

Vitamin D deficiency: A risk factor for myopia in children — a cross sectional study in a tertiary care centre

Sahira Aaraj,¹ Ayisha Kausar,² Sabeen Abid Khan³

Abstract

Objective: To study the role of vitamin D deficiency as a risk factor for myopia in children aged 5-15 years.

Method: The cross-sectional observational study was conducted from January to September 2019 at the Ophthalmology and Paediatric departments of Shifa Foundation Community Health Centre, Islamabad, Pakista. It comprised patients with suspected / symptomatic vitamin D deficiency who were enrolled from the paediatric outpatient department and referred to the ophthalmology clinics for eye exam. Apart from taking detailed ocular history, slit lamp examination, Snellen's distance visual acuity, auto-refraction to calculate spherical equivalent, and amplitude scan for measuring the axial length were performed. An average of 3 measurements was taken for both refraction and axial length calculation. Myopia was labelled if mean spherical equivalent was ≤ 0.25 dioptres. Vitamin D deficiency was defined as serum 25-hydroxyvitamin D level < 20 ng/ml. Data was analysed using SPSS 23.

Results: There were 72 subjects with a mean age of 10.11 ± 2.69 years; 37(51.4%) boys and 35(48.6%) girls. Myopia was seen in 40(55.6%) patients, while 32(44.4%) were emmetropic. The overall mean vitamin D level was 20.25 ± 12.18 ng/ml. There was no significant association between myopia and vitamin D deficiency ($p=0.115$). Significant associations were found between myopia and relatively older age ($p=0.005$), higher height ($p=0.001$), more weight ($p=0.001$) and higher body mass index value ($p=0.008$).

Conclusion: Low vitamin D levels were not significantly associated with myopia in children aged 5-15 years, but significant associations were found between myopia and relatively older age, and various anthropometric measures.

Keywords: Myopia, Vitamin D deficiency, Refractive error. (JPMA 72: 1075; 2022)

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Introduction

Visual impairment and blindness have an undesirable effect on learning abilities as well as on social and psychological health of a child.¹ Uncorrected refractive error (RE) is the commonest cause of visual impairment in children.² Population-based studies of uncorrected REs in different countries concluded a prevalence of 22.3% in China, 17.3% in Singapore and 17.1% in Malaysia.^{2,3}

Amongst all variations of REs, myopia is the commonest type.⁴ In Pakistan, a developing country with a population of over 220 million,⁵ the documented prevalence of myopia is 3.3% in children.⁶ Researchers have reported high incidence of myopia in children aged 5-15 years.⁴ In order to reduce the burden of RE associated visual disabilities in children, various treatable aspects contributing to the high incidence of myopia need to be investigated.

Over centuries epidemiological studies have recognised multiple potential risk factors for the development of myopia. Researchers have found low incidence of myopia

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^{1,3}Department of Pediatrics, ²Department of Ophthalmology, Shifa College of Medicine, Shifa Tameer-e-Millat University, Islamabad, Pakistan.

Correspondence: Sabeen Abid Khan. Email: sabeenk45@gmail.com

in those who spent more time outdoors compared to the ones more involved in indoor activities. This led to the concept that vitamin D might be a possible moderator of this association.⁷ All over the world, vitamin D deficiency and insufficiency have been found in all age groups, including children, belonging to different socioeconomic strata.⁸ Recent international studies reported an association with serum 25-hydroxy vitamin D 25OHD concentration and myopia.⁹⁻¹¹ Whether this reflects the association between outdoor exposure and myopia, or whether vitamin D itself plays a role in the pathophysiology is unclear.

The current study was conducted to investigate the association between the prevalence of myopia in children with suspected / symptomatic vitamin D deficiency aged 5-15 years.

Subjects and Methods

This cross-sectional observational study was conducted from January to September 2019 at the Ophthalmology and Paediatric departments of Shifa Foundation Community Health Centre, Islamabad, Pakistan. After approval from the institutional ethics review committee, the sample size was calculated using the OpenEpi calculator¹² with an expected frequency of children with

REs as $3.3 \pm 5\%$,⁶ confidence interval (CI) 99% and margin of error 5%.

The sample was raised using non-probability convenience sampling technique from the paediatric outpatient department (OPD) and were referred to the ophthalmology OPD for eye exam. Those included were paediatric patients of either gender aged 5-15 years with suspected / symptomatic vitamin D, deficiency. Children with vitamin D deficiency secondary to renal or hepatic issues and medications were excluded. Also excluded were those with ocular conditions other than REs, like congenital cataract, congenital glaucoma, ocular trauma and corneal dystrophy, and patients with myopia associated with connective tissue disorders, like the Marfan syndrome.

