Mean Posterior Corneal Power and Astigmatism in Normal Versus Keratoconic Eyes

Sepehr Feizi1, MD, MS; Siamak Delfazayebaher2, MD; Mohammad Ali Javadi2, MD; Farid Karimian2, MD; Vahid Ownagh2, MD; Fatemeh Sadeghpour2, MD

1Ocular Tissue Engineering Research Center, Shahid Beheshti University of Medical Sciences, Tehran, Iran
2Ophthalmic Research Center, Shahid Beheshti University of Medical Sciences, Tehran, Iran

Abstract

Purpose: To compare mean posterior corneal power and astigmatism in normal versus keratoconus affected eyes and determine the optimal cut-off points to maximize sensitivity and specificity in discriminating keratoconus from normal corneas.

Methods: A total of 204 normal eyes and 142 keratoconus affected eyes were enrolled in this prospective comparative study. Mean posterior corneal power and astigmatism were measured using a dual Scheimpflug camera. Correlation coefficients were calculated to assess the relationship between the magnitudes of keratometric and posterior corneal astigmatism in the study groups. Receiver operating characteristic curves were used to compare the sensitivity and specificity of the measured parameters and to identify the optimal cut-off points for discriminating keratoconus from normal corneas.

Results: The mean posterior corneal power was $-6.29 \pm 0.20$ D in the normal group and $-7.77 \pm 0.87$ D in the keratoconus group ($P < 0.001$). The mean magnitudes of the posterior corneal astigmatisms were $-0.32 \pm 0.15$ D and $-0.94 \pm 0.39$ D in the normal and keratoconus groups, respectively ($P < 0.001$). Significant correlations were found between the magnitudes of keratometric and posterior corneal astigmatism in the normal ($r=-0.76$, $P < 0.001$) and keratoconus ($r=-0.72$, $P < 0.001$) groups. The mean posterior corneal power and astigmatism were highly reliable characteristics that distinguished keratoconus from normal corneas (area under the curve, 0.99 and 0.95, respectively). The optimal cut-off points of mean posterior corneal power and astigmatism were $-6.70$ D and $-0.54$ D, respectively.

Conclusion: Mean posterior corneal power and astigmatism measured using a Galilei analyzer camera might have potential in diagnosing keratoconus. The cut-off points provided can be used for keratoconus screening.

Keywords: Keratoconus; Posterior Corneal Astigmatism; Posterior Corneal Power

INTRODUCTION

Keratoconus is a non-inflammatory and progressive corneal disease with unknown etiology. It leads to thinning and bulging of the cornea and, consequently, irregular...
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Recent studies have been oblique astigmatism on both corneal surfaces by increasing anterior against-the-rule (ATR) and posterior with keratoconus. Naderan et al and is correlated to total corneal astigmatism in eyes. Corneal astigmatism produces large, variable magnitudes correlation between anterior and posterior corneal astigmatism with disease progression and a significant They found a significant increase of posterior corneal astigmatism was approximately 1.0 D with a range of 0.0 to 2.90 D. They found no statistically significant increase in anterior, posterior, or total corneal astigmatism with the progressive stages of keratoconus. Shajari et al reported that the posterior corneal astigmatism was 0.69 ± 0.40 D in eyes across all keratoconus stages. They found a significant increase of posterior corneal astigmatism with disease progression and a significant correlation between anterior and posterior corneal astigmatism. Savini et al reported that posterior corneal astigmatism produces large, variable magnitudes and is correlated to total corneal astigmatism in eyes with keratoconus. Naderan et al reported a trend for increasing anterior against-the-rule (ATR) and posterior with-the-rule (WTR) astigmatism as well as decreasing oblique astigmatism on both corneal surfaces by increasing the keratoconus severity. They found a cut-off value of 0.4 D for posterior corneal astigmatism with 89.5% sensitivity and 85.0% specificity for discriminating keratoconus from normal corneas. The purpose of the present study was to measure and compare the magnitudes of mean posterior corneal power and astigmatism between normal and keratoconus corneas by using a dual Scheimpflug analyzer. Additionally, an attempt was made to identify optimal cut-off points to maximize sensitivity and specificity in discriminating keratoconus from normal corneas.

