Orthokeratology with increased compression factor in adolescent myopia control: a 2-year prospective randomized clinical trial

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Received: 2022-09-12 Accepted: 2023-03-10

Abstract

• **AIM:** To explore the long-term efficacy, safety, and optical mechanism of orthokeratology with increased compression factor in adolescent myopia control.

• METHODS: A prospective, double-masked, and randomized clinical trial was performed from May 2016 to June 2020. Subjects aged between 8 and 16y, with myopia (-5.00 to -1.00 D), low astigmatism (\geq -1.50 D) and anisometropia (≤1.00 D), were stratified into low (-2.75 to -1.00 D) and moderate (-5.00 to -3.00 D) myopia groups. Then they were randomly assigned to wear either increased compression factor (ICF; 1.75 D) orthokeratology or conventional compression factor (CCF; 0.75 D) orthokeratology. The data were recorded including axial length (AL), spherical equivalent (SE), best corrected visual acuity (BCVA), near visual acuity (NVA), corneal staining (using Efron grading scales), corneal hysteresis (CH), corneal resistance factor (CRF), higher-order aberrations (HOAs, expressed as root mean square, RMS_h), and subfoveal choroidal thickness (SFChT) in the 2-year followup period. Pearson's correlation coefficient was conducted to analyze the association between the changes in AL and RMS_h, SFChT.

• **RESULTS:** At the 2-year visit, there were no statistical differences in all the parameters between the ICF group and the CCF group in low myopia subjects (*P*>0.05). For

the moderate myopia subjects, the ICF group had shorter AL elongation (0.23 \pm 0.08 vs 0.30 \pm 0.11 mm, *P*=0.015), higher RMS_h (1.94 \pm 0.50 vs 1.65 \pm 0.51 µm, *P*=0.041), and higher SFChT (279.04 \pm 35.72 vs 254.08 \pm 29.60 µm, *P*=0.008) than those in CCF group. The change in AL was negatively correlated with RMS_h (*r*=-0.687, *P*<0.001) and SFChT (*r*=-0.464, *P*=0.013).

• **CONCLUSION:** ICF orthokeratology can control the progression of moderate myopia more effectively, which might be related to greater RMS_h and SFChT.

• **KEYWORDS:** orthokeratology; compression factor; myopia; axial length; higher-order aberrations; choroidal thickness

DOI:10.18240/ijo.2023.05.15

Citation: Tang WT, Zhang L, Zhang HD, Li SB, Liang H. Orthokeratology with increased compression factor in adolescent myopia control: a 2-year prospective randomized clinical trial. *Int J Ophthalmol* 2023;16(5):770-777

INTRODUCTION

O rthokeratology (ortho-k) is a popular and non-surgical method that can suppress the progression of myopia effectively in adolescents^[1-2]. Our previous study confirmed that wearing ortho-k for one year slowed down the axial elongation by 48.28%, compared with spectacles^[3]. However, the overnight ortho-k effect is temporary, and daytime refractive regression after lens removal has been reported frequently, especially in patients with moderate to high myopia. Refractive regression can be compensated by the compression factor (The extra power added onto the basic refractive error, also known as the Jessen factor)^[4].

Currently, many ortho-k lens manufacturers utilize conventional compression factor (CCF; 0.75 D). But there was still 0.25 D under-correction 4h after lens removal, and 0.50-0.75 D under-correction 8-10h after lens removal^[5-6]. Therefore, the conventional compression factor may not achieve the expected correction target. Chan *et al*^[7] analyzed 123 retrospective subjects and recommended that an extra 1.00 D should be

added to maintain satisfactory unaided vision. Several shortterm (4-7wk) studies^[8-10] have shown that the increased compression factor (ICF; 1.75 D) did not affect lens fitting, or ocular health, but did correct refractive error more rapidly. However, the lack of long-term and comprehensive studies made the conclusion and mechanism of myopia control equivocal. Short-term changes in higher-order aberrations (HOAs) and choroidal thickness are expected to be the efficacy predictors of ortho-k. Sun et al^[11] revealed that greater HOAs may relate to effective myopia control. Chen et al^[12] tried to alter the HOAs profile by modifying the lens design, but to date, no research has examined the effect of HOAs on the changes in axial length (AL) systematically. Changes in choroidal thickness after wearing ortho-k remain controversial^[13]. However, some of these studies did not consider the diurnal variation, age, and spherical equivalent (SE), which could affect choroidal thickness^[14].