After taking written informed consent from the parents or guardians, detailed dietary history, history of delayed dentition, delayed walking, pathological fracture, bone pains, backache, excessive sweating was taken and general physical examination was performed. At the ophthalmology OPD, detailed ocular history and examination was performed, including detailed slit lamp examination, Snellen's distance visual acuity (VA), auto-refraction to calculate spherical equivalent (SE) and amplitude scan (A-scan) for measuring the axial length. Cycloplegic refraction and fundoscopy was done in children with defective vision, and spectacles were prescribed accordingly. Axial lengths were measured using A-scan (Quantal Medical Axis-II). An average of 3 measurements was taken for both refraction and axial length calculation. Myopia was labelled as $SE \leq -0.25$ dioptres.

For vitamin D deficiency, blood sample was drawn through antecubital venipuncture and it was analysed through chemiluminescence. Vitamin D deficiency was defined as serum 25OHD level $<20\text{ng/ml}$. The demographic information and clinical features were noted on a specially-designed proforma.

Data was analyzed using SPSS 23. Mean \pm standard deviation values were calculated for quantitative variables, like age, height, weight, body mass index (BMI), vitamin D levels, axial length and SE. Frequencies and percentages were recorded for qualitative variables, like gender, parental myopia, time spent outdoors, dietary habits with focus on intake of milk, eggs and vitamin D supplements, birth order, and the presence or absence of myopia. Pearson's chi-square test was used to compare the difference in myopic and non-myopic population regarding various qualitative variables, and anthropometric measures including height, weight, BMI

and vitamin D levels. For numeric data, Shapiro-Wilk test was used to check data normality. Leven's test for equality of variance was used to measure homogeneity of variance between the groups. To compare means, variables with normal distribution were analysed with parametric independent sample t-test, and variables showing non normal distribution were analysed using Mann-Whitney U test. Confidence level was set at 95%. $P \leq 0.05$ was taken as statistically significant.

Results

Initially, 96 patients were enrolled, but comprehensive data was available for 72(75%) subjects; 37(51.4%) boys and 35(48.6%) girls. The overall mean age was 10.11 ± 2.69 years. Myopia was seen in 40(55.6%) patients, while 32(44.4%) were emmetropic. The overall mean vitamin D level was $20.25 \pm 12.18\text{ng/ml}$. There was no significant

Table-1: Demographic characteristics in the study groups.

	n72	Myopia n=40 (55.6%)	Emmetropes n=32 (44.4%)	P value
Gender	Male	20 (27.8%)	17 (23.6%)	0.792
	Female	20 (27.8%)	15 (20.8%)	
Residence	Urban	32 (44.4%)	32 (44.4%)	0.007
	Rural	8 (11.1%)	0	
Height Centile	< 5th centile	9 (12.5%)	15 (20.8%)	0.069
	5-10th centile	6 (8.3%)	5 (6.9%)	
	>10th centile	25 (34.7%)	12 (16.7%)	
Weight Centile	< 5th centile	10 (13.9%)	18 (25.0%)	0.009
	5-10th centile	8 (11.1%)	7 (9.7%)	
	>10th centile	22 (30.6%)	7 (9.7%)	
Parental Refractive Error	Both	11 (15.3%)	8 (11.1%)	0.638
	None	21 (29.2%)	17 (23.6%)	
Birth Order	Either parent	8 (11.1%)	7 (9.8%)	0.066
	Eldest	17 (23.6%)	14 (19.4%)	
	2nd born	9 (12.5%)	12 (16.7%)	
	3rd born	4 (5.6%)	6 (8.3%)	
	4th born	6 (8.3%)	0	
	5th born	2 (2.8%)	0	
Milk Intake	6th Born	2 (2.8%)	0	0.285
	Daily	15 (20.8%)	13 (18.1%)	
	1-2/week	14 (19.4%)	15 (20.8%)	
	3-4/week	2 (2.8%)	2 (2.8%)	
Egg consumption	None	9 (12.5%)	2 (2.8%)	0.151
	None	9 (12.5%)	7 (9.7%)	
	1-2/week	13 (18.1%)	17 (23.6%)	
Intake of Vitamin D Supplements	3-4/week	18 (25.0%)	8 (11.1%)	0.843
	Yes	7 (9.7%)	7 (9.7%)	
	No	20 (27.8%)	14 (19.4%)	
Time outdoor	Occasional	13 (18.1%)	11 (15.3%)	0.477
	> 3 hrs	3 (4.2%)	4 (5.6%)	
Serum Vitamin D levels	$\leq 3\text{hrs}$	37 (51.4%)	28 (38.9%)	0.49
	Deficient	26 (39.4%)	19 (28.8%)	
	Insufficient	6 (9.0%)	7 (10.6%)	
	Sufficient	3 (4.5%)	5 (7.6%)	

Table-2: The association of myopia with study variables.

