

# Intraoperative Swept-Source OCT–Based Corneal Topography for Measurement and Analysis of Stromal Surface After Epithelial Removal

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## ABSTRACT

**PURPOSE:** To assess intraoperative stromal topography measurements using swept-source optical coherence tomography (OCT)–based topography/tomography after epithelial removal and to analyze the epithelial contribution to the corneal topography and optics.

**METHODS:** This was a prospective series of 22 eyes of 19 patients referred to receive phototherapeutic keratotomy (PTK) for treatment of recurrent corneal erosion and a control group of 22 virgin eyes. Swept-source OCT corneal topography/tomography was obtained immediately before and immediately after mechanical deepithelialization before PTK. Epithelial thickness maps were obtained before the surgery using spectral-domain OCT in the control group and as a reference in the group with anterior basement membrane dystrophy. Topographic and optical characteristics, including the curvature, astigmatism, asphericity, and higher order aberrations of the cornea before and after deepithelialization were compared, and their differences correlated with the measurements derived from the epithelial thickness maps.

**RESULTS:** Stromal topography measurements after deepithelialization were easily obtained and showed excellent repeatability. Assessment of corneal edema induced by deepithelialization revealed that it did not significantly affect the measured parameters. The stromal surface was steeper by 1.28 diopters, had higher with-the-rule astigmatism by 0.41 diopters, was more prolate, and had more higher order aberrations compared to the intact epithelialized corneal surface. These differences correlated well with the parameters derived from epithelial thickness maps.

**CONCLUSIONS:** Measurement of stromal topography using swept-source OCT immediately after mechanical deepithelialization may be a viable method in therapeutic refractive surgery, where stromal topography-guided ablation is needed. A significant epithelial contribution to anterior corneal topography and optics was confirmed.

As the refractive medium with direct contact with the air, the corneal epithelium plays a significant role in the ocular refractive system. Even if the epithelium had a uniform thickness profile and a refractive index identical to that of the stroma, its sheer removal would increase the refractive power of the cornea due to the shortening of the radius of curvature.<sup>1</sup> In fact, the corneal epithelium has a higher refractive index (1.401) than the anterior stroma (1.380), a difference sufficient to affect overall corneal power.<sup>2</sup> Additionally, the epithelium has a non-uniform thickness profile due to the effect of eyelid blinking mechanics,<sup>3,4</sup> and the inherent behavior of epithelial cells to smooth the underlying stromal surface irregularities.<sup>4-8</sup> The epithelial thickness profile can produce myopic, hyperopic, astigmatic, and higher order aberration (HOA) shifts in the corneal optics.<sup>4,9</sup> The epithelial thickness profile can also be influenced by other factors, including anterior basement membrane dystrophy (ABMD)<sup>10</sup> and dry eye.<sup>11</sup> Epithelial remodeling may be sufficient to fully mask the stromal surface irregularity, as has been shown in the detection of keratoconus with normal topography.<sup>3,4,8,12</sup> This compensatory function means that anterior surface topography can be poorly representative of underlying stromal irregularity, resulting in poor outcomes of topography-guided therapeutic surgery and pointing to the need for transepithelial phototherapeutic keratotomy (PTK) and topography-guided ablation to achieve stromal regularization.<sup>9,13-16</sup> Reinstein et al<sup>17</sup> used epithelial thickness maps to generate stromal topography and performed the first stromal topography-guided treatment, whereas Vinciguerra et al<sup>18,19</sup> used intraoperative stromal topography to plan sequential stromal topography-guided treatment. Studies using either direct measurement of the corneal stromal surface immediately after epithelial debridement,<sup>1,20-23</sup> mathematically modeled stromal/Bowman's layer surface,<sup>24</sup> or Bowman's layer topography constructed by edge detection of the surface of Bowman's layer on 12 radial corneal swept-source optical coherence tomography (SS-OCT) B-scans<sup>25</sup> have also shown variation in epithelial and stromal topographic and optical characteristics.

The purpose of the current study was to assess the viability of SS-OCT technology for intraoperative measurement of stromal topography after epithelial removal. We also intended to compare the corneal anterior and stromal topographic and optical characteristics acquired by SS-OCT-based topography between the eyes with ABMD and the control group of virgin eyes, as well as to quantify the corneal epithelial contribution to the corneal topography and optics in the ABMD group.

## PATIENTS AND METHODS

This prospective study included 22 eyes of 19 patients with ABMD referred for PTK for treatment of recurrent corneal erosion at the Eye Department of the University Hospital of North Norway from April 2015 to March 2019. A control group of 22 healthy virgin eyes of 22 consecutive candidates for elective laser vision correction at the SynsLaser clinic in Tromsø, Norway.