Hence, this 2-year prospective, double-masked, and randomized study assessed the efficacy by AL, SE, best corrected visual acuity (BCVA), and assessed the safety by near visual acuity (NVA), corneal staining, corneal hysteresis (CH), corneal resistance factor (CRF), and explored the correlation of changes in AL with HOAs and subfoveal choroidal thickness (SFChT).

SUBJECTS AND METHODS

Ethical Approval This study was in accordance with the ethical standards formulated in the Helsinki Declaration and was approved by the Medical Ethics Committee of the First Affiliated Hospital of Chengdu Medical College (2020CYFYHEC-BA-30). All subjects and their parents had been told the complications and disadvantages of wearing ICF and CCF ortho-k in detail. Also, they were informed of the trial protocol, objectives, and procedures, then signed a written informed consent before commencement.

Subjects This randomized, double-blinded clinical trial enrolled 153 children who visited the First Affiliated Hospital of Chengdu Medical College for refraction correction from May 2016 to June 2020. The participants should conform to the criteria in Table 1.

Allocations and Treatments The subjects were divided into 2 groups according to the severity of myopia: low myopia group (-2.75 to -1.0 D) and moderate myopia group (-5.00 to -3.00 D). Then they were randomly (using a Microsoft Excel random number, and the numbers were sealed in opaque envelopes) divided into the experimental group and the control group. Patients and the examiner were both blinded to group assignments. We prepared a disclosure envelope for each randomization number in case emergency unblinding becomes necessary, such as serious adverse events (*e.g.*, infiltrates, pannus, microcysts, microbial keratitis). The experimental

Table 1 Inclusion and exclusion criteria

Inclusion criteria	Exclusion criteria						
8 to 16 years old (inclusive)	An experience of myopia control						
Myopia: -5.00 to -1.00 D	Ocular or systemic disease						
Anisometropia: ≤1.00 D	Contraindications for ortho-k lens						
Astigmatism: ≥-1.50 D	Poor compliance						
BCVA and NVA (logMAR) equal to or better than 0.00							

BCVA: Best corrected visual acuity; NVA: Near visual acuity.

group wore ICF ortho-k, and the control group wore CCF ortho-k. All subjects were fitted with the Mouldway ortho-k lenses (Autek China Inc., Sino-US Joint Venture) composed of Boston XO. The initial lens parameters including compression factor were inputted into the computer software provided by the manufacturer. All Subjects should perform an ideal lens fitting and they were instructed to wear ortho-k lenses every night for at least 8h.

Measurements All subjects should attend baseline and subsequent visits: 1d, 1wk, 1mo, 3mo, 6mo, 1y, 1.5y, and 2y after commencing lens wear. The aftercare visits were conducted from 8:00 to 10:00 *a.m.* and within 2h after lens removal. At the 1-year and 2-year visits, the subjects ceased lens wear for 3wk. Cycloplegic SE examination (using Tropiamide phenylephrine) was performed after the cessation period. Unscheduled consultations were provided if necessary to ensure the safety of lens wear.

