Research Article

Assessment of Visual Quality in Eyes with Forme Fruste Keratoconus and Mild and Moderate Keratoconus Based on Optical Quality Analysis System II Parameters

Zhonghao Ren,1 Liyan Xu,2 Qi Fan,2 Kaili Yang,2 Shengwei Ren,2 and Dongqing Zhao2

1Henan University People’s Hospital, Henan Provincial People’s Hospital, Henan Eye Hospital, Henan Eye Institute, Zhengzhou 450003, China
2Henan Provincial People’s Hospital, Henan Eye Hospital, Henan Eye Institute, People’s Hospital of Zhengzhou University, Henan University People’s Hospital, Zhengzhou 450003, China

Correspondence should be addressed to Shengwei Ren; shengweiren1984@163.com and Dongqing Zhao; ykszdq@163.com

Received 25 November 2019; Accepted 9 January 2020; Published 23 March 2020

Academic Editor: Ji-jing Pang

Copyright © 2020 Zhonghao Ren et al. This is an open access article distributed under the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Purpose. The study aimed to evaluate the visual quality of forme fruste keratoconus (FFK) and mild and moderate keratoconus by using an optical quality analysis system II (OQAS-II) and to explore the correlation between optical quality parameters and the disease progression. Methods. Twenty-one normal eyes, twenty-one FFK eyes, twenty-one mild keratoconus eyes, and twenty-one moderate keratoconus eyes were included in this prospective study. The optical quality parameters, such as object scatter index (OSI), modulation transfer function cutoff (MTF cutoff), strehl ratio (SR), and OQAS-II values at contrasts of 100% (OV-100), 20% (OV-20), and 9% (OV-9), were measured by OQAS-II. The repeatability of these parameters was analyzed by intraclass correlation coefficient (ICC), repeatability coefficient (RC), and coefficient of variation (CVw). Correlations between optical quality parameters and mean central keratometry readings ($K_{mean}$) were evaluated. The sensitivity and specificity of the parameters were analyzed using the receiver operating characteristic (ROC). Results. All the optical quality parameters among four groups showed good repeatability (all ICC $\geq 0.75$). The MTF cutoff, SR, OV-100, OV-20, OV-9 in FFK, mild and moderate keratoconus eyes were significantly lower than those in the normal group (all $P < 0.05$). The ROC analyses of the MTF cutoff, SR, OV-100, OV-20, and OV-9 showed significant area under the curve (AUC) in discriminating FFK from normal, mild keratoconus from FFK, and moderate keratoconus from FFK and mild keratoconus (all $P < 0.05$). The OSI in mild and moderate keratoconus eyes were significantly higher than that in FFK and normal group (all $P < 0.05$), while the OSI showed no significant difference between the FFK group and normal group ($P > 0.05$). The ROC analyses of the OSI showed significant AUC in discriminating mild keratoconus from FFK and moderate keratoconus from FFK (all $P < 0.05$). In addition, the MTF cutoff was closely correlated to $K_{mean}$ in keratoconus eyes ($r = -0.710$, $P < 0.001$). Conclusion. The repeatability of OQAS-II is good in measuring visual quality of normal as well as FFK, mild, and moderate keratoconus. The visual quality of the FFK, mild, and moderate keratoconus is worse than that in normal eyes. The OQAS-II has the potential value in screening FFK from normal eyes and might be a useful tool for evaluating the progression of keratoconus.

