# **Role of Vitamin D in Near Sightedness**

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#### ABSTRACT

**Purpose:** To determine the levels of Vitamin D in children with myopia and to compare them with age matched controls.

Study Design: Case control study.

**Place and Duration of Study:** The study was conducted in Naseer Memorial Hospital, Dadhyal Azad Kashmir from March 2016 to March 2017.

**Material and Methods:** Two hundred patients were selected using convenient sampling technique and were divided two groups (group I Myopic and group II control). Myopia was labeled if after subjective refraction a Spherical Equivalent (SE) of -0.50 diopters (D) or more was found. Vitamin D levels were measured by radioimmunoassay technique with Diasorin SR® kit following the user's manual. Vitamin D levels less than 20 ng/ml were considered Vitamin D deficient following the standards of American academy of pediatrics. The collected data was entered in the statistical package for social sciences (SPSS) version 21 for analysis. Independent t–test was used to determine the significant difference of means between controls and patients. P-values less than 0.05 were considered significant.

**Results:** Mean age of controls and myopes were  $10.65 \pm 3.9$  and  $10.20 \pm 2.5$  years respectively. Vitamin D levels in myopic children were found to be  $14.95 \pm 3.75$  ng/ml and there was no significant difference in mean values of Vitamin D levels in myopic and control group.

**Conclusion:** We found no difference in Vitamin D levels of myopic and non myopic children and concluded that Vitamin D has no role in development or progression of myopia.

Key Words: Vitamin D, Myopia, Refractive error, Objective refraction.

How to Cite this Article: Iqbal Y, Malik A, Shabbier R, Zaman A, Zia S, Talib M. Role of Vitamin D in Near Sightedness, Pak J Ophthalmol. 2020; **36 (2):** 137-141.

Doi: 10.36351/pjo.v36i2.990

#### **INTRODUCTION**

Types of refractive errors are myopia, hyperopia and astigmatism<sup>1</sup>. In Myopia, the person is able to see near objects and the distant ones are blurry hence the term near sightedness. It can occur at any age like in childhood or adolescence and even in old age. There is no gender predisposition and is affected globally<sup>2</sup>. It

Correspondence to: Yasir Iqbal Associate Professor of Ophthalmology, Watim Medical and Dental College, Rawat, Rawalpindi Email: yazeriqbal@gmail.com can be classified as physiologic and pathologic. The cause of physiological myopia can be high curvature of the cornea, nuclear sclerosis and elongated eyeball or combination of these factors with the absence of any other ocular pathology<sup>3</sup>. The cause of pathologic myopia is abnormal lengthening of the eyeball leading to thinning of the sclera wall and other complications. Another classification is based on age of onset. It tends to run in the families<sup>2</sup> and no occupational association has been found.

Myopia is prevalent in 20% of the population worldwide and is estimated that 12% children are

affected<sup>2</sup>. In Pakistan its prevalence is 6% in adults<sup>4</sup> and in children it is reported as high as 21%<sup>5</sup>. The figure is on the rise every year and is presenting as major global health problem<sup>2</sup>. Not only it has a social impact on the individual, making the person unable to perform the tasks of desire, it acts as a factor in adding individuals with less productivity<sup>6</sup>. Furthermore, myopia is a risk factor for other visually blinding diseases like glaucoma<sup>3</sup> and retinal detachment<sup>3</sup> which further implicates on the health system. For these reasons the pathophysiology of development of myopia remained a hot debate since ages.

The pathophysiology of myopia had been associated with many myths. Reading books in dim light or while lying on bed, watching too much television and even deficiency of Vitamin A had been proclaimed but no scientific reason was found. Some researchers advocate genetic predisposition<sup>7</sup> and have identified numerous genetic loci linked with myopia whereas others have attributed to less outdoor activity and Vitamin D deficiency<sup>8</sup>.

Vitamin D, once considered a Vitamin, is now being treated as a hormone. Many studies have demonstrated Vitamin D having effects on biological processes like calcium and phosphorus metabolism regulation as well as cell proliferation and immune differentiation, regulation and neurogenesis<sup>9,10</sup>. It is found to be associated with cardiovascular diseases, cancers, autoimmune and infectious diseases. About 90% of the Vitamin D is derived from the skin and around 10% from diet. There are two forms of Vitamin D, Vitamin D3 (cholecalciferol) and Vitamin D2 (ergocalciferol). Vitamin D3 is derived from the skin after exposure to ultraviolet light. After absorption from intestines and the synthesis by skin, Vitamin D is converted into 25 (OH) D in the liver. If the 25 (OH) D levels are greater than or equal to 30 ng/ml it is considered as normal and less than this is considered as Vitamin D deficiency<sup>11</sup>.