AL was measured by IOLMaster 500 (Carl Zeiss, Germany). Five mean values were recorded if each difference <0.02 mm and SNR>2.0. SE was measured by open-field WAM-5500 (Grand Seiko, Japan). The first three values with a difference ≤0.25 D were recorded. ETDRS charts 2000 (Precision Vision, USA) was used to measure low-contrast (10%) best corrected visual acuity (LC BCVA) and high-contrast (>90%) best corrected visual acuity (HC BCVA) successively. NVA was measured by Jaeger chart. Corneal staining was graded by the Efron grading system. CH and CRF were measured by Ocular Response Analyzer (ORA, Reichert Corporation, USA). Three records with each score \geq 5.0 were obtained. Ocular HOAs were measured by iTrace aberration analyzer (Tracey, USA) through the natural pupil in a dark room. The pupil diameter for analysis was 6 mm. The total HOAs were calculated by Zernike polynomial as the root mean square (RMS_b). SFChT was determined by Spectralis OCT (Heidelberg Engineering, Germany) with the linear scan pattern (average of the vertical and horizontal line) through the enhanced depth imaging (EDI) mode. Automatic registration aided to capture repeat scans, and follow-up mode Spectralis helped to ensure the same measured location at each visit.

Sample Size According to the randomized tests for two means (PASS11.0) and based on the previous research^[15], it



Figure 1 Study flowchart L-ICF: Low myopia subjects fitted with increased compression factor (1.75 D) ortho-k; L-CCF: Low myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with increased compression factor (1.75 D) ortho-k; M-CCF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-CCF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with increased compression factor (1.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k.

was estimated to induce at least a 0.17 mm difference in AL between the two groups over 2y, with a standard deviation (SD) of 0.21 mm. A sample size of 20 subjects in each group was required to achieve a power $(1-\beta)$ of 80% and a significance level (α) of 0.05. Assuming 30% dropouts, a minimum total sample size of 116 subjects (low myopia 29:29, moderate myopia 29:29) was allocated.

Statistical Analysis IBM SPSS Statistics 22.0 was applied for data analysis, and the data from right eyes were analyzed. Chi-squared test (or continuity-corrected Chi-square test, as appropriate) was used to compare the incidence of corneal staining and subjective discomforts between the two groups. The normality of the data was evaluated by Shapiro-Wilk tests. In the case of a normal distribution or approximate normal distribution, data were presented as mean±SD. The variance homogeneity was assessed by Levene's test. Then independent sample *t*-test was performed to assess the significant differences between the two groups. Pearson's correlation coefficient was conducted to calculate the correlation between the ocular biometrics (RMS_h, SFChT) and AL change. A *P* value <0.05 was considered statistically significant.

RESULTS

A total of 153 subjects were eligible to enroll in this research, of whom 131 subjects underwent fitting with the ortho-k lenses and proceeded to baseline data collection. A total of 59 low myopia subjects were randomly assigned to either ICF (L-ICF group, n=30) or CCF (L-CCF group, n=29) group, and 72 moderate myopia subjects were randomly assigned to either ICF (M-ICF group, n=37) or CCF (M-CCF group, n=35) group. Finally, a total of 97 (22 subjects in L-ICF group, 22 subjects in L-CCF group, 28 subjects in M-ICF group, and 25 subjects in M-CCF group) completed the 2-year follow-up (Figure 1).

Table 2 demonstrates the baseline biometric parameters of the analyzed subjects who completed the 2-year study. There were no statistical differences in baseline parameters between the ICF group and CCF group (all P>0.05). No serious adverse event was observed in the study period.

Changes in Axial Length The AL elongation in the M-ICF group was statistically slower than that in the M-CCF group. Over 2y, the AL elongation was 0.23 ± 0.08 mm in the M-ICF group and 0.30 ± 0.11 mm in the M-CCF group (*P*=0.015). The ICF group was 23.33% more effective in slowing moderate myopia AL elongation than the CCF group. No significant differences between the L-ICF group and the L-CCF group were observed in terms of AL elongation (Table 3).

