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# Hypovitaminosis D in 6 - 59-Month Children in the District of Parakou (Benin) in 2017

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#### **Abstract**

**Introduction:** Nutritional rickets is the visible tip of the iceberg formed by the diseases caused by hypovitaminosis. Vitamin D plasma is essentially influenced by the level of sunlight which is significant in Sub-Saharan Africa and in Benin. However, there are factors likely to impede endogenous production of that vitamin in children in our environment. This research work was conducted with the view to investigate hypovitaminosis D prevalence and associated factors in children aged 6 to 59 months in the District of Parakou in 2017. Methods: This research work was a cross-sectional study with prospective collection of data carried out in community in the District of Parakou in 2017. The study involved euthrophic and non-HIV infected children aged 6 to 59 months who did not benefit from vitamin D supplementation during the past three months. Vitamin D plasma concentration was obtained using "Radioimmunoassay" after cluster random sampling in accordance with WHO guidelines. According to Schwartz formula, nominal minimum sample size was 321. In this study, hypovitaminosis D was defined as a plasma level lower than 30 ng/l. Data processing was carried out using softwares such as EPI INFO version 7.2 and SPSS 21. Results: In total, 400 children were involved in the survey. Hypovitaminosis prevalence was 9.5% (38 out of 400 cases). In our research work, the factors associated with hypovitaminosis were child's age (p = 0.0470), mother's age (p = 0.0000), weaning age lower than 6 months (p = 0.0466) and mother's occupation out of direct sunlight (p = 0.0012). Conclusion: Nearly one out of ten children suffers from hypovitaminosis D in the District of Parakou in 2017. The associated factors were age of child and mother, early weaning and mother's professional occupation held out of direct sunlight. This research work should be continued and improved by a nationwide multicentre study. This could lead to the adoption of measures for

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hypovitaminosis D prevention among children in Benin.

# **Keywords**

Hypovitaminosis D, Associated Factors, Children, Benin

# 1. Introduction

Nutritional rickets is the visible tip of the iceberg formed by diseases caused by hypovitaminosis [1]. In addition to its function in bone health and immunomodulatory effects on the immune system, vitamin D also contributes to protect against cancers as well as auto-immune, cardiovascular, metabolic and infectious diseases [1] [2]. Vitamin D plasma concentration is influenced by level of sunlight which is significant in Africa. This may justify the little interest granted to that micronutrient by the African scientists. In our environment, there are factors that impede the endogenous production of vitamin D in children. These are skin color, level of sun exposure, dress style and nutrition. In a study conducted in a temperate zone (New Zeland) among children in 2005, Rockell et al. had reported a prevalence of 35% [3]. Hypovitaminosis D is common among children both in temperate and tropical regions. For instance, in Nigeria in 2000, Tom et al. had reported a prevalence of 37% [4]. Besides, the relation between hypovitaminosis D and infant and child mortality in Africa should be considered. In Benin, a study conducted in Parakou on the plasma vitamin D profile in children hospitalized for severe malaria showed a frequency of 83.8% hypovitaminosis [5]. This raises the question of the interaction between this micronutrient and malaria on the one hand, and then others infectious diseases on the other hand in a country without any policy of Vitamin D supplementation under the pretext of a high level of sunshine. The purpose of this research work was to investigate hypovitaminosis D prevalence and its associated factors among children aged 6 to 59 months in the District of Parakou in 2017.

#### 1.1. Methods

This research work was a cross-sectional study with prospective data collection performed within the community in the District of Parakou (Northern region of Benin). The study was carried out during the period running from May to July 2017. The study population consisted of all children aged 6 to 59 months and living in the District of Parakou for at least 6 months. On the other hand, the study excluded children suffering from acute malnutrition (Mild upper arm circumference (MUAC) < 125 mm or weight for height score lower than –2 standard deviation (SD) or nutritional edema). In addition, the study did not involve children with vitamin D supplementation underway or in the three past months. Eventually, this study did not include children tested as HIV positive. Moreover, those whose parents did not give their consent to participate to the study were

