

Published in final edited form as:

J Biomed Opt. 2009 ; 14(5): 050501. doi:10.1117/1.3213569.

Extended *in vivo* anterior eye-segment imaging with full-range complex spectral domain optical coherence tomography

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Abstract

We demonstrate the capability of full-range complex (FRC) spectral domain optical coherence tomography (SD-OCT) to image the anterior eye segment from the cornea to the posterior surface of the lens. With an adapted spectrometer design, we developed a SD-OCT system with an extended normal (single half-space) depth range of 7mm (in air). This OCT-intrinsic depth range was doubled with a FRC technique. We demonstrate the performance of our OCT system by imaging the whole anterior segment of a healthy human eye *in vivo*.

Keywords

biomedical optics; ophthalmology; imaging systems

Optical coherence tomography (OCT) is an imaging modality that enables high-resolution cross-sectional imaging of biological tissues and translucent materials.^{1,2} Spectral domain (SD) OCT is a high-speed and high-sensitivity variant of OCT that largely replaced the older time domain variant in the recent years.

In SD-OCT, the depth-resolved information can be reconstructed by Fourier transform of the cross spectral density measured with a spectrometer located in the detection arm of an interferometer.^{3,4} However, SD-OCT suffers from two drawbacks that restrict its measurement range.

First, the Fourier transform of the real-valued cross-spectral density is symmetrical about the zero path difference. Therefore, one cannot distinguish between positive and negative optical path differences with respect to the reference mirror. This effect particularly concerns imaging of objects with larger depth extensions, such as the anterior eye segment, where measurement ranges of the order of 10 mm are needed. In order to suppress the mirror images, the so-called full-range complex (FRC) technique was introduced.⁵ In addition to the amplitude of the spectral interferogram, its phase is measured to reconstruct the full complex spectral interferogram—the analytic function. Inverse Fourier transformation of the analytic function directly yields the true object structures without any mirror terms. Several FRC approaches were developed that differ in the way of generating the $\pi/2$ phase-shifted quadrature function of the spectral interferometric signal.^{6–8} An elegant method uses the phase modulation that is introduced by off-pivot-point illumination of the galvanometer scanner mirror.^{9–11} A second drawback of SD-OCT is the spectrometer-intrinsic depth

range, which is limited by its spectral resolution. Until now, the depth range of standard SD-OCT systems is ~ 3 mm, which can be doubled by applying FRC methods. However, this axial measurement range is not sufficient to cover the entire anterior eye segment.

In this letter, we demonstrate an SD-OCT system with a modified spectrometer design combined with a FRC measurement range doubling that achieves an imaging depth of 14 mm, sufficient to cover the human anterior eye segment from the cornea to the posterior surface of the lens.

Figure 1 shows a schematic diagram of our system. A superluminescent diode (Superlum, Moscow) with a center wavelength of 835 nm and a bandwidth (FWHM) of 18 nm was used as the light source. The round trip coherence length was $17 \mu\text{m}$. The collimated beam (1.5 mm diam) was divided by a 50/50 nonpolarizing beamsplitter into a reference and a sample beam. A variable neutral density filter was mounted in the reference arm to adjust the light power so that the line-scan camera of the spectrometer is operated close to the saturation limit to get maximum sensitivity. In the sample arm, a galvanometer scanner was mounted on an x - y translation stage for correct adjusting of the scanner position to achieve a $\pi/2$ phase shift between adjacent A scans.⁹ The scanner was driven by a saw-tooth voltage generated by a DAQ-card (National Instruments, PCI 6110, Austin, TX). An achromatic object lens with a focal length of 80 mm was used to focus the beam onto the sample. This provided a transversal resolution of $\sim 57 \mu\text{m}$ with a confocal range of 6 mm. Sample and reference beams were recombined at the 50/50 splitter, coupled into a single mode fiber and guided to the spectrometer where the real part of the spectral interferogram $\mathcal{S}(x,\lambda)$ was recorded.

