Commentary: Linear discriminant score for differentiating early primary open angle glaucoma from suspects

Diagnosing glaucoma suspects is a challenge and it is important to be able to recognize the ones most likely to progress to glaucoma. To this end, there has been a search for appropriate tools to be able to discriminate between glaucoma suspects who may actually be early glaucoma and those who may be normal. Because no one parameter is often adequate, there have been attempts to combine parameters in an attempt to provide greater discriminative capability.

The advent of spectral-domain optical coherence tomography (SD-OCT) technology has allowed advanced macular imaging protocols to play an important role in the diagnosis and monitoring of glaucoma.^[1-4] The ganglion cell analysis (GCA) obtained by the Cirrus HD-OCT[®] system (Zeiss) segments and measures the thickness of the ganglion cell-inner plexiform layer (GC-IPL), thereby potentially increasing its diagnostic accuracy compared to conventional peripapillary retinal nerve fiber layer (RNFL) thickness measurement.

Published studies have reported comparable values for area under receiver operating characteristic curves (AROCs) of GCA and RNFL thickness for discriminating early glaucoma from normal.^[2-5] Once there is a visual field defect, such a differentiation is not difficult. The challenge comes in differentiating a glaucoma suspect from normal in the presence of normal visual fields.

In this article,^[6] the authors have used the SD-OCT to study the RNFL thickness and GCA and used a linear discriminant score as a discriminator between glaucoma suspects and early glaucoma patients. They found better accuracy than individual ganglion cell complex (GCC) and optic nerve head (ONH) and RNFL parameters and deduced that it could be used for the diagnosis of glaucoma.

One explanation for the lower AROC of GCA could be that the GCA in its present form only measures an annulus of 4.3 × 4.0 mm around the macula. Though this region contains 50% of retinal ganglion cells (RGCs),^[7] the loss that is picked up depends upon the location of the RNFL defect. Recent studies have pointed out that the topographic profiles of RNFL defects affect the diagnostic performance of macular scans in

preperimetric glaucoma.^[8] Because of the arrangement of the arcuate fibres, in cases of RNFL defects which are closer to the poles, RNFL atrophy may not be reflected by RGC loss in the GCA in the machine simply because it is further away from the measurement circle. The diagnostic ability of macular ganglion cel inner plexiform layer (GCIPL) parameters was comparable to that of peripapillary RNFL and ONH parameters in preperimetric glaucoma (PPG). The inner directional angle of RNFL defects, but not the angular width, affects the diagnostic sensitivity of macular GCIPL parameters.

As for all technology, the GCA analysis also needs to be interpreted with caution, keeping the clinical setting in mind. In a cohort of healthy eyes, Kim *et al.*^[9] reported abnormal diagnostic classifications in 40.4% and 30.8% on GCA and RNFL maps, respectively, especially in eyes with long axial lengths, large fovea-disc angles, and small optic discs. It is also important to keep in mind that conditions such as diabetic macular edema and age-related macular degeneration, which are common comorbidities in the age group of glaucoma patients, may affect the macular RGC thickness.

At present, the GCA measurements do not appear to outperform average RNFL measurements to discriminate between glaucoma suspects and established early glaucoma or normal subjects. In that sense the linear discriminant score appears to be a viable additional tool in our armamentarium of imaging. However, applicability in the clinical setting and easy incorporation in commercially available machines would be required before it could be widely used in clinical practice.

Sushmita Kaushik

Glaucoma Services, Advanced Eye Centre, Postgraduate Institute of Medical Education and Research, Chandigarh, India

> Correspondence to: Prof. Sushmita Kaushik, Glaucoma Services, Advanced Eye Centre, Postgraduate Institute of Medical Education and Research, Chandigarh, India. E-mail: sushmita_kaushik@yahoo.com

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