Check for updates

OPEN ACCESS

EDITED BY Xiuming Jin, Zhejiang University, China

REVIEWED BY Sebastião Cronemberger, Federal University of Minas Gerais, Brazil Raul Martin, IOBA - Universidad de Valladolid, Spain

*CORRESPONDENCE Aicun Fu Maicun2019@qq.com

RECEIVED 24 July 2023 ACCEPTED 13 October 2023 PUBLISHED 02 November 2023

CITATION

Qin J, Qing H, Ji N, Lyu T, Ma H, Shi M, Yu S, Ma C and Fu A (2023) Changes in axial length in anisometropic children wearing orthokeratology lenses. *Front. Med.* 10:1266354. doi: 10.3389/fmed.2023.1266354

COPYRIGHT

© 2023 Qin, Qing, Ji, Lyu, Ma, Shi, Yu, Ma and Fu. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.

Changes in axial length in anisometropic children wearing orthokeratology lenses

Jian Qin¹, Huiling Qing¹, Na Ji², Tianbin Lyu¹, Hui Ma¹, Menghai Shi¹, Shiao Yu³, Conghui Ma¹ and Aicun Fu^{3*}

¹Department of Ophthalmology, Henan Provincial People's Hospital, Henan Eye Hospital, Henan Eye Institute, People's Hospital of Zhengzhou University, Zhengzhou, China, ²Department of Ophthalmology, The Affiliated Eye Hospital of Suzhou Vocational Health College, Suzhou, China, ³Department of Ophthalmology, The First Affiliated Hospital of Zhengzhou University, Zhengzhou, China

Purpose: There is a particular anisometropia occurring in one eye with myopia, while the other eye has very low myopia, emmetropia, or very low hyperopia. It is unclear how the binocular axial length changes when these children wear unilateral OK lenses only in the more myopic eyes. This study investigates the changes in the axial elongation of both eyes.

Methods: This is a 1-year retrospective study. In total, 148 children with myopic anisometropia were included. The more myopic eyes were wearing orthokeratology lenses (treated eyes), whereas the contralateral eyes were not indicated for visual correction (untreated eyes). The untreated eyes were classified into three subgroups based on the spherical equivalent refraction (SER): low myopia (\leq -0.50 D, n = 37), emmetropia (+0.49 to -0.49 D, n = 76), and low hyperopia (\geq 0.50 D, n = 35). Changes in the axial length (AL) were compared between the untreated and treated eyes and among the three subgroups.

Results: The axial elongation was 0.14 \pm 0.18 mm and 0.39 \pm 0.27 mm in all treated and untreated eyes, respectively (p < 0.001). The interocular AL difference decreased significantly from 1.09 \pm 0.45 mm at the baseline to 0.84 \pm 0.52 mm at 1 year (p < 0.001). The baseline median (Q1, Q3) SER of the untreated eyes were -0.75 D (-0.56, -0.88 D), 0.00 D (0.00, -0.25 D), and +0.75 D (+1.00, +0.62 D) in low myopia, emmetropia, and low hyperopia subgroups, respectively. The axial elongation was 0.14 \pm 0.18 mm, 0.15 \pm 0.17 mm, and 0.13 \pm 0.21 mm (p = 0.92) in the treated eyes and 0.44 \pm 0.25 mm, 0.35 \pm 0.24 mm, and 0.41 \pm 0.33 mm in the untreated eyes (p = 0.11) after 1 year. Multivariate linear regression analyses only showed significant differences in axial elongation between the emmetropia and low myopia subgroups of untreated eyes (p = 0.04; p > 0.05 between other subgroups).

Conclusion: Unilateral orthokeratology lenses effectively reduced axial elongation in the more myopic eyes and reduced interocular AL differences in children with myopic anisometropia. The refractive state of the untreated eyes did not affect the axial elongation of the more myopic eye wearing the orthokeratology lens. In the untreated eyes, AL increased faster in the low myopia subgroup than in the emmetropia subgroup.

KEYWORDS

anisometropia, myopia, unilateral orthokeratology lens, axial length, refractive state

1. Introduction

Anisometropia is defined as the difference of \geq 1.00 D between the spherical equivalents of the two eyes (1). Anisometropia >2.00 D against the rule cylinder and 3.00 D sphere (2) may cause strabismus, amblyopia, binocular vision dysfunction, and asthenopia (3-5). Various measures such as spectacles and contact lenses with different designs were used to correct anisometropia. However, due to the lens effect-related aniseikonia, many anisometropic patients may not tolerate spectacles (6). Traditional rigid gas-permeable contact lenses (7) and singlevision soft contact lenses (8) can decrease aniseikonia and improve binocular vision but cannot prevent the progression of myopia (7, 9). Additionally, some children cannot tolerate the foreignbody sensation caused by daily wearing rigid gas-permeable contact lenses. These are not ideal methods for controlling myopia progression and decreasing anisometropia in children with myopic anisometropia (6-9). Orthokeratology (OK) lenses are specifically designed gas-permeable contact lenses that can not only improve the unaided visual acuity (VA) but also reduce anisometropia and prevent myopia progression in anisometropic patients (10-19). Two 2-year (11, 15) and three 1-year (12-14) retrospective studies found a significant reduction in interocular axial length (AL) difference of unilateral myopic anisometropia (UMA, one eye is myopic and the other eye is emmetropic) children wearing a unilateral OK lens.

