# ANALYSIS OF CLINICAL RELATIONSHIP OF VISUAL ACUITY WITH OPTICAL COHERENCE TOMOGRAPHY AND PERIMETRY PARAMETERS IN PRIMARY OPEN-ANGLE GLAUCOMA

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## ABSTRACT

Aims: This study aims to analyze the clinical relationship of visual acuity with optical coherence tomography and perimetry in primary open-angle glaucoma. Methods: This retrospective cross-sectional study was conducted with patients who were diagnosed with primary open-angle glaucoma in the ophthalmology department of Trakva University School of Medicine between January 2010 and January 2021. Perimetric data of patients such as visual field index, mean defect, pattern standard deviation, short-term fluctuation, corrected pattern standard deviation; results of optical coherence tomography such as average retinal nerve fiber layer thickness, average C/D ratio, and thickness of retinal nerve fiber layer quadrants, and records of examinational findings were statistically tested to evaluate the statistical relationship. Numbers, percentages, mean and standard deviation were used as the descriptive statistics. Results: Initially 80 eyes of 49 patients diagnosed with primary open-angle glaucoma met the inclusion criteria. 22 patients were female and 27 patients were male. Thinner mean retinal nerve fiber layer, inferior and superior quadrants were observed in eyes with lower visual acuity. It was observed that eyes with lower visual acuity had a worse visual field index and mean defect. Conclusion: In tertiary clinics that receive various numbers of patient referrals, perimetry can be seen as more useful and accurate in primary open-angle glaucoma detection and monitoring. It provides us with better and more accurate results for glaucoma management in the later stages. It is recommended that both optical coherence tomography and perimetry be used for disease monitoring, as this allows doctors to better monitor disease progression. In addition, since optical coherence tomography is an objective test and is less likely to require a patient response, it should be kept in mind that advanced and severe glaucoma may occur when performing perimetry, especially in patients with poor cooperation. Keywords: Glaucoma, perimetry, optical coherence tomography, visual field, epidemiology

# **INTRODUCTION**

Primary open-angle glaucoma (POAG) is a chronic, degenerative optic neuropathy that causes selective loss of retinal ganglion cells, and is one of the leading causes of blindness worldwide. Even though POAG often occurs with elevated intraocular pressure (IOP), many POAG cases were reported to progress with a normal range of IOP values (1, 2). The condition of retinal ganglion cell loss leads to progressive enlargement of the optic nerve cup and thinning of the neural rim (3). Glaucoma is the leading cause of blindness, seen in 60 million people around the world (3). According to the survey data of 2013, POAG accounts for 44 million people across the world and is estimated to hit 53 million people by the year 2020 (4). According to recent studies in the United Kingdom, POAG is reported to be the most common type of glaucoma (5). Studies suggest that POAG is the second most common cause of blindness in the United States, approximately affecting 2.5 million people (6).

Optic disc findings frequently are detected before the deficit of the visual field appears. An enlarged cup-to-disc (C/D) ratio greater than 0.5, C/D ratio asymmetry of 0.2 or more between two eyes or a high asymmetry of the cup in one eye are considered to be diagnostic findings. The destructions caused by glaucoma are clinically revealed by OCT and perimetry tests (6). Perimetry, also known as visual field testing, is a crucial diagnostic tool that maps out the visual field of the patient which makes it necessary for the diagnosis and management of POAG (7). Typical visual field changes in POAG include asymmetry of visual field between two eyes, temporal wedge, paracentral scotoma, general constriction of peripheral field, and increased variability of responses in an area which later developed field defects (7). The perimetry threshold for the detection of visual loss is estimated to reliably stand at 40-50% ganglion cell loss. Thus, in POAG, typical optic nerve head structural changes occur earlier than functional change (visual field loss) (7). It should be noted that there is the phase of characteristic glaucomatous changes and increased tendency of damages to the retinal nerve fiber layer (RNFL) that comes before the perimetrical defects are detectable (7).

This retrospective study aims to analyze the clinical relationship of best-corrected visual acuity (BCVA) with optical coherence tomography and perimetry on patients with POAG in the ophthalmology department of Trakya University School of Medicine.

# MATERIAL AND METHODS

This study was approved by the Scientific Research Ethics Committee of Trakya University School of Medicine (Protocol Code: TÜTF-BAEK 2021/104). This retrospective cross-sectional study analyzed 80 eyes of 49 patients who were diagnosed with POAG

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in the Ophthalmology Department of Trakya University Hospital between January 2010 and January 2021. The study was carried out under the tenets of the Declaration of Helsinki. Written informed consent was obtained from all the participants.

