Ultra-high-Resolution Measurement by Optical Coherence Tomography of Dynamic Tear Film Changes on Contact Lenses

Qi Chen, Jianhua Wang, Atizhu Tao, Meixiao Shen, Shuliang Jiao, and Fan Lu

PURPOSE. To determine the dynamic pre- and postlens tear film (PLTF and PoLTF) thicknesses by using optical coherence tomography (OCT).

METHODS. Ultra-high-resolution OCT was used to image the tear film of 22 subjects before and after contact lens wear. A soft lens with 1 drop of artificial tears on its concave surface was inserted onto one randomly selected eye. OCT images were taken before insertion, immediately afterward, and every 2 minutes for 10 minutes. For the contralateral eye, the lens inserted was not prewetted on the concave surface. OCT images were taken before insertion, immediately afterward, and at 3 minutes. Then another drop was instilled, and images were taken immediately afterward and every 2 minutes for 10 minutes. Images were processed by custom software to yield tear film thickness.

RESULTS. The thickness of precorneal tear film (PCTF) was 1.9 ± 0.9 μm. The PoLTF was visualized clearly in all cases immediately after lens insertion, with 1 drop on the lens concave surface. Through the first 6 minutes after insertion, the PoLTF was greater than the PCTF. The PLTF (n = 12) and PoLTF (n = 9) were visualized immediately after lens insertion. After 3 minutes, the PLTF in most subjects and PoLTF in all subjects were invisible. The thickness of the PLTF increased after the instillation of artificial tears, whereas the PoLTF did not at any checkpoint for 10 minutes.

CONCLUSIONS. Ultra-high-resolution OCT is a promising tool for measuring the tear film. The PoLTF did not increase after instillation of artificial tears. (Invest Ophthalmol Vis Sci. 2010; 51:1988–1993) DOI:10.1167/iovs.09-4389

Determining the thickness of the precorneal, prelens, and postlens tear films (PCTF, PLTF, and PoLTF) is indispensable for understanding the normal physiology of the tear film and the etiology of dry eye associated with contact lens wear. More than 50% of contact lens wearers experience ocular dryness and discomfort, and some of them abandon lens wear, as intolerable dry eye symptoms affect daily life. The continuous exchange of the tears underneath the contact lens is needed for maintaining ocular health. Accordingly, the PoLTF is considered to be largely responsible for contact lens–associated dry eye, although the PLTF also contributes to it. Therefore, the measurement of the tear film thickness on and underneath the contact lens should be addressed. Some instruments can measure the thickness of the PCTF, PLTF, and PoLTF with good precision. However, direct cross-sectional visualization of the tear film may provide a better understanding of its role in contact lens fitting and interaction of the lens with the ocular surface.

Technological advancements in optical coherence tomography (OCT) have now made it possible to use ultra-high-resolution (~5 μm) spectral-domain OCT (SD-OCT) to image the tear film on contact lenses in situ. Thus, any tear film thicker than 3 μm theoretically can be detected. Although change in tear film are important in evaluating the fit of the contact lens on the cornea, not much information is available regarding the dynamic changes in the PLTF and PoLTF. Bruce et al., King-Smith et al., and Nichols et al. investigated the PCTF, PLTF, and PoLTF using interferometric methods. However, these methodologies cannot capture two-dimensional images of the anterior segment of the eye, nor can they directly visualize the tear layers on and underneath the contact lens. The goals of the present study were to use direct or indirect measurements by ultra-high-resolution OCT to visualize the PCTF, PLTF, and PoLTF and to investigate the dynamic changes in thickness after the instillation of artificial tears.

METHODS

Subjects

This prospective study was approved by the research review board of the University of Miami. Informed consent was obtained from each subject in accordance with the tenets of the Declaration of Helsinki. During screening, slit lamp evaluation after fitting a study lens (~3.00 D; base curve: 8.6 mm; silicone hydrogel contact lenses, PureVision; Bausch & Lomb, Rochester, NY) was performed by one of the investigators (CQ) to confirm the lens fitting with a centration of less than 1 mm. After a screening test, 22 subjects (14 men and 8 women; mean ± SD age, 31.4 ± 6.6 years; range, 24–46) were selected. Twelve did not wear contact lenses, and 10 wore soft lenses. None of the subjects had a history of previously diagnosed dry eye or any current ocular or systemic diseases.