Changes in Spherical Equivalent The increases of SE (measured after lens cessation for 3wk) in the M-ICF group were lower than that in the M-CCF group (1-year cessation for 3wk: P=0.008, 2-year cessation for 3wk: P=0.023). However, there were no significant differences observed between the L-ICF group and the L-CCF group at all the yearly washout visits (Table 4).

Int J Ophthalmol, Vol. 16, No. 5, May 18, 2023 www.ijo.cn Tel: 8629-82245172 8629-82210956 Email: ijopress@163.com

Table 2 Baseline data mea									
Parameters	L-ICF group (n=22)	L-CCF group (n=22)	t	Р	M-ICF group (n=28)	M-CCF group (n=25)	t	Р	
Age (y)	11.32±1.78	10.50±1.65	1.578	0.122	12.04±2.36	12.52±2.16	-0.775	0.442	
BCVA (logMAR)	-0.05±0.04	-0.04±0.03	-0.654	0.517	-0.05±0.05	-0.05±0.04	-0.412	0.682	
SE (D)	-2.08±0.53	-1.89±0.51	-1.147	0.258	-4.15±0.74	-4.33±0.70	0.900	0.372	
AL (mm)	23.97±0.63	23.95±0.61	0.126	0.901	24.80±0.82	25.02±0.96	-0.869	0.389	
RMS _h (µm)	0.66±0.31	0.57±0.42	0.801	0.427	0.75±0.31	0.82±0.37	-0.768	0.446	
SFChT (µm)	263.05±23.59	259.00±16.15	0.664	0.511	229.07±27.54	224.16±25.59	0.670	0.506	

L-ICF: Low myopia subjects fitted with increased compression factor (1.75 D) ortho-k; L-CCF: Low myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with increased compression factor (1.75 D) ortho-k; M-CCF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; BCVA: Best corrected visual acuity; SE: Spherical equivalent; AL: Axial length; RMS_h: Higher-order aberrations expressed as root mean square; SFChT: Subfoveal choroidal thickness.

Table 3 Chang	es in axial length in d	ifferent groups					mean	±SD, mm
Follow-ups	L-ICF group (<i>n</i> =22)	L-CCF group (<i>n</i> =22)	t	Р	M-ICF group (<i>n</i> =28)	M-CCF group (<i>n</i> =25)	t	Р
6mo	0.10±0.05	0.09±0.04	0.183	0.855	0.05±0.03	0.08±0.03	-2.218	0.031
1y	0.17±0.06	0.18±0.06	-0.126	0.900	0.12±0.05	0.17±0.05	-3.253	0.002
1.5y	0.24±0.08	0.27±0.10	-0.916	0.365	0.20±0.08	0.25±0.08	-2.364	0.022
2у	0.28±0.10	0.31±0.10	-0.815	0.420	0.23±0.08	0.30±0.11	-2.524	0.015

L-ICF: Low myopia subjects fitted with increased compression factor (1.75 D) ortho-k; L-CCF: Low myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with increased compression factor (1.75 D) ortho-k; M-CCF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k.

Table 4 Chang	es in spherical equival	lent after lens cessation	on for 3w	k in differ	ent groups		me	an±SD, D
Follow-ups	L-ICF group (n=22)	L-CCF group (<i>n</i> =22)	t	Р	M-ICF group (<i>n</i> =28)	M-CCF group (n=25)	t	Р
1y	-0.36±0.28	-0.43±0.33	0.744	0.461	-0.22±0.20	-0.42±0.29	2.794	0.008
2у	-0.55±0.36	-0.63±0.45	0.662	0.512	-0.35±0.28	-0.57±0.40	2.355	0.023

L-ICF: Low myopia subjects fitted with increased compression factor (1.75 D) ortho-k; L-CCF: Low myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with increased compression factor (1.75 D) ortho-k; M-CCF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k.