Only eyes with intact epithelium on slit-lamp examination, normal epithelial distribution on epithelial thickness map, and no history of corneal erosion at least 1 month before the PTK surgery were included. Patients presenting with ocular pathology other than the history of recurrent corneal erosions and eyes that had undergone previous ocular surgery were excluded from the study. Soft contact lens users were instructed to stop using the lenses 1 week before the measurements; no hard contact lens users were included. The study was approved by the Norwegian Regional Committee for Medical & Health Research Ethics (REK-Nord 2014/1401). All patients provided informed consent for the anonymous use of their data in scientific analyses and publications. Standard preoperative ocular examination, including slit-lamp biomicroscopy, visual acuity/subjective manifest refraction, and fundus examinations were performed in all eyes. An SS-OCT-based topographer/tomographer (Casia SS-1000; Tomey Corporation) was used for corneal topography and a spectral-domain OCT (SD-OCT) scanner (RTVue-100; Optovue, Inc) was used for the epithelial thickness maps. To limit the influence of overnight swelling,<sup>26</sup> the measurements were performed at least 4 hours after the patients awoke.

The Casia SS-1000 was located approximately 1 meter away from the surgery bed. Initial topography with SS-OCT was obtained before the application of 0.5% proparacaine anesthetic drops (Alcaine; Alcon

Laboratories, Inc). The patients were asked to blink three times before each of the three acquisitions, and the average value was used for further analyses. The epithelium was then removed completely over the central 9 mm of the corneal surface using an electric epithelial scrubber (Amoils brush; Innovative Excimer Solutions). The surgeon exercised special care in removing the epithelium in the gentlest but most thorough manner to leave an intact and smooth Bowman's membrane. The corneal surface was rinsed with a balanced salt solution (BSS; Alcon Laboratories, Inc) and inspected under the microscope to ensure that all epithelial debris had been removed. After the deepithelialization and before the final PTK procedure, the patients were asked to move to the Casia SS-1000 again to undergo another three acquisitions of the topographies. The patients were asked to blink before each acquisition. No artificial tears were necessary to enhance the examination and the corneas were not dried before the acquisition. The raw OCT data and the instrument's edge-detection processing of each image were checked for artifacts. A bandage contact lens (Acuvue Oasys; Johnson & Johnson) was placed over the eye after the PTK procedure had been completed.

## Measurements

The Casia SS-1000 automatically moves the chinrest until the patient's eye reaches the correct position and initiates 0.34-second long acquisition, with the reference axis of the measurements aligned with the corneal vertex.<sup>27,28</sup> The analysis software identifies the anterior and posterior corneal surfaces and generates the corneal pachymetry, elevation, curvature, and refractive maps. We compared the following topographic and optical parameters of the anterior corneal (epithelium-covered) surface and the stromal (deepithelialized) surface: (1) radius of the best-fit sphere (BFS) at the 3.5-, 6-, and 9-mm zones; (2) radius of the axial curvature and its orientation along the steepest and flattest meridians at the 3.5-mm zone; (3) corneal asphericity (mean Q-value) measured over the central 6-mm zone; and (4) Fourier HOAs measured over the central 3- and 6-mm zones.

In addition, the BFS radius of the posterior corneal surface at the 3.5-, 6-, and 9-mm zones, and the pachymetry at the corneal vertex before and after deepithelialization were also analyzed. The amount of edema induced by the deepithelialization was expressed as the difference between the reduced pachymetry after deepithelialization and the epithelial thickness measured by SD OCT preoperatively (amount of central edema = preoperative central epithelial thickness – [central pachymetry<sub>epi-on</sub> – central pachymetry<sub>epi-off</sub>]).

Preoperative epithelial thickness map measurements by SD-OCT, centered on the corneal vertex, covering the central 6-mm diameter zone, were also performed. Three consecutive acquisitions were performed for each eye. Auto-generated averaged regional epithelial thickness was recorded for the following areas: central 2 mm and paracentral 2- to 5-mm ring divided into superior, superior temporal, temporal, temporal inferior, inferior, inferior nasal, nasal, and superior nasal sectors.

## Contribution of the Epithelium to the Refractive Power

To evaluate the dioptric refractive power contribution of the epithelium, radii of the anterior cornea with epithelium-on and the stromal surface after epithelial removal were converted to power. According to the law of Gaussian optics, the following formula was used for the two interfaces:  $\text{Power} = (n' - n)/r$ , where  $n$  and  $n'$  are the refractive index of the media in front and behind the refractive interface, respectively, and  $r$  is the radius of the interface curvature measured by the Casia SS-1000. The refractive indexes used for the epithelium and anterior stroma were 1.401 and 1.380, respectively.<sup>2</sup> Considering the epithelium as a thin refractive lens with two refractive interfaces (air–epithelium interface and epithelium–stroma interface), the power of the “epithelium lens” residing between air and the stroma is represented by  $\text{power}_{\text{air-epithelium}} + \text{power}_{\text{epithelium-stroma}} - \text{power}_{\text{air-stroma}}$ .