1. Introduction

Keratoconus is a degenerative disorder characterized by corneal thinning and conical-shaped protrusion of the cornea [1]. It usually happens in adolescence and progresses until the third or fourth decade of life [2]. The progressive corneal protrusion and thinning would induce irregular astigmatism, leading to the impairment of visual function [3]. The individuals with mild keratoconus usually have their vision corrected with contact lenses or spectacles [4]. Nevertheless, the majority of keratoconus patients still have ocular discomfort and poor vision quality [5].
Several studies have found that the keratoconus patients have bad visual quality by questionnaire investigations [6,7]. Furthermore, it has been identified that the mild and moderate keratoconus had lower grades than normal people according to the National Eye Institute Visual Function Questionnaire-25 [8]. In addition, several studies have shown that both ocular and corneal aberrations are significantly higher in keratoconic eyes than those in normal eyes [9,10]. With the continuous development of digital image and computer processing, some new quantitative evaluation technology of the visual quality appeared. The optical quality analysis system (OQAS-II) is a double-pass (DP) system that could provide an objective clinical evaluation of the visual quality and has been successfully used to objectively classify the maturity of cataracts [11]. As far as we know, there are few studies aiming to evaluate the visual quality in keratoconus using OQAS-II. Ye et al. [12] found that the modulation transfer function cutoff (MTF cutoff), Strehl ratio (SR), OQAS values at contrasts of 100% (OV-100), OQAS values at contrasts of 20% (OV-20), and OQAS values at contrasts of 9% (OV-9) had significant differences between the normal and forme fruste keratoconus (FFK) patients. Leonard et al. [13] reported that the OSI was increased in keratoconic eyes and could be considered as a clinically significant parameter to stage keratoconus by directly evaluating visual quality. However, to the best of our knowledge, comparisons of all the parameters among FFK, mild keratoconus, moderate keratoconus patients, and normal have not yet been reported.

Therefore, the present study aimed to investigate the measurement variability of the OQAS-II and evaluate the visual quality of FFK, mild, and moderate keratoconus by OQAS-II. Additionally, we investigated the diagnostic ability of OQAS-II in screening FFK from normal eyes and explored the correlation between optical quality parameters and the disease progression.

2. Materials and Methods

2.1. Study Subjects. This study was conducted according to the Declaration of Helsinki guidelines, and all procedures involving human subjects were approved by the Institutional Review Board of Henan Eye Hospital. Written informed consents were obtained from all subjects.

Twenty-one FFK patients, 21 mild keratoconus patients and 21 moderate keratoconus patients were recruited in the refractive surgery center of Henan Eye Hospital from March 2018 to March 2019. Twenty-one simple refractive errors patients with matched age and gender were enrolled as the normal group. The FFK group consisted of 21 topographically normal eyes of patients with KC in the other eye [14]. The diagnosis of keratoconus was based on clinical examinations and the presence of characteristic corneal topographic features [15], in which the patients were presented with the eccentric steepening keratometry and anterior and the posterior elevation patterns such as I-S asymmetry, as well as at least one of the clinical diagnostic signs such as Fleischer ring, Vogt’s striae, and corneal thinning by means of slit-lamp biomicroscopy. According to the Amsler-Krumeich scales (Supplemental Table 1), stage 1 of the Amsler-Krumeich scales (AK1) and stage 2 of the Amsler-Krumeich scales (AK2) were defined as the mild and moderate keratoconus, respectively [15]. In patients with FFK, the fellow eye with keratoconus was excluded from the mild and moderate group. Eyes with previous ocular surgery, corneal haze, scar, cataract, vitreous opacity or aqueous humor opacity, rigid contact lens wears within 4 weeks, soft contact lens wears within 2 weeks, and severe keratoconus were excluded.

2.2. Clinical Examination. For each participant, a complete eye examination was performed, including best-corrected visual acuity (BCVA, logMAR), manifest refraction, silt-lamp and fundus examination, Visante Omni anterior segment OCT (Carl Zeiss Jena GmbH, Germany), corneal thickness in the thinnest point, intraocular pressure, and the axial length.

OQAS-II (Visiometrics, Spain) was used to measure the optical quality parameters of all subjects. The light source of the system is a 780 nm laser diode which is fully filtered and collimated. The point light source is imaged on the retina. After retina reflection, light passes twice through ocular media. Then, the HD Analyzer analyses the size and the shape of the reflected light spot. The measurement was performed in a darkroom to avoid the effects of spherical aberration and astigmatism. The subjects adapt to the dark environment for 10 minutes to acquire the largest pupil in the natural state after wearing the corrective lenses. The subjects were instructed to remain stationary after being positioned on the chin rest of the instrument. It was made sure that the subjects continued to fixate on the target and blink prior to each measurement to maintain a good tear film during the examination. The pupil diameter was set at 4 mm, and the spherical refraction errors were corrected by an incorporated optometer in the DP system (+5D∼−8D). For subjects with more than 0.5 D cylindrical refraction errors, the astigmatism was corrected with an external cylindrical lens [16]. All subjects underwent 3 consecutive tests and were measured by the same investigator with 5 minutes interval. The average value was used in the current analysis.