Just like UMA children, anisometropic children have more myopia in one eye and very low myopia or very low hyperopia in the other eye. They are common that UMA children wear the OK lenses just in the more myopic eye, while the contralateral untreated eye serve as control. In order to understand the change in AL of both eyes in these children and find reasonable treatments, this study compared the interocular differences in AL and the AL changes in the more myopic eyes wearing the OK lenses during different refractive states of the contralateral untreated eyes. AL changes in the contralateral untreated eyes in three different refractive states were also compared.

2. Methods

2.1. Participants

Data obtained from the patients' clinical records of 148 children with UMA who visited *** hospital: the First Affiliated Hospital of Zhengzhou University between January 2017 and September 2019 and met the following criteria were retrospectively collected. The inclusion criteria were as follows: patients aged 8–14 years, cycloplegic spherical equivalent refraction (SER) of -1.00 D to -4.50 D in the more myopic eye and +2.00 D to -1.25 D in the contralateral eye, anisometropia of \geq 1.0 D, astigmatism of <2.00 D in both the eyes, monocular best corrected VA of 0 (LogMAR charts) or better, intraocular pressure of <21 mmHg, no history of use of medications or contact lenses to control the progression of myopia, no manifest strabismus, no history of other eye diseases and surgery, and the decentration of treatment zone <1 mm during the wearing of OK lenses.

2.2. Study design

The more myopic eyes were wearing the OK lenses (treated eyes), whereas and the contralateral eyes were not indicated for visual correction (untreated eyes). The untreated eyes were classified into three subgroups according to their refractive states: low myopia (SER: \leq -0.50 D), emmetropia (SER: +0.49 D to -0.49 D), and low hyperopia (SER: \geq 0.50 D) (20, 21). Therefore, the treated eyes were classified into three subgroups according to the refractive state of the untreated eyes (low myopia, emmetropia, or low hyperopia). Four-zone five-curve reverse geometry OK lenses [Boston XO material; oxygen permeability 100×10^{-11} (cm²/s) (mlO²/ml × mmHg); Autek Corporation, Hefei, China] were used. All patients wore the OK lenses continuously for 8–10 h during sleep every day. Routine follow-ups were arranged, and additional unscheduled visits were arranged if necessary.

AL, corneal power, corneal astigmatism, and anterior chamber depth (ACD) were measured using a non-contact partial coherence interferometer (IOLMaster-500, Carl Zeiss, Jena, Germany). Three drops of 1% cyclopentolate (Alcon, Puurs, Belgium) were instilled with 10-min intervals between consecutive administrations. Three autorefraction measurements (ARK-1, Nidek, Japan) were performed 10 min later. The mean values indicated the degree of myopia, which is expressed as SER. OK lenses were replaced at approximately every 1.5 years. However, the OK lens was reordered when the residual SER was > -0.50 D at any visit after treatment stabilization.

Corneal topography with the Topographic Modeling System (TMS-4, Tomey Corporation, Tokyo, Japan) was used to measure the corneal parameters by the same professional technician. A different tangential curvature corneal topography was established by subtracting the pre-OK map from the 1-month post-OK in this study. We adjusted the refractive power change around the corneal vertex to 1.00 D aiming to keep the color of the optical area within two colors. The central blue zone is the treatment zone, and the surrounding red zone is the reverse curve zone. The center of the treatment zone and degree of decentration on the tangential difference map were analyzed with MATLAB. The distance from the pupil center to the center of the treatment zone was considered to be a decentration distance (22).

2.3. Statistical analysis

Normal-distributed (mean \pm SD) and skew-distributed continuous variables [medians with first (Q1) and third quartiles (Q3)] of baseline parameters were compared using a *t*-test and non-parametric rank-sum test, respectively, and the chi-squared test was used to evaluated categorical variables. The independent sample *t*-test was conducted to assess differences in axial elongation after 1 year between treated and untreated eyes and interocular difference in AL between baseline and 1 year. Analysis of variance (ANOVA) was used to assess the axial elongation after 1 year among different subgroups of the treated and untreated eyes. Furthermore, multivariate linear regression analyses were applied to estimate the axial elongation between untreated and treated eyes, the changes in AL among different subgroups of the treated and untreated eyes, and the reduction in the interocular AL difference. All statistical analyses in this study were performed in software Empower (R; www.empowerstats.com, X & Y Solutions Inc., Boston, MA, USA) and R (http://www.R-project.org, the R foundation, Austria, Vienna).

3. Results

Table 1 shows the baseline data of the patients. No significant differences were observed in corneal power and thickness, ACD, and intraocular pressure (all p > 0.11); however, significant differences were observed in SER, AL, and corneal astigmatism (all p < 0.001) between the untreated and treated eyes. Excluding the anisometropia value and interocular AL difference (both p < 0.001) among the three groups in the treatment eyes, other parameters including SER, AL, corneal power, thickness and astigmatism, ACD, and intraocular pressure did not show a significant difference (all p > 0.29). Excluding SER, AL, corneal astigmatism, anisometropia value, and interocular AL difference (all p < 0.001 except p = 0.01 for astigmatism) among the three groups in the untreated eyes, the other parameters did not show a significant difference (all p > 0.21).