## Evaluation of Astigmatism

The axial radii and their orientation along the steepest and flattest meridians within the anterior diameter of 3.5 mm, with and without epithelium, were recorded for evaluating astigmatism. The radii were converted to power values using the law of Gaussian optics. The axis and power of astigmatism were calculated for the air–epithelium interface, epithelium–stroma interface, and air–stroma interface. The astigmatism values were then decomposed into two Jackson cross-cylinder lenses, one with power  $J_0 = (-C/2) \cos(2a)$  at axis  $a = 0 = 180$  degrees, and the other with power  $J_{45} = (-C/2) \sin(2a)$  at axis  $a = 45$  degrees, for further comparison.<sup>29</sup> In this representation, an increase in  $J_0$  indicates an increased component of with-the-rule (WTR) astigmatism or decreased component of against-the-rule (ATR) astigmatism, and vice versa. A change in  $J_{45}$  indicates a change in oblique astigmatism. The contribution of the epithelium to astigmatism was calculated as:

$$\sqrt{(J_0 + J_0' - J_0'')^2 + (J_{45} + J_{45}' - J_{45}'')^2}$$

where  $J_0$  and  $J_{45}$ ,  $J_0'$  and  $J_{45}'$ , and  $J_0''$  and  $J_{45}''$  represent the Jackson cross-cylinder components of astigmatism for the air–epithelium, epithelium–stroma, and air–stroma interface, respectively. For vector summation, air–epithelium and air–stroma interface astigmatism were defined using the negative magnitude and flat axis. Because the epithelium–stroma interface has a divergent optical effect, epithelium–stroma interface astigmatism was also defined with a negative magnitude but using the steep axis for vector addition.

### Statistical Analysis

Statistical analysis was performed using SPSS Statistics 24 (IBM Corporation). The normal distribution of continuous variables was confirmed using the Kolmogorov-Smirnov test and Q-Q plot. The paired *t* test and related-samples Wilcoxon signed-rank test were used for comparing the epithelium-on and epithelium-off parameters with normal and non-normal distribution, respectively. The Pearson correlation and Spearman rank correlation were used for defining the correlation between parameters with normal and non-normal distribution, respectively. A *P* value of less than .05 was considered statistically significant.

## RESULTS

The 22 topographies before and after deepithelialization, and the preoperative epithelial thickness maps, were from 7 women (36.8%) and 12 men (63.2%). The mean patient age was  $46.21 \pm 13.18$  years (range: 25 to 70 years). The measured parameters and calculated values for the epithelial and stromal surfaces are listed in Table 1. The patients' demographic and topographic data were comparable to those of the 22 normal virgin eyes in the control group (Table A, available in the online version of this article). The correlation between the deepithelialization-induced changes and the epithelial thickness in different areas are listed in Table B (available in the online version of this article). Preoperative epithelial thickness distribution over the central 5-mm diameter of the ABMD and control groups is shown in Figure 1. The average epithelial thickness profile was similar between groups, with both showing thicker epithelium inferiorly compared to superiorly. This difference was greater in the ABMD group (6.00  $\mu\text{m}$ ) compared to the control group (2.08  $\mu\text{m}$ ). The epithelial thickness was thicker in the ABMD group in the central and inferior zones, as shown in Figure 2.

The best-fit sphere (BFS) radius of the anterior corneal surface (epithelium-covered) was significantly larger at both the 3.5- and 6-mm diameter zone compared to that of the stromal surface (measured after deepithelialization), whereas the opposite was true at the 9-mm diameter zone. The difference in the BFS radius between the two surfaces at the 3.5-mm diameter showed a statistically significant correlation with the paracentral 2 to 5 mm epithelial thickness ( $P = .011$ ,  $R = 0.571$ ). Compared to the stromal surface, the anterior corneal surface had lower prolateness at the 6-mm diameter zone, and lower HOAs at both the 3- and 6-mm diameter zone. Asphericity between the epithelial and stromal surfaces was correlated with paracentral epithelial thickness ( $P = .039$ ,  $R = 0.477$ ). The difference in HOAs at the 3-mm diameter zone was correlated with the standard deviation of the epithelial thickness profile ( $P = .041$ ,  $R = -0.472$ ).

There was no statistically significant change in the BFS radius of the posterior corneal surface at the 3.5- and 6-mm diameter zone after epithelial debridement, whereas the BFS radius at the 9-mm diameter zone was significantly increased after deepithelialization ( $P < .001$ ).