OQAS-II can provide six optical quality parameters, including OSI, MTF cut off, SR, OV-100, OV-20, and OV-9. The OSI quantifies intraocular scattered light and is defined as the ratio of the light of peripherally annular area versus that of the central peak in the acquired system. Small value of OSI is usually linked to eye with low scattering. The MTF cutoff is set as the cutoff point of the MTF curve on the x-axis and is calculated from the point spread function directly. The cutoff value represents the highest spatial frequency at which the MTF reaches the lowest contrast of 1%. The SR is defined as the ratio of the area under the MTF curve of the measured eye to that of the ideal aberration-free eye. The three OVs are normalized values of three spatial frequencies that correspond to the MTF values for three contrast conditions: 100 percent (OV-100), 20 percent (OV-20), and 9 percent (OV-9).
2.3. Statistical Analysis. All statistical analyses were performed by the SPSS 22.0 package (SPSS Inc., Chicago, IL, USA). Quantitative variables were expressed with mean ± standard deviation (SD) and analyzed by one-way analysis of variance with least significance difference (LSD-t) corrections. The qualitative variables were expressed with percentage and analyzed by Pearson’s chi squared test. The measurement repeatability of optical quality parameters was assessed through three indicators, which include intraclass correlation coefficient (ICC), repeatability coefficient (RC), and coefficient of variation ($CV_w$). For calculation, $RC$ was calculated by $(CV_w)^2$ and then divided by the overall mean. $RC$ showed excellent repeatability (ICC $> 0.75$) in AK2. All the parameters with statistically significant difference, receiver operating characteristic (ROC) analyses were performed to demonstrate the accuracy of the parameters in distinguishing FFK from normal eyes, AK1 from FFK, and AK2 from AK1. $P$ values less than 0.05 were considered statistically significant.

3. Results

3.1. Demographic Data of Subjects. The demographic data of the four groups are shown in Table 1. The results showed no significant difference in gender, age, and the axial length among the 4 groups (all $P > 0.05$). The spherical equivalent in the AK2 group was significantly higher than that in the normal group ($P < 0.05$), while the spherical equivalent in AK1 group showed no significant difference with the FFK group as well as the normal group (all $P > 0.05$). The astigmatism, steep keratometry ($K_s$), flat keratometry ($K_f$), mean central keratometry readings ($K_{mean}$), and BCVA (logMAR) in the AK1 group and AK2 group were significantly higher than those in the normal group (all $P < 0.05$), while the corneal thickness in the thinnest point and intraocular pressure in the AK1 group and AK2 group were significantly lower than those in the normal group ($P < 0.05$).

3.2. Repeatability of Optical Quality Parameters. Table 2 summarized the repeatability values of the OQAS-II parameters. In the normal group, 5 of 6 parameters (83.33%) showed excellent repeatability (ICC $> 0.90$) and 1 parameter (16.67%) showed good repeatability (ICC $> 0.75$). Similarly, 5 of 6 parameters (83.33%) showed excellent repeatability (ICC $> 0.90$) and 1 parameter (16.67%) showed good repeatability (ICC $> 0.75$) in FFK. All the parameters (100%) showed excellent repeatability (ICC $> 0.90$) in AK1, while only 1 parameter (16.67%) showed excellent repeatability (ICC $> 0.75$) and 5 parameters (83.33%) showed good repeatability (ICC $> 0.75$) in AK2.

3.3. Comparison of Optical Quality Parameters. The comparison of optical quality parameters among the four groups is shown in Table 3. The MTF cutoff, SR, OV-100, OV-20, OV-9 in FFK, AK1, and AK2 were significantly lower than those in the normal group (all $P < 0.05$). Similarly, the MTF...
### Table 2: Measurement of repeatability of OQAS-II parameters for the normal group, FFK, AK1, and AK2.