Ultra-high-Resolution Spectral Domain OCT

To detect the micrometer-thin tear layers on the cornea and the contact lens, we constructed a custom-built, high-speed, ultra-high-resolution SD-OCT and used it in the present study. Briefly, the image capture rate was 48 frames per second when a light source was used that had a center wavelength of 840 nm and a broadband width at 100
nm. The source was connected with a telecentric light delivery system driven by an X-Y galvanometer scanner. The power of the incident light delivered into the anterior segment was adjusted to 750 μW. The scan width was up to 15 mm with a depth of 3 mm. In theory, this OCT system has a ~3-μm depth resolution in tears or tissues of the eye. The repeatability of the system was tested for the measurement of soft contact lenses. Five soft contact lenses with the same parameters as those for the lenses used in this study were immersed in a wet cell and imaged by OCT two times, with approximately a 5-hour interval between the two measurements. Three measurements of the central thickness were averaged. The repeatability was defined as the SD of the differences between the two measurement sessions in each lens. The repeatability was 0.92 μm. The images captured by the camera were delivered to a computer workstation for processing and display. The light delivery system was mounted on a standard slit lamp that also incorporated a digital video system in the viewer. The subjects were asked to sit in front of the slit lamp and look forward horizontally. The room light was dimmed to avoid reflex tearing that might influence the results.

**Experimental Procedure**

The study was conducted in a consulting room with controlled temperature (15–25°C) and humidity (30%–50%). All subjects were scheduled after 10 AM to avoid the edematous cornea and the alteration of the tear film induced by sleep that might affect the result of the study.16,17 The corneal image, including the micrometer-thin tear film, of a randomly chosen initial eye was taken as the baseline in an 8-mm-width scan on the horizontal meridian with ultrahigh-resolution SD-OCT. Before insertion of a soft contact lens, 1 drop (35 μL) of artificial tears (Refresh Liquigel; Allergan, Irvine, CA) was instilled on the concave surface, and the lens was inserted on the eye. The anterior segment of the eye with the contact lens was imaged immediately after lens insertion and centration verification. It was then imaged again every 2 minutes for 10 minutes. Each image was taken after good centration verification. After that, the contact lens was removed, and 1 drop of artificial tears was instilled on the cornea so that it could be reimaged to obtain the true corneal thickness. This value was used to indirectly calculate the thickness of the PCTF and PoLTF. The subject was permitted to have a 5-minute rest to reduce the risk that manipulation of the initial eye would influence the contralateral eye.

After the 5-minute rest period, the baseline image of the contralateral eye was taken. Another silicone hydrogel contact lens with the same specification was then inserted onto this eye; however, for the second eye, no drop of artificial tears was placed on the concave surface of the lens before insertion onto the eye. The OCT images were taken immediately after instillation and every 2 minutes for 10 minutes. The lens was then removed, and the same procedure that was performed on the initial eye was again used to obtain the true corneal thickness of the contralateral eye.

After the lenses were removed, they were soaked in the contact lens solution and imaged by OCT to obtain the lens thickness. This value was used to calculate the PLTF indirectly.7

**PCTF, PLTF, and PoLTF Measurements**

Commercial software (MATLAB; ver. 7.1; The MathWorks, Inc., Natick, MA) was used to perform the image processing and obtain the thickness of a randomly chosen initial eye was taken as the baseline in an 8-mm-width scan on the horizontal meridian with ultrahigh-resolution SD-OCT. Before insertion of a soft contact lens, 1 drop (35 μL) of artificial tears (Refresh Liquigel; Allergan, Irvine, CA) was instilled on the concave surface, and the lens was inserted on the eye. The anterior segment of the eye with the contact lens was imaged immediately after lens insertion and centration verification. It was then imaged again every 2 minutes for 10 minutes. Each image was taken after good centration verification. After that, the contact lens was removed, and 1 drop of artificial tears was instilled on the cornea so that it could be reimaged to obtain the true corneal thickness. This value was used to indirectly calculate the thickness of the PCTF and PoLTF. The subject was permitted to have a 5-minute rest to reduce the risk that manipulation of the initial eye would influence the contralateral eye.