The astigmatism contribution of the “epithelium lens” between the air and the stroma was  $-0.25 \text{ D} \times 7$ , whereas the astigmatism for the air–stroma interface after deepithelialization was  $-1.04 \text{ D} \times 178$ , yielding epithelium-induced astigmatism of  $-0.76 \text{ D} \times 85$  (Figure 3). The average central epithelial thickness before debridement was  $57.42 \pm 3.55 \text{ }\mu\text{m}$ , whereas the mean variation of central corneal thickness before and after deepithelialization was  $37.95 \pm 7.95 \text{ }\mu\text{m}$ , indicating an average central stromal thickening of  $19.47 \pm 7.09 \text{ }\mu\text{m}$ , which was interpreted as stromal edema. The stromal edema was not correlated with any of the measured parameters on the anterior cornea, whereas the amount of flattening of the posterior cornea at the 9-mm diameter zone after deepithelialization showed a significant correlation with the amount of stromal edema ( $P = .027$ ,  $R = 0.482$ ).

The intraclass correlation coefficients of three consecutive measurements for each anterior curvature parameter before and after epithelial removal were high, ranging between 0.993 and 0.999, and are summarized in Table C (available in the online version of this article).

## DISCUSSION

We found SS-OCT technology capable of producing measurements of stromal topography after epithelial removal, as well as repeatable and convenient for use in therapeutic refractive surgery in cases where intraoperative stromal mapping after deepithelialization is necessary. We also looked at the epithelial contribution to the corneal optics by studying SS-OCT topographies before and after deepithelialization in eyes with ABMD. We found that the epithelial contribution to the total corneal power consisted of negative optical power, positive asphericity, and HOAs that are inverse to the stromal aberrations. These findings are in accord with the literature,<sup>22,23</sup> as is the comparison of the epithelial thickness maps between eyes with ABMD and virgin eyes, showing thicker central and inferior epithelium in the ABMD group.<sup>10</sup>

As Reinstein et al<sup>4</sup> described in 2008, epithelium in normal eyes is not a layer with uniform thickness and there is also a variation in thickness distribution between two eyes of the same person. The epithelial thickness was found to be distributed according to the stromal surface curvature gradient and the template provided by the eyelid.<sup>4,13,15,16,30-32</sup> The superior-inferior asymmetry in the average epithelial thickness profile found in the current study was similar to that previously reported and the thinner superior epithelium is most likely due to the influence of the upper eyelid.

To study the topographic and optical contribution of the epithelium, we used direct stromal topography obtained immediately after deepithelialization and correlated it to the anterior corneal topography obtained by the same device right before deepithelialization. It is difficult to accurately measure the stromal topography of newly deepithelialized cornea via Placido-based, scanning-slit-based,<sup>33</sup> or PAR Corneal Topography,<sup>34,35</sup> The reason may be that these technologies use an intense visible examination light, which causes photophobia, involuntary ocular movements, and hence improper patient fixation after deepithelialization. In the current study, the deepithelialized corneas were exposed to SS-OCT infrared light of 1,300-nm wavelength, which is not visible to the patient. It also allows better penetration into an opaque cornea<sup>36</sup> compared to visible light. The short topography acquisition time additionally reduces patient discomfort and may contribute to minimizing motion artifacts resulting from ocular movement. Finally, the instrument uses a quick software-controlled alignment system, which automatically moves the unit to the measurement position according to the corneal reflex,<sup>28</sup> making examination possible even under the most difficult conditions. It should be noted that the tear film, which is normally registered as the first corneal surface by OCT-based topography, has a smoothing effect on the corneal curvature, introducing a potential measurement error, especially in highly irregular cases.

In 1993, Simon et al<sup>1</sup> published an evaluation of surface topography before and after deepithelialization to determine epithelial power in human cadaver eyes. Measurement of corneal topographic changes after epithelial debridement in normal myopic eyes or keratoconic eyes has also been reported based on the use of either high-frequency digital ultrasound,<sup>24</sup> anterior elevation,<sup>17</sup> or Placido topography.<sup>20-22</sup> The Triton SS-OCT scanner (Topcon, Inc) has also been used for noncontact quantification of topography of the anterior corneal surface and Bowman's layer.<sup>25</sup> Most of these studies concluded that the epithelium tends to reduce corneal curvature,<sup>1,21,22,24,25</sup> magnitude of astigmatism,<sup>20,22,25</sup> irregularities,<sup>20,21</sup> and prolateness.<sup>1,20,22</sup> Our results concur, demonstrating that the epithelium-on corneal surface was significantly flatter, and had lower magnitude of astigmatism and decreased HOAs and prolateness compared to the measurements after deepithelialization. These differences correlated well with the parameters derived from epithelial thickness maps in the same group of eyes obtained before deepithelialization (Table B). Figure A (available in the online version of this article) shows the difference between the elevation topographies before and after deepithelialization alongside the epithelial thickness map of the same eye before deepithelialization. The epithelial profile can be seen to account for the asymmetric difference in elevation topography. Our general conclusions concerning the stromal topography and epithelial contribution to the corneal optics for both groups in the current study were in agreement and comparable with previously published similar studies.<sup>1,20-22,24</sup>