The spectrometer consisted of a transmission diffraction grating (Wasatch Photonics, Logan, Utah) with 1500 lines/mm, an achromatic camera lens with a focal length of 300 mm, and a 2048 pixel line-scan camera (Atmel, Aviva M2 CL 2014, San Jose, CA) with $14 \times 14 \mu\text{m}^2$ pixel size. In our configuration, the spectrometer resolution was 25 pm. The SD-OCT intrinsic depth range, which is limited by the Nyquist sampling theorem, was measured with 7 mm (in air).⁶ After applying the full-range algorithm, the depth range was doubled to an axial imaging range of 14 mm. The probing beam power on the cornea was 2 mW, which is well below the safety limits.¹² The integration time per A scan was set to $50 \mu\text{s}$ to optimize the trade-off between mirror term suppression and sensitivity. With this setting, the sensitivity of the system was measured with ~ 106 dB near zero position. The sensitivity decrease was ~ 17 dB over three-quarters of the depth range. The system worked with an A-scan rate of 20 kHz. A single B scan with 2048 A-scans was recorded in ~ 100 ms. The acquisition time for a 3-D scan with 120 B scans was ~ 15 s. The scanning range covered the whole transversal width of the anterior eye segment (14 mm for a single B scan and $14 \times 14 \text{ mm}^2$ for a 3-D scan).

In conventional SD-OCT, the intensity distribution $I(x,z)$ (which represents the depth profiles) is retrieved by inverse Fourier transform of the recorded spectral interferogram $I(x,z) = \text{FT}^{-1}_{k \rightarrow z} \{ \mathcal{S}(x,k) \}$, where x is the transversal scanning direction, z the axial depth range, and k the wavenumber. Prior to the Fourier transform, fixed pattern noise and dc term were removed by subtracting a mean spectrum (averaged over 2048 A scans) from each spectral data set, followed by rescaling the spectral data from λ space into k space.

As mentioned above, the Fourier transform of a real valued function is Hermitian, and therefore, the depth profile is symmetrical about zero path difference, giving rise to mirror artifacts in conventional SD-OCT. To suppress these mirror artifacts and double the measurement range, we used a phase shift introduced by the x scanning mirror.⁹⁻¹¹ In brief, the sample beam hits the fast scanning mirror slightly away from the scanner pivot axis.

Therefore, during the transversal scan, the optical path length is changed. A constant phase shift between adjacent A scans is generated that depends on the mirror axis offset. If the phase difference is set appropriately ($\pi/2$), one can reconstruct the complex spectral interferogram by Hilbert transform of $S(x,k)$ along the transverse coordinate x for each wavenumber k . Finally, an inverse Fourier transform of each complex spectral A scan yields the depth profiles with suppressed mirror images.

The measured spectral interferograms were transferred via camera link and a high-speed frame grabber card (National Instruments, PCI 1428, Austin, Texas) to a personal computer, where the data were stored and postprocessed.

We demonstrate the performance of our system by 2-D and 3-D imaging of the human anterior eye segment *in vivo*. Figure 2 shows a B scan. The signal intensity was plotted on a logarithmic gray scale and covers an image size of $14(x) \times 14(z)$ (z, optical distance) mm^2 . Figure 2(a) shows a SD-OCT image obtained by inverse fast Fourier transform (FFT) of the spectral interferogram without applying FRC reconstruction. The object structure is disturbed by overlapping of mirror images. Figure 2(b) shows the same data set with FRC post-processing. Note that the imaging depth ranges from the front surface of the cornea to the posterior surface of the lens. Even the epithelium of the cornea and backscattered intensity within the lens can be observed, as well as the lens capsule. However, there are still some residual mirror artifacts from highly backscattering structures, such as the iris.

To quantify the mirror term suppression of our system, the extinction ratio was measured within a highly backscattering structure (iris *in vivo*) and at a weakly scattering black synthetic (nonmoving) surface.⁹ Only pixels with an intensity level above a certain threshold (noise level) in both imaging regions (corresponding to the positive and negative frequencies) were used to calculate the extinction ratio. With this convention, the extinction ratio was measured from Fig. 2(b) with 22 dB in contrast to the nonmoving sample with 36 dB. The reduced extinction ratio might be caused by sample motion that leads to phase instabilities and loss in mirror term suppression efficiency. Axial sample motion of >2 mm/s would invert the suppression effect (suppressing the real image and enhancing the mirror image).⁹

Figure 3 shows the anterior eye segment during the accommodation from the far point (a) to the near point (b). The bright lines in both figures indicate the lens surfaces and demonstrate the thickness-change of the lens. Video 1 shows a 3-D scan of the anterior eye segment. It contains 120 sequenced B scans, which were recorded during one sweep of the y scanner mirror. The size of the 3-D scan is approximately $14(x) \times 14(y) \times 14(z)$ mm^3 . Video 2 shows the same 3-D data set in the *en face* plane.

One drawback of this method is that only a fixed scanning pattern (transversal range and speed) can be used for a specific scanning-mirror pivot-point offset. To overcome this problem, a motorized mount for the transverse scanner can be used. However, this would result in a more expensive and more complex system.