Among the untreated eyes, the uncorrected VA of the low hyperopia and emmetropia subgroups was 0 (LogMAR charts) or better, and 12 eyes (32.4%) in the low myopia subgroup exhibited an uncorrected VA > 0.1, with an average VA of 0.25 \pm 0.17. Among the 37 eyes in the low myopia subgroup, 13 eyes had a sphere diopter between -0.50 and -1.00 D and a cylinder diopter between 0 and -0.75 D, 10 eyes had a sphere diopter of -0.25D and a cylinder diopter between -0.50 and -1.25 D, 13 eyes had a sphere diopter of 0 D and a cylinder diopter between -1.00and -2.00 D, and one eye had a sphere diopter of +0.50 D and a cylinder diopter of -2.00 D. The average SER [median (Q1, Q3)] was -2.50 D (-1.75 D, -3.00 D) and 0.00 D (+0.38 D, -0.41 D) in the treated and untreated eyes, respectively (p < 0.001; Table 1). In the untreated eyes, the average SER values were -0.75 D (-0.56D, -0.88 D), 0.00 D (0.00 D, -0.25 D), and +0.75 D (+1.00 D, +0.62 D) in low myopia (n = 37), emmetropia (n = 76), and low hyperopia (n = 35) subgroups; respectively.

After 1-year follow-up, axial elongation was 0.14 ± 0.18 mm and 0.39 \pm 0.27 mm in all treated and untreated eyes, respectively (independent sample *t*-test, p < 0.001). The interocular difference in AL decreased from 1.09 \pm 0.45 mm at baseline to 0.84 \pm 0.52 mm at 1 year (independent sample *t*-test, p < 0.001) and the reduction in the interocular AL difference (equal to binocular AL difference at baseline minus binocular AL difference at 1 year after treatment) was $0.25 \pm 0.30 \,\mathrm{mm}$ (Table 2). After univariate analysis, the reduction in the interocular AL difference was correlated with age ($\beta = -0.02$, p = 0.04), anisometropia value ($\beta = 0.05$, p =0.03), and refractive state of the untreated eyes (all p < 0.05), but independent of sex (p = 0.76). In the untreated eyes, AL changes in low myopia, emmetropia, and low hyperopia subgroups were $0.44 \pm 0.25 \,\mathrm{mm}, 0.35 \pm 0.24 \,\mathrm{mm}$, and $0.41 \pm 0.33 \,\mathrm{mm}$, respectively (ANOVA, p = 0.11). AL changes in the corresponding eyes from the treated eyes were 0.14 ± 0.18 mm, 0.15 ± 0.17 mm, and 0.13 ± 0.21 mm (ANOVA, p = 0.92).

Multivariate linear regression analyses showed no significant differences in axial elongation in the treated eyes when the corresponding eyes from the untreated eyes had low myopia, emmetropia, or low hyperopia (all p > 0.05; adjusting for age, sex, and anisometropia values) and significant differences in axial elongation only between low myopia and emmetropia subgroups of the untreated eyes (β = 0.09, p = 0.04; p > 0.05 between other subgroups; adjusting for age, sex, corneal astigmatism, and anisometropia). The axial elongation in the low myopia subgroup was 0.09 mm greater than that in the emmetropia subgroup of the untreated eyes (Table 3). Moreover, the axial elongation in the untreated eyes was greater than that in the treated eyes ($\beta = 0.23$, p < 0.001; adjusting for baseline AL, corneal power, and anterior chamber depth). The reduction in the interocular AL difference was significantly associated with age ($\beta = -0.04$, p = 0.009), anisometropia value ($\beta = 0.09$, p = 0.003), and refractive state of the untreated eyes ($\beta_{\text{low myopia vs. emmetropia subgroups}} = -0.12$, p =0.045; p > 0.05 between other subgroups) after multivariate linear regression analyses.

Of all children, 15 children exhibited slight peripheral corneal epithelial staining, but no obvious symptoms of irritation in the early days of wearing the OK lens. However, the eyes showed complete recovery several days after discontinuing the OK lens, with or without 0.1% sodium hyaluronate eye drops. The eyes wearing the OK lens showed no other serious complications, such as corneal stromal infiltration or corneal ulceration.

4. Discussion

This retrospective cohort study included myopic anisometropic children who wore the OK lens only in the more myopic eye. It was observed that axial elongation in the eyes wearing the OK lenses (treated eyes) was significantly slower than that in the uncorrected contralateral eyes (untreated eyes), and the interocular AL differences decreased after 1-year follow-up. The refractive state (low myopia, emmetropia, and low hyperopia) of the untreated contralateral eyes did not affect the axial elongation of the myopic eyes wearing the OK lens. The AL of the untreated contralateral eyes increased faster in the low myopia subgroup than in the emmetropia subgroup.