METHODS

In this prospective comparative study, the right eyes of 204 normal (90 male subjects) and 142 eyes of 98 patients with keratoconus (56 male patients) were enrolled. The study was approved by the Institutional Review Board of the Ophthalmic Research Center, which is affiliated with Shahid Beheshti University of Medical Sciences, Tehran, Iran, and followed the tenets of the Declaration of Helsinki. Informed consent, signed by the participants, was obtained after the purpose of the study was thoroughly explained.

In the normal group, the only ocular problem was refractive error (sphere between −13.0 and +0.75 D and refractive astigmatism lower than 6.5 D), and a history of diagnosis of ocular pathology such as dry eye, keratoconus, glaucoma, retinal disease, or any previous ocular surgery led to subject exclusion. The keratoconus group consisted of patients with clinical keratoconus. Eyes with previous acute corneal hydrops, stromal scar, other corneal and ocular diseases, or a history of any ocular surgery were excluded from the keratoconus group. Contact lens users were asked to discontinue the use of lenses for at least 2 weeks or 1 month before the preoperative examination for soft and gas permeable contact lenses, respectively.

A complete ocular examination, including slit-lamp biomicroscopy, manifest refraction, uncorrected and best-spectacle corrected visual acuity (UCVA and BSCVA, respectively) measurement using a Snellen acuity chart, keratometry, intraocular pressure measurement, and dilated funduscopy was performed. The diagnosis of keratoconus was based on clinical slit-lamp findings (stromal thinning, conical protrusion, Fleischer ring, and Vogt striae) and associated characteristics of Placido disk-based (TMS-2, Tomey, Nagoya, Japan) topographic patterns. Keratoconic eyes were divided into the following three groups per mean keratometry (K) readings: mild (K < 48.0 D), moderate (48.0 D ≤ K ≤ 52.0 D), and severe (K > 52.0 D).

For measurements with the Galilei dual Scheimpflug analyzer (software version 5.2; Ziemer Ophthalmic System AG), the participants were seated with their chinrest and forehead against the forehead strap while focusing on the target. The alignment of the scan...
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Statistical Analysis

Considering an α error of 5.0%, study power of 80%, difference in posterior corneal astigmatism between the two study groups (d) of 0.2 D, and standard deviation of difference in posterior corneal astigmatism of 0.3 D, the sample size was calculated to be 135 patients in each group. Data were analyzed using Statistical Package for the Social Sciences version 21 (IBM Corp., Armonk, NY, USA). Values indicating means and standard deviations, ranges, frequencies, and percentages were used to express the data. The normal distribution of continuous variables was verified using a Kolmogorov–Smirnov test and Q-Q plot. The chi-square test was used for comparing percentages between the groups. Generalized estimating equation models were used to compare the study groups to prevent the confounding effects of potential correlations between measurements from both corneas of the same patient. The Pearson or partial correlation coefficient was calculated in the normal and keratoconus groups, respectively, to assess the relationship between the magnitudes of keratometric and posterior corneal astigmatism. The latter coefficient was also used to determine the correlation between the two eyes of each patient in the keratoconus group. To distinguish clinical keratoconus from normal eyes, receiver operating characteristic (ROC) curves were calculated and quantified using the area under the curve (AUC) for the mean corneal power and astigmatism of each component. The AUC describes the predictive accuracy of the indices; an area of 1 implies that the test perfectly discriminates between the study groups. Optimal cut-off points to distinguish keratoconus from normal corneas were calculated for the measured parameters. The positive likelihood ratio [sensitivity/(1‑specificity)] and negative likelihood ratio [(1-sensitivity)/specificity] for these cut-off points were determined. A P value of less than 0.05 was considered statistically significant and all P values were two sided.