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**PCTF, PLTF, and PoLTF Measurements**

Commercial software (MATLAB; ver. 7.1; The MathWorks, Inc., Natick, MA) was used to perform the image processing and obtain the thickness of the central specular hyperreflective reflex were removed. Ten axial 50 axial scans (0.30 mm width) of the central specular hyperreflective reflex were removed. Ten axial scans on each side of the specular reflex were then used to yield OCT longitudinal reflectivity profiles from the anterior surface of the cornea to the posterior surface of the PLTF. (C) Processing of the image produced peaks a, the posterior surface of the cornea; b, the anterior surface of cornea or posterior surface of PoLTF; c, the anterior surface of the PoLTF or the posterior surface of the contact lens; d, the anterior surface of the contact lens or the posterior surface of PLTF; and e, the anterior surface of the PLTF. The thickness of each layer was calculated by determining the number of pixels between each two peaks and then converting them to micrometers by the factor 1.13.
CL Insertion with Drop

FIGURE 2. Changes in PLTF and PoLTF after lens insertion with 1 drop of artificial tears on the concave side. (A) A soft contact lens (CL) was inserted onto the cornea (CO) with 1 drop (35 μL) of artificial tears on the concave surface of the lens. Both the PLTF (green *) and PoLTF (red #) were clearly visualized by SD-OCT immediately afterward. (B–F) After that, the PLTF and PoLTF gradually decreased. (F) At 10 minutes after lens wear, both the PLTF and PoLTF were still visible.

### Table 1. PCTF, PLTF, and PoLTF Thicknesses at Each Time Point

<table>
<thead>
<tr>
<th>Eye/Time Point</th>
<th>PCTF</th>
<th>PLTF</th>
<th>PoLTF</th>
</tr>
</thead>
<tbody>
<tr>
<td>Initial eye*</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Baseline</td>
<td>1.7 ± 1.5</td>
<td>15.4 ± 9.0</td>
<td>7.9 ± 8.7</td>
</tr>
<tr>
<td>CL insertion with drop</td>
<td>18.5 ± 9.5</td>
<td>15.4 ± 9.0</td>
<td>7.9 ± 8.7</td>
</tr>
<tr>
<td>2 min</td>
<td>12.6 ± 7.2</td>
<td>8.7 ± 4.8</td>
<td>4.6 ± 2.4</td>
</tr>
<tr>
<td>4 min</td>
<td>8.2 ± 5.5</td>
<td>4.8 ± 2.9</td>
<td>3.2 ± 1.7</td>
</tr>
<tr>
<td>6 min</td>
<td>6.0 ± 4.5</td>
<td>3.6 ± 1.9</td>
<td>3.2 ± 1.7</td>
</tr>
<tr>
<td>8 min</td>
<td>4.6 ± 3.4</td>
<td>2.5 ± 2.1</td>
<td>2.0 ± 2.2</td>
</tr>
<tr>
<td>10 min</td>
<td>3.2 ± 3.7</td>
<td>2.0 ± 2.2</td>
<td>2.0 ± 2.2</td>
</tr>
<tr>
<td>Contralateral eye†</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Baseline</td>
<td>2.0 ± 1.5</td>
<td>3.0 ± 2.1</td>
<td>1.1 ± 1.1</td>
</tr>
<tr>
<td>CL insertion</td>
<td>6.0 ± 5.2</td>
<td>3.0 ± 2.1</td>
<td>1.1 ± 1.1</td>
</tr>
<tr>
<td>3 min</td>
<td>2.3 ± 1.7</td>
<td>1.4 ± 1.5</td>
<td>1.1 ± 1.1</td>
</tr>
<tr>
<td>Drop instillation</td>
<td>29.5 ± 12.5</td>
<td>1.0 ± 1.2</td>
<td>1.1 ± 1.1</td>
</tr>
<tr>
<td>2 min</td>
<td>15.1 ± 7.9</td>
<td>1.4 ± 1.4</td>
<td>1.1 ± 1.1</td>
</tr>
<tr>
<td>4 min</td>
<td>9.5 ± 5.2</td>
<td>0.9 ± 1.2</td>
<td>1.1 ± 1.1</td>
</tr>
<tr>
<td>6 min</td>
<td>7.3 ± 5.4</td>
<td>0.9 ± 1.1</td>
<td>1.1 ± 1.1</td>
</tr>
<tr>
<td>8 min</td>
<td>5.3 ± 4.1</td>
<td>1.1 ± 1.1</td>
<td>1.1 ± 1.1</td>
</tr>
<tr>
<td>10 min</td>
<td>4.3 ± 4.1</td>
<td>1.1 ± 1.0</td>
<td>1.1 ± 1.0</td>
</tr>
</tbody>
</table>

Data are expressed as mean micrometers ± SD.