Changes in Root Mean Square Figure 2 represents changes and differences in RMS_h values between the ICF group and CCF group during the study course. At the 3-month visit, RMS_h differed significantly between the M-ICF group (1.96±0.49 µm) and the M-CCF group (1.61±0.46 µm, *P*=0.011). After 3mo of lens wear, M-ICF showed a significantly greater increase in RMS_h (all *P*<0.05). No significant differences between the L-ICF group and the L-CCF group were observed in RMS_h values.

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Changes in Subfoveal Choroidal Thickness Changes and differences in SFChT for CCF and ICF groups are shown in Figure 3. SFChT in the M-ICF group was significantly thicker than that in the M-CCF group from the 6-month visit onward (P<0.05), except for the follow-up at 1.5y (P=0.086). However, there were no significant differences observed in terms of SFChT between the L-ICF group and the L-CCF group at all visits.

Correlation Analysis for Root Mean Square and Subfoveal Choroidal Thickness Since AL elongation differed statistically only between the M-ICF group and M-CCF group, and was shorter in the M-ICF group, correlation analysis



Figure 2 the changes and differences in RMS_h L-ICF: Low myopia subjects fitted with increased compression factor (1.75 D) ortho-k; L-CCF: Low myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with increased compression factor (1.75 D) ortho-k; M-CCF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; RMS_h: Higher-order aberrations expressed as root mean square. ^a*P*<0.05, ICF group *vs* CCF group by independent sample *t*-test. Error bars represent standard deviation.

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Table 5 Comparisons of DCVA, NVA, CH, CKF, Comean staining, and subjective disconnorts over	Table	5 Compa	arisons of I	BCVA, NV	Ά, CH,	CRF,	corneal staining	, and sub	jective (discomforts	over
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Parameters L-ICF group (n=22	L-CCF group (<i>n</i> =22) -0.04±0.06	t/χ^2 -0.028	P 0.978	M-ICF group (<i>n</i> =28)	M-CCF group (<i>n</i> =25)	t/χ^2	Р
	-0.04±0.06	-0.028	0.978				
HC BCVA (IOgIVIAR) -0.05±0.05	0 20+0 14		0.070	-0.06±0.05	-0.05±0.05	-0.800	0.427
LC BCVA (logMAR) 0.24±0.13	0.20 ± 0.14	1.124	0.268	0.23±0.19	0.31±0.19	-1.729	0.088
NVA (logMAR) 0.03±0.02	0.03±0.03	-0.437	0.664	0.06±0.03	0.04±0.05	1.602	0.118
CH (mm Hg) 11.15±1.72	10.86±1.80	0.544	0.590	10.94±1.69	10.91±1.71	0.048	0.962
CRF (mm Hg) 11.04±2.04	11.22±1.79	-0.319	0.751	10.82±1.68	10.79±1.69	0.056	0.956
Corneal staining (G1, %) 18.18	13.64	0.000	>0.999	21.43	20.00	0.016	0.898
Corneal staining (G2, %) 9.09	18.18	0.193	0.660	14.29	16.00	0.000	>0.999
Glare, ghosting (%) 27.27	18.18	0.518	0.472	17.86	12.00	0.044	0.833
Visual fluctuation (%) 4.55	13.64	0.275	0.600	3.57	8.00	0.010	0.919

L-ICF: Low myopia subjects fitted with increased compression factor (1.75 D) ortho-k; L-CCF: Low myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with increased compression factor (1.75 D) ortho-k; M-CCF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; LC BCVA: Low-contrast best corrected visual acuity; HC BCVA: High-contrast best corrected visual acuity; NVA: Near visual acuity; CH: Corneal hysteresis; CRF: Corneal resistance factor.