Recently time spent outdoors has become area of interest in myopia research. Researchers have found low incidence of myopia in those who spent more time outdoors as compared to the ones who are more involved in indoor activities, which led to the concept that Vitamin D might be a possible moderator of this association<sup>8</sup>. Researchers have reported high incidence of myopia in children aged 5 to 15 years<sup>12</sup>. Therefore, in this study we compared Vitamin D levels of myopic and non-myopic children in order to determine

whether Vitamin D has a role in myopia pathophysiology or not.

## MATERIAL AND METHODS

It was an observational case-control study conducted at Naseer Memorial Hospital, Dadyal Azad Kashmir, from March 2017 to March 2019. The study was conducted according to the guidelines of Declaration of Helsinki. A formal verbal consent from the children and their parents was taken before the commencement of the study. After fulfilling the inclusion and exclusion criteria, patients were selected using convenient non-probability sampling technique and were divided two groups (group I Myopic and group II control). Selection criteria for group I was; children of any gender with age between 5 to 15 years and diagnosed with physiological myopia. Group II were age-matched controls. Subjects with history of ocular surgery, eye diseases like glaucoma, uveitis, retinal disease, cataract, systemic disease, any therapeutic regimen or steroid use were excluded from the study. Demographic information i.e., age, gender and history of systemic disorders were recorded. Children underwent complete ocular examination including visual acuity, detailed slit lamp examination, intraocular pressure measurement and ophthalmoscopy before cycloplegic retinoscopy was done. Myopia was labeled if after subjective refraction a Spherical Equivalent (SE) of -0.50 diopters (D) or more was found (Mild myopia if SE less than -3.0 D; moderate myopia if greater than or equal to -3.0 D; and high myopia was defined as more than or equal to -6.0 D). Children with myopia were included in group I whereas those having astigmatism and hyperopia was excluded. Children with no refractive error were included in the control group.

Informed consent was taken from the parents of all participants. For assessment of Vitamin D levels, 2cc of venous blood was collected and after centrifugation serum was stored at  $-20^{\circ}$ C temperature in laboratory freezer for further analysis. Vitamin D levels were measured by radioimmunoassay technique with Diasorin SR® kit following the user's manual. Vitamin D levels less than 20 ng/ml were considered Vitamin D deficiency following the standards of American academy of pediatrics.

The collected data was entered in the statistical package for social sciences (SPSS) version 21 for analysis. Gender was expressed as percentages and frequency whereas numerical variables like age and Vitamin D levels were expressed as mean and standard deviation. Independent t-test was used to determine the significant difference of means between controls and patients. P values less than 0.05 was considered as significant.

#### RESULTS

A total of 1587 children in between the age of 5 - 15years were examined during the study period. After fulfilling the inclusion /exclusion criteria, 100 children were selected by simple convenient sampling method for each group. Group I consisted of 47.18% males and 52.11% females whereas in group II were 55.82% males and 44.17% females. The mean ages of controls and myopics were  $10.65 \pm 3.9$  and  $10.20 \pm 2.5$  years respectively. The age difference between the two groups was statistically insignificant (Table 1). In group I, 79.4% had mild myopia, 19.6% had moderate myopia and 0.6% had high myopia. The Vitamin D levels in myopic children were found to be 14.95  $\pm$ 3.75 ng/ml and there was no significant difference in mean values of Vitamin D levels in myopic and control group.

Table 1:	Descriptive	<b>Statistics</b>	for the	Myopia and	d Control.

		Group I (Myopia) n = 100	Group II (Control) n = 100	P-value
Mean Age (years ± SD)		$10.20\pm2.5$	$10.65\pm3.9$	0.789
Gender	Male Female	47.18% 52.11%	55.82% 44.17%	0.675 0.647
Serum Vitamin D levels (ng/ml)		$15.95\pm3.75$	$16.02\pm5.11$	0.625
Spherical equivalent refractive error (D)		$-3.08 \pm 2.45$	$+0.25\pm0.26$	Not applicable

#### DISCUSSION

The pathogenesis of myopia has been a hot debate since ages. Its association with excessive near work and less outdoor activity has been advocated by many<sup>13</sup>. Atta Z et al<sup>14</sup> found myopia in 52.6% of the children studying in madrasas and proposed less outdoor activity to be associated with myopia. Similarly, Pan CW<sup>15</sup> found that children involved more in outdoor activity have less chances of myopia. Similar idea was advocated by Tideman JW et al<sup>16</sup>. But the question remained unanswered that how outdoor activity affects refractive error development. It was proposed that better quality of retinal image is attained while viewing distant objects with a small pupil size and accommodative errors may be inhibiting ocular growth and decreasing the risk of myopia<sup>17</sup>. However, evidence from animal models did not support this hypothesis. Another proposed hypothesis was an increase in retinal dopamine secretion in response to bright light during outdoors suppresses axial elongation but again no scientific evidence was presented.