	Study Cohort	Myopia	Emmetropia	p- value
Mean Age (years \pm SD)	10.11 \pm 2.69	11.12 \pm 2.50	9.35 \pm 2.62	0.005
Height (cm)	132.80 \pm 15.88	138.28 \pm 15.61	126.13 \pm 14.55	0.001
Weight (kg)	27.86 \pm 10.42	32.48 \pm 10.39	23.88 \pm 9.04	0.001
BMI	15.49 \pm 3.25	16.60 \pm 3.06	14.72 \pm 2.90	0.008
Serum Vitamin D levels (ng/ml)	20.25 \pm 12.18	16.11 \pm 7.38	24.46 \pm 17.64	0.115
Axial length (mm)	22.97 \pm 0.83	23.35 \pm 0.79	22.59 \pm 0.76	0.001
Spherical equivalent (Diopters Sphere)	-0.31 \pm 1.42	-1.21 \pm 1.09	0.77 \pm 1.29	<0.000

SD: Standard deviation, BMI: Body mass index.

Table-3: Association of vitamin D levels with various variables.

Clinical parameters	p- value
Age	0.469
Height (cm)	0.277
Height centile	0.707
Weight centile	0.719
Weight (kg)	0.533
BMI	0.967
Urban/ Rural residence	0.503
Birth order	0.889
Intake of dairy products	0.536
Consumption of eggs	0.637
Intake of Vitamin D supplements	0.011
Time spent in sunlight	0.970

BMI: Body mass index.

difference between the two groups in terms of gender, birth order, parental myopia, time spent outdoors, serum vitamin D levels and intake of vitamin D supplements, milk and eggs ($p > 0.05$). Myopia was more common in children from rural areas ($p = 0.007$) and weight was a significant factor between the groups (Table-1).

Significant associations were found between myopia and relatively older age ($p = 0.005$), higher height ($p = 0.001$), more weight ($p = 0.001$) and higher body mass index value ($p = 0.008$) as well in terms of axial length ($p = 0.001$ and SE (Table-2). Regarding vitamin D levels and myopia, there was no significant association ($p = 0.115$).

The intake of vitamin D supplements significantly affected serum vitamin D levels ($p = 0.011$), but no significant association was found between serum vitamin D levels and other dietary / environmental factors (Table-3).

Discussion

The current study established that serum Vitamin D levels had no association with myopia after adjusting for confounding factors. Vitamin D is currently perceived as a hormone which affects almost all body systems apart from calcium homeostasis and bone health.⁸ Most of the study population had low vitamin D level in both groups, but low

serum vitamin D levels were not found to have any significant association with myopia. A study in Pakistan reported 58.17% of the population being deficient in vitamin D and 26.65% having insufficient vitamin D.¹³ Similar observations were reported in another local study in Azad Kashmir.¹⁴ Comparable results were reported among adults in a multi-country European study.¹⁵ No convincing evidence for a direct role of vitamin D in myopia risk was found. In contrast, certain international studies have reported that low vitamin D levels were inversely proportional to the risk of myopia in children.^{10,16-19} Myopia is prevalent in the Asian population.¹ At present, recommended methods for preventing myopia that are supported by strong evidence are time spent outdoors, bifocal lenses, and a drop of atropine.¹⁹ Studies on outdoor activities are a bit controversial as some have indicated no association,¹⁹ while others reported that outdoor time was protective against myopia.²⁰⁻²² The current study did not find significant association of myopia with time spent outdoor. A systematic review and meta-analysis from China demonstrated the protective effect of outdoor time against myopia onset, but not for myopic progression.⁷ The exact mechanism of outdoor light in preventing myopia is yet to be determined.^{7,20}

Parental myopia is thought to be associated with a greater risk of early-onset myopia in the Asian population,^{23,24} and some studies have found parental myopia as a risk factor for early-onset myopia.^{24,25} In contrast, the current study could not find a significant association between single or both parent RE and children myopia ($p = 0.638$). Different population cohorts or limited sample size could be the possible reason for this.

The current study found equal gender distribution in myopic patients at a M:F ratio of 1:1. Similar finding were reported earlier,²⁶ While other studies reported a male predominance.²⁷

Age was significantly associated with myopia, as highlighted by the fact that the axial length increased

with age, and the mean age was higher in the myopic patients compared to the non-myopes ($p=0.005$). In literature, myopia prevalence has been reported to be significantly higher in children aged 12-13 years ($p<0.001$).²⁸

Regarding vitamin D supplementation, most patients in both groups in the current study were not on regular vitamin D supplements, and their consumption of dairy products was inadequate which explains deficient and insufficient vitamin D status in these subjects. Lack of awareness and access to these food items can be responsible for the low levels.