Conclusion

In conclusion, we have developed a SD-OCT system with an extended depth range. With the high sensitivity and high speed of the instrument, the whole anterior eye segment from the cornea to the posterior surface of the lens could be imaged *in vivo* with a recording time of ~ 100 ms per B scan at an A scan rate of 20 kHz; 120 sequenced B scans formed a 3-D scan, which was recorded in ~ 15 s. Possible clinical applications of our system may be in cataract and glaucoma (chamber angle) diagnostics, as well as in accommodation studies.

Acknowledgments

Technical support by C. Wölfl and financial support by the Austrian Science Fund (FWF Grant No. P 19624-B02) are gratefully acknowledged.

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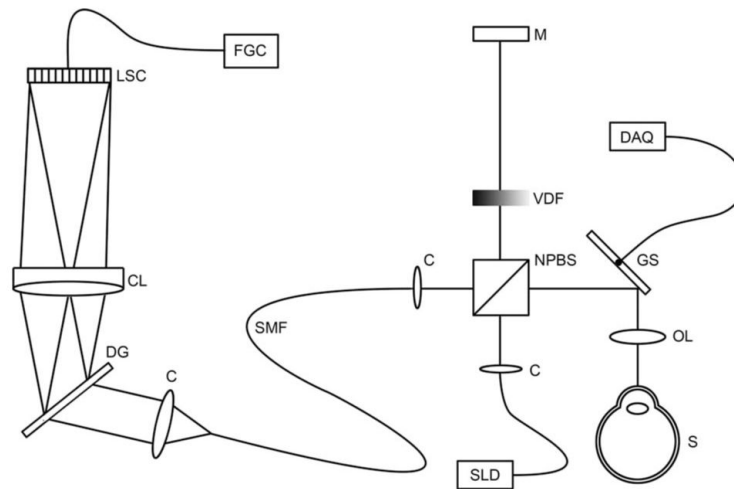


Fig. 1. Schematic setup of the FRC-SD-OCT system. SLD, super luminescent diode; C, collimator; NPBS, nonpolarizing beamsplitter; VDF, variable density filter; M, reference mirror; GS, x - y galvanometer scanner; OL, object lens; S, sample; DAQ, data acquisition card; SMF, single-mode fiber; DG, diffraction grating; CL, camera lens; LSC, line scan camera; and FGC, frame grabber card.

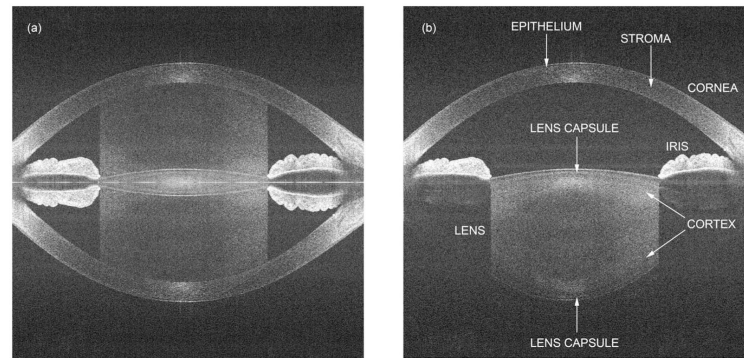


Fig. 2.

In vivo measurements of human anterior eye segment: (a) B scan with 2048 A scans obtained by standard SD-OCT processing; (b) full-range reconstruction by FFT of the complex spectral interferometric signal; image range from cornea to back surface of the lens. Extinction ratio at the iris: 22 dB; image size: $14 \times 14 \text{ mm}^2$; and Dynamic range 50 dB.