|        | ICC   | RC   | CV_w (%) | ICC   | RC   | CV_w (%) | ICC   | RC   | CV_w (%) | ICC   | RC   | CV_w (%) |
|--------|-------|------|----------|-------|------|----------|-------|------|----------|-------|------|----------|
| OSI    | 0.973 | 0.152 | 12.448   | 0.968 | 0.303 | 18.053   | 0.994 | 0.971 | 12.306   | 0.934 | 0.470 | 17.319   |
| MTF cutoff (c/d) | 0.965 | 5.472 | 4.231    | 0.915 | 12.289 | 11.432   | 0.953 | 10.879 | 17.877   | 0.792 | 9.840 | 14.131   |
| SR     | 0.868 | 0.088 | 11.712   | 0.887 | 0.088 | 14.011   | 0.950 | 0.088 | 24.325   | 0.770 | 0.088 | 52.705   |
| OV-100 | 0.966 | 0.175 | 4.054    | 0.915 | 0.411 | 11.471   | 0.953 | 0.361 | 17.861   | 0.791 | 0.328 | 49.301   |
| OV-20  | 0.957 | 0.232 | 6.915    | 0.910 | 0.361 | 13.725   | 0.963 | 0.248 | 18.254   | 0.779 | 0.232 | 52.291   |
| OV-9   | 0.928 | 0.215 | 10.328   | 0.900 | 0.248 | 15.227   | 0.947 | 0.175 | 21.809   | 0.785 | 0.152 | 54.772   |

**Table 3: OQAS-II parameters among the normal, FFK, AK1, and AK2 group.**

|        | NL (n = 21) | FFK (n = 21) | AK1 (n = 21) | AK2 (n = 21) | FFK/ NL | AK1/ NL | AK2/ NL | FFK/ AK1 | FFK/ AK2 | AK1/ AK2 |
|--------|-------------|--------------|--------------|--------------|--------|--------|--------|---------|---------|---------|
| OSI    | 0.44 ± 0.18 | 0.61 ± 0.37  | 2.85 ± 2.62  | 9.88 ± 3.86  | 0.815  | 0.990  | 0.000  | 0.000  | 0.000  | 0.000   |
| MTF cutoff (c/d) | 46.69 ± 6.43 | 38.81 ± 8.93 | 21.97 ± 10.59 | 7.23 ± 4.51 | 0.002* | 0.000* | 0.000* | 0.000* | 0.000* | 0.000* |
| SR     | 0.27 ± 0.05 | 0.23 ± 0.06  | 0.13 ± 0.05  | 0.06 ± 0.02  | 0.002* | 0.000* | 0.000* | 0.000* | 0.000* | 0.000* |
| OV-100 | 1.56 ± 0.21 | 1.29 ± 0.30  | 0.73 ± 0.35  | 0.24 ± 0.15  | 0.002* | 0.000* | 0.000* | 0.000* | 0.000* | 0.000* |
| OV-20  | 1.21 ± 0.22 | 0.95 ± 0.26  | 0.49 ± 0.27  | 0.16 ± 0.10  | 0.000* | 0.000* | 0.000* | 0.000* | 0.000* | 0.000* |
| OV-9   | 0.75 ± 0.16 | 0.59 ± 0.17  | 0.29 ± 0.16  | 0.10 ± 0.07  | 0.000* | 0.000* | 0.000* | 0.000* | 0.000* | 0.000* |

**3.4. The Relationship between the Optical Quality Parameters and K_mean.** Figure 1 showed the MTF cutoff was closely correlated to \( K_{\text{mean}} \) in keratoconus eyes (AK1 and AK2) \((r = -0.710, P < 0.05)\), while the MTF cutoff showed no significant correlation to \( K_{\text{mean}} \) in the normal group \((r = 0.004, P = 0.987)\) and FFK group \((r = -0.335, P = 0.138)\).

The relationships between other optical quality parameters and \( K_{\text{mean}} \) are shown in Supplemental Table 2.

**3.5. ROC Curve Analyses.** Then, we sought to evaluate the optical quality parameters as adjunctive diagnostic indicators by using ROC analysis (Figures 2–4). Comparing the cutoff, SR, OV-100, OV-20, and OV-9 values in the AK1 group and AK2 group were significantly lower than those in the FFK group (all \( P < 0.05 \)). The OSI values in the AK1 group and AK2 group were significantly higher than those in the FFK group and normal group (\( P < 0.05 \)), while the OSI values showed no significant difference between the FFK group and normal group.