* The concave surface of the contact lens contained a drop (35 μL) of artificial tears before lens insertion.

† The contact lens was inserted onto the contralateral eye with no drop. Three minutes after lens insertion, 1 drop (35 μL) of artificial tears was instilled on the convex surface of the lens.

The thickness of each layer equaled the distance in pixels between the corresponding two peaks. As shown in the magnified image (Fig. 1B), the PLTF, PoLTF, soft contact lens, and cornea were visualized clearly after lens insertion with 35 μL artificial tears on the concave surface. In the reflectivity profile, five sharp peaks were obtained, which are respectively labeled a through e, from left to right (Fig. 1C). Thicknesses of the PLTF, PoLTF, soft contact lens, and cornea, measured in pixels, were easily obtained. The tear film thickness in micrometers was then calculated by using a factor of 1.13 with the refractive index of tears as 1.343.18

However, at some points, the tear film on the cornea and the contact lens could not be visualized, because it was too thin to detect. To obtain the thickness of these invisible tear layers that actually existed, we used an indirect calculation. Before lens insertion, the thickness of central cornea plus the PCTF (Ct) was measured. Then, the instillation of 1 drop of artificial tears outlined the cornea in images obtained by SD-OCT, so that the true central cornea thickness (Ct) was obtained. Thus, the PCTF thickness was calculated as Ct − C2. The calculations for the PLTF and the PoLTF thicknesses were based on the same principle. The true thickness of the contact lens (Lc) was determined from the image taken after the lens was soaked in the contact lens solution. This thickness was subtracted from the central lens thickness (L1) that included the PLTF measured during lens wear, giving the PLTF thickness as PLTF = L1 − L2. The thickness of true cornea plus the PoLTF (Ct) was measured in vivo. As with the direct calculations of thicknesses, indirectly calculated thicknesses were initially measured in pixels and then converted to micrometers by the multiplication factor of 1.13.

### Data Analysis

Paired t-tests were used for the comparison of tear film thickness between the right and left eyes at baseline. Repeated-measurements analysis of variance (Re-ANOVA) was performed to determine whether there were differences in the tear film thicknesses among the different time points. Post hoc tests were used to compare the tear film thick-
before insertion. The PLTF and PoLTF were immediately increased above baseline between non–contact lens wearers and experienced wearers. There were no significant differences at any of the 2-minute time points. Data are presented as the mean ± SD for all the variables at each time point. P < 0.05 was considered significant (all analyses: SPSS ver. 13.0; SPSS Inc., Chicago, IL).

**RESULTS**

At baseline, there were no significant differences between the PCTF thickness in the right and left eyes (paired t-tests, P > 0.05); therefore, we averaged both PCTF thicknesses for each subject. The mean thickness of the PCTF at baseline was 1.9 ± 0.9 µm (range, 0.2–3.7 µm). After insertion of the contact lens, there were no significant differences at any of the 2-minute time points over 10 minutes in PLTF and PoLTF or the PCTF at baseline between non–contact lens wearers and experienced contact lens wearers (Re-ANOVA, P > 0.05).