Demographic data such as age and gender, accompanying systemic comorbidities such as hypertension and diabetes, risk factors such as high IOP levels, and the duration of onset POAG, the usage of topical prostaglandins, beta-blockers, carbonic anhydrase inhibitors and alpha agonists, initial and following clinical findings were obtained from the medical records of the patients. All patients underwent complete ophthalmologic examination at each visit including BCVA determined by the Snellen chart, anterior segment biomicroscopic examination, IOP measurement with Goldmann applanation tonometer, and detailed fundus examination obtained with 78 diopters non-contact lens.

The following findings were documented at the clinical visits: visual field parameters such as VFI, mean deviation, pattern standard deviation, optical coherence tomography (OCT) results such as mean RNFL thickness, mean C/D ratio, thicknesses of RNFL quadrants, surgical procedures that the patients underwent, the usage of topical anti-glaucomatous agents and types of anti-glaucomatous agents that have been used.

Initially, 80 eyes of 49 patients diagnosed with POAG met the inclusion criteria of having BCVA with positive light perception and above, absence of corneal pathologies, ocular-vascular diseases (central retinal artery occlusion, retinal venous occlusion), accompanying macular pathologies such as a macular hole and macular degeneration. POAG patients who have poorer BCVA than positive light perception, onset corneal pathologies, ocular-vascular diseases, and macular pathologies which could cause unreliable fundus examination results were excluded from this cross-sectional study.

Perimetric data of patients such as visual field index (VFI), mean defect (MD), pattern standard deviation (PSD), short-term fluctuation (SF), corrected pattern standard deviation (CPSD); results of OCT such as mean RNFL thickness, mean C/D ratio, and thickness of RNFL quadrants, and records of examinational findings were statistically tested to evaluate the statistical relationship.

Numbers, percentages, mean and standard deviation were used as the descriptive statistics. The variables were tested for normal distribution by the One-sample Kolmogorov-Smirnov test. The Chi-square test was used for qualitative comparison. Quantitative data were compared with the Independent Sample t-Test. The bivariate Pearson Correlation test was performed to measure the strength and direction of linear relationships between pairs of continuous variables. A p-value of <0.05 was set for statistical significance. The data were analyzed with IBM SPSS version 20.

#### RESULTS

In our study, 22 (44.9%) patients were female and 27 (55.1%) patients were male. The summary of patients' characteristics (mean age, duration of onset glaucoma, gender, accompanying comorbidity, and the number of the right or left eye that was affected) is shown in Table 1.

The mean BCVA for all eyes was  $0.86 \pm 0.26$  Snellen decimals (ranging from 0.0080 to 1.0 Snellen decimals). It was found that bigger C/D ratios due to fundus examination led to eyes in worsened BCVA (p=0.018, r=-0.263, Pearson Correlation Test). Mean IOP was observed to be  $16.3 \pm 3.5$  mm Hg (ranging from 6 to 23 mm Hg) and was not observed to have a noticeable correlation with BCVA (p=0.228). The summary of examinational findings (BCVA,

#### Table 1: Patient characteristics.

	Number of Patients [n (%)]
Age (years)*	63.1 ± 10.2 (43-84)
Duration of onset glaucoma (months) $^*$	55.5 ± 45.3 (1-204)
Gender	
Female	22 (44.9)
Male	27 (55.1)
Affected eye	
Right	40 (50.0)
Left	40 (50.0)
Comorbidity	
Hypertension	15 (18.8)
Diabetes Mellitus	12 (16.3)
Hyperthyroidism	2 (2.5)
None	49 (59.9)

\*Data were expressed as mean ± SD (min-max).

#### Table 2: Examinational findings.

	Number of Patients [n (%)]
BCVA (Snellen Unit)*	$0.86 \pm 0.26 \ (0.0080 - 1.0)$
IOP (mm Hg)*	16.3 ± 3.5 (6-23)
Biomicroscopic findings	
Cataract	3 (3.9)
Nepheline	2 (2.5)
Nevus Filament	1 (1.3)
Pterygium	2 (2.5)
Biomicroscopic C/D ratio*	$0.6 \pm 0.3 \ (0.1 \text{-} 1.0)$
Lens statement	
Phakic	60 (75)
Pseudophakic	20 (25)

**BCVA:** Best corrected visual acuity, **IOP:** Intraocular pressure, **C/D:** Cup-to-disc

\*Data were expressed as mean ± standard deviation (min-max).

IOP, biomicroscopic findings, lens status, and biomicroscopic C/D ratio) was shown in Table 2.

In terms of perimetry and OCT results, the mean VFI was revealed as 74.2  $\pm$  34.6 % (ranging from 0 to 100%), and the mean thickness of RNFL was found out to be 76.9  $\pm$  17.5  $\mu$ m. The summary of perimetry and OCT results (VFI, MD, PSD, SF, CPSD, mean C/D ratio, mean RNFL thickness) is shown in Table 3.