![Figure 1: Correlation between the MTF cutoff and \( K_{\text{mean}} \) among normal, FFK, and keratoconus eyes (AK1 and AK2). A graph showing a significant correlation between the MTF cutoff and \( K_{\text{mean}} \) in keratoconus eyes \((r = -0.710, P < 0.001)\). No significant correlation was found between the MTF cutoff and \( K_{\text{mean}} \) in the normal group \((r = 0.004, P = 0.987)\) and FFK group \((r = -0.335, P = 0.138)\).](attachment:image.png)
MTF cutoff, SR, OV-100, OV-20, and OV-9 in normal eyes to FFK, the areas under the curve (AUC) were 0.760, 0.740, 0.761, 0.781, and 0.765, respectively (all \( P < 0.001 \)) (Figure 2). Moreover, the OSI, MTF cutoff, SR, OV-100, OV-20, and OV-9 showed a significant ability to discern AK1 from FFK with the AUC of 0.889, 0.893, 0.905, 0.893, 0.901, and 0.907, respectively (all \( P < 0.001 \)) (Figure 3). In addition, the OSI, MTF cutoff, SR, OV-100, OV-20, and OV-9 also demonstrated a significant ability to discern AK2 from AK1 with the AUC of 0.948, 0.909, 0.908, 0.910, 0.916, and 0.908, respectively (all \( P < 0.001 \)) (Figure 4). The cutoff value, sensitivity, specificity, and Youden index of these parameters are shown in Supplemental Tables 3–5, respectively.

**4. Discussion**

Our findings showed that OQAS-II had good repeatability in measuring visual quality of normal, FFK, AK1, and AK2. The visual quality in FFK, AK1, and AK2 was inferior to that in normal. Our results showed that the MTF cutoff was significantly associated with mean in keratoconus eyes. In addition, our results demonstrated that the OQAS-II might help clinicians to better understand the visual quality in keratoconus and could be a useful tool for detecting FFK and monitoring its progression.

Keratoconus is a progressive corneal ectasia characterized by localized corneal thinning which leads to the protrusion of cornea [2]. Corneal tomography is currently the most widely available method to diagnose early keratoconus. Although the changes on the topography of cornea could be obviously detected before the clinical signs of keratoconus, these changes do not correlate with the visual acuity [18]. As far as we know, the wavefront sensors and OQAS-II were both objective evaluating devices of visual quality. While in eyes where scattered light and aberrations are prominent, wavefront sensors may overestimate image quality [19]. In contrast, the OQAS-II can reflect a more accurate description of the visual quality and has been widely used for clinical application [20–22]. OQAS-II images contained all the information about the visual quality of the eye including all the higher-order aberrations and scattered light, being both generally missed by most aberrometric techniques. In addition, the OQAS-II can also help clinicians explain why some patients have good BCVA, but the subjective visual disturbance is obvious. Therefore, it has been considered as a convenient and objective method for visual quality assessment.
In order to assess the visual quality and explore the potential diagnostic value of OQAS-II in keratoconus patients, the measurement repeatability of OQAS-II needs to be explored to make a reliable clinical judgement. Several studies have identified a good measurement repeatability of OQAS-II. Xu et al. [23] measured 119 healthy eyes with OQAS-II and concluded that the OQAS-II showed excellent repeatability for objective measurements of overall visual quality in clinic. Iijima et al. [24] also reported a good repeatability of OQAS in healthy adults. Furthermore, studies have also shown that the measurement repeatability of the DP system was good in FFK [12], which was in accordance with our current findings. To our knowledge, there are no studies on the measurement repeatability of OQAS-II in mild and moderate keratoconus. In clinic, ICC $\geq 0.75$ indicates good to excellent repeatability, and ICC $\geq 0.90$ means the device has excellent repeatability [25]. Our study showed that the repeatability of all the optical quality parameters detected by OQAS-II was excellent in AK1, while some parameters including MTF cutoff, SR, OV-100, OV-20, and OV-9 in AK2 showed good measurement repeatability. This might be attributed to the obvious irregular corneal distortion in AK2, consequently degrading the retinal image quality [26]. Our findings support that OQAS-II measurements are reliable in evaluating the visual quality changes in eyes with FFK, mild, and moderate keratoconus. Further studies on different stages of keratoconus should be performed to confirm our findings.