**Tear Film Thickness after Lens Insertion with 1 Drop of Artificial Tears Instilled on the Concave Surface**

Both the PLTF and PoLTF were clearly evident and visualized immediately after lens insertion with 1 drop of artificial tears instilled on the concave surface in all subjects (Fig. 2A). Over the 10-minute period of observation, the PLTF decreased gradually with blinking (post hoc test, P < 0.05; Table 1, Figs. 2B–F, 3A). At each time point after lens insertion, except at 10 minutes, it was significantly greater than PCTF at baseline (post hoc test, P < 0.05; Table 1, Fig. 3A). Similarly, the PoLTF decreased continuously for 8 minutes after insertion (post hoc test, P < 0.05; Table 1, Figs. 2B–F, 3A). It was greater than the PCTF through the first 6 minutes after insertion (post hoc test, P < 0.05; Table 1; Fig. 3A). In six of the 22 subjects, the PoLTF was not detected after 2 minutes. In the others, it was visible for 4 (n = 5), 6 (n = 6), 8 (n = 2), and 10 (n = 3) minutes.

**Tear Film Thickness after Lens Insertion with 1 Drop of Artificial Tears Instilled on the Convex Surface after 5 Minutes of Lens Wear**

In the second eye in which the concave surface of the lens was not prewetted with artificial tears, the PLTF (n = 12) and PoLTF (n = 9) were visualized immediately on lens insertion on the cornea (Table 1, Figs. 3B, 4A). Three minutes later, the PLTF was still apparent in only two subjects, whereas the PoLTF was not apparent in any of the subjects, because it became too thin (Table 1, Figs. 3B, 4B). After instillation of a single drop of artificial tears, the PLTF clearly increased in thickness (post hoc test, P < 0.05; Table 1, Figs. 3B, 4C), but the PoLTF did not (post hoc test, P > 0.05; Table 1 Figs. 3B, 4D). After that, the PLTF gradually decreased in the following 8 minutes (post hoc test, P < 0.05; Table 1, Figs. 3B, 4D, 4E), but the PoLTF did not increase and remained undetectable (Re-ANOVA, P > 0.05; Table 1, Figs. 3B, 4D, 4E). The thinnest tear film directly detected by ultrahigh-resolution SD-OCT used in this study was 3.2 µm, which was very close to the 3-µm theoretical resolution in tears by the instrument.

**DISCUSSION**

How to best measure the thickness of the PCTF, PLTF, and PoLTF is of considerable interest to clinicians and researchers; however, no agreement has been achieved regarding the optimal method. Doane7 developed a tear film interferometer that applied a contour plot to the measurement of the PLTF thickness. Fogt et al.10 used interferometry of spectral oscillations to determine the PLTF thickness. Bruce and Brennan11 and Bruce and Mainstone19 used biomicroscopy to observe the PoLTF in specular reflection. Prydal et al.12 found the PCTF thickness to be 34 to 45 µm by noninvasive interferometry and 41 to 46 µm by confocal microscopy. Wong et al.14 and Creech et al.15 obtained theoretical predictions of PCTF thickness as 8.0 and 10.4 µm, respectively. King-Smith et al.17 reported that the average thickness of the PCTF was 2.7 µm. Nichols and King-Smith22 found both PLTF and PoLTF to be 2.3 µm, when measured by interferometry. In contrast, Lin et al.23 reported a value of 11.5 µm for the PoLTF when determined by optical pachometry. Although the technologies and methods described above can be used to estimate tear film thickness with higher resolution than OCT, they cannot rapidly acquire large numbers of ultrahigh-resolution images that are necessary to precisely evaluate tear dynamics.

Commercial and custom-made OCT devices have been used to study many aspects of the ocular anterior segment such as the whole cornea, the epithelium, anterior chamber width, depth and angle, the flap after laser and others.24–28 The high resolution of the time domain OCT instrument, approximately 10 µm, made it useful to investigate the tear menisci and tear film.8,29 Wang et al.30 presented distinct images of the cornea, contact lenses, and tear film and reported PLTF, PLTF, and the PoLTF thicknesses of approximately 3.3, 3.8, and 4.6 µm, as
determined by indirect calculation. Recent studies demonstrated that a high-resolution SD-OCT or an ultrahigh-resolution SD-OCT could obtain higher quality images and detect more details of the ocular anterior segment. Kaluzny et al. indicated that the high-resolution OCT instrument, with a 4- to 6-μm longitudinal resolution, was a promising device in contact lens research and practice. However, the tears on the contact lens surface were not included in their study, and tear films were not evident in their images. In a previous study, we used a custom-built, ultrahigh-resolution (~3 μm) OCT instrument to precisely estimate tear menisci and tear films on and underneath contact lenses and to record the interaction between the lens edge and the ocular surface. In addition, we compared the fitting characteristics of two lenses with different materials, back curves, and edge designs and suggested that rational lens design might improve the fitting characteristics by improving tear exchanges around the lens edge. Therefore, the SD-OCT instrument was considered to be a promising tool for studying tear dynamics on the lens. However, because of the small subject sample in that study, conclusions regarding the true PLTF and PoLTF thickness were not drawn.