When mean C/D ratio and other quadrants of RNFL were observed, mean C/D ratio, nasal and temporal quadrants did not have a significant correlation (p=0.082 for mean C/D ratio, p=0.237 for nasal quadrant, p=0.211 for temporal quadrant). The inferior quadrant thickness had a negative correlation with age (p=0.01, r=-0.273).



The mean thickness of the RNFL and BCVA had a statistically significant linear relationship (p<0.001, r=0.418). Thinner RNFL was also observed in eyes with lower BCVA, indicating that they are positively correlated. VFI and BCVA also had a statistically significant linear relationship (p=0.001, r=0.441). It was observed that eyes with lower BCVA had worsened VFI. Additionally, MD had a statistically significant linear relationship with BCVA (p<0.001, r=0.545). RNFL inferior quadrant (p<0.001, r=0.457) and RNFL superior quadrant (p=0.006, r=0.303) had a moderate positive correlation with BCVA. All of the correlations are presented in Table 4 and Table 5.

The percentages of anti-glaucomatous agents' usage were observed to be 67.5% for prostaglandins, 68.8% for beta-blockers, 61.3% for carbonic anhydrase inhibitors, and 47.5% for alpha agonists. Regarding the operative approach, 11 (13.8%) eyes underwent trabeculectomy, 8 (10.0%) eyes underwent phacoemulsification cataract extraction and 1 (1.3%) eye underwent vitrectomy. Table 3: Perimetry and optical coherence tomography results.

	$Mean \pm SD (min-max)$
Perimetry	
VFI (%)	74.2 ± 34.6 (0 – 100)
MD (dB)	-9.6 ± 9.1 (-30.9 – 1.1)
PSD (dB)	5.6 ± 3.1 (1.5 – 14.4)
SF (dB)	$2.0 \pm 1.0 \ (0 - 5.7)$
CPSD (dB)	4.8 ± 3.3 (0 – 14.1)
<b>Optical Coherence Tomography</b>	
Mean C/D Ratio	$0.7 \pm 0.2 \ (0.1 - 1.1)$
Mean RNFL Thickness (µm)	76.9 ± 17.5 (40 – 115)

VFI: Visual field index, MD: Mean defect, PSD: Pattern standard deviation, SF: Short-term fluctuation, CPSD: Corrected pattern standard deviation, C/D: Cup-to-disc, RNFL: Retinal nerve fiber layer

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Continuous variables	r	P-value	r	P-value	Z	P-value	Z	P-value	Z	P-value	Z	P-value
VFI	-0.141	0.213	0.084	0.0460	-0.526	0.599	-1.510	0.131	-0.927	0.354	-2.531	0.011
MD	-0.165	0.144	0.117	0.300	-0.706	0.480	-1.639	0.101	-0.801	0.423	-2.895	0.004
PSD	-0.118	0.299	-0.002	0.98	-0.140	0.889	-0.523	0.601	-1.998	0.046	-1.600	0.110
The mean thickness of RNFL	-0.176	0.118	0.157	0.165	-0.706	0.480	-0.835	0.404	-1.937	0.053	-2.008	0.003
I Quadrant	-0.273	0.014	0.044	0.696	-0.457	0.648	-0.790	0.429	-1.264	0.206	-2.082	0.037
S Quadrant	-0.122	0.282	0.110	0.333	-0.280	0.779	-0.741	0.131	-1.408	0.159	-3.011	0.003
N Quadrant	-0.185	0.101	-0.048	0.672	-0.336	0.736	-0.702	0.483	-1.347	0.178	-1.572	0.116
T Quadrant	-0.207	0.066	-0.040	0.725	-0.519	0.480	-0.672	0.502	-0.457	0.647	-0.800	0.424
Mean C/D Ratio	-0.135	0.233	0.113	0.319	-1.288	0.198	-1.655	0.098	-0.915	0.360	-4.583	< 0.001

VFI: Visual field index, MD: Mean defect, PSD: Pattern standard deviation, RNFL: Retinal nerve fiber layer, I Quadrant: Inferior quadrant, S Quadrant: Superior quadrant, N quadrant: Nasal quadrant, T Quadrant: Temporal quadrant, BB: Beta blocker, AA: Alpha agonist, CAI: Carbonic anhydrase inhibitor

# Table 5: The strength and direction of linear relationships between pairs of continuous variables.

	BCVA				
Continuous variables	r	P-value			
VFI	0.441	0.001			
MD	0.545	< 0.001			
The mean thickness of RNFL	0.418	< 0.001			
I Quadrant	0.547	< 0.001			
S Quadrant	0.303	0.006			
N Quadrant	134	0.237			
T Quadrant	0.141	0.211			
Mean C/D Ratio	-0.196	0.082			
Biomicroscopic C/D Ratio	-0.263	0.018			
IOP	0.136	0.228			