The results that unilateral OK lenses effectively suppressed axial elongation in the more myopic eyes and reduced interocular AL differences in children with myopic anisometropia were consistent with those of five other studies (Table 4) (11-15). The reduction in interocular AL differences in anisometropic children following treatment with monocular OK lenses may be due to the influence of peripheral defocus induced by OK wear (23) rather than the natural development laws of anisometropia. In a 1-year retrospective study, Long et al. (13) divided patients with UMA into the treated eyes (wearing OK lens only in the myopic eyes) group and the spectacle group, which was matched for age, refraction, and AL. They observed that changes in AL in the myopic eyes (treated eyes) were lower than that in the non-myopic eyes (treated eyes) and both the eyes from the spectacle group. However, there was no statistically significant difference in change in AL between the non-myopic eyes of the two groups and between the two eyes in the spectacle group. The most probable mechanism of action TABLE 1 Baseline characteristics of the study participants grouped by the SER of untreated eyes [mean ± SD or median (Q1, Q3)].

	Treated eyes	Untreated eyes	<i>p</i> -value		Treated	eyes		Untreated eyes				
	All	All		Low myopia in the untreated eyes	Emmetropia in the untreated eyes	Low hyperopia in the untreated eyes	p-value	Low myopia	Emmetropia	Low hyperopia	p-value	
Number of eyes (<i>N</i>)	148	148		37	76	35		37	76	35		
Age (year)	11.5 ± 1.9			11.1 ± 2.2	11.6 ± 1.9	11.6 ± 1.8	0.40					
Sex (male, <i>n</i> and %)	65 (43.9%)			18 (48.6%)	35 (46.1%)	12 (34.3%)	0.41					
SER(D)	-2.50 (-1.75, -3.00)	0.00 (+0.38, -0.41)	<0.001	-2.70 ± 0.94	-2.41 ± 0.95	-2.46 ± 0.78	0.29	-0.75 (-0.56, -0.88)	0.00 (0.00, -0.25)	+0.75 (+1.00,+0.62)	<0.001	
AL (mm)	24.67 ± 0.73	23.58±0.67	< 0.001	24.76 ± 0.76	24.65 ± 0.76	24.62 ± 0.66	0.70	$23.90 \ \pm 0.62$	23.57 ± 0.67	23.28 ± 0.59	<0.001	
Anisometropia value (D)	2.45 ± 1.04			1.90 ± 0.89	2.36 ± 0.95	3.34 ± 0.87	< 0.001					
Interocular AL difference (mm)	1.09 ± 0.45			0.86 ± 0.39	1.08 ± 0.45	1.35 ± 0.38	<0.001					
Corneal power (D)	42.55 ± 1.38	42.45 ± 1.33	0.50	42.36 ± 1.56	42.69 ± 1.41	42.45 ± 1.07	0.53	42.20 ± 1.40	42.62 ± 1.40	42.35 ± 1.04	0.21	
Corneal astigmatism (D)	1.17 ± 0.42	1.45 ± 0.61	<0.001	1.27 ± 0.47	1.15 ± 0.41	1.13 ± 0.40	0.36	1.71 ± 0.68	1.39 ± 0.48	1.25 ± 0.67	0.01	
Central corneal thickness (µm)	534.1 ± 37.0	531.8 ± 36.3	0.63	534.4 ± 42.8	531.1 ± 31.2	537.8 ± 39.6	0.92	528.7 ± 38.5	531.9 ± 30.9	535.1 ± 42.2	0.74	
Intraocular pressure (mmHg)	15.72 ± 2.61	16.04 ± 2.90	0.45	15.50 ± 2.52	15.91 ± 2.80	15.58 ± 2.35	0.82	15.88 ± 2.75	16.30 ± 3.06	15.72 ± 2.78	0.75	
Anterior chamber depth (mm)	3.72 ± 0.27	3.67 ± 0.26	0.11	3.72 ± 0.21	3.75 ± 0.23	3.67 ± 0.36	0.80	3.67 ± 0.25	3.66 ± 0.24	3.60 ± 0.25	0.75	

SER, spherical equivalent refraction; AL, axial length.

TABLE 2 Change in AL of the two eyes in the three subgroups [mean \pm SD or median (Q1, Q3)].

	Treated eyes	Untreated eyes	<i>p</i> -value		Treated	eyes		Untreated eyes				
	All	All		Low myopia in the untreated eyes	Emmetropia in the untreated eyes	Low hyperopia in the untreated eyes	p-value	Low myopia	Emmetropia	Low hyperopia	<i>p</i> -value	
Number of eyes (N)	148	148		37	76	35		37	76	35		
SER at baseline (D)	-2.50 (-1.75, -3.00)	0.00 (+0.38,-0.41)	< 0.001	-2.70 ± 0.94	-2.41 ± 0.95	-2.46 ± 0.78	0.29	-0.75 (-0.56, -0.88)	0.00 (0.00, -0.25)	+0.75 (+1.00,+0.62)	<0.001	
AL at baseline (mm)	24.67 ± 0.73	23.58 ± 0.67	< 0.001	24.76 ± 0.76	24.65 ± 0.76	24.62 ± 0.66	0.70	23.90 ± 0.62	23.57± 0.67	23.28 ± 0.59	<0.001	
AL at 1 year (mm)	24.81 ± 0.72	23.97 ± 0.73	< 0.001	24.89 ± 0.78	24.80 ± 0.75	24.75 ± 0.63	0.69	24.34 ± 0.72	$23.92{\pm}0.71$	23.69 ± 0.65	< 0.001	
Change in AL at 1 year (mm)	0.14 ± 0.18	0.39 ± 0.27	< 0.001	0.14 ± 0.18	0.15 ± 0.17	0.13 ± 0.21	0.92	0.44 ± 0.25	0.35 ± 0.24	0.41 ± 0.33	0.11	
AL difference (treated minus untreated eyes at baseline)#	1.09 ± 0.45			0.86 ± 0.39	1.08 ± 0.45	1.35 ± 0.38	<0.001					
AL difference (treated minus untreated eyes at 1 year)%	0.84 ± 0.52			0.55 ± 0.47	0.88 ± 0.50	1.06 ± 0.51	<0.001					
Reduction of AL difference (# minus*)	0.25 ± 0.30			0.30 ± 0.25	0.20 ± 0.28	0.28 ± 0.37	0.17					

