Bony manifestation of rickets in a sunny city - a case report from Yazd, Iran

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Abstract:
Rickets is disease that occurs in growing bones in which defective mineralization occurs in both the bone and the cartilage of the epiphyseal growth plate, resulting in the retardation of growth and skeletal deformities. Rickets is more common in areas with less sunlight. However, this case report presents a case of the bony manifestation of rickets with the intake of vitamin D supplements in Yazd, a city in central Iran that has sunshine almost every day. A patient was referred to an outpatient general pediatric clinic for deformities of the legs and growth disturbance, with his height far below the normal range. The changes that were most evident in his X-rays were the bowing of the long bones of the legs and forearms and the cupping of the wrist metaphyseal region. In summary, we present a patient with bony manifestation of rickets despite living in a sunny area and taking vitamin D supplements. Thus, it is important to remember that rickets is still a common disease among children in Iran. More studies of this issue should be conducted, including the identification of abnormal cases and rescheduling vitamin D supplementation programs.

Keywords: rickets, vitamin D deficiency, dietary supplement

1. Introduction
Rickets is the bony manifestation of altered vitamin D, calcium, and phosphorus metabolism in a child. The causes of rickets are varied and include nutritional deficiencies, especially poor dietary intake of vitamin D and calcium. There is another type of rickets known as non-nutritional rickets (1). The earliest reports describing the syndrome of rickets appeared in the English literature around 1650, suggesting the causes of rickets, osteomalacia, and the various metabolic defects in rickets and osteomalacia (2). By definition, rickets occurs only in children whose growth plates have not closed. Children with rickets are described as apathetic and irritable, often with a short attention span, and they are seemingly indifferent. They tend to sit for long periods of time due to the deformity associated with the bowing of the lower extremities (2, 3). Permanent skeletal deformities may appear in patients who have rickets. The relevant clinical findings are rachitic rosary (enlarged ends of the ribs), scoliosis, skull-craneotabes, bowing of the long bones (caused by loading), thickening of the wrists and ankles, greenstick fractures, genu varum, genu valgum, and laxity of the ligaments (2, 4).

Plain radiographic findings include physeal cupping and widening, fraying of the metaphysis, flattening of the skull and dorsal kyphosis (2, 5, and 6). Vitamin D deficiency is one of most common causes of rickets, and this deficiency
may be due to a lack of sun exposure and inadequate intake of the vitamin in food. In some cultures, whole body clothing prevents adequate sun exposure, e.g., the purdah system in Muslim communities. Socioeconomic status also may predispose people to vitamin D deficiency, and, thus, to having rickets due to many contributing factors, including the frequency and number of labor hours, malnourished infants, the economic situation of the family, living in cities where there is air pollution, living in high altitude areas, and the nutritional status of the mother. Vitamin D deficiency also is common at the end of the winter due to less sun exposure (7). Dietary vitamin D deficiencies can occur in children to different extents depending on ethnicity, skin pigmentation, and variations in the intake of supplements (8).

A Caucasian infant's vitamin D requirements are met by exposure to sunlight for 30 minutes per week, clothed only in a diaper, or for two hours per week when fully clothed with no hat. Asians require approximately threefold longer periods of exposure to sunlight because of the protective pigmentation in their skin, and Africans need six times the exposure of Caucasians (9). Although there are many osseous diseases that can occur in children, the goal of this research was to demonstrate that rickets continues to be a formidable, yet treatable, disease that should be considered in the differential diagnosis of children even though they are taking vitamin D supplements and getting adequate sun exposure.

2. Case Presentation
2.1. Clinical presentation
A young boy, 18 months old, was referred to an outpatient clinic because his parents had noticed that the child, at the age of 10 months, had knock knees and growth abnormalities.

![Figure 1. Genus varum](image)

2.2. Past history
The child was healthy at birth, and his birth weight was 2900 g. His occipitofrontal circumference (OFC) was 47 cm (25th percentile for his age). He was short in comparison to the norm, measuring just 73 cm in height (less than the 5th percentile for his age), and he weighed 12 kg. He was breastfed for the first six months, after which formula was added to his diet. He had received vitamin D supplements since he was born. The parents were of normal stature and build, and there was no consanguinity of rickets in the family history. The patient had no history of or features of malabsorption, chronic renal disease, or hepatic disease.
2.3. Physical examination
Upon examination, the patient had a bilateral genu varum (Figure 1), widened wrists, bowing of the arms (Figure 2), and short stature; however, his face, skull, spinal column, diaphragmatic insertion, dentition, and costochondral junctions were normal. His hearing, vision, mental acumen, and motor development were normal.

2.4. Laboratory findings
The results of biochemical investigations are summarized in (Tables 1, 2, and 3). His alkaline phosphatase, parathyroid hormone (PTH) was elevated. Calcium, phosphorus, serum BUN, vitamin D₃, and urine calcium, phosphorus and creatinine results were normal.
Table 1. Hematologic index

| Lab test | Patient value         | Normal value for age             |
|----------|-----------------------|----------------------------------|
| WBC      | 7×10^6 / lit          | 4.1-10.9×10^6 / lit              |
| RBC      | 4.48 mil/UL           | 3.6-6.1 mil/UL                   |
| Neutrophils | 36%                  | 35-80%                           |
| Lymphocytes | 22%                  | 20-50%                           |
| Hgb      | 11.7 g/dl             | 10.5-14 g/dl                     |
| MCV      | 82 fl                 | 78-101 fl (Female)               |
| Platelets | 305 ×10^3 / µL        | 140-450×10^3/µL                  |