Figure 3 The changes and differences in SFChT L-ICF: Low myopia subjects fitted with increased compression factor (1.75 D) ortho-k; L-CCF: Low myopia subjects fitted with conventional compression factor (0.75 D) ortho-k; M-ICF: Moderate myopia subjects fitted with increased compression factor (1.75 D) ortho-k; M-CCF: Moderate myopia subjects fitted with conventional compression factor (0.75 D) ortho-k. SFChT: Subfoveal choroidal thickness. ^aP<0.05, ^bP<0.01, ICF group vs CCF group by independent sample *t*-test. Error bars represent standard deviation.

was performed for RMS_h and SFChT in the M-ICF group. Examined data from the M-ICF group revealed that change in AL over 2y had a negative association with the change in RMS_h (r=-0.687, P<0.001; Figure 4A) and SFChT (r=-0.464, P=0.013; Figure 4B).

Comparisons of Other Measurements There were no statistical differences between the ICF group and the CCF group for each parameter over 2y (all *P*>0.05; Table 5). No G3,



Figure 4 Scatterplots exhibit the correlation between the change in RMS_h (A), SFChT (B) and the change in AL in the M-ICF group over 2y Regression lines indicate the coefficients of the Pearson correlation in the M-ICF group. AL: Axial length; RMS_h: Higherorder aberrations expressed as root mean square; SFChT: Subfoveal choroidal thickness.

G4 corneal staining (Efron's Grading Scale), poor near visual acuity, or poor dark adaptation was observed at any visit in all subjects.

DISCUSSION

As the most important parameter for ortho-k lens design, the back optic zone radius (BOZR) was determined based on the Jessen formula. That formula is BOZR=Kf-T-CF, while Kf represents the flat keratometry, T represents the target myopic reduction and CF represents the compression factor. Lee *et al*^[16] inferred that ICF ortho-k might steepen the mid-peripheral cornea and induce more peripheral myopic defocus to control myopia progression.

To the best of our knowledge, this is the first randomized trial to explore the efficacy and the optical mechanism of ICF ortho-k for myopic children stratified by myopia degrees over 2y. Table 6 illustrates the comparisons of current studies about ICF ortho-k.

Our results demonstrate that ICF ortho-k can correct lowmoderate myopia in adolescents safely, which are basically consistent with conclusions drawn by He *et al*^[17] and Ren *et al*^[18].

Table 6	Compariso	ns of prev	ious studie	s and our stu	dy			
Ctudy	Age range	Sample	Follow-up	Myopia	Initial	SE (D)	Change ir	n AL (mm)
Study	(y)	size	time	range (D)	ICF	CCF	ICF	CCF
Lau ^[8-9]	7.8-11	28	4-7wk	-4.0 to -0.5	-2.30±1.03	-2.27±0.99	-0.011	0.002
Wan ^[10]	6-10	25	4wk	-4.0 to -0.5	-2.08±0.97	-2.11±0.97	-	
He ^[17]	18-38	44	12mo	-5.0 to -0.75	-3.46±0.90	-3.84±0.77	-	
Ren ^[18]	20-36	50	1mo	-5.0 to -0.75	-3.58±0.90	-3.88±0.75	-	
Wan ^[19]	6-11	69	4wk	-4.0 to -0.5	-2.12±0.86	-2.17±0.81	-0.031	0.003
Present	8-16	97	2у	-5.0 to -1.0	-2.08±0.53 (L); -4.15±0.74 (M)	-1.89±0.51 (L); -4.33±0.70 (M)	0.28 (L); 0.23 (M)	0.31 (L); 0.30 (M)

ICF: Increased compression factor (1.75 D) ortho-k; CCF: Conventional compression factor (0.75 D) ortho-k; SE: Spherical equivalent; L: Low myopia subjects; M: Moderate myopia.

