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RESEARCH ARTICLE

THE FREQUENCY OF VITAMIN D DEFICIENCY AMONG IRANIAN CHILDREN

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ABSTRACT

Background

Iran is a tropical country and yet there is widespread vitamin D deficiency among our infants and children. Our study was conducted to assess the Vitamin D status in children in and around Iran.

Methods

This cross sectional study was conducted in 196 children from the newborn period to 18 years of age. Under strict aseptic precautions 2 ml of blood was taken from each child and serum levels of 25(OH)D levels were measured by the direct ELISA method.

Results

Median age was 59 months (range 2–161); 3.1% were vitamin D deficient, 19.4% insufficient. There was no significant difference in mean 25 (OH)D level between Indigenous (93.2, standard deviation (SD) 21.9, n = 82) and non-Indigenous (97.3, SD 27.9, n = 112) children (P = 0.32). Median number of hospitalizations/year were similar (P = 0.319) between vitamin D sufficient (0.34, range 0–12, n = 152) and insufficient (0.22, 0–6, n = 44) children. There was no significant difference between the number of infective admissions per year between vitamin D sufficient/insufficient groups (P = 0.119).

Conclusion

Suboptimal vitamin D status is common among otherwise healthy young children. Predictors of vitamin D status vary in infants vs toddlers, information that is important to consider in the care of these young patients. More information is needed about the optimum level of vitamin D for non-bone-related health in children.

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INTRODUCTION

It is now generally accepted that vitamin D deficiency is a worldwide health problem (Hosseini-Nezhad et al., 2013). It has only recently been recognized that vitamin D plays an important role in up regulating immunity (Kamen and Tangpricha, 2010). Spanning many continents and including all ages, genders and racial/ethnic groups (Lappe, 2011) and are highly prevalent among children worldwide (Thornton et al., 2013).

It has been reported to have spread in many Asian countries (Fraser, 2004) and even, to have increased in its prevalence in North America, Europe, Australia, and New Zealand (Au et al., 2012). Several studies have shown a high prevalence of vitamin D deficiency in Middle East countries (Fields et al., 2011); the figures for suboptimal levels from several countries in this region were reported between 20% and 80% (El-Hajj Fuleihan, 2009). In order to obtain a better estimation of the vitamin D status in Iran, studies encompassing different populations in various age and sex groups are required. Few studies have been performed on vitamin D status in this country, focused mainly on pregnant or elderly women (Maghbooli et al., 2007). The prevalence of vitamin D deficiency is reported to be higher in

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school age children. In Iran, different studies have shown a high prevalence of vitamin D deficiency, which is present in 86% of school-age children in Tehran (Neyestani *et al.*, 2012) and in 46.2% between 14 - 18 years old children in Isfahan (Moussavi *et al.*, 2005). Since it is possible to easily prevent complications arising from vitamin D deficiency by nutritionally enriched vitamin D supplements. The present study was carried out to assess the level of vitamin D in the newborn period to 18 years of age in Iran in 2015.

METHODS

Detection of 25 OH vitamin D by ELISA

Vitamin D level was determined in serum by Enzyme Linked Immune Sorbent Technique (ELISA) using OH vitamin D ELISA kit; Catalog No.EIA-4696 (DRG International, Inc.), CA, U.S.A. the status of 25OH vitamin D was evaluated as follows: According to Endocrine Society Clinical Practice Guidelines, vitamin D deficiency was defined as a 25 (OH)D < 20ng/L (50nmol/L), and insufficiency as a 25 (OH)D between 21 and 29ng/L (52.5 and 72.5nmol/L), and 25(OH) D level ≥ 30 ng/L (75nmol/L) as the optimal level (Holick *et al.*, 2012).

Definition of vitamin D deficiency

Mild, moderate and severe vitamin D deficiencies were defined as 25-OHD values of 20-30 ng/ml, 10-20 ng/ml, and <10 ng/ml respectively. There is also another classification for vitamin D deficiency in the literature. In this classification, the combination of moderate and severe vitamin D deficiencies are considered vitamin D deficiency (25-OHD <20 ng/mL) and mild vitamin D deficiency (25-OHD 20-30 ng/mL) as vitamin D insufficiency (Holick *et al.*, 2009). We used the first classification. However, the second classification was used after stating its usage.

Statistical analyses

The results were analysed using Microsoft Excel (Microsoft Corp., Redmond, WA, USA) and SPSS 18.0 (PASW Statistics 18; SPSS Inc., Chicago, IL, USA). Data conforming to a normal distribution were assessed for significance using a t-test; other data were analysed using a Kruskal-Wallis test, two-tailed P-value of <0.05 were considered significant.