In the present study, we used ultrahigh-resolution SD-OCT and confirmed its theoretical depth resolution of 3 μm by visualizing a PLTF of 3.2 μm in one subject. As long as the tear film was thicker than 3.2 μm, it was detectable and calculated directly. In addition, the PoLTF was visualized and calculated directly immediately after lens insertion in some subjects, as suggested by Wang et al. To the best of our knowledge, this is the first time the PoLTF has been quantified directly from images without the aid of artificial tears. The increased tear film is mainly the result of reflex tearing, as we avoided applying soaking solution to the eye when the lens was inserted. In some subjects, there was no detectable increase in the PoLTF. This means only that the PoLTF did not exceed the 3-μm limit of resolution. It is possible that these subjects had poor sensitivity or good adaptability to soft contact lens wear and did not have any significant reflex tearing. Immediately after lens insertion or eye drop instillation, the PLTF and PoLTF were readily visualized. However, after 3 minutes of lens wear, they were not visible in most subjects. Thus, normally the PLTF and PoLTF during lens wear are too thin to be visible by ultrahigh-resolution OCT. The central thicknesses of the PCTF, PLTF, and the PoLTF were close to but a little thinner than those reported by King-Smith et al., Nichols and King-Smith, and Wang et al. This slight difference may be due to the differences in sample size, the race of the subjects recruited, and the evident improvement of the longitudinal resolution of the instrument.

To investigate the effect of 1 drop on the PLTF and the PoLTF, we monitored these variables for 10 minutes after instillation on the lens. The results indicated that the lubricating drops used to improve ocular comfort during lens wear do not flow around the lens edge into the space between lens and cornea. Thus, the drops may relieve only the friction between eyelid and contact lens by increasing the PLTF. When the artificial tears were placed on the concave surface of lens before insertion, the PoLTF was readily apparent. However, it was quickly extruded by blinking. Thus, the PoLTF cannot be maintained during lens wear, even when extra tears are added.

The SD-OCT instrument used in the present study has opened a new era in the evaluation of the different fitting characteristics of various lens designs and materials. Our results suggest that soft contact lens design and materials must be improved so that they hold the PoLTF and enhance the rate of tear exchange beneath the lens. Only in this way can the dry eye symptoms induced by contact lens wear be alleviated and the health of the ocular surface be sustained.
contact lens and the cornea used in the indirect calculations of tear films did not change during the approximately 10 minutes of lens wear. Any thickness changes in the lens and cornea that did occur could introduce measurement errors for the indirect calculations. In addition, lens decentration during imaging may be another concern. After fitting the lens on the eye, we used a slit lamp biomicroscope to evaluate the fitting, especially the centration. Good centration of less than 1 mm was achieved on each eye. Although we were aware of variation of the lens fitting during blinking, the images were taken during the first gaze. Slight decentration appeared not to have an impact on the direct visualization of the tear film; however, some variation may have been introduced into the indirect calculation. Using test lenses, we measured the variation of lens thickness. The variation of the central 1-mm zone was approximately 1.5 μm between the thinnest and thickest points. In an attempt to offset the systematic error due to decentration of the lens, we processed multiple acquisitions along both sides of the apex to arrive at the results. Last, in this one-visit study, we defined normal healthy subjects as individuals without any previously diagnosed dry eye. We did not perform any tear function tests, like tear break-up time or the Schirmer test. The absence of testing may have allowed some undetected variation, if some of our subjects had subclinical dry eye. Further studies are needed to correlate the measurements with these tests.

In summary, the ultrahigh-resolution SD-OCT instrument is a novel and promising tool for evaluating the tear dynamics during contact lens wear. With this instrument, we have shown that the tears cannot be held underneath the soft contact lens, and extra lubricating drops used clinically for improving ocular comfort did not alter the PoLTF.

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References