also excluded. Sampling was probabilistic; it was carried out using a two-stage cluster sampling in accordance with WHO guidelines. Cluster unit consisted of one group of children aged 6 to 59 months living in an area of the District of Parakou. Statistical unit was represented by one child aged 6 to 59 months who meets the inclusion criteria. Areas were randomly selected; and cluster interval, number of clusters as well as number of children by cluster were calculated. Sample minimum size expected in the study was 321 (according to Schwartz formula), with an 83.3% prevalence of a study conducted in Tanzania in 2000 [6]. For each child involved, a "face-to-face" interview (interviewer/respondent) was performed to collect data among mothers or child minders using a survey form. The main variable of this study was hypovitaminosis D. Vitamin D level was defined as normal when it oscillated between 30 and 150 ng/ml. A child was considered with hypovitaminosis when vitamin D concentration was lower than 30 ng/ml. Vitamin D deficiency was retained when concentration was lower than 20.8 ng/ml; deficiency was classified in two categories: moderate (between 12 and 20.8 ng/ml) and severe (lower than 12 ng/ml). A child was with hypervitaminosis D when vitamin D level was higher than 150 ng/ml [7]. Vitamin D plasma concentration was obtained through "radioimmunoassay" on plasma after blood collection among selected children. The blood thus taken into a dry tube was centrifuged to collect plasma which was stored at -18°C during 1 to 2 months before being handled. Actual vitamin D screening test by means of radioimmunoassay was implemented in four phases which are extraction with acetonitrile, incubation, counting and finally reading of test results. The independent variables were economic variables, gynecological and obstetrical histories among mothers as well as sociodemographic variables, child care data, history of diseases and clinical data among the children. Entry and processing of data were carried out using softwares such as EPI INFO version 7.2 and SPSS 21. Measures of central tendencies and dispersion (Mode, Average, Standard deviation) served to describe quantitative variables. The ratios and their confidence interval were used for the description of qualitative variables. Ratios were compared using Karl Pearson's Chi-square test with a significance threshold estimated at 0.05.

# 1.2. Ethical and Professional Considerations

The city council of Parakou, the heads of boroughs and chiefs of areas granted written authorizations for the conduct of this research work. The ethics committee of the Faculty of Medicine of the University of Parakou has approved it. Mothers and minders of the involved children had given their oral free and informed consent before any participation. The biological tests done within the framework of the study were fully free of charge for the participants. The refusal of a child mother or minder to participate to the study had no discriminatory effect. Children with hypovitaminosis D were treated free with an Ergocalciferol alcohol solution at a loading dose of 200,000 IU, followed by a maintenance dose of 1600 IU per day. Cases of hypervitaminosis have not been treated.

#### 2. Results

A total of 502 children were shortlisted and four hundred (400) were finally involved in this study. One hundred and two (102) shortlisted children were not included in the study for many reasons: forty (40) were malnourished, twenty-one (21) were on poly-vitamin supplementation and forty-one (41) refused to participate to the study. None of the children was tested as HIV/AIDS infected. **Figure 1** outlines the flow chart showing the source population and the study population.

# 3. Population Description

#### 3.1. Data Related to the Children

Among the 400 children investigated, 192 were female (48.0%) and 208 were male (52.00%) *i.e.* a sex ratio of 1.08. Mean age was  $36.22 \pm 15.72$  months. The BCG vaccination coverage was 92.75% (29/400). Exclusively breastfed children were 283 *i.e.* 70.75%. In this research work, weaning started before the age of six months in 203 children (50.70%). Children weighed on average  $12.8 \pm 3.09$  kg with extremes ranging from 7 to 23 kg. The average size of the children investigated was  $87.89 \pm 12.51$  cm with extremes oscillating between 55 and 115 cm. In this study, weight for height (WFH) oscillated between -2 and 0 Z-score in

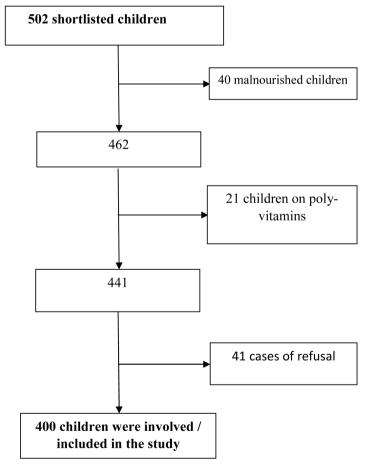


Figure 1. Flow chart showing the study population in Parakou in 2017.

