Accommodation-induced variations in retinal thickness measured by spectral domain optical coherence tomography

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1 Introduction

With the increasing prevalence of presbyopia and myopia worldwide, research into understanding the accommodation mechanism and its association with presbyopia and myopia has attracted great interest among ophthalmic research communities. Accommodation is defined as a dioptric change in the power of the eye to focus a near object onto the retina. When the eye accommodates, the ocular dimensions undergo a series of changes: an increase in lens thickness, a reduction of the lens’s equatorial diameter, a forward movement of the lens’s anterior pole, a backward movement of the lens’s posterior pole, and a contraction of the ciliary body with a reduction in the ciliary ring diameter. These changes lead to a concomitant decrease of the anterior chamber depth (measured from the cornea to the anterior lens surface) and an increase in the anterior segment length (measured from the cornea to the posterior lens surface).

In addition to the dimensional changes in the anterior segment, changes in the posterior segment of the eye during accommodation also have been widely investigated, because retinal stretching or distortion with accommodation may be associated with accommodative micropia and other accommodation-dependent perceptual phenomena. However, most reported results because retinal stretching or distortion with accommodation were based on psychophysics and visual optics methods. Direct measurements of morphological changes of the human retina during accommodation using high-resolution three-dimensional (3-D) imaging techniques, such as optical coherence tomography (OCT), have not been reported before. This type of measurement is able to provide direct evidence and quantification of retinal stretching or distortion.

In previous studies, we investigated dimensional changes of the anterior segment in different accommodated states using a dual-channel spectral domain OCT (SD-OCT) system, the results of which were in agreement with those of other
In this paper, we report on measurements of retinal thickness (RT) of the macular area using the same OCT system to investigate accommodation-induced morphological changes of the central retina. This study takes advantage of the imaging system’s capability of simultaneously measuring ocular dimensions, including RT and axial length. With the knowledge of axial length, correction for optical magnification of the eye can be performed to ensure a more objective comparison of RT between unaccommodated and accommodated states.

2 Methods

2.1 Experimental System and Performance

The dual-channel SD-OCT system used in this study consists of two fiber-based Michelson interferometers (Fig. 1). The collimated light beams in the sample arm were combined by a beam-splitter cube. The combined probing beams were coaxial. By placing a lens after one of the collimated light beams, the divergence of the two probing beams was turned to make them focus at different depths in the eye. The light beam responsible for imaging the cornea [OCT-1, consisting of super luminescent diode 1 (SLD 1) light source, the optics of channel 1, and spectrometer 1] was focused at the anterior segment, while the other (OCT-2, consisting of SLD 2 light source, the optics of channel 2, and spectrometer 2) was focused at the retina. Two SLD light sources with the same full width at half maximum (FWHM) of 45 nm (IPSDD0807, Inphenix, Livermore, California) were used for two OCT subsystems. The linear CCD cameras (Aviiva-SM2-CL-2014, 2048 pixels with 14 micron pixel size operating in 12-bit mode, e2V, Chelmsford, Essex, England) in the two spectrometers were synchronized and operating at a rate of 24,000 A-lines per second. The calibrated imaging depths in air of OCT-1 and OCT-2 were 7.28 and 5.16 mm, respectively. The total exposure power in front of the corneal surface was 1.96 mW, which was safe for long-term exposure to the eye according to ANSI Z136.1. A Badal system was integrated to compensate for ocular refractive error and induce accommodation in each eye. Other technical details of the imaging system can be found in our previous publication.

2.2 Subjects

Twenty-three subjects (right eye, mean spherical error: $-3.62 \pm 2.36$ D; astigmatism < 0.5 D; 10 men and 13 women) aged from 22 to 30 years (mean age: 25.8 ± 2.6 years) were recruited from among students of Shanghai Jiao Tong University for this study. The maximum amplitude of accommodation of each subject was measured with the push-up method (mean maximum amplitude of accommodation: $8.02 \pm 0.19$ D). All subjects had no abnormal ophthalmological findings and no history of ocular disease, surgery, or trauma. The study was performed in accordance with the tenets of the Declaration of Helsinki. The subjects understood the nature and possible consequences of this research and informed consent was obtained from all volunteers in this study.