SER, spherical equivalent refraction; AL, axial length. *.* The symbol for the horizontal column title. This makes the bottom column of the table more concise when calculating the difference between the two variables.

	Subgroups	Non-adjusted model	ed model	Sex and age-adjusted model	ijusted model	Sex, age, corneal astigmatism, and anisometropia value-adjusted model	cigmatism, and adjusted model
		eta, 95% Cl	<i>p</i> -value	eta, 95% CI	<i>p</i> -value	eta, 95% CI	<i>p</i> -value
Treated eyes	Emmetropia in the untreated eyes	0		0			
	Low myopia in the untreated eyes	$-0.02 \ (-0.10, 0.05)$	0.54	$-0.03 \ (-0.10, 0.04)$	0.44	$-0.01 \ (-0.08, 0.07)$	0.86
	Low hyperopia in the untreated eyes	-0.01 (-0.09, 0.06)	0.72	$-0.03 \left(-0.10, 0.03\right)$	0.33	$-0.04 \left(-0.11, 0.03\right)$	0.25
Untreated eyes	Emmetropia	0		0		0	
	Low myopia	0.09 (-0.02, 0.19)	0.10	0.08 (0.01, 0.15)	0.04	0.09 (0.01, 0.17)	0.04
	Low hyperopia	0.06 (-0.04, 0.17)	0.25	$0.04 \left(-0.06, 0.14\right)$	0.41	0.06 (-0.04, 0.16)	0.22
SER, spherical equivalent refraction.	refraction.						

TABLE 3 Multivariate linear regression analyses of change in axial length in the three subgroups based on the SER of untreated eyes

of the OK lens was increased myopic defocus in the peripheral retina, thereby decreasing the tendency of axial elongation (23). Although these studies and the present study reported that AL differences between the uncorrected contralateral eyes and the eyes wearing the OK lenses decreased in children with UMA, reduction in the binocular AL differences from baseline to 1 year after treatment were slightly different among various studies [0.15 mm (11), 0.31 mm (12), 0.29 mm (13), 0.19 mm (14), 0.36 mm (15), and 0.25 mm in the present study]. These differences in the values may be attributed to the differences in study populations, baseline characteristics, anisometropia values, genetic factors such as the number of myopic parents (24), and other possible factors. This study observed that younger children with low anisometropia values in the low myopia subgroup exhibited a greater reduction in the binocular AL differences.

In the present study, the refractive state (low myopia, emmetropia, and low hyperopia) of the untreated contralateral eyes did not affect the change in AL of the more myopic eyes wearing the OK lens. Similar results have not been reported in any of the previous studies. Clinically, regardless of the refractive state of the untreated contralateral eye (low myopia, emmetropia, or low hyperopia), wearing the OK lens was a good option after the diagnosis of unilateral myopia since it could provide optical correction and also inhibit axial elongation in the myopic eyes, resulting in reduced AL difference between the two eyes.

The current study observed that AL in low-myopic eyes grew faster than emmetropic eyes in untreated eyes. The difference in axial elongation among the untreated contralateral eyes at different refractive states when the myopic eye was wearing the OK lens may relate to several factors. First, it is not related to the OK lens but may relate to the change in peripheral refraction (myopic refraction) induced by midperipheral corneal steeping induced by OK wear (23). Long et al. (13) reported no significant difference in axial elongation between emmetropic and low hyperopic untreated contralateral eyes regardless of whether the myopic eye had OK lenses or spectacles. However, there was a slight difference in the SER range in untreated contralateral eyes between the study by Long et al. (13) (+1.50 D to -0.50 D) and the present study (+1.88 D to -1.13 D). Second, the differences in peripheral refraction (between myopic, emmetropic, or hyperopic eyes of the untreated contralateral eyes) could be related to AL change (25-27). Third, there may be a sympathetic effect between the eyes of the same patient, as seen with other eye conditions. Finally, among the 37 eyes in the low myopia subgroup of the untreated eyes (SER: -0.50 D to -1.13 D), 13 eyes exhibited a sphere diopter of more than -0.50 D, and 19 cases exhibited a cylinder diopter of more than -1.00 D. The uncorrected VA of the untreated eyes was higher than 0 (LogMAR charts). However, these children and their parents or guardians were reluctant to receive any form of intervention and gave consent only for the OK lens in the more myopic eye. Thus, the untreated eyes were in a state of uncorrected myopia or minus astigmatism during the follow-up period. Uncorrected myopia (28, 29) and astigmatism (30, 31) have been shown to hasten the progression of myopia. Therefore, for children with UMA, methods of correcting the eyes with lower myopia such as wearing the OK lens should be adopted to slow down axial elongation and balance the binocular VA when