Stromal edema was seen to occur after epithelial debridement in all of our cases, with a mean central increase in stromal thickness value of  $19.47 \pm 7.09 \mu\text{m}$ . Gatinel et al<sup>20</sup> reported a central corneal thickness difference of  $37.84 \pm 9.82 \mu\text{m}$  after deepithelialization, which amounts to approximately  $15 \mu\text{m}$  of stromal edema, if subtracted from the average epithelial thickness of  $53 \mu\text{m}$ , as published by Reinstein et al<sup>3</sup> using the gold standard of very high-frequency digital ultrasound. Potential measurement errors in curvature measurements caused by stromal edema were a concern, but we found that changes in central stromal thickness were not correlated with the measured curvature parameters of the anterior cornea; However, we did find a significant correlation with the amount of posterior corneal flattening at the 9-mm diameter zone. This result is in accordance with the findings of Meek et al,<sup>37</sup> who reported that fluid entering the cornea causes more swelling in the posterior lamellae than in the anterior lamellae, explained by the lamellar interweaving and their insertion into Bowman's layer, as well as the differences in the glycosaminoglycans in the anterior and posterior stroma. In addition, *in vitro* studies have shown that substantial corneal hydration may cause statistically significant but clinically minimal change in the anterior corneal curvature.<sup>38,39</sup>

One inclusion criterion in the ABMD group was a healthy epithelium on slit-lamp examination and normal epithelial thickness distribution on epithelial thickness map. The recurrent erosion syndrome group had symptoms of recurrent corneal erosion without signs. Our control group of healthy virgin eyes with no symptoms of recurrent erosion was recruited from candidates for elective laser vision correction. Table 2 shows no significant difference between the groups in terms of preoperative central pachymetry, radius of anterior BFS, and anterior asphericity. However, the group representing eyes with ABMD showed thicker epithelium generally and a greater inferior-superior difference, as shown on Figures 1-2. Buffault et al<sup>10</sup> recently published similar findings. This observation is as expected: ABMD results in duplication of the basement membrane of the epithelium with abnormalities and alteration in the healthy columnar organization from cuboidal basal cells to wing-cells and then flattened squamous cells. It is thus expected that the epithelial thickness profile of eyes with ABMD will be different from that of a normal cornea.

Current surface ablation treatments are performed on the deepithelialized stromal surface, whereas the ablation parameters have been based on the corneal topography and refractive measurements before deepithelialization. The differences in topography of the two surfaces and the optical influence of the epithelium have been modelled to result in ablation errors, the average of which is nomogram-compensated in normal regular corneas.<sup>40,41</sup> In the case of irregular corneas, the epithelium will be even more irregular, explaining why transepithelial topography-guided procedures are more effective for repairing irregular corneas than stromal ablations based on topography alone.<sup>14-16,42</sup> In transepithelial topography-guided surgery, the transepithelial PTK component treats the irregularity masked by the epithelium and the topography-guided

component treats the remainder. Also, the empirically derived preset ablation rate and epithelial thickness compensations may be a source of ablation depth errors<sup>43</sup> in irregular cases. Transepithelial topography-and epithelial thickness map-guided PRK may provide improved accuracy, although there will be a high demand on the treatment laser's scanning resolution and its tracking precision that may be beyond the currently available technology. Transepithelial PTK does not require this level of tracking accuracy. Alternatively, direct stromal topography-guided treatments have been demonstrated. In the first, Reinstein et al<sup>17</sup> obtained stromal topography maps by subtracting the very high-frequency digital ultrasound-derived epithelial thickness profile from the corneal front surface elevation data, derived from scanning slit tomography (Orbscan; Bausch & Lomb), and used this to perform a custom stromal surface topography-guided laser in situ keratomileusis (LASIK) enhancement following a short flap LASIK complication. The second method described by Vinciguerra et al<sup>19</sup> used intraoperative stromal topography measurements after deepithelialization to perform stromal topography-based wavefront-guided ablation.

Anterior segment OCT technology has lately undergone rapid development.<sup>44</sup> Current SD-OCT instruments produce good epithelial thickness maps, but not topography, whereas the SS-OCT instruments produce excellent topography, but not epithelial thickness maps. Currently, the only hybrid topographer/tomographer is the MS-39 (CSO). The MS39 obtains the front surface topography by combining the instantaneous Placido ring-derived surface with auto-correlated rotational corneal SD-OCT scanning technology, allowing the back surface to be calculated by using the highly accurate thickness measurements "hanging" from the aforementioned anterior surface. A second device, an ultra-thin blue laser scanner imaging topographer/tomographer (Precisio2; iVIS), is currently able to deliver both topography and epithelial thickness mapping. These devices may be in an excellent position to allow for stromal topography-guided ablation planning.