Height, weight and BMI in the current study were positively associated with myopia. Previous reports have shown that height is associated with increased axial length. Changes in axial length may involve remodelling of the scleral extracellular matrix, which would increase the axial length because of the lengthening of the vitreous chamber. Similar findings were observed in a study which found that taller children and those with higher BMI were more likely to develop myopia.²⁸⁻³⁰ However, one study showed that myopic RE was not associated with weight or BMI.³¹ These inconsistencies may result from ethnic and demographic differences.

Children living in rural areas were seen to have myopia more frequently in the current study, which is a finding different from others who found more myopia in urban population.²⁸ The difference in finding may be because of the fact that the current study had no patient from the rural population in its emmetropic group which might be a confounding factor.

The prevalence of myopia was numerically higher in first-born child in the current study (23.6%) as has been reported earlier.³²

Regarding various dietary and lifestyle factors noted in the current study, no significant association was between serum vitamin D level and dietary and lifestyle factors because the study population had low vitamin D levels uniformly with similar dietary and lifestyle patterns.

The current study has certain epidemiological limitations, as, being a cross-sectional study, it inherently lacked the effect of temporality. Also, it could not take into account the effect of seasonal variations on serum vitamin D levels. Data was collected in resource-limited conditions and the study used convenience sampling. The sample size was also relatively small.

Conclusion

The mean serum vitamin D levels were lower in myopic

children, but the difference was not significant compared to the non-myopic patients. Myopia was more common in relatively older children living in rural areas and having higher height, weight and BMI values.

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Conflict of Interest: None.

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References

1. Abbas H, Awais M, Naimat K. Prevalence and pattern of refractive errors in school going children of Mangla Cantonment. *Pak Armed Forces Med J.* 2019; 69:1125-28.
2. World Health Organization. Visual Impairment and Blindness. [Online] [Cited 2021 October 25]. Available from: URL: <http://www.who.int/mediacentre/factsheets/fs282/en/>.
3. Wu PC, Huang HM, Yu HJ, Fang PC, Chen CT. Epidemiology of myopia. *Asia Pac J Ophthalmol.* 2016; 5:386-93.
4. Bodla MA, Bodla AA, Moazzam A, Tariq N. Correlation between changes in intraocular pressure and refractive error indices post Cataract Surgery. *Pak J Med Sci.* 2020; 36:574-7.
5. Bukhari MH, Mahmood K, Zahra SA. Over view for the truth of COVID-19 pandemic: A guide for the Pathologists, Health care workers and community'. *Pak J Med Sci.* 2020; 36:COVID19-5111-4.
6. Gull A, Raza A. Visual screening and refractive errors among school aged children. *J Rawalpindi Med Coll (JRMC).* 2014; 18:97-100.
7. Xiong S, Sankaridurg P, Naduvilath T, Zang J, Zou H, Zhu J, et al. Time spent in outdoor activities in relation to myopia prevention and control: a meta-analysis and systematic review *Acta Ophthalmol.* 2017; 95:551-66.
8. Moorani KN, Mustufa MA, Hasan SF, Kubar N. Vitamin D status in under five children in diverse communities of Karachi. *Pak J Med Sci.* 2019; 35:414-9.
9. Yazar S, Hewitt AW, Black LJ, McKnight CM, Mountain JA, Sherwin JC, et al. Myopia is associated with lower vitamin D status in young adults. *Invest Ophthalmol Vis Sci.* 2014; 55:4552-9.
10. Williams KM, Bentham GCG, Young IS, McGinty A, McKay GJ, Hogg R, et al. Association between Myopia, Ultraviolet B Radiation Exposure, Serum Vitamin D Concentration and Genetic Polymorphism in Vitamin D Metabolic Pathways in a Multicountry European Study. *JAMA Ophthalmol.* 2017; 135:47-53.
11. Tang SM, Lau T, Rong SS, Yazar S, Chen LJ, Mackey DA, et al. Vitamin D and its pathway genes in myopia: systematic review and meta-analysis. *Br J Ophthalmol.* 2019; 103:8-17.
12. Dean AG, Sullivan KM, Soe MM. Open Epi: Open Source Epidemiologic Statistics for Public Health, Version. [Online] [Cited 2021 November 4]. Available from: URL: www.OpenEpi.com
13. Syed F, Latif MSZ, Ahmed I, Bibi S, Ullah S, Khalid N. Vitamin D deficiency in Pakistani population: critical overview from 2008 to 2018. *Nut Food Sci.* 2020; 50:105-15.
14. Iqbal Y, Malik A, Shabbier R, Zaman A, Zia S, Talib M. Role of Vitamin D in Near Sightedness, *Pak J Ophthalmol.* 2020; 36:137-41.
15. Williams KM, Bentham GCG, Young IS, McGinty A, McKay GJ, Hogg R, et al. Association Between Myopia, Ultraviolet B Radiation Exposure, Serum Vitamin D Concentrations, and Genetic Polymorphisms in Vitamin D Metabolic Pathways in a Multicountry European Study. *JAMA Ophthalmol.* 2017; 135:47-53.
16. Jung BJ, Jee D. Association between serum 25-hydroxyvitamin D