TABLE 4 Change in AL of both eyes in anisometropia children who wore OK lenses in the more myopic eyes in different studies.
--

References	Number	Follow-up time (year)	Age (year)	OK lens g	group (treated	d eyes)	Control gr	oup (untreate	Baseline SER difference	Reduction of AL difference (mm/year)	
				Baseline SER (D)	Baseline AL (mm)	AL elongation rate (mm/year)	Baseline SER (D)	Baseline AL (mm)	AL elongation rate (mm/year)		
Chen et al. (12)	25	1.93	11.2 ± 1.9 (9-15)	-2.32 ± 0.94 (-1.00 to -4.00)	24.56 ± 0.72	0.08 ± 0.15	$+0.24 \pm 0.50$ (+1.75 to -0.50)	23.47 ± 0.65	0.39 ± 0.32	2.56	0.31
Long et al. (13)	79	1.0	11.46 ± 1.76 (8-14)	-2.60 ± 1.14	24.79 ± 0.90	0.05 ± 0.19	$+0.10 \pm 0.42$ (+1.50 to -0.50)	23.64 ± 0.78	0.34 ± 0.21	2.70	0.29
Fu et al. (14)	27	1.0	12.41 ± 2.32 (8-18)	-2.83 ± 1.37 (-1.25 to -6.50)	24.68 ± 0.70	0.11 ± 0.19	-0.07 ± 0.35 (+0.50 to -0.50)	23.51 ± 0.72	0.30 ± 0.28	2.76	0.19
Tsai et al. (11)	31	2.01	12.32 ± 3.07 (>8)	-2.73 ± 0.95	24.15 ± 1.13	-	-0.37 ± 0.50 (+0.50 to -0.50)	23.32 ± 1.04	-	2.36	0.15 (1st year) 0.09 (2nd year)
Xu et al. (15)	47	1.28	12.35 ± 2.37 (8-16)	-2.31 ± 1.16 (-1.00 to -6.00)	24.54±0.94	About 0.11	$0.15 \pm 0.49 (+1.50)$ to -0.50	23.46±0.74	About 0.35	2.46	0.36 (1st year) 0.14 (2nd year)
Current study	1										
All	148	1.18	11.5 ± 1.9 (8-14)	-2.50 ± 0.91 (-1.00 to -4.38)	24.67 ± 0.73	0.14 ± 0.18	-0.05 ± 0.65 (+1.88 to -1.13)	23.58 ± 0.67	0.39 ± 0.27	2.45	0.25
Low myopia	37	1.11	11.1 ± 2.2 (8-14)	-2.70 ± 0.94 (-1.00 to -4.38)	24.76 ± 0.76	0.14 ± 0.18	-0.80 ± 0.20 (-0.50 to -1.13)	23.90 ± 0.62	0.44 ± 0.25	1.90	0.30
Emmetropia	76	1.22	11.6 ± 1.9 (8-14)	-2.41 ± 0.95 (-1.00 to -4.25)	24.65 ± 0.76	0.15 ± 0.17	-0.05 ± 0.20 (+0.38 to -0.38)	23.57 ± 0.67	0.35 ± 0.24	2.36	0.20
Low hyperopia	35	1.02	11.6 ± 1.8 (8-14)	-2.46 ± 0.78 (-1.00 to -4.10)	24.62 ± 0.66	0.13 ± 0.21	$0.88 \pm 0.37 (+1.88$ to +0.50)	23.28 ± 0.59	0.41 ± 0.33	3.34	0.28

SER, spherical equivalent refraction; AL, axial length.

the OK lens is used in the more myopic eye. Chen et al. (12) observed that in children with UMA, the OK lens could also slow down axial elongation in the initially non-myopic eye that developed myopia subsequently. Therefore, further prospective studies should include peripheral refraction assessment (at baseline, with and without OK lens wear, and after treatment) between other clinical variables.

The present study has some limitations. The sample sizes of the three subgroups were unequal, with more cases in the emmetropia subgroup than in the low hyperopia and low myopia subgroups. However, all patients who met the inclusion criteria at the hospital were included in the study. Moreover, the current study evaluated the effect of the uncorrected eye in monocular OK lens wearing. The uncorrected eyes usually have a low degree of refractive error which may result in small differences of refractive error in the three subgroups and play little impact on axial elongation or change in interocular AL differences. Other limitations were the lack of a spectacle-wearing control group, not collecting genetic factors (number of myopic parents) from the children of participants, not measuring and analyzing the differences in peripheral refraction, the retrospective nature of the study, and the short follow-up time. Further prospective investigations with long-term followup and studies including control group design and peripheral refraction analysis should be conducted to confirm the findings of the present study.