BCVA: Best-corrected visual acuity, r: Pearson correlation coefficient, VFI: Visual field index, MD: Mean defect, RNFL: Retinal nerve fiber layer, I quadrant: Inferior quadrant, S quadrant: Superior quadrant, N quadrant: Nasal quadrant, T quadrant: Temporal quadrant, C/D: Cup-to-disc, IOP: Intraocular pressure

#### DISCUSSION

Primary open-angle glaucoma is an asymptomatic, progressive, and degenerative optic neuropathy characterized by enlarging optic disc and loss of visual field. Patients at high risk for POAG include black people older than 40 years, white people older than 65 years, and people with a family history of glaucoma or people with a history of diabetes or severe myopia (-6.00 diopters or more, i.e., farther from 0.00) (6). In our study, the mean age of people with POAG was 63.1 years, many other studies confirming an older age prevalence as well (1, 6, 8-10). It was also seen that hypertension and diabetes were two of the major risk factors for POAG. Other studies with different ethnicities have revealed that blood pressure changes and onset diabetes played a major role in increasing the risk of POAG (11, 12). Therefore, adequate management of blood pressure, blood sugar levels, and HbA1c are crucial for decreasing the risk of POAG.

In our study, mean BCVA results were seen as 0.86, with mean IOP being in the normal range. Our study did not reveal a statistically significant linear relationship between IOP and BCVA. The fact that POAG could occur and progress with normal IOP should not be overlooked. The study with 229 eyes conducted by Omodaka et al. (13) confirmed that IOP had a weak correlation with BCVA,



which is in line with the results of our study. According to the study of Leung et al. (14), better BCVA is more correlated with normal RNFL thickness, especially with the thickness of the temporal quadrant RNFL. On the contrary, our study revealed the temporal quadrant of RNFL to not have a correlation with BCVA, unlike superior and inferior quadrants which have strong correlations with BCVA. This situation could be explained by the earlier and more destructive effects of POAG on inferior and superior quadrants. In addition, it was seen that older patients had thinner inferior RNFL quadrant. It is believed to occur due to the age-related changes in the retinal layer of the globe. Moreover, the mean thickness of the RNFL was within the normal range in our study, which could explain the reason for the better BCVA results.

Visual field loss is also one of the other causes of POAG. Peripheral vision loss is usually the initial defect and is followed by central vision loss (blindness) in POAG. Perimetry should be done in patients with POAG to detect visual field defects. Perimetry is the gold standard investigation tool in which defects are detectable when 40-50% loss of ganglion cells occurs. Therefore, it is believed that structural tests can identify progression in the early stages better while perimetry is more useful in the later stages (15-20). Thus, it is believed that perimetry is more relevant in the later stages of POAG (18). On the contrary, OCT detects even the slightest defects on the optic disc head which gives a higher detection chance of disease progression in the early stages and is still indeed a handy tool for POAG and examination of worsened BCVA. Nonetheless, the fact that moderate BCVA could be gotten despite OCT results even with minimal defects should be noted (7). Therefore, examination of peripapillary RNFL thickness will provide a better chance of detecting early glaucoma than that of perimetry (14, 18). In our study, it was observed that perimetry parameters were more accurate in POAG than the results of OCT in which VFI and MD had better strength to reveal the defects than the parameters of OCT such as mean RNFL thickness, superior and inferior quadrants of RNFL. These results could be attributed to the fact that the patient group in our study was referred to our tertiary clinic from another one. Additionally, the study of Banegas et al. (16) supported the functionality of OCT in progression detection of pre-perimetric glaucoma, while perimetry is more reliable in later stages. Given these facts, we believe that worsened BCVA is better explained with perimetry in the tertiary clinics.

The main limitation of our study is that our clinic is considered a tertiary center which meant that the majority of our patients were already on anti-glaucomatous medications at the time of their referral, indicating that their POAG was relatively under control. This left us with better parameters of POAG such as BCVA, IOP, RNFL thickness, C/D ratio, which were used for investigation and management.

In conclusion, OCT has higher sensitivity in the detection of progression than perimetry in the early stages, which gives OCT the ability of progression detection within a shorter follow-up time in early POAG. On the other hand, perimetry provides us better and more accurate results for POAG management in the later stages. Therefore, our study has revealed that in tertiary clinics that receive various numbers of patient referrals, perimetry is more useful and accurate in POAG detection and monitoring. Using both OCT and perimetry for disease monitoring is advisable, as this provides us better disease progression tracking with a better chance than using one method without the other. *Ethics Committee Approval:* This study was approved by the Scientific Research Ethics Committee of Trakya University School of Medicine (Protocol Code: TÜTF-BAEK 2021/104).

*Informed Consent:* Informed consent was obtained from the participants of this study.

Conflict of Interest: The authors declared no conflict of interest.

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