Ptychographic optical coherence tomography

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Ptychography is a robust computational imaging technique that can reconstruct complex light fields beyond conventional hardware limits. However, for many wide-field computational imaging techniques, including ptychography, depth sectioning remains a challenge. Here we demonstrate a high-resolution three-dimensional (3D) computational imaging approach, which combines ptychography with spectral-domain imaging, inspired by optical coherence tomography (OCT). This results in a flexible imaging system with the main advantages of OCT, such as depth-sectioning without sample rotation, decoupling of transverse and axial resolution, and a high axial resolution only determined by the source bandwidth. The interferometric reference needed in OCT is replaced by computational methods, simplifying hardware requirements. As ptychography is capable of deconvolving the illumination contributions in the observed signal, speckle-free images are obtained. We demonstrate the capabilities of ptychographic optical coherence tomography (POCT) by imaging an axially discrete lithographic sample and an axially continuous mouse brain sample. © 2021 Optical Society of America under the terms of the OSA Open Access Publishing Agreement

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Ptychography [1,2] has been successfully applied for two-dimensional (2D) imaging using various radiation sources, e.g., optical, x ray, and electron sources [3–6], and various methods have been investigated to extend its applicability to three dimensions (3D). The 2D ptychographic forward model assumes a thin object such that the interaction between the illumination and object can be factorized into a simple multiplication [5]. For thicker samples that do not satisfy this multiplication condition, multislice ptychography [7–11] computationally divides a sample into a series of thin slices, applying the 2D forward model from each slice to the next to reconstruct a 3D image. However, the depth resolution of multislice ptychography is limited to the order of the depth-of-field (DoF) of the imaging system. Another approach to extend conventional ptychography is to combine it with other 3D imaging modalities. In the x ray regime, techniques termed ptychographic tomography (PT) [12–14] and ptychographic laminography [15] reconstruct a 3D image from 2D projections at different object viewing angles. Recently, multislice ptychographic tomography (MSPT) [16] has been introduced to extend the imaging depth of the technique while preserving the depth resolution. But in general, a drawback of tomographic techniques is the requirement for dense, time consuming angular scans. In the optical regime, optical coherence tomography (OCT) has become a prominent 3D imaging method [17–19]. OCT obtains 3D information by measuring interference between light backscattered from an object and an external reference by taking advantage of the low temporal coherence of a broadband light source. OCT has also been extended and combined with other computational imaging techniques to account for refractive distortion [20] and diffraction effects inside the imaged 3D volume [21–23]. An OCT system requires the use of a stable interferometric reference arm to encode temporal phase. Furthermore, OCT suffers from speckle noise [19] intrinsic to most coherent imaging methods.

Here we present a new technique that enables high-resolution 3D imaging in a reflection geometry, without the need for object rotation or interferometry, by combining ptychography with swept-source OCT. We demonstrate this ptychographic optical coherence tomography (POCT) technique using a near-infrared swept-source laser in a lensless ptychography setup, where a fast wavelength scan is performed at each transverse scan position. Similar to OCT, the axial resolution in POCT is decoupled from the lateral resolution and is only limited by the source spectral bandwidth. This feature enables fine depth sectioning, not limited by the DoF of the imaging setup that would limit a multislice ptychography approach. The electronically controlled wavelength scan offers a speed advantage over angular scans needed in tomographic approaches. Furthermore, by using ptychography’s ability to retrieve the complex-valued contributions in the observed signal, speckle artefacts that are common in OCT. As the phase is retrieved through interferometric reference arm can be eliminated from the setup, resulting in a simple, compact, and robust layout. We show POCT reconstructions of both an axially discrete two-layer lithographic sample and an axially continuous mouse brain sample.

Figure 1 shows the schematic of the POCT experimental setup where a swept-source is used in a reflection ptychography
configuration. Details of the swept-source system are described in [24]. In the reflection ptychography part, the sample is translated relative to the stationary probe beam with overlapping areas by a 2D motorized stage. A Fermat’s spiral pattern [25] is used as the scan grid. At each transverse scan position, an equidistant frequency scan is performed, and monochromatic ($\lambda/1000$) backscattered intensity patterns are recorded by the camera at each frequency. Lenses L1, L2, L3 and the iris are used to create and control a localized probe for ptychography.

After the data acquisition, single-slice ptychographic reconstructions are performed at each wavelength. Theoretically, the multiplicative approximation in single-slice ptychography holds when the sample thickness $T$ satisfies $T \leq (\Delta r)^2/\lambda$, where $\lambda$ is the wavelength and $\Delta r = \lambda/2N_A$ is the lateral half-pitch resolution. Under the multiplicative approximation, single-slice ptychography offers quantitative object reconstruction, with which high-resolution surface profilometry has been successfully demonstrated [3,27]. By measuring semi-transparent samples and using multiple-wavelength measurements, POCT aims at volumetric imaging.