In conclusion, among children with myopic anisometropia aged 8-14 years, wearing OK lenses in the more myopic eyes without any intervention in the contralateral eyes (SER: +2.00 to -1.25 D) could effectively slow down axial elongation in more myopic eyes, thereby reducing the interocular AL difference. The refractive state (low myopia, emmetropia, or low hyperopia) of the untreated eyes did not affect the axial elongation of more myopic eyes wearing the OK lens. In the untreated eyes, AL increased faster in the low myopia subgroup than in the emmetropia subgroup. Clinically, one of the reasonable treatment methods for children with UMA is wearing the OK lenses in the more myopic eyes and a regular follow-up of the untreated eyes if they have emmetropia or very low hyperopia. Appropriate treatment measures employed when the untreated eyes become myopic can balance the VA in both eyes, control axial elongation, and reduce the interocular AL difference. These could reduce the incidence of strabismus and amblyopia and could better establish binocular visual function.

Data availability statement

The raw data supporting the conclusions of this article will be made available by the authors, without undue reservation.

References

Ethics statement

The studies involving humans were approved by Medical Ethics Committee of the First Affiliated Hospital of Zhengzhou University. The studies were conducted in accordance with the local legislation and institutional requirements. Written informed consent for participation in this study was provided by the participants' legal guardians/next of kin.

Author contributions

JQ: Conceptualization, Data curation, Writing—review & editing, Investigation, Project administration, Writing original draft. HQ: Investigation, Writing—review & editing, Data curation, Supervision. NJ: Writing—review & editing. TL: Writing—review & editing, Data curation. HM: Data curation, Writing—review & editing. MS: Data curation, Writing—review & editing. SY: Data curation, Writing—review & editing. CM: Data curation, Writing—review & editing, Formal analysis. AF: Data curation, Formal analysis, Writing—review & editing, Conceptualization, Methodology.

Funding

The author(s) declare financial support was received for the research, authorship, and/or publication of this article. This study was supported by the Medical Science and Technology Research Project of Henan Health Commission (LHGJ20200065), Suzhou Vocational Health College (szwzy201905), and Key Scientific Research Project of Universities of Henan Education Department (22A320024).

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

3. Barrett BT, Bradley A, Candy TR. The relationship between anisometropia and amblyopia. *Prog Retin Eye Res.* (2013) 36:120–58. doi: 10.1016/j.preteyeres.2013.05.001

^{1.} O'Donoghue L, McClelland JF, Logan NS, Rudnicka AR, Owen CG, Saunders KJ. Profile of anisometropia and aniso-astigmatism in children: prevalence and association with age, ocular biometric measures, and refractive state. *Invest Ophthalmol Vis Sci.* (2013) 54:602–8. doi: 10.1167/iovs.12-11066

^{2.} Gawecki M. Threshold values of myopic anisometropia causing loss of stereopsis. *J Ophthalmol.* (2019) 2019:1–8. doi: 10.1155/2019/2654170

4. Weakley DR Jr. The association between nonstrabismic anisometropia, amblyopia, and subnormal binocularity. *Ophthalmology.* (2001) 108:163–71. doi: 10.1016/S0161-6420(00)00425-5

5. Wang HY, Zhao KX. The effects of myopic anisometropia on binocular vision function. *Chin J Exp Ophthalmol.* (2013) 31:559–63. doi: 10.3760/cma.j.issn.2095-0160.2013.06.009

6. Karimian F, Ownagh V, Amiri MA, Tabatabaee SM, Dadbin N. Stereoacuity after wavefront- guided photorefractive keratectomy in anisometropia. *J Ophthalmic Vis Res.* (2017) 12:265–9. doi: 10.4103/jovr.jovr_138_16

7. Katz J, Schein OD, Levy B, Cruiscullo T, Saw SM, Rajan U, et al. A randomized trial of rigid gas permeable contact lenses to reduce progression of children's myopia. *Am J Ophthalmol.* (2003) 136:82–90. doi: 10.1016/S0002-9394(03) 00106-5

8. Walline JJ, Jones LA, Sinnott L, Manny RE, Gaume A, Rah MJ, et al. A randomized trial of the effect of soft contact lenses on myopia progression in children. *Invest Ophthalmol Vis Sci.* (2008) 49:4702–6. doi: 10.1167/iovs.08-2067

9. Horner DG, Soni PS, Salmon TO, Swartz TS. Myopia progression in adolescent wearers of soft contact lenses and spectacles. *Optom Vis Sci.* (1999) 76:474–9. doi: 10.1097/00006324-199907000-00023

10. Kakita T, Hiraoka T, Oshika T. Influence of overnight orthokeratology on axial elongation in childhood myopia. *Invest Ophthalmol Vis Sci.* (2011) 52:2170-4. doi: 10.1167/iovs.10-5485

11. Tsai WS, Wang JH, Lee YC, Chiu CJ. Assessing the change of anisometropia in unilateral myopic children receiving monocular orthokeratology treatment. *J Formos Med Assoc.* (2019) 118:1122–8. doi: 10.1016/j.jfma.2019.02.001

12. Chen Z, Zhou J, Qu X, Zhou X, Xue F. Effects of orthokeratology on axial length growth in myopic anisometropes. *Cont Lens Anterior Eye.* (2018) 41:263–6. doi: 10.1016/j.clae.2017.10.014