Noncontact quantification of topography of Bowman's layer on the intact corneas using SS-OCT<sup>25</sup> and SD-OCT<sup>45</sup> has already been reported. We have demonstrated the viability of stromal topography measurements with SS-OCT measured intraoperatively on deepithelialized stroma, a method appropriate for use in therapeutic refractive surgery in cases with significant epithelial opacity and/or defects where deepithelialization is planned as part of the treatment procedure.

### AUTHOR CONTRIBUTIONS

Study concept and design (WZ, XC, TPU, AS); data collection (WZ, YF); analysis and interpretation of data (WZ, DZR, TJA, AS); writing the manuscript (WZ, YF); critical revision of the manuscript (WZ, XC, DZR, TJA, TPU); statistical expertise (WZ, DZR, TJA); supervision (XC, TPU, AS)

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doi:10.3928/1081597X-20200122-01

TABLE 1

| Parameter                                      | Difference in Measurements Before and After Epithelial Removal |  | P      |
|--|--|--|--------|
|  | Before Epithelial Removal, Mean<br>± SD                        | After Epithelial Removal, Mean<br>± SD |        |
| Radius of BFS of anterior surface (mm)         |  |  |        |
| 3.5 mm   | 7.70 ± 0.39  | 7.58 ± 0.37                            | < .001 |
| 6 mm   | 7.82 ± 0.34  | 7.76 ± 0.36                            | < .001 |
| 9 mm   | 7.93 ± 0.34  | 8.01 ± 0.34                            | < .001 |
| Radius of BFS of posterior surface (mm)        |  |  |        |
| 3.5 mm   | 6.59 ± 0.35  | 6.58 ± 0.32                            | .687   |
| 6 mm   | 6.63 ± 0.31  | 6.63 ± 0.30                            | .702   |
| 9 mm   | 6.77 ± 0.30  | 6.80 ± 0.29                            | < .001 |
| Fourier higher order aberrations (μm)          |  |  |        |
| 3 mm   | 0.16 ± 0.07  | 0.27 ± 0.15                            | < .001 |
| 6 mm   | 0.18 ± 0.07  | 0.33 ± 0.19                            | < .001 |
| Components of anterior corneal astigmatism (D) |  |  |        |
| J0   | 0.12 ± 0.47  | 0.52 ± 0.7                             | .001   |
| J45  | 0.03 ± 0.43  | -0.03 ± 0.52                           | .281   |
| Anterior asphericity (6 mm)                    | -0.25 ± 0.35   | -0.63 ± 0.45                           | .006   |

SD = standard deviation; BFS = best-fit sphere

TABLE A

Demographic and Topographic/Tomographic Data in Main and Control Group

| Parameter                      | Main Group                         |               | Control Group                  |                | P     |
|--------------------------------|------------------------------------|---------------|--------------------------------|----------------|-------|
|                                | Mean ± SD                          | Range         | Mean ± SD                      | Range          |       |
| Age (y)                        | 46.21 ± 13.18                      | 25 to 70      | 30.23 ± 5.85                   | 20 to 39       | < .05 |
| Gender                         | 36.8% female and 63.2% male        |               | 40.91% female and 59.09% male  |                |       |
| Eye                            | 59.1% right eye and 40.9% left eye |               | 50% right eye and 50% left eye |                |       |
| Spherical equivalent (D)       | -0.55 ± 1.35                       | -4.38 to 1.38 | -3.00 ± 1.25                   | -5.13 to -1.00 | < .05 |
| Central pachymetry (μm)        | 545.06 ± 35.46                     | 478 to 596    | 535.32 ±<br>27.82              | 489 to 599     | .32   |
| Radius of anterior BFS at 9 mm | 7.93 ± 0.34                        | 7.26 to 8.71  | 7.81 ± 0.27                    | 7.42 to 8.42   | .22   |
| Anterior asphericity (6 mm)    | -0.25 ± 0.35                       | -0.90 to 0.69 | -0.07 ± 0.46                   | -0.57 to 0.73  | .15   |