Vitamin D is a vital element for absorption of calcium in the intestines and plays a significant role in the growth of bones along with mineral and Calcium homeostasis9. The commonly used indicator of Vitamin D status is serum 25 (OH) D concentrations. Researchers have found a strong association between serum 25 (OH) D concentrations and myopia advocating greater time spent outdoors reduces the risk of myopia. Sherwin JC et al<sup>8</sup> reported low serum concentrations of Vitamin D in myopes. Similarly, low Vitamin D concentration was reported by Tideman JW et al<sup>16</sup> in patients with high axial length. How Vitamin D prevents myopia remains to be answered. One theory is that, as deficiency of Vitamin D leads to alteration of intracellular Ca level and subsequently causes impaired contraction and relaxation of the ciliary muscles thus leading to myopia<sup>18</sup>. Another theory is Calcium deficiency secondary to lack of adequate Vitamin D levels leading to head and orbit deformity and consequently myopia of prematurity but against this was the finding of no change in refractive status of low birth weight infants after extra-enteral Ca supplementation<sup>18</sup>. Vitamin D is also thought to affect pathological scleral growth and myopia via retinoic acid. Retinoic acid and Vitamin D receptors form heterodimers which participate in signaling and cellcycle regulation but data based studies are lacking<sup>17</sup>.

In our study, we found no difference in Vitamin D levels of myopics and age matched control. Same finding was reported by another researcher, according to whom myopes prefer to stay indoors and consequently have low 25 (OH) D3 levels therefore serum 25 (OH) D3 is simply a biomarker of sun exposure<sup>19</sup>. Hence outdoor activity might be mediating myopia prevention via some other pathway instead of Vitamin D. Analysis done by Williams KM et al<sup>20</sup> also negated the hypothesis that outdoor activity protects against myopia by Vitamin D levels. Furthermore, Williams KM et al<sup>20</sup> objected that if outdoor activity is protective against myopia it should slow the progression in those who already have myopia. These questions are yet to be answered.

In our study the participants had below normal mean 25 [OH] D concentrations and were below the accepted normal value of 20 ng/ml. This is in contrast to other studies<sup>15,16</sup> in which only myopes had low levels of Vitamin D. This might be due to other factors like ethnicity, outdoor activity and diet. Vitamin D deficiency is reported more in African Americans compared to Caucasians and more in the regions of Middle East, China, Mongolia, and the Indian subcontinent<sup>21</sup>. The confounder of ethnicity was not addressed in our study. Likewise, high Vitamin D levels and less incidence of myopia is advocated to be associated with more outdoor activity<sup>15</sup> and to higher dietary intake of Vitamin D<sup>16</sup> but Mutti et al<sup>17</sup> found no association. According to Mutti et al<sup>17</sup> the prevalence of myopia is on the rise in Asian population despite the Vitamin D rich fish diet. Therefore, further researchis needed to identify relevant biological connections between Vitamin D and myopia. The diet confounder was also not taken into account in our study.

Limitation of the study is that it was an observational study not a randomized trial. The confounding factors like time spent outdoors, seasonal variation of measurement and sunlight exposure, dietary habits and demographic variables were not taken into account.

## CONCLUSION

We found no difference in Vitamin D levels of myopic and non myopic children and concluded that Vitamin D has no role in development or progression of myopia.

## **Ethical Approval**

The study was approved by the Institutional review board/Ethical review board.

## **Conflict of Interest**

Authors declared no conflict of interest

#### Authors' Designation and Contribution

Yasir Iqbal; Associate Professor: Study design, Data Collection, Manuscript Writing, Final review

Aqsa Malik; Assistant Professor: Study design, Manuscript Writing, Final review.

below the Massoma Talib: Assistant Professor: Study

Masooma Talib; Assistant Professor: Study design, Final review.

Rabbia Shabbier; Lecturer: Study design, Final review.

Atteava Zaman, Assistant Professor: Study design,

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Final review.

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