WBC: white blood cell; RBC: red blood cell; Hgb: hemoglobin; MCV: mean corpuscular volume

Table 2. Biochemical study

| Lab Tests | Patient | Normal range |
|-----------|---------|--------------|
| AST       | 37 IU/L | 9-80 IU/L    |
| ALT       | 23 IU/L | 5-45 IU/L    |
| ALKP      | 2260 IU/L | 150-420 IU/L |
| GGT       | 13.9 IU/L | 5-32 IU/L    |
| Ca (Total)| 9.4    | 9-11 mg/dL   |
| P         | 4.9    | 4-6.5 mg/dL  |
| PTH       | 75 pg/mL | 10-65 pg/mL  |
| Vitamin D₃| 37.8 ng/mL | 4.8-52.8 ng/mL |
| Na        | 137 mEq/L | 135-147 mEq/L |
| K         | 4.4 mEq/L | 4.1-5.3 mEq/L |
| INR       | 1.1     |              |

AST: aspartate aminotransferase; ALT: alanine aminotransferase; ALKP: alkaline phosphatase; TG: triglyceride; GGT: gamma-glutamyl transferase; Ca: calcium; P: phosphorus; PTH: parathyroid hormone; Na: sodium; K: potassium INR: international normalized ratio

Table 3. Urine analysis

| Lab test | Patient value | Normal value for age |
|----------|---------------|----------------------|
| Ca       | 2 mg/dl       | Up to 4 mg/dl/24h    |
| Cr       | 77.25 mg/dl   | 20-300 mg/dl         |

2.5. Imaging
X-rays showed widening, cupping, and severe fraying of the metaphysical regions, knock knees with the patient in the standing position, and the cupping of the metaphysis were calcified to a greater extent than the metaphysis (osteopenia). These findings were suggestive of the bony manifestations of rickets.

2.6. Treatment
Treatment consisted of 10,000 IU of vitamin D3 (cholecalciferol) daily and 600 mg of elemental calcium per day in three equal doses.

2.7. Patient follow-up
The treatment resulted in resolution of the biochemical and radiological abnormalities within three months. Skeletal deformities regressed completely after medical therapy.

2.8. Ethical consideration
Informed consent was obtained directly and in writing from the patient’s parents for publication of this manuscript and any accompanying images. Consent for the recommended clinical course of action and research participation was freely given based on full prior knowledge of potential benefits and risks. This personal ‘story’ does not belong to the caregiver. It is the individual subject who possesses the original ‘copyright.’ This premise is reflected throughout our medical care system by carefully established privacy controls that protect against dissemination of
patients’ information to third parties without the consent of the affected individuals (10). The subject of a case report must be aware of both the premise and the undertaking whenever possible and feasible. A corollary to the ‘copyright’ was obtained to preserve confidentiality. Wherever possible, the case report could be reviewed by the patient’s parents to permit editing or removal of any confidential or compromising material, thus overtly respecting the ‘copyright.’ The caregiver and/or author were constantly aware of the need for informed consent and the protection of confidentiality, and this awareness provided an additional level of protection. The Medical Ethics Committee of Shohadaye Kargar Hospital approved the study protocol.

3. Discussion
Tests for rickets should continue to be important elements of the differential diagnosis in children who present for evaluation with a bowed extremity, gait disturbances, diminished height, and failure to thrive. Radiography, radio nucleotide imaging, and measurements of blood urea nitrogen (BUN), creatinine, calcium, phosphorus, alkaline phosphatase, vitamin D₃, PTH, and a variety of urinary measurements, including calcium, are extremely useful aids in establishing the diagnosis of rickets and in elucidating the etiology of the disorder (11, 12).

Children with rickets as a result of vitamin D deficiency typically present with bowing of the lower extremities related to weight bearing when the child begins to walk. They have genu varum, genu valgum, coxavara, and short stature. Our patient presented with similar features. Cupping, flaying, fraying of metaphysis, and generalized osteopenia are characteristic features that are observed in a roentgenographic study. X-rays of the wrists of our patient had all of these features, along with delay in bone age. Vitamin D deficiency can occur even with adequate sun exposure (13). It can occur in people who consume foods that are not fortified with vitamin D or if there is intestinal malabsorption of vitamin D. There are very few foods that naturally contain vitamin D, and most of these are meat or fish, so they may not be acceptable in some cultures. Even though vitamin D is essential for bone health, especially in children, few foods are fortified with vitamin D. The fortification of milk with vitamin D is highly recommended in lactating and pregnant women, and, in developed countries where this process is performed, it has been successful in controlling this condition. Also, such countries have conducted many investigative studies of this issue, which we have not done throughout our country to educate people about preventive public health strategies and to document the use of vitamin D supplementation in conjunction with breast milk. So, routine fortification of milk and other food products with vitamin D should be considered. Also, we should ensure that healthcare professionals are aware of the risk of vitamin D deficiency and its consequences in children.

4. Conclusions
We presented a patient with bony manifestation of rickets despite vitamin D supplementation and living in a sunny area. We must remember that rickets is still a common disease in children in our country. Even though vitamin D supplementation has been recommended for all infants in Iran, especially those who are breastfed, this is not done in all parts of the country. Given that there are special genetic predispositions to have rickets, we recommend more studies of patients who have overt symptoms of vitamin D deficiency. We also recommend a new policy that includes improved fortification of foods with vitamin D and requires screenings of children in order to prevent the adverse consequences of low vitamin D, such bone deformities.

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Conflict of Interest:
There is no conflict of interest to be declared.

Authors’ contributions:
Both of authors contributed to this project and article equally. Both authors read and approved the final manuscript.

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