In order to assess the visual quality and explore the potential diagnostic value of OQAS-II in keratoconus patients, the measurement repeatability of OQAS-II needs to be explored to make a reliable clinical judgement. Several studies have identified a good measurement repeatability of OQAS-II. Xu et al. [23] measured 119 healthy eyes with OQAS-II and concluded that the OQAS-II showed excellent repeatability for objective measurements of overall visual quality in clinic. Iijima et al. [24] also reported a good repeatability of OQAS in healthy adults. Furthermore, studies have also shown that the measurement repeatability of the DP system was good in FFK [12], which was in accordance with our current findings. To our knowledge, there are no studies on the measurement repeatability of OQAS-II in mild and moderate keratoconus. In clinic, ICC $\geq 0.75$ indicates good to excellent repeatability, and ICC $\geq 0.90$ means the device has excellent repeatability [25]. Our study showed that the repeatability of all the optical quality parameters detected by OQAS-II was excellent in AK1, while some parameters including MTF cutoff, SR, OV-100, OV-20, and OV-9 in AK2 showed good measurement repeatability. This might be attributed to the obvious irregular corneal distortion in AK2, consequently degrading the retinal image quality [26]. Our findings support that OQAS-II measurements are reliable in evaluating the visual quality changes in eyes with FFK, mild, and moderate keratoconus. Further studies on different stages of keratoconus should be performed to confirm our findings.

Then, we comparatively evaluated the visual quality in FFK, mild, and moderate keratoconus patients. The significant upward trend in OSI and downward trend in MTF cutoff, SR, OV-100, OV-20, and OV-9 observed from normal to moderate keratoconus indicated that as the disease continued to advance, the visual quality in keratoconus declined gradually. Our results showed that the MTF cutoff, SR, OV-100, OV-20, and OV-9 in FFK were significantly lower than those in the normal group (all $P < 0.05$). These results were consistent with those given by Ye et al. [12], but they did not evaluate the OSI value. Leonard et al. [13] compared the OSI values between the keratoconus eyes and normal eyes and found statistically significant increments of OSI scores in the AK1 and AK2 group, which was also consistent with our results. However, our results further showed the OSI had no significant difference between the FFK and normal group. Miháltz et al. [27] also reported that visual quality in terms of the Strehl ratio and the spot radius

![Figure 3: Comparisons between FFK and AK1. Receiver operating characteristic (ROC) curves were constructed by plotting sensitivity versus 100% specificity at varying cutoff values of the OSI (a), MTF cut off (b), SR (c), and OV-100 (d), OV-20 (e), OV-9 (f), respectively. Area under the curve (AUC) and 95% CI are noted at the bottom right of each graph. For further information about cut-off values, sensitivity, specificity, and Youden index, see Supplemental Table 4.](image-url)
was degraded in the subclinical keratoconus and keratoconus group compared with that in the control group; although related, the Strehl ratio parameter described in their study was different from our results. Moreover, our results showed that the MTF cutoff was significantly associated with $K_{\text{mean}}$ in keratoconus eyes. Further studies with different populations should be conducted to confirm the findings.

ROC curve analysis could illustrate the diagnostic ability of a binary classifier system [28]. To evaluate the potential diagnostic value of OQAS-II parameters in FFK, mild, and moderate keratoconus, we made the ROC curve analyses of these parameters in our study. And we found some optical quality parameters except OSI displayed a significant ability to discern AK1 from FFK and AK2 from AK1, which indicated that the optical quality parameters could help to evaluate the progression of keratoconus and detect the early stage of the disease. The OV-20 had the largest area under the curve (AUC = 0.781) in FFK compared to the normal group, followed by the OV-9, while the OV-9 had the largest area under the curve (AUC = 0.907) in identifying AK1 from FFK. The OSI had the largest area under the curve (AUC = 0.948) in identifying AK2 from AK1.

Regardless of any correlation between keratoconus age and stage, the significant upward trend in OSI and downward trend in MTF cutoff, SR, OV-100, OV-20, and OV-9 observed from normal to AK2 and significant AUC by ROC analysis (Figures 2–4) indicated that the optical quality parameters measured by OQAS-II may be helpful to monitoring the progression of keratoconus.

Several limitations of our study need to be addressed. Firstly, the sample size of the present study was small, which might affect the validity of our results. Secondly, we did not include the severe keratoconus eyes in our study since it is difficult to measure eyes with high astigmatism, which might get some bias of the visual quality through an external cylindrical lens. Thirdly, the mean OSI value recorded over 19.5 seconds without blinking was not included in current study, which reflects the tear film dynamic alterations and might be different among normal, FFK, mild, and moderate keratoconus. A further multicenter study should be conducted later.