SD = standard deviation; BFS = best-fit sphere

TABLE B

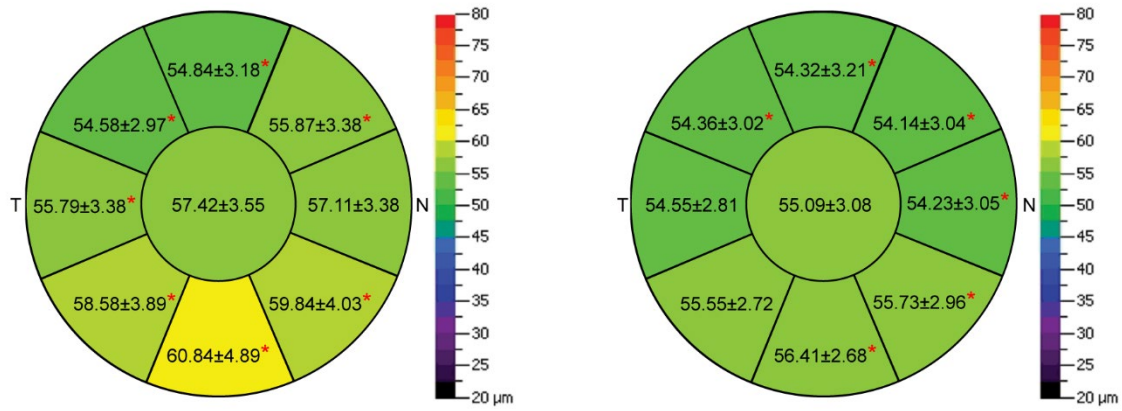
Correlation Between the Epithelium Removal-Induced Changes and the Epithelial Thickness in Different Areas

| Parameter                        | Epithelial Thickness |   |   |                   |                   |
|----------------------------------|----------------------|---|---|-------------------|-------------------|
|                                  | Central 2 mm         | Paracentral 2-5 mm                          | V-Ha                                    | SD of Epib        | Edema             |
| Radius of anterior BFS at 3.5 mm | P = .061             | P = .011, R = 0.571, R <sup>2</sup> = 0.326 | P = .38                                 | P = .184          | P = .305          |
| Radius of anterior BFS at 6 mm   | 0.569                | 0.435                                       | 0.200                                   | 0.765             | 0.827             |
| Radius of anterior BFS at 9 mm   | 0.425                | 0.534                                       | 0.289                                   | 0.262             | 0.426             |
| Anterior asphericity             | 0.131                | 0.039, R = 0.477, R <sup>2</sup> = 0.228    | 0.625                                   | 0.237             | 0.433             |
| Radius of posterior BFS at 9 mm  | 0.331                | 0.240                                       | 0.969                                   | 0.171             | 0.027, R = -0.482 |
| J0                               | 0.757                | 0.608                                       | 0.02, R = -0.529, R <sup>2</sup> = 0.28 | 0.842             | 0.948             |
| HOAs at 3 mm                     | 0.558                | 0.283                                       | 0.657                                   | 0.041, R = -0.472 | 0.219             |
| HOAs at 6 mm                     | 0.751                | 0.994                                       | 0.369                                   | 0.136             | 0.782             |

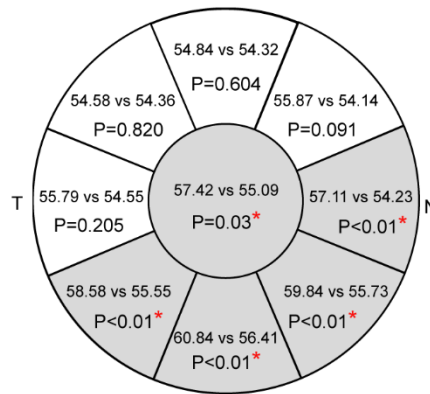
BFS = best-fit sphere; HOAs = higher order aberrations; Difference in average epithelial thickness along vertical and horizontal meridian within 5-mm diameter. bStandard deviation of central 5-mm epithelial thickness.

| Parameter                                      | Epithelium-on |                | Epithelium-off |                |
|--|---------------|----------------|----------------|----------------|
|  | ICC           | 95% CI         | ICC            | 95% CI         |
| Radius of anterior BFS at 3.5 mm               | 0.993         | 0.985 to 0.997 | 0.996          | 0.988 to 0.999 |
| Radius of anterior BFS at 6 mm                 | 0.998         | 0.988 to 0.999 | 0.998          | 0.995 to 0.999 |
| Radius of anterior BFS at 9 mm                 | 0.999         | 0.997 to 0.999 | 0.999          | 0.998 to 0.999 |
| Radius of posterior BFS at 3.5 mm              | 0.994         | 0.988 to 0.998 | 0.995          | 0.989 to 0.998 |
| Radius of posterior BFS at 6 mm                | 0.999         | 0.998 to 1.00  | 0.999          | 0.997 to 0.999 |
| Radius of posterior BFS at 9 mm                | 0.999         | 0.999 to 1.000 | 0.998          | 0.997 to 1.000 |
| Anterior asphericity                           | 0.902         | 0.997 to 1.000 | 0.871          | 0.756 to 0.941 |
| HOAs at 3 mm                                   | 0.84          | 0.707 to 0.924 | 0.788          | 0.602 to 0.902 |
| HOAs at 6 mm                                   | 0.874         | 0.768 to 0.942 | 0.917          | 0.836 to 0.963 |
| Vertex pachymetry                              | 0.813         | 0.662 to 0.910 | 0.995          | 0.90 to 0.998  |
| Axial radius along steepest meridian at 3.5 mm | 0.997         | 0.993 to 0.999 | 0.992          | 0.984 to 0.997 |
| Axial radius along flattest meridian at 3.5 mm | 0.994         | 0.987 to 0.997 | 0.992          | 0.982 to 0.996 |
| Axis of steepest meridian at 3.5 mm            | 0.967         | 0.922 to 0.986 | 0.884          | 0.768 to 0.950 |