2.3 Imaging Procedure

All imaging procedures were performed by the same experienced operator. All eyes were scanned with a 512 (horizontal) × 128 (vertical) raster pattern in unaccommodated and maximally accommodated states successively. The crosshair fixation target was uniformly illuminated by a white light-emitting diode lamp. The subjects were instructed to keep the target in sharp focus throughout the imaging procedures.
During the experiment, the subject was asked to sit in front of a modified slit-lamp and fixate at the crosshair target through a Badal system. The other eye was covered throughout the experiment to ensure reliable fixation of the test eye. The fixation target was first adjusted to the optical far point of the eye to compensate for its refractive error and maintain an unaccommodated state as much as possible. The central scanning line was adjusted to cover the foveal center. The OCT images were acquired when the cornea and retina were clearly visualized and a specular reflex appeared in the corneal image. Then, the operator moved the target to induce accommodation until the subject reported blurring of the target. At this point, subjective maximum accommodation was induced. Two repeated measurements were performed in unaccommodated and maximally accommodated states during each test session. Two independent test sessions were performed for each eye with a 15-min break to test the reproducibility of the measurements.

2.4 Data Analysis

We developed a computer program to segment the layers of the internal limiting membrane (ILM) and RPE automatically using the acquired 3-D volumetric data covering a 6 mm × 6 mm macular area. The segmented ILM and RPE boundaries from each cross-sectional image in the volumetric OCT data were manually examined and corrected to ensure accuracy by a professional operator who was blinded to the subjects’ information. The RT was determined by calculating the distance between the ILM and RPE boundaries. The refractive index used in the RT calculation was 1.380. The retinal volume (RV) is the sum of the areas nasal, temporal, superior, and inferior from the foveal center [the area marked with a red square in Figs. 2(b) and 2(d)]. Several factors may influence the comparison, e.g., eye movements during OCT scanning and/or accommodation, nodal point shift during accommodation, eye AL, and so on. To ensure that the calculations of the RT change were not affected by these factors, registration of the tested area together with a correction for optical magnification between unaccommodated and maximally accommodated states was performed.

First, the OCT cross-sectional image that contained the deepest foveal pit (defined as the foveal center) was manually identified from the corresponding 128 cross-sectional images obtained from the unaccommodated and accommodated states, respectively [Figs. 2(c) and 2(e)]. Then, the position of the A-scan that corresponded to the smallest RT could be identified. Hence, the exact position of the foveal center can be obtained from each OCT volumetric dataset.

Second, in consideration of the optical magnification caused by accommodation and various ALs, the actual scan distance at the retina and the corresponding scan resolution of each eye needed to be corrected individually. A schematic illustrating the correction is shown in Fig. 3. The actual distance being scanned at the retina is equal to the preset distance (6 mm) only for a schematic eye (AL: 23.9 mm; nodal position in an unaccommodated state: 7.4 mm) with no refractive error. The actual scan distance at the retina of each eye can be calculated based on geometric optics with a preset scan length (L = 6 mm) in a schematic eye.

\[ A_{\text{unacc}} = \frac{[(AL_{\text{unacc}} - N_{\text{unacc}})/(23.9 - 7.4)] \times L}{A_{\text{acc}}} = \frac{[(AL_{\text{acc}} - N_{\text{acc}})/(AL_{\text{unacc}} - N_{\text{unacc}})] \times A_{\text{unacc}}}{(AL_{\text{unacc}} - N_{\text{unacc}}) / (AL_{\text{acc}} - N_{\text{acc}})}, \]  

where \( A_{\text{unacc}}, A_{\text{acc}}, AL_{\text{unacc}}, AL_{\text{acc}}, N_{\text{unacc}}, \) and \( N_{\text{acc}} \) are the actual scan distance, the corresponding AL, and the nodal positions in unaccommodated and accommodated states, respectively. For example, under the same preset scan length, the actual scan length in an eye with a longer AL is larger than that in the schematic eye in either unaccommodated (A1 > A0) or accommodated states (A5 > A3). In addition, the actual scan length in an eye in an accommodated state is larger than that in an unaccommodated state (A3 > A0, A5 > A1). Then, the corresponding scan resolution of each eye can be obtained according to geometric optics.
to the respective corrected scan length (i.e., the horizontal or vertical scan length/its containing A-scan number). With individually corrected scan resolutions, the number of A-scans contained in horizontal and vertical directions of 4 mm can be accurately calculated.