Our data analysis pipeline is summarized in Fig. 2. At the starting wavelength, we use an auto-focus algorithm specifically tailored to ptychography [28] to calibrate the sample-detector distance. Using this calibration, a momentum-accelerated ePIE algorithm [29,30] is used to reconstructed the object and probe. Thanks to the small step size in the wavelength scan, the previously reconstructed probe and object can be used as initial guesses for the following reconstruction to speed up the convergence rate. In this way, ptychographic reconstructions are performed wavelength by wavelength.

Although quantitative object reconstructions are obtained at each wavelength, initially they appear uncorrelated because of several reasons. First, the reconstructed object pixel size is wavelength-dependent when using Fraunhofer propagation [31]. Second, the absolute lateral object position is one of the ambiguities of ptychography, causing independent reconstructions to contain arbitrary spatial shifts as indicated in Fig. 2. Last, there is a random constant phase term for each object reconstruction due to the non-uniqueness of a constant phase offset in ptychography. For lateral alignment, all estimated objects are scaled and linearly interpolated to the reconstruction with the smallest pixel size, and then registered with sub-pixel precision [32]. The phase relation between the different-wavelength images can be determined from the object itself, thereby replacing the external interferometric reference surface in swept-source OCT, provided the signal is dominated by a single depth reflection at one object location. In contrast to common-path OCT [33], no flat reference layer is required as phase shifts in one layer are identified by the ptychographic reconstruction upon phase synchronization at a single object location (pixel). Once a reference position is chosen, we synchronize the constant phase offsets of all wavelengths at this position. Finally, a 1D Fourier transform along the wavelength axis reveals the depth information of the object. Given by the Fourier transform relation, the optical sectioning resolution is

$$\Delta z_{\text{opt}} = \frac{\lambda_1 \lambda_N}{|\lambda_1 - \lambda_N|}. \quad (1)$$

Taking into account the refractive index $n$ of the object and the double pass in the reflection geometry, the physical depth resolution is given by $\Delta z_{\text{phy}} = \Delta z_{\text{opt}}/2n$.

As a first demonstration of POCT, a custom-made two-layer sample has been measured using $N = 51$ wavelengths ranging from 708.8 nm to 802.8 nm at $M = 201$ transverse scan positions. The sample consists of semi-transparent metal structures (50 nm thick) axially separated by a layer of transparent polymer. Fabrication details of the sample can be found in [24]. A 3D representation of the sample is shown as part of the setup schematic in Fig. 1, and Fig. 3(a) shows the optical microscope image of the sample. The sample-detector distance calibrated by zPIE [28] is 38.6 mm. At the shortest wavelength of 708.8 nm and with the detection NA of 0.085, the lateral resolution is $\Delta r = \lambda_{\text{max}}/2N_A = 4.7 \mu m$, and the DoF is 240 $\mu m$. The optical sectioning resolution $\Delta z_{\text{opt}}$ based on Eq. (1) is 6 $\mu m$, resulting in a depth resolution of 2 $\mu m$, an improvement over conventional ptychography by 2 orders of magnitude. See Supplement 1 for intermediate results. The resolved tomograms of the top and the bottom layers are shown in Figs. 3(b) and 3(c), respectively. Figure 3(d) shows three depth plots at three locations marked in Figs. 3(b) and 3(c), respectively. The phase synchronization was performed at location 1. The width of the peaks shows that the expected depth resolution of 2 $\mu m$ is indeed achieved. We do observe a slight broadening of the peak at location 2, which is caused by strongly dispersive spacer layer as also observed in OCT measurements [24]. Phase synchronization at location 2 would have led to a spectrum-limited resolution at this depth, and more advanced numerical dispersion compensation should enable depth-independent resolution.
The lateral resolution is evaluated by calculating the Fourier ring correlation (FRC) [34] of two object intensities reconstructed from the shortest (708.8 nm) and longest wavelengths (803.8 nm), and the 1/2 bit resolution criteria are used. In Fig. 3(e), the object correlation curve (solid line) is above the 1/2 bit threshold (dashed line) across the detected spatial frequency spectrum, meaning that the two independent reconstructions are consistent up to the Nyquist frequency. Thus, the measured half-period lateral resolution in our lensless setup is confirmed to be at the diffraction limit of 4.7 µm. A 3D rendering of the sample is shown in Fig. 3(f). As depicted in Fig. 1, the top layer is the metal/air or polymer/air interface, and the bottom layer has metal structures at the polymer/glass interface with matching refractive index. A 3D visualization video is available online (Visualization 1).