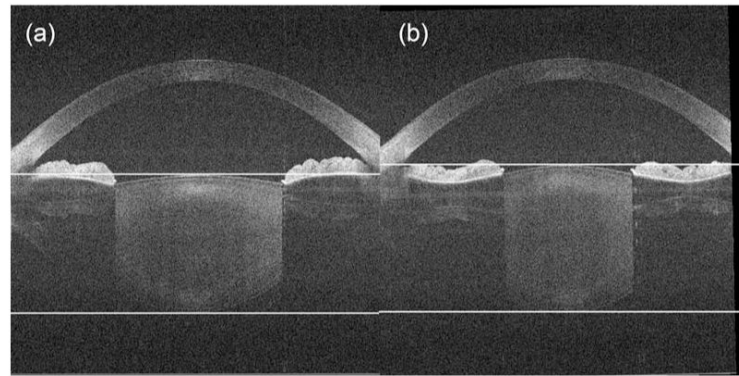
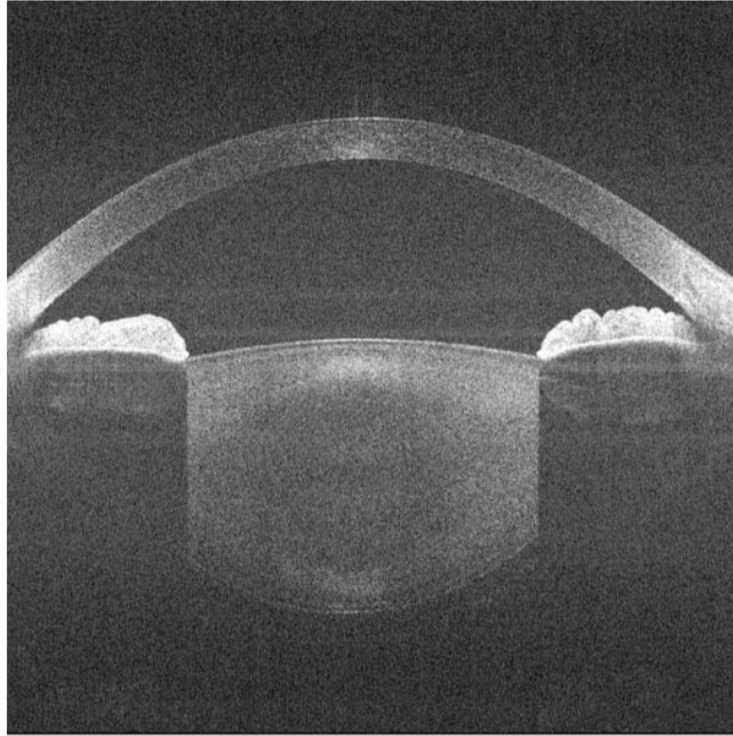
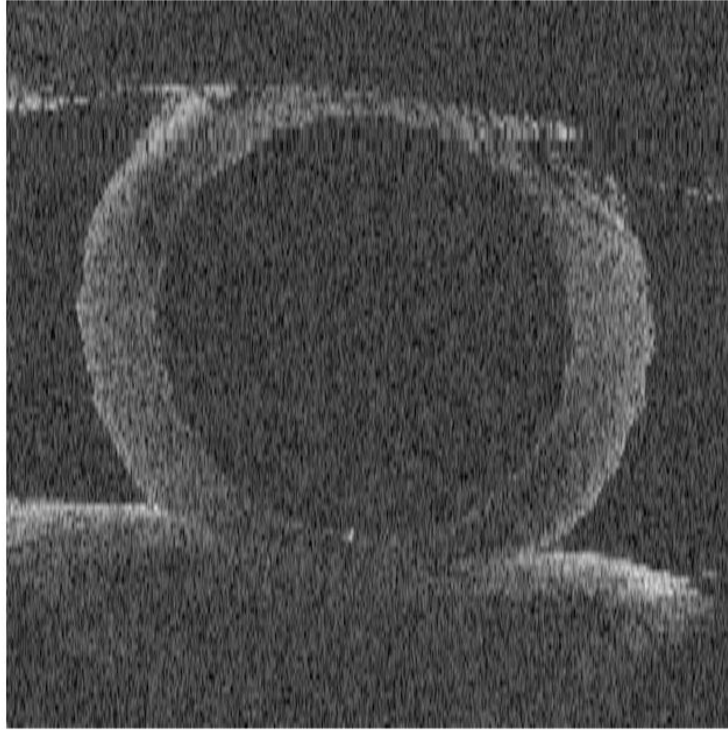


Fig. 3. OCT B-scans during the accommodation of the eye from (a) the far point to (b) the near point. Changes of the anterior eye segment, especially the thickness of the lens, are visible. The white lines are for better visualization of the lens surface position changes during accommodation.

**Video 1.**

3-D scan of the anterior eye segment; movie contains sequence of 120 B scans that were recorded in ~15 s. Size: $14 \times 14 \times 14 \text{ mm}^3$ (QuickTime, 4.5 MB). [URL: <http://dx.doi.org/10.1117/1.3213569.1>].

**Video 2.**

Video derived by the same data set showing the *en face* cross sections of the anterior eye segment from the cornea via the iris to the lens. Size: $14 \times 14 \times 14 \text{ mm}^3$ (QuickTime, 2.6 MB). [URL: <http://dx.doi.org/10.1117/1.3213569.2>].