RESULTS

Table 1 presents and compares demographic and refractive data between the two study groups. In the keratoconus group, mild keratoconus was observed in 41 eyes (28.9%), whereas 46 eyes (32.4%) and 55 eyes (38.7%) had moderate and severe keratoconus, respectively. Table 2 lists and compares the simulated keratometric, posterior, and mean total corneal power and astigmatism between the study groups. Comparisons of the magnitudes of keratometric, posterior, and mean total corneal power and astigmatism among the different keratoconus severity subgroups are presented in Table 3. The table shows that the mean corneal power and astigmatism of the different components were significantly higher in the keratoconus group than in the normal group and increased with the disease stage.

Significant correlations were found between the magnitudes of keratometric and posterior corneal astigmatism in the normal (r=−0.76, P < 0.001) and keratoconus (r=−0.72, P < 0.001) groups. Posterior corneal astigmatism could be predicted from keratometric astigmatism in the normal and keratoconus groups by using the following linear regression equations: posterior corneal astigmatism = −0.171 – 0.129 × keratometric astigmatism and posterior corneal astigmatism = −0.363 – 0.105 × keratometric astigmatism, respectively. In the subgroup analysis, the association between keratometric and posterior corneal astigmatism was weak in the mild keratoconus subgroup (r = −0.38, P = 0.02). However, this association was strong in the moderate (r = −0.87, P < 0.001) and severe (r = −0.86, P < 0.001) keratoconus subgroups.
In the normal group, WTR, ATR, and oblique keratometric astigmatism were found in 156 eyes (76.5%), 20 eyes (9.8%), and 28 eyes (13.7%), respectively. Corresponding figures in the keratoconus group were 103 (72.0%), 5 (3.5%), and 35 (24.5%), respectively ($P = 0.01$). WTR, ATR, and oblique astigmatism of the posterior corneal surface were found in 2 eyes (1.0%), 198 eyes (97.0%), and 4 eyes (2.0%) in the normal group and in 4 eyes (2.8%), 114 eyes (79.7%), and 25 eyes (17.5%) in the keratoconus group, respectively ($P < 0.001$). These values indicate that more eyes showed oblique astigmatism of both the anterior ($P = 0.02$) and posterior ($P < 0.001$) corneal surfaces in the keratoconus group than in the normal group. The subgroup analysis revealed that the axis orientation of the keratometric and posterior corneal surfaces remained unchanged regardless of the disease stage ($P = 0.81$ and 0.92, respectively) [Table 4].

Table 1. Comparisons of demographic and refractive data between normal eyes and eyes with keratoconus.

|                      | Normal group | Keratoconus group | $P$   |
|----------------------|--------------|-------------------|-------|
| Age (years)          | 29.9±5.9 (20 to 54) | 28.7±9.6 (16 to 66) | 0.24  |
| Sex (Female/Male)    | 114/90       | 68/74             | 0.51  |
| Eye (Right/Left)     | 204/0        | 69/73             | <0.001|
| Spherical equivalent refraction (D) | -2.69±1.80 (-13.88 to +0.5) | -6.69±4.77 (-22.5 to -0.75) | <0.001 |
| Refractive astigmatism (D) | -0.77±0.51 (-6.25 to -0.0) | -4.82±2.52 (-12.0 to 0.0) | <0.001 |

Table 2. Comparisons of keratometric, posterior and total corneal mean power and astigmatism (mean±standard deviation; range) between normal eyes and eyes with keratoconus.

|                      | Normal group | Keratoconus group | $P$   |
|----------------------|--------------|-------------------|-------|
| Mean power (D)       | Simulated keratometry 43.73±1.14 (41.09 to 45.98) | 51.27±4.63 (44.90 to 73.30) | <0.001 |
|                      | Posterior    -6.29±0.20 (-6.84 to -5.83) | -7.77±0.87 (-10.33 to -6.34) | <0.001 |
|                      | Total        43.73±1.12 (40.66 to 45.74) | 50.45±4.51 (43.39 to 75.0) | <0.001 |
| Astigmatism (D)      | Simulated keratometry 1.13±0.86 (0.06 to 4.45) | 5.44±2.71 (0.38 to 15.70) | <0.001 |
|                      | Posterior    -0.32±0.15 (-0.82 to -0.08) | -0.94±0.39 (-2.94 to -0.16) | <0.001 |
|                      | Total        1.03±0.82 (0.06 to 4.29) | 5.34±2.87 (0.37 to 20.90) | <0.001 |