47.50% of cases. **Table 1** shows the distribution of children according to type of breastfeeding, weaning age and weight for height score.

# 3.2. Data Related to Mothers and Child Minders

Mothers' mean age was  $28.20 \pm 6.13$  years with extremes ranging from 17 to 50 years. The predominant age group was the one varying between 20 and 29 years (63.75%). Among them, 200 were uneducated (50.00%). Two hundred and fifteen mothers worked in the sun (56.25%) and 225 had a low socioeconomic status (56.25%). **Table 2** shows the distribution of mothers of involved children according to educational attainment, work place and socioeconomic status.

**Table 1.** Distribution of children involved in the study according to type of breastfeeding, weaning age and weight for height score in the District of Parakou in 2017.

	Population size	Ratios	
Type of breastfeeding			
Exclusive breastfeeding	283	70.75	
Milk substitute only	11	2.75	
Mixed	105	26.50	
Weaning age (months)			
<6	203	50.75	
6 - 11	165	41.25	
12 - 17	20	5.00	
18	12	3.00	
Weight for height (z-score)			
[-2 to 0]	190	47.50	
[0 to 2]	210	52.50	

**Table 2.** Distribution of mothers of involved children according to educational attainment, workplace and socioeconomic status in the District of Parakou in 2017.

	Population size	Ratios
Educational attainment		
Uneducated	200	50.00
Literate	4	1.00
Primary school	80	20.00
Secondary school	106	26.50
Higher education	10	2.50
Workplace		
In the sunlight	215	53.75
Out of the sunlight	185	46.25
Socioeconomic status		
Low	225	56.25
Average	173	43.25
High	2	0.50

#### 3.3. Vitamin D Plasma Level

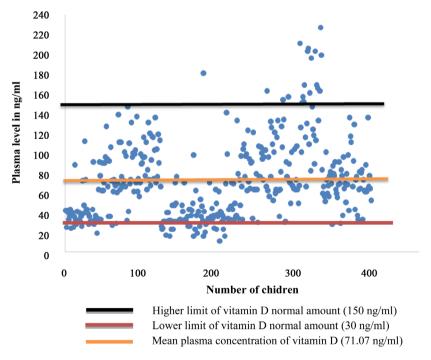
Among the 400 children involved in the study, 38 children suffer from hypovitaminosis D. Average plasma vitamin D concentration in investigated children was  $71.07 \pm 39.43$  ng/ml with extremes ranging from 11 to 227 ng/ml. The average concentration of children with hypovitaminosis D was  $24.05 \pm 4.86$  ng/ml with extremes ranging from 11 to 29.90 ng/ml. Two hundred and thirty-four children (58.50%) had a vitamin D plasma concentration below the average (71.07 ng/ml). **Figure 2** shows vitamin D plasma level in investigated children.

# 3.4. Hypovitaminosis D Prevalence in Children Aged 6 to 59 Months

Thirty-eight out of the four hundred investigated children had hypovitaminosis D *i.e.* a prevalence of 9.50%. Among those 38 cases of hypovitaminosis D, there were 29 cases of vitamin D deficiency (7.25%), eight children (2%) had moderate vitamin D deficiency and one child (0.25%) had severe deficiency. Besides, considering the threshold that defines vitamin D function outside the bones (50 ng/ml or higher), the ratio of children unlikely to have the benefits of that micronutrient was 37.25%. **Figure 3** shows distribution of involved children according to their vitamin D status.

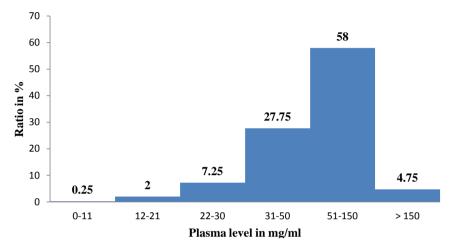
### 3.5. Factors Associated with Hypovitaminosis D

In this research work, child age and weaning age were statistically associated with hypovitaminosis D, with respectively p = 0.047 and p = 0.000. Mothers' age



**Figure 2.** Vitamin D plasma concentration among 6 - 59-month children in the District of Parakou in 2017.

and workplace were also associated with hypovitaminosis D, with p=0.000 and p=0.001. On the contrary, the socioeconomic status and the weight-for-height score were not associated with hypovitaminosis. **Table 3** indicates the relationship between hypovitaminosis D and sociodemographic data, data related to weaning age and nutritional status.