In a second experiment, a cryo-cut unstained slice of cerebellar mouse brain tissue with a nominal thickness of 40 µm has been measured using POCT. This mouse brain sample has continuous axial structures. A low-resolution, bright-field, reflection optical microscope image of the sample is shown in Fig. 4(a). A high-NA objective is used in the microscope to focus on the top and the bottom layer of the sample, images of which are shown in Figs. 4(b) and 4(c). In the experiment, the sample is measured with \( N = 49 \) wavelengths ranging from 708.8 nm to 851.3 nm at \( M = 201 \) scan positions. The optical sectional resolution \( \Delta x_{\text{opt}} \) is 4.3 µm. The calibrated propagation distance from the object to the detector is 25.4 mm, resulting in a detection NA of 0.12, a lateral half-period resolution of 3 µm, and a DoF of 106 µm. Figure 4(d) shows the reconstructed tomogram of the deepest layer of the mouse brain tissue, where the bright areas indicate the air/glass substrate interface and the dark areas indicate the brain/glass interface. A tomogram of an upper layer of the sample is shown in Fig. 4(e), where the bright areas are ridges of the mouse brain tissue that correspond to the sharp structures in Fig. 4(b). Using an averaged value of \( n = 1.38 \) for the refractive index of mouse brain tissue across the full wavelength range [35], we obtain a value for the thickness of the mouse brain of 20 µm with a depth resolution of 1.6 µm. Figure 4(f) shows a color-coded height plot of the reconstructed sample, where for each pixel the height is chosen at the maximum intensity location along the z direction. Figure 4(g) shows a 3D rendering of the reconstructed mouse brain sample. See Visualization 2 for a 3D rotating projection.

With the two-layer and mouse brain samples, we demonstrate that POCT successfully performs depth-resolved imaging, where the lateral and axial resolutions are decoupled. This shows that POCT has a great potential for 3D high-resolution imaging. Compared to OCT, POCT does not rely on an external reference, and it deconvolves the illumination from the object reconstruction. Therefore, POCT does not suffer from reference- and illumination-induced artefacts. Additionally, the current setup only contains a beam splitter between the object and the camera; thus, the reconstruction results [Figs. 3(b) and 3(c) and Figs. 4(d) and 4(e)] are almost free of typical coherent imaging artefacts (“speckle”) caused by imperfections in optical systems.

One advantage of POCT over CT-based 3D imaging techniques [12–14,16] is speed. First, wavelength scanning is faster than mechanical stage rotation, as our acousto-optic scanning method enables sub-ms wavelength switching times, and fast cameras achieve multi-kHz update rates. Second, computational complexity is reduced, as POCT only requires a series
of 2D reconstructions plus a phase synchronisation step, and the correlation between different-wavelength images provides a strong initial guess for both object and probe after a first reconstruction at a single wavelength, ensuring fast convergence. Furthermore, no image stitching algorithms are needed. Lastly, as the phase synchronisation is achieved numerically, no phase-stable optical path is needed. In addition to the present swept-source implementation of POCT, the concept can be extended to other methods in which wavelength-resolved diffraction data is measured, such as Fourier-transform-based diffractive imaging [36,37] or computational time-domain OCT [24].

Going beyond the present proof-of-concept, various improvements can still be envisaged. One clear limitation of POCT is the absence of coherent amplification by a strong reference wave. We, therefore, envision the main applications of POCT in material science and lithographic device inspection, where photon flux on sample is not the limiting factor, or in situations where the generation of a reference beam is challenging such as with extreme ultraviolet radiation. While in the lensless geometry the NA is limited by the sample-detector distance and detector size, the use of high-NA optics can yield a significant transverse resolution improvement. In addition, the axial imaging range is currently limited to the single-layer projection limit in ptychography, which is typically in the order of tens or hundreds microns at optical wavelengths. In principle, multi-layer ptychography can handle multiple scattering to a certain degree, more sophisticated forward modeling in ptychography is required to apply POCT on highly scattering samples in reflection.

In conclusion, we have introduced POCT as a new concept for high-resolution 3D optical imaging. Further improvements to this new imaging modality are clearly possible, and we believe that the POCT concept has great potential as a simple yet robust 3D imaging method, especially on wafer-based planar nanostructures commonly found in modern nanotechnology.

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Supplemental document. See Supplement 1 for supporting content.

REFERENCES