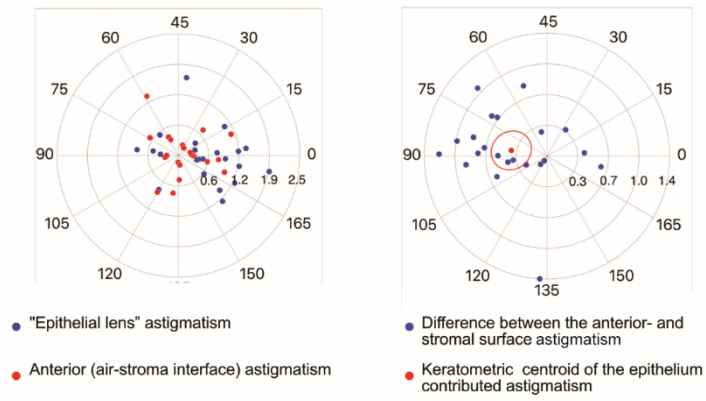
*ICC = intraclass correlation coefficient; OCT = optical coherence tomography;  
BFS = best-fit sphere; HOAs = higher order aberrations*



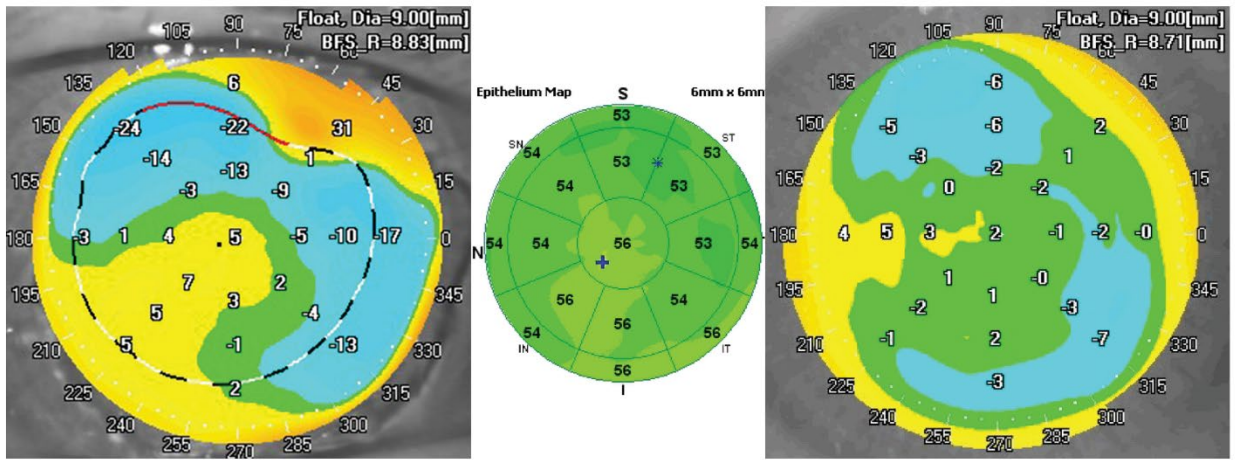
**Figure 1.** Preoperative epithelial thickness distribution over the central 5-mm diameter in the (Left) main group and (Right) control group.



**Figure 2.** Mean epithelial thickness in eyes with anterior basement membrane dystrophy versus virgin eyes (22 eyes each) with *P* values (red asterisks denote the sections where epithelium in the eyes with anterior basement membrane dystrophy is significantly thicker).



**Figure 3.** Double-angle plots representing “epithelial lens” astigmatism and stromal surface astigmatism (left) and difference between corneal anterior and stromal surface astigmatism with keratometric centroid of the “epithelial lens” astigmatism (right).



**Figure A.** Corneal elevation topographies of the same eye before (left) and after (right) epithelial removal, as well as the corneal epithelial thickness map (center) taken before the epithelial removal