Table 3. Comparisons of keratometric, posterior and total corneal mean power and astigmatism (mean±standard deviation; range) among different subgroups of keratoconus severity.

|                      | Mild keratoconus subgroup | Moderate keratoconus subgroup | Severe keratoconus subgroup | $P$   |
|----------------------|---------------------------|-------------------------------|-----------------------------|-------|
| Mean power (D)       | Simulated keratometry 46.49±0.92 (44.90 to 47.89) | 49.92±1.23 (48.0 to 51.90) | 55.96±3.58 (52.10 to 73.30) | <0.001 |
|                      | Posterior                -6.90±0.33 (-8.38 to -6.34) | -7.48±0.33 (-8.25 to -6.57) | -8.65±0.61 (-10.33 to -7.60) | <0.001 |
|                      | Total                    45.90±1.05 (43.39 to 47.75) | 49.24±1.14 (47.30 to 51.17) | 54.84±3.84 (50.40 to 75.0) | <0.001 |
| Astigmatism (D)      | Simulated keratometry -0.81±0.44 (-2.94 to -0.16) | -0.92±0.29 (-1.58 to -0.18) | -1.05±0.40 (-2.12 to -0.23) | 0.02  |
|                      | Posterior                4.01±2.02 (0.37 to 8.10) | 5.03±2.12 (0.39 to 10.40) | 6.58±3.43 (1.96 to 20.90) | <0.001 |

D: diopter

Table 4. Comparison of axis orientations of keratometric and posterior corneal astigmatism among different subgroups of keratoconus severity.

|                      | WTR astigmatism | ATR astigmatism | Oblique astigmatism |
|----------------------|-----------------|-----------------|---------------------|
|                      | N (%)           | N (%)           | N (%)               |
| Keratometric astigmatism |                 |                 |                     |
| Mild keratoconus subgroup | 28 (68.3%) | 1 (2.4%) | 12 (29.3%) |
| Moderate keratoconus subgroup | 35 (76.1%) | 1 (2.2%) | 10 (21.7%) |
| Severe keratoconus subgroup | 40 (72.7%) | 3 (5.5%) | 12 (21.8%) |
| Posterior astigmatism |                 |                 |                     |
| Mild keratoconus subgroup | 1 (2.4%) | 31 (75.6%) | 9 (22.0%) |
| Moderate keratoconus subgroup | 1 (2.2%) | 38 (82.6%) | 7 (15.2%) |
| Severe keratoconus subgroup | 2 (3.6%) | 44 (80.0%) | 9 (16.4%) |

WTR: with the rule; ATR: against the rule; N: number
Figure 1 compares the results of the ROC curve analyses, and Table 5 presents the AUC of keratometric, posterior, and mean total corneal power and astigmatism for the keratoconus group versus the normal group. The AUC was close to 1.0 for all measurements, indicating that they can effectively differentiate keratoconus from normal corneas. The posterior corneal steep meridian had the strongest power to distinguish keratoconus from normal corneas. Table 6 lists the optimal cut-off points and their sensitivity, specificity, and positive and negative likelihood ratios for keratometric, posterior, and mean total corneal power and astigmatism to distinguish keratoconus from normal eyes. As demonstrated, the cut-off points of all parameters except for the flat meridian of total corneal power had high sensitivity and specificity for identifying keratoconus.