Finally, according to the A-scan number corresponding to the position of the foveal center, the region of interest (i.e., a 4-mm² macular area centered at the foveal center) can be determined. A spline interpolation method was applied to guarantee the same transverse resolution of the tested area.

By applying the registration process to the area of interest, the influence of eye position displacement on RT comparisons between unaccommodated and accommodated states could be minimized.

2.5 Statistical Analysis

SPSS v. 16.0 (SPSS Inc., Chicago, Illinois, USA) was used for the statistical analysis. The intraclass correlation coefficient (ICC) with a one-way random-effect model was used to evaluate measurement repeatability. A paired t test was applied to analyze the statistical significance of the RT difference between unaccommodated and accommodated status. A one-way analysis of variance (ANOVA) was used to analyze quadrant differences in RV in the same state (unaccommodated or accommodated) in a 4-mm² macular area, as well as accommodation-induced RV changes. P < 0.05 was considered statistically significant.

3 Results

The right eyes of all subjects were imaged at unaccommodated and maximally accommodated states in the study. The RT and AL measured with the dual-channel SD-OCT system showed good reproducibility between the two independent test sessions in the same state (P < 0.001, ICC > 0.9, one-way random ICC). The measured ALs ranged from 23.21 to 27.65 mm (mean AL: 25.10 ± 1.28 mm) and from 23.22 to 27.66 mm (mean AL: 25.11 ± 1.27 mm) at unaccommodated and maximally accommodated states, respectively.

The RT of the horizontal cross-section passing through the foveal center (lateral length: 4 mm, centered at the fovea) was calculated based on the corrected cross-sectional OCT image. The result showed a small but statistically significant decrease in RT in the maximally accommodated state compared to that in the unaccommodated state (paired t test, p < 0.001). The corresponding average RT at the unaccommodated and maximally accommodated states and their difference (mean and standard deviation) are shown in Fig. 4 (mean of all subjects).

The RT of the 4 mm × 4 mm macular area centered at the fovea was also calculated based on corrected OCT volumetric data. The maps of average RT at unaccommodated and
maximally accommodated states and their differences are shown in Fig. 5 (mean of all subjects). There is an obvious RT decrease in the macular area with maximum accommodation [Fig. 5(c)].

In addition, the mean RV of all subjects in unaccommodated and accommodated states and their differences in the superior nasal, inferior nasal, superior temporal, and inferior temporal quadrants of the macular area are shown in Table 1. There was no significant difference in the mean RV among the four quadrants of the 4-mm² macular area (p > 0.05, one-way ANOVA). A significant decrease in the RV of the 4 mm × 4 mm macular area around the fovea occurred with maximum accommodation (p < 0.05, paired t test). The variation in RV induced by maximum accommodation was not significantly different among the four quadrants (p > 0.05, one-way ANOVA).

4 Discussion

Using a dual-channel SD-OCT system manufactured in-house, we demonstrated that RT in the 4-mm² macular area surrounding the fovea underwent a small but significant change during maximum accommodation in a population of young adults. The retinal thinning observed during maximum accommodation may imply retinal stretching or distortion from another perspective, which is in line with previous reports. However, our results showed no significant difference in the RT change induced by maximum accommodation among the four quadrants of the macular area (p = 0.958, one-way ANOVA), as well as between the nasal and temporal area (p = 0.656, one-way ANOVA), which was inconsistent with that proposed by Blank and Enoch. The asymmetry of retinal distortion on the horizontal meridian during marked accommodation may be introduced because of the location of the optic nerve head. Hence, this inconsistency may result from the different areas tested in the studies (the tested area in our study was a 4-mm² macular area centered at the fovea, while the tested area in the study by Blank and Enoch included the whole retina on the horizontal meridian).

Taking optical magnification of the eye into account, the corrections for AL and refractive power were applied to compensate for the difference in transverse image resolution before averaging and comparing the RT values. In this study, we...
used the nodal point position of the schematic eye instead of the individual nodal point position for the correction procedure, which may have caused a small calculation error in the retinal change. In order to assess this calculation error, we also calculated the change in RT without any correction with refractive power and AL. The results also demonstrated quadrant-independent retinal thinning with an RV decrease of 0.216 mm$^3$ in a 4-mm$^2$ macular area (the mean RV change was $\sim$7455 $\mu$m$^3$/A-scan and the mean RT change was $\sim$13.50 $\mu$m/A-scan during maximum accommodation, which is similar, but $\sim$1.23% smaller than those with correction (with correction, the decrease in RV was 0.219 mm$^3$, i.e., the mean RV change was $\sim$7547 $\mu$m$^3$/A-scan and the mean RT change was $\sim$13.67 $\mu$m/A-scan). Hence, the possible calculation error caused by the small difference between the individual and schematic eye’s nodal point positions is unlikely to influence the conclusions in this work.