RESULTS

Over the 24-month period, 200 children were recruited, and sufficient blood for vitamin D analysis was obtained in 196 children. No children were excluded as none had previously been diagnosed with vitamin D deficiency or were on vitamin D supplements. The median age of the cohort was 56 months (range 2-161). Other demographics of the children are outlined in Table 1. Of the 196 children in whom sufficient blood was collected for analysis, three children (3.1%) were found to be vitamin D deficient. Of these, two were mildly deficient (25-50 nmol/L), and one severely deficient (<12.5 nmol/L). Two of the three children identified as being deficient had risk factors for vitamin D deficiency; the severely vitamin D deficient child was a 10-year-old boy admitted with nephrotic syndrome. Of the mildness vitamin D deficient children, one was an Indigenous child hospitalised with sepsis requiring an intensive care unit admission, and the second child had no known risk factors for vitamin D deficiency and had a normal level on repeat testing. Nineteen children (19.4%) were vitamin D 'insufficient' (50-

75 nmol/L), and the rest ($n = 152$, 77.6%) were deemed 'sufficient'. Of the 38 children who were vitamin D insufficiency, 48% were Indigenous and 52% non-Indigenous.

Table 1. Socio-demographics of the children ($n = 200$)

Ethnicity	Non-Indigenous	57%
	Indigenous	43%
Sex	Male	65%
	Female	35%
Age group	Infant	8%
	1-5 years	43%
	5-18 years	48%
Gestation	Term	76%
	Preterm	7%
	Unknown	17%
Place of recruitment	Wards/ED	38%
	Daycase procedure unit	33%
	HRCT/bronchoscopy list	16%
	Healthy blood drive	13%
Mean weight z-scores ($n = 132$)	Indigenous	-0.6
	Non-Indigenous	0.2
	Overall	-0.3

There was no significant difference ($P = 0.33$) in the mean 25 (OH)D (nmol/L) among Indigenous children ($n = 84$, Mean 92.2) and non-Indigenous children ($n = 128$, mean 97.3).

DISCUSSION

This study shows vitamin D level of serum in children, which is low and particularly critical in 0 - 18 years old girls. Where the prevalence of vitamin D deficiency was found to be 91.7% in 9 - 2 years old students in Tehran. In another study in 7 - 18 years old students in Tehran, 52% were vitamin D deficient, of whom 26% suffered vitamin D insufficiency (Moussavi *et al.*, 2005). Also, vitamin D deficiency was 65.2% between 14 - 18 years old students in a similar study from Isfahan (Rabbani *et al.*, 2009). Bener A *et al* revealed that the prevalence of vitamin D deficiency in Qatari children <5 years of age was 9.5%, 5-10 years was 28.9%, 11-16 years was 61.6% (Maghbooli *et al.*, 2007). Large prevalence study of children aged 1-11 years was carried out in the USA, showing that <1% of children had levels <25 nmol/L, 18% <50 nmol/L and 95% had levels <75 nmol/L. A similar study in the UK found that 35.1% of children aged 4-17 years had 25(OH)D levels <50 nmol/L. As the main source of vitamin D is sunlight, the most likely major reason for the lower prevalence of vitD deficiency /insufficiency in Darwin is its higher average number of hours per year of sunlight; in Darwin it is 8.4 h per Day (approximately 3102 h per year) (AGBM, 2011). We found no significant difference between the numbers of acute or infective hospitalisations per year depending on vitamin D status.

There have been multiple studies documenting possible associations between clinical rickets and a predisposition to infection (Yamshchikov *et al.*, 2009). This study was performed during cold seasons in which vitamin deficiency is highly critical. An inverse correlation was found between the levels of 25 (OH) and seasons of respiratory tract infection (Li *et al.*, 2015). There are some reports of a higher occurrence of type one diabetes mellitus T1DM and multiple sclerosis MS during the cold seasons, which was related to the lower vitamin D status (Amital *et al.*, 2010). Vitamin D deficiency is reported to be associated with increased incidence of ear and lung infections in children. Severity of various diseases currently

has a high profile in the literature, leading to increased vitamin D testing (and thus increased cost) worldwide. In Australia, the cost of vitamin D testing has increased by nearly a hundredfold since the year 2000 (Bilinski *et al.*, 2012). Further, others have expressed concern over the mismatch between high quality evidence and causation. (Harvey, 2012) Our pilot study contributes to this debate as our data do not support the routine screening of children in the NT as it is very unlikely that vitamin D deficiency contributes to their high burden of infectious disease in our setting. Our study does not support aiming for a vitamin D level of >75 nmol/L in children, we recommend continuing to follow the APEG guideline, aiming for vitamin D levels >50 nmol/L (Munns *et al.*, 2006).

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