- levels and myopia in general Korean adults. *Indian J Ophthalmol.* 2020; 68:15-22.
17. Tideman JW, Polling JR, Voortman T, Jaddoe VW, Uitterlinden AG, Hofman A et al. Low serum vitamin D is associated with axial length and risk of myopia in young children. *Eur J Epidemiol.* 2016; 31:491-9.
 18. Tang SM, Lau T, Rong SS, YazarS, ChenLJ, Mackey DA, et al. Vitamin D and its pathway genes in myopia: systematic review and meta-analysis. *Br J Ophthalmol.* 2019; 103:8-17.
 19. Jin-woo K, Jin AC, Yoon LT. Serum 25-hydroxyvitamin D level is associated with myopia in the Korea national health and nutrition examination survey, *Med. Medicine (Baltimore).* 2016; 95:e5012.
 20. Pan CW, Dirani M, Cheng CY, Wong TY, Saw SM. The Age-Specific Prevalence of Myopia in Asia. *Optom Vis Sci.* 2015; 92:258-66.
 21. Theophanous C, Modjtahedi BS, Batech M, Marlin DS, Luong TQ, Fong DS. Myopia prevalence and risk factors in children. *Clin Ophthalmol.* 2018; 12:1581-7.
 22. Guo Y, Liu LJ, Tang P, Lv YY, FengY, XuL, et al. Outdoor activity and myopia progression in 4-year follow-up of Chinese primary school children: The Beijing Children Eye Study. *PLoS One.* 2017; 12.
 23. Ansar A, MR Robina, Nawaz MA, KK Maheen, Waqas A. Risk factors for refractive errors among school going children in wah cantonment, Rawalpindi. *RMJ.* 2016; 41:432-6.
 24. Jiang X, Hornoch KT, Cotter SA, Matsumura S, Mitchell P, Rose KA, et al. Association of Parental Myopia With Higher Risk of Myopia Among Multiethnic Children Before School Age. *JAMA Ophthalmol.* 2020; 138:501-9.
 25. Jiang D, Lin H, Li C, Liu L, Xiao H, Lin Y, et al. Longitudinal association between myopia and parental myopia and outdoor time among students in Wenzhou: a 2.5-year longitudinal cohort study. *BMC Ophthalmology.* 2021; 21:11.
 26. Latif MZ, Khan MA, Afzal S, Gillani SA, Chouhadry MA. Prevalence of refractive errors; an evidence from the public high schools of Lahore, Pakistan. *J Pak Med Assoc.* 2019; 69:464-7.
 27. Wajuihian SO, Mashige KP. Gender and age distribution of refractive errors in an optometric clinical population. *J Optom.* 2021; 14:315-27.
 28. Harrington SC, Stack J, Dwyer VO. Risk factors associated with myopia in school children in Ireland. *Br J Ophthalmol.* 2019; 103:1803-9.
 29. Lee DC, Lee SY, Kim YC. An epidemiological study of the risk factors associated with myopia in young adult men in Korea. *Sci Rep.* 2018; 8:511.
 30. Kim H, Seo JS, Yoo WS, Kim GN, Kim RB, Chae AG, et al. Factors associated with myopia in Korean children: Korea National Health and nutrition examination survey 2016–2017 (KNHANES VII). *BMC Ophthalmol.* 2020; 20:31.
 31. Jung SK, Lee JH, Kakizaki H, Jee D. Prevalence of myopia and its association with body stature and educational level in 19-year-old male conscripts in Seoul, South Korea. *Invest Ophthalmol Vis Sci.* 2012; 53: 5579-83.
 32. Guggenheim JA, McMahon G, Northstone K, Mandel Y, Kaiserman I, Stone RA, et al. Birth order and myopia. *Ophthalmic Epidemiol.* 2013; 20:375-84.
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