The optical magnification associated with the change in refractive power of the eye influences not only the scan resolution of the tested area, but also the RT estimation, although the optical path length measured through the retina (ILM to RPE) should not be affected by an alteration in lens thickness induced by accommodation. To investigate the potential influences on intraocular distances caused by magnification, we performed additional measurements with a model eye. SD-OCT measurements on the ocular distances of a model eye with or without a $+8\,\text{D}$ lens placed in front of it showed no difference, which suggested that the RT measurement was unlikely to be influenced by the magnification associated with 8 D of accommodation.

The hypothesis for retinal thinning during accommodation is likely to be explained by the forward movement of the ora serrata caused by an accommodation-induced ciliary muscle contraction. However, the relationship and mechanism between RT variation and accommodation is complicated and still unknown. For example, whether retinal thinning is a passive response induced by accommodation or an active motion to enhance or even induce accommodation needs to be investigated further.

Because of limitations in the imaging system, we did not investigate changes in the choroid and sclera induced by accommodation. It will be further studied with our newly developing experimental system with a 1050-nm light source in the future. In addition, the AL used for the correction of optical magnification was measured by a dual-channel SD-OCT with an average refractive index of 1.353, which may have slightly affected the accuracy of the true AL. Additionally, the maximally accommodated amplitude of the subjects varied by a small amplitude. Both factors will influence the correction for optical magnification and then affect the precision of the retinal change calculation between unaccommodated and accommodated states. Nevertheless, as mentioned above, the difference between accommodation-induced retinal changes with and without correction was only $\sim$1.23%. Therefore, the calculation error was too small to influence retinal change consequences during maximum accommodation. Besides, as mentioned in Sec. 2.4, a registration procedure was performed to minimize the potential influence on the retinal change in the tested area between unaccommodated and accommodated states caused by the changes in fixation or by cyclo-rotational changes in eye position or by any tilts in the OCT scans.

In conclusion, we demonstrate that the macular thickness between the ILM and RPE surrounding the fovea had a slight but significant decrease during maximum accommodation, and
the change was not quadrant-dependent, which will provide new evidence for retinal stretching or distortion. This finding may be explained by an accommodation-induced ciliary muscle contraction, but the intrinsic mechanism is complicated and needs to be investigated further.

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References


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Table 1 Retinal volume of the four quadrants around the fovea (4 mm × 4 mm) in unaccommodated and maximally accommodated states.

<table>
<thead>
<tr>
<th></th>
<th>IN</th>
<th>IT</th>
<th>SN</th>
<th>ST</th>
<th>P value&lt;sup&gt;c&lt;/sup&gt;</th>
</tr>
</thead>
<tbody>
<tr>
<td>Retinal volume (mm³)</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Unaccommodated</td>
<td>1.299 ± 0.087</td>
<td>1.266 ± 0.080</td>
<td>1.325 ± 0.082</td>
<td>1.284 ± 0.080</td>
<td>0.109</td>
</tr>
<tr>
<td>Maximally accommodated</td>
<td>1.243 ± 0.087</td>
<td>1.211 ± 0.082</td>
<td>1.268 ± 0.081</td>
<td>1.229 ± 0.079</td>
<td>0.127</td>
</tr>
<tr>
<td>Difference&lt;sup&gt;b&lt;/sup&gt;</td>
<td>−0.056 ± 0.010</td>
<td>−0.055 ± 0.015</td>
<td>−0.056 ± 0.011</td>
<td>−0.055 ± 0.012</td>
<td>0.958</td>
</tr>
</tbody>
</table>

<sup>a</sup>Difference = retinal volume (maximally accommodated)—retinal volume (unaccommodated). <sup>b</sup>The p value indicates the statistical significance of the difference in retinal volume in the same quadrant between relaxed and maximally accommodated states with a paired t test. <sup>c</sup>The p value indicates the statistical significance of quadrant differences in retinal volume at the same state (unaccommodated or maximally accommodated) by a one-way analysis of variance.
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