**Figure 3.** Distribution of involved children according to their vitamin D status in 2017.

**Table 3.** Relationship between hypovitaminosis D and sociodemographic data, data related to weaning age and nutritional status in 6 - 59-month children in the District of Parakou in 2017.

	Hypovitaminosis D					
	Total	n	%	RP	95% CI	P
Children' age (months)						0.047
6 - 24	119	6	5.04	1		
≥25	281	32	11.38	2.41	[1.02 - 6.50]	
Weaning age (months)						0.0000
6 - 11	368	29	7.88	1		
12 - 18	32	9	25.71	4.5	[1.9 - 9.7]	
Mothers' age (years)						0.0000
17 - 29	262	15	5.72	1		
≥30	138	23	16.66	0.30	[0.1 - 0.6]	
Educational attainment						0.066
Uneducated	204	14	6.86	1		
Educated	196	24	12.24	0.52	[0.2 - 1.1]	
Workplace						0.001
Exposed to sunshine	215	11	5.12	1		
Out of the sunlight	185	27	14.59	3.16	1.5 - 6.6	
Weight-for-height score						0.090
-2 to 0	190	23	12.11	1.7	[0.9 - 3.5]	
0 to 2	210	15	7.14	1		

#### 4. Discussion

The fact of not taking into account some diseases such as tuberculosis and malaria which are infections often associated with hypovitaminosis D, may be a limitation of this study. However, its prospective character is a factor likely to limit missing data during collection. Blood collection and plasma sample conservation were performed under the conditions required for such a survey. Conservation time was short; this may help avoid a possible degradation of vitamin D. Vitamin D screening test was performed using radioimmunoassay. This technique has the advantage of being sensitive, specific, quick and quantitative. Its performance on children randomly selected within the community may help extrapolate the results on all the children aged 6 to 59 months in the District of Parakou.

# 4.1. Prevalence of Hypovitaminosis

Thirty-eight out of the 400 children involved in the study were suffering from hypovitaminosis D. Mean plasma vitamin D concentration in deficient children was 24.05 ± 4.86 ng/ml versus mean plasma concentration of all investigated children estimated at  $71.07 \pm 39.43$  ng/ml. The mean vitamin D plasma concentration in deficient children found out in this study is similar to the one reported by Cusick et al. (25.3 ± 7.10 ng/ml) in Uganda in 2014 among children aged 18 months to 12 years [8]. In Tanzania, Rwebembera et al. have reported in their study on 6-month children in 2013 a mean plasma concentration of 23.70  $\pm$  8.10 ng/ml similar to the one identified in this research work [6]. It is lower than the one reported by Nabeta et al. in 2015 (32.20 ± 10.90 ng/ml) in a study carried out in Uganda among 6 - 24-month children [9]. The prevalence of hypovitaminosis D in 2017 among 6 - 59-month children in the District of Parakou amounted to 9.50%. A similar prevalence has been reported by Bener et al. in a study conducted in 2009 in Qatar among under 5-year children seen in community [10]. This prevalence is lower than the one reported in 5 - 35-month children (37%) suspected of rickets by Thacher et al. in 2000 in Nigeria and by Rwebembera in 2013 in Tanzania among 6-month children exposed to HIV [6] [11]. Normally, given the high level of sunlight in the District of Parakou, hypovitaminosis D ratio among children should be very low and even null. This points out that there are factors which predispose that group of child population to hypovitaminosis D. These predisposing factors may be their dress style, history of early births, long hospitalizations, eating habits, duration of sun exposure and particularly skin pigmentation. As a matter of fact, in the study of Guardia et al. conducted in 2008 in the United States of America, the ratio of people presenting with vitamin D deficiency was significantly higher among children with black skin than children with fair skin [12]. Although it is a protective factor against sunburn, skin pigmentation is a major cause of vitamin D deficiency in children.