In conclusion, the repeatability of OQAS-II was good in normal as well as FFK, mild, and moderate keratoconus eyes. And the FFK, mild, and moderate keratoconus patients had worse visual quality compared with that in normal eyes. Furthermore, the OQAS-II might be a new method in
detecting FFK and a useful tool for objectively evaluating the progression of keratoconus.

Data Availability

The data used to support the findings of this study are available from the corresponding author upon request.

Conflicts of Interest

The authors declare that there are no conflicts of interest regarding the publication of the paper.

Acknowledgments

This study was supported by Henan Natural Science Foundation (no. 182300410362) and Open Program of Shandong Provincial Key Laboratory of Ophthalmology (no. 2018-04).

Supplementary Materials

Supplemental Table 1: Amsler-Krumeich classification for keratoconus severity. Supplemental Table 2: relationship between $K_{\text{mean}}$ and OQAS-II parameters. Supplemental Table 3: the ROC analysis of OQAS-II parameters between FFK and NL. Supplemental Table 4: the ROC analysis of OQAS-II parameters between FFK and AK1. Supplemental Table 5: the ROC analysis of OQAS-II parameters between AK1 and AK2. (Supplementary Materials)

References

[1] Y. S. Rabinowitz, "Keratoconus," *Survey of Ophthalmology*, vol. 42, no. 4, pp. 297–319, 1998.
[2] M. Romero-Jiménez, J. Santodomingo-Rubido, and J. S. Wolfohn, "Keratoconus: a review," *Contact Lens and Anterior Eye*, vol. 33, no. 4, pp. 157–166, 2010.
[3] S. Basu and J. Vazirani, "Keratoconus: current perspectives," *Clinical Ophthalmology*, vol. 7, no. 7, pp. 2019–2030, 2013.
[4] A. Suzuki, N. Maeda, M. Fuchihata, S. Koh, K. Nishida, and T. Fujikado, "Visual performance and optical quality of standardized asymmetric soft contact lenses in patients with keratoconus," *Investigative Ophthalmology & Visual Science*, vol. 58, no. 7, pp. 2899–2905, 2017.
[5] S. Lee, G. Jung, and H. K. Lee, "Comparison of contact lens corrected quality of vision and life of keratoconus and myopic patients," *Korean Journal of Ophthalmology*, vol. 31, no. 6, pp. 489–496, 2017.
[6] V. Saunier, A.-E. Mercier, T. Gaboriau et al., "Vision-related quality of life and dependency in French keratoconus patients: impact study," *Journal of Cataract & Refractive Surgery*, vol. 43, no. 12, pp. 1582–1590, 2017.
[7] H. Mahdaviazad, S. Bamdad, N. Roustaei, and S. Mohaghegh, "Vision-related quality of life in Iranian patients with keratoconus," *Eye & Contact Lens: Science & Clinical Practice*, vol. 44, no. 2, pp. S350–S354, 2018.
[8] S. Aydin Kurna, A. Altun, T. Gencaga, S. Akkaya, and T. Sengor, "Vision related quality of life in patients with keratoconus," *Journal of Ophthalmology*, vol. 2014, Article ID 694542, 7 pages, 2014.
[9] B. Jafri, X. Li, H. Yang, and Y. S Rabinowitz, "Higher order wavefront aberrations and topography in early and suspected keratoconus," *Journal of Refractive Surgery*, vol. 23, no. 8, pp. 774–781, 2007.
[10] Z. Schlegel, Y. Lefé, H. S. Bains, and D. Gatinel, "Total, corneal, and internal ocular optical aberrations in patients with keratoconus," *Journal of Refractive Surgery*, vol. 25, no. 10, pp. S951–S957, 2009.
[11] P. Artal, A. Benito, G. M. Perez et al., "An objective scatter index based on double-pass retinal images of a point source to classify cataracts," *PLoS One*, vol. 6, no. 2, Article ID e16823, 2011.
[12] C. Ye, P. K.-F. Ng, and V. Jhanji, "Optical quality assessment in normal and forme fruste keratoconus eyes with a double-pass system: a comparison and variability study," *British Journal of Ophthalmology*, vol. 98, no. 11, pp. 1478–1483, 2014.
