and oncological diseases.
4. The role of vitamin D in the prevention of complications in postmenopausal age requires further study on the basis of more clinical material.

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