**DISCUSSION**

The current study reported the characteristics of the posterior corneal surface measured with a dual Scheimpflug topographer and compared the results of eyes with keratoconus and those with normal corneas. Previous studies have shown that the Galilei Scheimpflug analyzer has excellent repeatability for corneal curvature measurements. Our results indicated that the magnitude of mean posterior corneal power and astigmatism was significantly higher in keratoconus corneas than in normal corneas. A recent study evaluated the posterior corneal astigmatism in keratoconic corneas by using Pentacam.[16] It reported that the mean magnitude of posterior astigmatism was approximately 1.0 D with a range of 0.0–2.90 D, which corresponds to the range of values reported in the present study. Kamiya et al[16] found no statistically significant increase in anterior, posterior, or total corneal astigmatism with the progression of keratoconus. Their finding contrasts with our results, which indicated that the magnitude of the mean posterior corneal power and astigmatism significantly increased with disease progression. This difference can be explained by the fact that Kamiya et al[16] used a different classification of keratoconus severity that accounted for astigmatism, corneal power, corneal transparency, and corneal thickness. It is unclear which is the more reliable method for classification of keratoconus severity.

We found a significantly higher prevalence of oblique astigmatism of both the anterior and posterior corneal surfaces in the keratoconus group than in the normal group. However, in most keratoconic eyes, the axis orientations of the keratometric and posterior corneal astigmatism were WTR and ATR, respectively, and did not change as the disease severity increased. These data suggest that posterior corneal astigmatism provides a
compensatory effect for the astigmatism of the anterior corneal surface in keratoconus eyes irrespective of the stage of the disease.

Kamiya et al\[16\] reported a significant correlation between the magnitudes of anterior and posterior corneal astigmatism in keratoconus. Coupled with these findings, the results of the current study show that the magnitude of posterior corneal astigmatism is significantly related to the magnitude of keratometric astigmatism at the different stages of keratoconus. This association is weak in mild keratoconus but strong in moderate and severe keratoconus. This observation can be attributed to the fact that manifestations of keratoconus occur at the posterior corneal surface in early stages of the disease, when the anterior surface demonstrates subtle topographic changes.\[8-10\] With the progression of the disease, however, an increase in the anterior corneal curvature occurs in parallel with the posterior corneal curvature, which results in a linear relationship between the two surfaces at a constant ratio. The significant correlation between the keratometric and posterior corneal astigmatism is of clinical importance; this correlation allows the estimation of total corneal astigmatism from the keratometric value when the posterior corneal measurement is unavailable.\[22\] Toric intraocular lenses have been developed for the correction of astigmatic errors in normal and keratoconic eyes with cataracts.\[23-26\] The magnitude of corneal astigmatism power determined using anterior corneal measurements should be adjusted based on the posterior corneal astigmatism to improve the accuracy of predicted corneal astigmatism correction achieved by these lenses, particularly in keratoconus eyes in which the role of the posterior cornea is magnified.\[27\] Devices that measure the posterior corneal surface are not available in all eye clinics; the intercept and slope parameters reported in the current study can be used in such situations to predict posterior corneal astigmatism.

The ability of the mean power and astigmatism of the posterior corneal surface to discriminate keratoconus from normal eyes was investigated using the area under the ROC curves. The results indicated that mean posterior corneal power and astigmatism are strong enough to distinguish eyes with keratoconus from normal eyes and the measurements of the posterior corneal steep meridian were the coefficients with the highest discriminative ability. This is congruent with previous studies that have demonstrated that manifestations of keratoconus occur at the posterior corneal surface at early stages of the disease.\[8-10\] Furthermore, the current study attempted to define the cut-off points for posterior corneal surface measurements

### Table 5. Area under the receiver operating characteristics curve (AUC) of keratometric, posterior, total corneal mean power and astigmatism for keratoconus versus normal corneas

|                  | AUC  | 95% confidence interval | P    |
|------------------|------|-------------------------|------|
| **Simulated keratometry** |      |                         |      |
| Mean power       | 0.992 | 0.987 to 0.998           | <0.001|
| Flat meridian    | 0.934 | 0.906 to 0.962           | <0.001|
| Steep meridian   | 0.987 | 0.973 to 1.0             | <0.001|
| Astigmatism      | 0.949 | 0.923 to 0.976           | <0.001|
| **Posterior**    |      |                         |      |
| Mean power       | 0.990 | 0.981 to 0.999           | <0.001|
| Flat meridian    | 0.960 | 0.937 to 0.982           | <0.001|
| Steep meridian   | 0.995 | 0.991 to 1.0             | <0.001|
| Astigmatism      | 0.950 | 0.925 to 0.976           | <0.001|
| **Total**        |      |                         |      |
| Mean power       | 0.986 | 0.975 to 0.996           | <0.001|
| Flat meridian    | 0.915 | 0.881 to 0.948           | <0.001|
| Steep meridian   | 0.994 | 0.989 to 0.999           | <0.001|
| Astigmatism      | 0.951 | 0.925 to 0.977           | <0.001|