[13] A. P. Leonard, S. D. Gardner, K. M. Rocha, E. R. Zeldin, D. M. Tremblay, and G. O. Waring, "Double-pass retinal point imaging for the evaluation of optical light scatter, retinal image quality, and staging of keratoconus," *Journal of Refractive Surgery*, vol. 32, no. 11, pp. 760–765, 2016.
[14] A. Saad and D. Gatinel, "Topographic and tomographic properties of forme fruste keratoconus corneas," *Investigative Ophthalmology & Visual Science*, vol. 51, no. 11, pp. 5546–5555, 2010.
[15] V. Mas Tur, C. MacGregor, R. Jayaswal, D. O’Brart, and N. Maycock, "A review of keratoconus: diagnosis, pathophysiology, and genetics," *Survey of Ophthalmology*, vol. 62, no. 6, pp. 770–783, 2017.
[16] M. Tian, H. Miao, Y. Shen, J. Gao, X. Mo, and X. Zhou, "Intra- and inter-session repeatability of an optical quality and intraocular scattering measurement system in children," *PLoS One*, vol. 10, no. 11, Article ID e0142189, 2015.
[17] J. M. Bland and D. G. Altman, "Statistics notes: measurement error," *BMJ*, vol. 313, no. 7059, p. 744, 1996.
[18] M.-R. Nilforoushan, M. Speaker, M. Marmor et al., "Comparative evaluation of refractive surgery candidates with Placido topography, Orbscan II, Pentacam, and wavefront analysis," *Journal of Cataract & Refractive Surgery*, vol. 34, no. 4, pp. 623–631, 2008.
[19] F. Diaz-Douton, A. Benito, J. Pujol, M. Arjona, J. L. Guell, and P. Artal, "Comparison of the retinal image quality with a Hartmann-Shack wavefront sensor and a double-pass instrument," *Investigative Ophthalmology & Visual Science*, vol. 47, no. 4, pp. 1710–1716, 2006.
[20] M. Vilaseca, M. J. Romero, M. Arjona et al., "Grading nuclear, cortical and posterior subcapsular cataracts using an objective scatter index measured with a double-pass system," *British Journal of Ophthalmology*, vol. 96, no. 9, pp. 1204–1210, 2012.
[21] R. G. Anera, J. J. Castro, J. R. Jiménez, C. Villa, and A. Alarcón, "Optical quality and visual discrimination capacity after myopic LASIK with a standard and aspheric ablation profile," *Journal of Refractive Surgery*, vol. 27, no. 8, pp. 597–601, 2011.
[22] M. Vilaseca, A. Padilla, J. Pujol, J. C. Ondategui, P. Artal, and J. L. Gáell, "Optical quality one month after verisyse and Verilux phakic IOL implantation and Zeiss MEL 80 LASIK for myopia from 5.00 to 16.50 diopters," *Journal of Refractive Surgery*, vol. 25, no. 8, pp. 689–698, 2009.
[23] C. C. Xu, T. Xue, Q. M. Wang, Y. N. Zhou, J. H. Huang, and A. Y. Yu, "Repeatability and reproducibility of a double-pass optical quality analysis device," *PLoS One*, vol. 10, no. 2, Article ID e0117587, 2015.
[24] A. Iijima, K. Shimizu, H. Kobashi, A. Saito, and K. Kamiya, “Repeatability, reproducibility, and comparability of subjective and objective measurements of intraocular forward scattering in healthy subjects,” BioMed Research International, vol. 2015, Article ID 925217, 6 pages, 2015.

[25] Association JS, Accuracy (Trueness and Precision) of Measurement Methods and Results—Part 6: Use in Practice of Accuracy Values, 1994.

[26] A. Jinabhai, W. Neil Charman, C. O’Donnell, and H. Radhakrishnan, “Optical quality for keratoconic eyes with conventional RGP lens and simulated, customised contact lens corrections: a comparison,” Ophthalmic and Physiological Optics, vol. 32, no. 3, pp. 200–212, 2012.

[27] K. Miháltz, I. Kovács, K. Kránitz, G. Erdei, J. Németh, and Z. Z. Nagy, “Mechanism of aberration balance and the effect on retinal image quality in keratoconus: optical and visual characteristics of keratoconus,” Journal of Cataract & Refractive Surgery, vol. 37, no. 5, pp. 914–922, 2011.

[28] A. N. Kamarudin, T. Cox, and R. Kolamunnage-Dona, “Time-dependent ROC curve analysis in medical research: current methods and applications,” BMC Medical Research Methodology, vol. 17, no. 1, p. 53, 2017.