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# En face OCT angiography demonstrates flow in early type 3 neovascularization (retinal angiomatous proliferation)

## Abstract

*Introduction* The characteristics of type 3 neovascularization (NV), also known as retinal angiomatous proliferation, have been well described clinically, as well as with fluorescein angiography (FA), indocyanine green angiography, and optical coherence tomography (OCT). OCT angiography (OCT-A) is a novel and non-invasive technique for imaging retinal microvasculature by detecting changes, with respect to time, in reflectivity related to blood flow.

Method In this case series, we describe two patients who presented with type 3 NV and underwent clinical examination and multimodal imaging, including OCT-A. *Results* In the first patient, OCT-A demonstrated flow within two separate lesions in the same eye, one of which was only weakly detected by FA. In the second patient, sequential OCT-A demonstrated a reduction in intralesional flow following intravitreal therapy. *Conclusions* OCT-A may have a role in the early diagnosis of type 3 NV and in assessing the response to treatment. Further studies are needed to determine sensitivity and specificity.

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#### Introduction

Type 3 neovascularization (NV), also known as retinal angiomatous proliferation, is thought to arise predominantly from the deep capillary plexus of the retina and grow toward the RPE, forming anastomoses with type 1 (sub-RPE) NV.<sup>1–4</sup> Fluorescein angiography (FA) reveals, at the site of the intraretinal vascular lesion, a

hyperfluorescent focus that typically appears first during dye transit, followed by late leakage. Optical coherence tomography (OCT) localizes the lesion to the deeper layers of the retina.

The advent of OCT angiography (OCT-A) enables retinal microvascular flow to be imaged in the clinical setting without the injection of intravenous fluorescent dye.

#### Case report 1

An 88-year-old female with a 3-year history of NV age-related macular degeneration (NV-AMD) in her right eye presented for a maintenance injection of intravitreal bevacizumab. Fundus examination showed a small new intraretinal hemorrhage in her left eye. Spectral domain OCT (SD-OCT; OCT +HRA, Heidelberg Engineering) and FA confirmed the presence of two separate type 3 NV lesions (Figures 1a and e), the more superior lesion being only weakly detectable on FA.

The patient also underwent OCT-A (Avanti, Optovue, Fremont, CA, USA), demonstrating flow within the active lesions (Figures 1f and g). Intravitreal ranibizumab was administered to the left eye to initiate a course of therapy.

#### Case report 2

A 70-year-old female with a prior history of type 3 NV in her left eye presented for follow up. Assessment of the previously non-NV right eye confirmed a new type 3 NV lesion as documented by color photography, FA/ICG, and SD-OCT (Figures 2a and d). The patient underwent swept-source OCT and OCT-A (Topcon DRI OCT-1, Topcon Corporation, Tokyo, Japan), following which the volume data <sup>1</sup>Vitreous Retina Macula Consultants of New York, New York, NY, USA

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CASE SERIES



**Figure 1** Multimodal imaging of the left eye of patient 1 showing two type 3 NV lesions bordering the foveal avascular zone. Yellow arrows: high-flow lesion; green arrows: second, low-flow lesion. (a) Color photograph; (b, c) mid- and late-venous phases of fluorescein angiogram; (d, e) SD-OCT through high- and low-flow lesions, respectively; (f, g) en face OCT-A segments of superficial and deep capillary plexus, respectively.

was processed by Topcon using a proprietary angiography algorithm. The resulting images revealed a hyperintense flow signal at the inferior border of the foveal avascular zone correlating with the location of the type 3 NV (Figure 2e). The patient began treatment with intravitreal ranibizumab in the right eye on a monthly regimen. After two consecutive treatments, SD-OCT showed a reduction in the retinal thickness with partial resolution of the retinal cyst and intraretinal hemorrhage. Repeat OCT-A showed a reduction in the brightness of the type 3 lesion (Figure 2f); it also revealed a region of supranormal flow, inferotemporal to the main lesion (Figure 2f, black arrow).

## Discussion

Type 3 NV may be difficult to diagnose in the early stages, when it may manifest as intraretinal hyper-reflective features on OCT, without retinal thickening or obvious RPE disruption;<sup>5</sup> serial FA may be necessary for early detection.

OCT-A relies on the detection of changes—in amplitude, phase, or other physical characteristics of the reflected light—that result from the interaction with the components of the flowing blood.<sup>6</sup> These are rendered as two-dimensional en face maps in which the pixel brightness represents blood flow.<sup>7</sup> The relative absence of stationary or time-static structures from OCT-A maps distinguishes these images from those rendered by conventional en face OCT.

Our first patient had two foci of type 3 NV. The lesion at the superior border of the foveal avascular zone was seen on SD-OCT but was only detected weakly by FA. OCT-A demonstrated both the lesions, one with a stronger flow signal than the other (the inferior, brighter lesion saturating the detection capabilities of the device). En face segmentation enhanced the sensitivity of OCT-A to detect these lesions, by excluding the flow signals from the inner retinal capillaries. Segmentation also localized the lesions to the deep capillary plexus, consistent with type 3 NV.

In the second patient, OCT-A demonstrated a single type 3 NV lesion in the deep capillary plexus, communicating with a sub-RPE component. Sequential OCT-A suggested a reduction in the intralesional flow as a result of the treatment with 2 anti-VEGF injections as represented by a reduction in brightness. It also revealed a complex of ectatic capillaries, inferior to the main lesion, with supranormal flow, not detectable by FA or ICGA in our patient. Telangiectasis adjacent to type 3 NV has recently been described as an early feature of type 3 NV.<sup>8</sup>

The technology to perform OCT-A in the clinical setting is now commercially available. OCT-A detects, and may quantify, flow in early or partially treated type 3 NV lesions, and has the potential to increase our understanding of the pathogenesis. To the best of our knowledge, this is the first published report of OCT-A of



**Figure 2** Multimodal imaging of the right eye of patient 2 showing a type 3 NV lesion (yellow arrow) at the inferior border of the foveal avascular zone, and accompanying intraretinal hemorrhage that is adjacent and superior (red arrow). (a) Color photograph; (b) fluorescein angiogram; (c) indocyanine green angiogram; (d) SD-OCT showing a hyper-reflective intraretinal lesion communicating with a localized pigment epithelial detachment at its outer aspect and an adjacent hemorrhage within a cystic space at its inner aspect; (e) baseline OCT angiogram; (f) OCT angiogram after two intravitreal treatments. Supranormal flow is seen in a region of the capillary plexus inferior to the type 3 lesion (black arrow).

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type 3 NV. Larger studies are needed to determine the sensitivity and specificity of OCT-A in the early diagnosis of type 3 NV, and its reliability in determining lesion activity, through quantification of microvascular flow.

#### Summary

What was known before

• Type 3 NV is well described on the basis of clinical findings and multimodal imaging including FA and cross-sectional OCT.

What this study adds

• Our cases show that OCT-A is capable of demonstrating flow in early type 3 NV (that is only weakly detectable on FA), and that OCT-A is capable of quantifying a reduction in flow following intravitreal therapy.

# **Conflict of interest**

KBF is a consultant for Heidelberg Engineering.

# Acknowledgements

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