# 4.2. Factors Associated with Hypovitaminosis D

The study focused on 6 - 59-month children. Children's mean age was 36.22  $\pm$ 

15.72 months. The age group predominantly vitamin D deficient was the one of children whose age was higher or equal to 25 months. As a result, the older children's age is the higher risk of hypovitaminosis D is; this may be due to susceptibility to infectious diseases and hygiene conditions which become poorer. In addition, as from that age group skin becomes darker and may thus represent a protective barrier against UV radiation necessary for endogenous synthesis of vitamin D. The difference observed was statistically significant (p = 0.047). The same remark has been made by Fasih in a study on the assessment of hypovitaminosis D frequency among 1 - 5-year children [13]. On the contrary, in a cross-sectional study conducted in Argentina on 290 children, Hirschler et al. have highlighted lack of correlation between vitamin D serum level and child age (p = 0.14) [14]. If children aged 25 months and above are more exposed to hypovitaminosis D contrary to the WHO data which suggest that infants are particularly exposed to that phenomenon, therefore vitamin D endogenous production is difficult among them due to skin color, dress style and other situations which are likely to impede UV wavelength through the skin [15].

In this research work, mother's occupation out of sunlight was associated with hypovitaminosis D (p = 0.001). The same remark has been made by other authors who pointed out a significant relationship between the duration of sun exposure and hypovitaminosis D. For instance, Bener *et al.* have found out in 2009 in Qatar among under 5-year children a significant correlation (p < 0.05) between hypovitaminosis D and duration of sun exposure [10]. On the contrary, in the same country in 2016, Almehmadi *et al.* identified a lack of correlation between duration of sun exposure and hypovitaminosis D in a study on children, with an average duration estimated at 4 hours (p = 0.48) [16]. The significant relationship between mothers' professional occupation out of sunlight and hypovitaminosis D identified in this study may be due to the cultural behavior based on the fact that mothers carry their children on their backs while working.

The old age of mother or child minder higher or equal to 30 years was a significant determinant of hypovitaminosis D in this research work (p = 0.002). This interaction involves many questions, especially whether old mothers provide more protection to their children against UV rays or practice a diet low in vitamin D.

The results of this study indicate that early weaned children were more prone to have hypovitaminosis D, with a significant difference (p = 0.046). This may mislead into assuming that breast milk is a protective factor against hypovitaminosis D. This is not the case since breast milk is vitamin D deficient. Better still, risk for hypovitaminosis is four times higher in children weaned at 24 months of age than those weaned at 12 months of age. In addition, no child on milk substitute fortified with vitamin D had developed hypovitaminosis D. Without wishing to make the promotion of milk substitutes regarding their vitamin D content, it may be suggested, subject to further studies, to consider implementing vitamin D supplementation among children on exclusive breast-

feeding. This finding of our study is similar to the one reported by Atiq *et al.* in Pakistan in 1998 among 6-year children weaned at the age of 4 months, with significant statistical difference (p = 0.01) [17]. This relataionship existing between early weaning age and hypovitaminosis D may be due to the fact that weaning food were vitamin D deficient.

# 5. Conclusion

Nearly one out of ten children suffers from hypovitaminosis D in the District of Parakou in 2017. The associated factors were age of child and mother, early weaning and mother's professional occupation held out of sunlight. These results indicate that despite the high level of sunshine in the Northern regions of Benin, hypovitaminosis D among children is a critical issue of great concern. Therefore, other nationwide surveys should be conducted in order to capitalize on and show the benefits of vitamin D in the national health policy. Moreover, those further nationwide studies will help adopt new preventive measures against hypovitaminosis D among children in Benin.

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- The corresponding author.

#### **Conflict of Interest**

The authors have not disclosed any conflict of interest concerning this research work implementation and the process of publication of this paper.

# **Authors' Contribution**

- All the authors have proofread this paper;
- Alphonse Noudamadjo has initiated this research work and was responsible for the protocol drafting and paper writing;
- Bakiou Babatoundé participated to the protocol drafting, data collection and report writing;
- Joseph Agossou was involved in the protocol drafting and paper proofreading;
- Julien Didier Adédémy participated to the protocol drafting and paper proofreading;
- Falilatou Mohamed Agbeille participated to the protocol drafting and paper proofreading;
- Gérard Kpanidja participated to the protocol drafting and paper proofreading;
- Abogbo Gibril participated to vitamin D screening test;
- Marcellin Kuassi Amoussou-Guenou participated to vitamin D screening test.

# **Authors' Agreement**

All the authors have agreed to submit this paper for publication.

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