Our most intriguing finding is that ICF ortho-k can prevent myopia progression more effectively in moderate myopia adolescents, as assessed by 2 primary endpoints (change in AL and SE). In the moderate myopia group, the ICF group was 23.33% more effective in slowing axial elongation than CCF group after 2-year therapy. The increase of SE in the M-ICF group manifested 0.22 D less than in the M-ICF group after 2-year therapy. Wan et al^[19] reported that the axial elongation in ICF group (-0.031±0.06 mm) was significantly slower than that in CCF group (0.003±0.038 mm) after 4-week therapy, which was in agreement with our study. Nevertheless, Lau et al^[8] concluded that there was no statistical difference in AL and SE between ICF group and CCF group. That might be caused by the different follow-up times, sample sizes, and baseline SE. While ICF ortho-k was invalid for more myopic control in low myopia adolescents. This was in accordance with the study proved by Lau *et al*^[9]. However, we have to acknowledge that it was not so appropriate to compare these studies with ours simply because of differences in follow-up times, sample sizes, and demographic characteristics.

Our findings revealed that two factors might be responsible for slower axial growth in M-ICF group: greater higher-order aberrations (HOAs) or thicker choroidal thickness. On the one hand, this long-term study confirmed that ICF ortho-k led to a significant increase in HOAs and RMS_h metrics, particularly after 3months. Recent studies investigated that children with higher levels of HOAs in the root-mean-square (RMS) values exhibited slower axial growth^[20-22]. As the corneal shape is modified during ortho-k treatment, ocular aberrations, primarily HOAs, significantly increase^[23-25]. The increase in HOAs may usually be asymmetric. This can provide directional cues to the retina to respond to different visual stimuli rapidly and accurately. This compensatory optimization of image quality can suppress axial elongation by improving accommodative facility^[26-27]. In this study, HOAs increased significantly, demonstrating theoretically that the visual quality in moderate myopic children worsened after ICF ortho-k treatment. However, we found no difference in LC BCVA or subjective discomforts between the two groups. Therefore,

we speculate that children with ICF ortho-k may experience a higher visual compensation process or a blur adaptation induced by increased HOAs and visual fluctuation^[28-29]. Yet, we need to weigh the pros and cons of visual quality and myopia control.

On the other hand, this research also investigated that there was a negative correlation between the change in SFChT and AL at 2y in the M-ICF group. Therefore, it showed that choroidal thickening had a certain protective effect (r=-0.464) on axial growth. This was consistent with the results drawn by Lau et $al^{[8]}$ (r=-0.86), and Jin et $al^{[30]}$ (r=-0.4). However, Ding et al^[31] held a different view that the change in SFChT was irrelevant to the change in SE. The cause might be different analysis indicators (SE vs AL), and different measurement times (10:00-15:00 vs 8:00-10:00 a.m.). Nevertheless, the mechanism remains unclear. We speculate that improvement in optical performance and steady-state structure may lead to an increase in choroidal blood circulation and thickness^[32]. Choroidal thickening could change the optical imaging flat of the retina^[33], or generate certain chemical substances (NO, etc.) that induce a signal of scleral remodeling to slowed axial elongation^[34]. In recent years, EDI-SD OCT has been widely used in clinical practice to observe the choroidal tomographic structure and measure the thickness of the choroid due to its non-invasive, intuitive, and repeatable advantages^[35]. However, the change of choroidal thickness after wearing ortho-k was slight (usually measured in tens of μ m)^[36]. We utilized automatic registration and follow-up mode Spectralis to guarantee the precision and identical location at each visit.

Two main limitations should be noticed. First, the sample size of this study was still insufficient to group by age or lens design. Second, we did not control for confounding factors such as genetic and environmental factors.

In conclusion, ortho-k with increased compression factor (an extra 1 D) demonstrated safe and effective myopia correction in adolescents during the observation period, and it could control moderate myopia progression more effectively, which might be related to increasing RMS_h and SFChT. This study may provide a novel and effective ortho-k lens design for patients with moderate myopia.

ACKNOWLEDGEMENTS

Foundation: Supported by Education Department Foundation of Sichuan Province (No.15ZA0262).

Conflicts of Interest: Tang WT, None; Zhang L, None; Zhang HD, None; Li SB, None; Liang H, None.

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