### Table 6. Cut-off points of keratometric, posterior and total corneal mean power and astigmatism and their sensitivity, specificity, and positive and negative likelihood ratios (LR) to distinguish eyes with keratoconus from controls

|                  | Cut-off point (D) | Sensitivity (%) | Specificity (%) | LR+   | LR−   |
|------------------|------------------|----------------|----------------|-------|-------|
| **Simulated keratometry** |      |                |                |       |       |
| Mean power       | 45.88            | 92.3%          | 98.5%          | 62.73 | 0.08  |
| Flat meridian    | 44.64            | 85.2%          | 89.7%          | 9.28  | 0.16  |
| Steep meridian   | 46.69            | 96.5%          | 97.5%          | 39.36 | 0.04  |
| Astigmatism      | 2.62             | 88.7%          | 93.1%          | 12.93 | 0.12  |
| **Posterior**    |      |                |                |       |       |
| Mean power       | -6.70            | 93.7%          | 99.5%          | 191.07| 0.06  |
| Flat meridian    | -6.48            | 85.9%          | 96.6%          | 25.04 | 0.15  |
| Steep meridian   | -6.85            | 97.2%          | 97.5%          | 39.65 | 0.03  |
| Astigmatism      | -0.54            | 88.0%          | 91.7%          | 10.56 | 0.13  |
| **Total**        |      |                |                |       |       |
| Mean power       | 45.69            | 89.4%          | 99.5%          | 182.45| 0.11  |
| Flat meridian    | 45.04            | 73.2%          | 98.0%          | 37.35 | 0.27  |
| Steep meridian   | 46.47            | 94.4%          | 99.0%          | 96.25 | 0.06  |
| Astigmatism      | 2.67             | 88.0%          | 94.6%          | 16.33 | 0.13  |
to differentiate keratoconus and normal eyes. The values provided in this study can be used in clinical settings, particularly for the screening of keratoconus in refractive surgery candidates. For example, we found that posterior corneal astigmatism >0.54 D indicates keratoconus with a sensitivity and specificity of 88% and 92%, respectively. Naderan et al.\(^9\) reported a cut-off value of 0.4 D for posterior corneal astigmatism with 89.5% sensitivity and 85.0% specificity for discriminating keratoconus from normal corneas. Posterior corneal astigmatism >0.50 D has been reported in 9% and 12.5% of normal corneas.\(^{28,29}\)

In the current study, this rate was 11.3% in the normal group and 88.7% in the keratoconus group.

There were two limitations in this study. First, two eyes of the same patient were evaluated in some cases in the keratoconus group. We attempted to exclude the confounding effects of potential correlations between measurements from both corneas of the same patient by using generalized estimating equation models and partial correlation coefficients. Second, this study included patients with clinical keratoconus and excluded patients with subclinical forms. A method of distinguishing keratoconus from normal eyes is more valuable when it can also identify subclinical forms with acceptable sensitivity and specificity. Additionally, the threshold of cut-off points may vary if subclinical forms of keratoconus are considered.

CONCLUSION

In conclusion, since posterior corneal imaging became possible, it has received increasing clinical interest, particularly for the accurate correction of astigmatic errors using toric intraocular lenses. Posterior corneal measurements using a dual Scheimpflug camera seem to have potential in diagnosing keratoconus patients. Additional studies are advisable to confirm the usefulness of the power of posterior corneal surface measurements for the identification of keratoconus and subclinical keratoconus.

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Conflicts of Interest
There are no conflicts of interest.

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