13. Long W, Li Z, Hu Y, Cui D, Zhai Z, Yang X. Pattern of axial length growth in children myopic anisometropes with orthokeratology treatment. *Curr Eye Res.* (2020) 45:834–8. doi: 10.1080/02713683.2019.1701685

14. Fu AC, Qin J, Rong JB, Ji N, Wang WQ, Zhao BX, et al. Effects of orthokeratology lens on axial length elongation in unilateral myopia and bilateral myopia with anisometropia children. *Cont Lens Anterior Eye.* (2020) 43:73–7. doi: 10.1016/j.clae.2019.12.001

15. Xu J, Gao B, Tian Q, Wu Q, Zhang X, Lin X, et al. Effects oforthokeratology on axial length elongation in anisometropes. *Ophthalmic Res.* (2021) 64:991-1001. doi: 10.1159/000516907

16. Zhang Y, Chen Y. Effect of orthokeratology on axial length elongation in anisomyopic children. *Optom Vis Sci.* (2019) 96:43–7. doi: 10.1097/OPX.00000000001315

17. Zhong Y, Ke L, Qiong W, Liu F. Orthokeratology lens for management of myopia in anisometropic children: a contralateral study. *Cont Lens Anterior Eye.* (2020) 43:40–3. doi: 10.1016/j.clae.2019.03.003

18. Tsai WS, Wang JH, Chiu CJ. A comparative study of orthokeratology and low-dose atropine for the treatment of anisomyopia in children. *Sci Rep.* (2020) 10:14176. doi: 10.1038/s41598-020-71142-3

19. Tsai HR, Wang JH, Chiu CJ. Effect of orthokeratology on anisometropia control: a meta -analysis. *Meta-Analysis J Formos Med Assoc.* (2021) 120:2120-7. doi: 10.1016/j.jfma.2021.05.024

20. French AN, Morgan IG, Burlutsky G, Mitchell P, Rose KA. Prevalence and 5- to 6-year incidence and progression of myopia and hyperopia in Australian schoolchildren. *Ophthalmology*. (2013) 120:1482–91. doi: 10.1016/j.ophtha.2012.12.018

21. Galvis V, Tello A, Otero J, Serrano AA, Gómez LM, Camacho PA, et al. Prevalence of refractive errors in Colombia: MIOPUR study. *Br J Ophthalmol.* (2018) 102:1320–3. doi: 10.1136/bjophthalmol-2018-312149

22. Li T, Zuo X, Zhang T, Liu L, Wang Z, Han L, et al. Patients with intermittent exotropia and exophoria exhibit non-aggravated lens decentration after orthokeratology application: the Nanjing Strabismus Cohort. *Ophthalmol Ther.* (2023) 12:1535–45. doi: 10.1007/s40123-023-00685-1

23. Li X, Friedman IB, Medow NB, Zhang C. Update on orthokeratology in managing progressive myopia in children: efficacy, mechanisms, and concerns. J Pediatr Ophthalmol Strabismus. (2017) 54:142–8. doi: 10.3928/01913913-20170106-01

24. Xiang F, He M, Morgan IG. The impact of parental myopia on myopia in Chinese children: population-based evidence. *Optom Vis Sci.* (2012) 89:1487– 96. doi: 10.1097/OPX.0b013e31826912e0

25. Sng CC, Lin XY, Gazzard G, Chang B, Dirani M, Chia A, et al. Peripheral refraction and refractive error in Singapore Chinese children. *Invest Ophthalmol Vis Sci.* (2011) 52:1181–90. doi: 10.1167/iovs.10-5601

26. Mutti DO, Hayes JR, Mitchell GL, Jones LA, Moeschberger ML, Cotter SA, et al. Refractive error, axial length, and relative peripheral refractive error before and after the onset of myopia. *Invest Ophthalmol Vis Sci.* (2007) 48:2510–9. doi: 10.1167/iovs.06-0562

27. Chen X, Sankaridurg P, Donovan L, Lin Z, Li L, Martinez A, et al. Characteristics of peripheral refractive errors of myopic and non-myopic Chinese eyes. *Vision Res.* (2010) 50:31–5. doi: 10.1016/j.visres.2009.10.004

28. Chung K, Mohidin N, O'Leary DJ. Under-correction of myopia enhances rather than inhibits myopia progression. *Vision Res.* (2002) 42:2555–9. doi: 10.1016/S0042-6989(02)00258-4

29. Adler D, Millodot M. The possible effect of under-correction on myopic progression in children. *Clin Exp Optom.* (2006) 89:315–21. doi: 10.1111/j.1444-0938.2006.00055.x

30. Tong L, Saw SM, Lin Y, Chia KS, Koh D, Tan D. Incidence, and progression of astigmatism in Singaporean children. *Invest Ophthalmol Vis Sci.* (2004) 45:3914–8. doi: 10.1167/iovs.04-0492

31. Fan DS, Rao SK, Cheung EY, Islam M, Chew S, Lam DS. Astigmatism in Chinese preschool children: prevalence, change, and effect on refractive development. *Br J Ophthalmol.* (2004) 88:938–41. doi: 10.1136/bjo.2003.030338