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#### ORIGINAL RESEARCH

Comparison of Posterior Hyaloid Assessment Using Preoperative Optical Coherence Tomography and Intraoperative Triamcinolone Acetonide Staining During Vitrectomy

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Purpose: To compare the specificity of diagnosing posterior vitreous detachment (PVD) using preoperative optical coherence tomography (OCT) versus intraoperative triamcinolone acetonide (TA) staining in patients undergoing vitrectomy.

Patients and Methods: This retrospective cohort study included patients undergoing pars plana vitrectomy for diverse retinal pathologies. Intraoperatively, surgeons evaluated the posterior hyaloid status with TA staining and compared it with preoperative OCT findings. Results: One hundred six patients underwent intraoperative assessments of posterior hyaloid

status, with 72% (76/106) of the eyes showing positive staining. Sixty-two patients had also undergone preoperative OCT. Of the patients diagnosed with PVD on preoperative OCT, 50% (15/30) showed positive TA staining intraoperatively. The sensitivity of preoperative OCT assessment was 83.3%, and its specificity was 65.9%.

Conclusion: Preoperative OCT imaging is associated with lower sensitivity and specificity for diagnosing PVD when compared to intraoperative TA staining.

Keywords: optical coherence tomography, posterior hyaloid, posterior vitreous detachment, vitrectomy, vitreoschisis, triamcinolone acetonide

#### Introduction

The vitreous, a colorless, gel-like substance, is an extracellular matrix that plays a key role in ocular disease and health.<sup>1</sup> In young individuals,<sup>2</sup> the vitreous has a gel-like consistency, with strong overall attachment to the retina.<sup>3</sup> As an individual ages, however, alterations in macromolecular interactions within the vitreous create a more liquid consistency. This liquefaction, known as syneresis, causes destabilization of the vitreous and leads to vitreous collapse and consequent posterior vitreous detachment (PVD).<sup>4</sup>

Preoperative recognition of the posterior hyaloid status helps the surgeon to better understand the surgical anatomy and allows a more precise planning of the surgical approach.<sup>5</sup> In addition, the use of colored stains and particulate suspensions (eg, triamcinolone acetonide [TA]) intraoperatively can facilitate visualization of the multilamellar nature of the vitreous.<sup>6</sup> Intraoperative identification and removal of the posterior hyaloid may mitigate the risk of postoperative complications, including recurrence of retinal detachment and formation of epiretinal membrane (ERM). This study aimed to compare intraoperative assessments of the posterior hyaloid presence by surgeons after triamcinolone staining with the preoperative diagnosis of PVD using spectral-domain optical coherence tomography (OCT).

## **Patients and Methods**

This retrospective cohort study compared OCT-based preoperative diagnosis of PVD with intraoperative TA-based assessments of the presence or absence of residual vitreous or posterior hyaloid after a full vitrectomy. The study sample consisted of 106 eyes of 106 patients who had undergone intraoperative TA staining, of which preoperative OCT data of sufficient quality were available for 62 eyes. All patients were evaluated and assessed at King Khaled Eve Specialist Hospital (KKESH), Rivadh, Saudi Arabia, from January 2018 to March 2020. Patients with a diagnosis of tractional retinal detachment, vitreous hemorrhage (VH), macular hole (MH), ERM, vitreomacular traction (VMT) and those with rhegmatogenous retinal detachment (RRD) for no longer than 1 month were included. Exclusion criteria included media opacities (eg, cataract or corneal opacity), recurrent retinal detachment, and previous surgical interventions for any retinal pathology. This study adhered to the tenets of the Declaration of Helsinki. The ethics review board at KKESH granted IRB approval for this study. Informed consent was not required due to the retrospective nature of the study.

Preoperatively, patients underwent a clinical assessment that included measurements of best-corrected visual acuity using a Snellen chart, fundus examination, and applanation tonometry. Retinal structure and vitreoretinal interface were analyzed qualitatively and quantitatively using transforeal horizontal spectral-domain OCT scans (Spectralis OCT; Heidelberg Engineering, Inc., Heidelberg, Germany) and macular volume raster scans consisting of 19 horizontal 6mm high-resolution horizontal line scans obtained within 90 days before the surgery. A trained technician performed the OCT acquisition for all eyes and two masked ophthalmologists (VAS and WAA) assessed the presence or absence of PVD from the OCT results. Disagreements between the assessors were resolved with joint review of the scans. Staging was based on a previously published OCT staging system, where stages 0, 1, 2, and 3 were jointly classified as attached vitreous and stage 4 was categorized as total PVD.<sup>7</sup>

Surgery was performed using a standard three-port full vitrectomy (23-gauge). Next, intravitreal TA was administered (KENACORT OR KENALOG 40 mg/mL diluted 1:4), and surgeons evaluated the presence or absence of a posterior vitreous detachment. After TA administration,

the surgeon's preferred surgical approach was used for the procedure. Any intraoperative complications were documented.

Data were collected in an Excel spreadsheet (v. 2010; Microsoft Corp., Redmond, WA, USA). Data were then cleaned, managed, and coded in Excel. Statistical analyses were performed with SPSS (v. 19.0, IBM Corp., Armonk, NY, USA). Confidence intervals were set as 95%, with a corresponding p-value less than 0.05 was considered statistically significant. Descriptive analysis was done where categorical variables were presented as frequency (%), while continuous variables were presented as mean (SD). Inferential analysis was then done, where intraoperative findings were correlated to the diagnoses from preoperative OCT imaging. Sensitivity and specificity analysis was conducted considering Triamcinolone staining as the gold standard. Associated sensitivity, specificity in addition to both positive and negative predictive values were calculated.

### Results

The study sample included 55 males (52%) and 51 females (48%). At presentation, the mean age of the patients was 53 years (Table 1). The retinal pathologies requiring vitrectomy in our patients are described in Table 2. Preoperative OCT data were available for 62 patients. Among these, 32 eyes had an attached vitreous: 6 were classified as stage 0, 14 as stage 1, 9 as stage 2, and 3 as stage 3; the 30 remaining eyes were categorized as stage 4 with a total PVD. We excluded 44/106 eyes from OCT analysis because of inadequate quality of OCT imaging (Table 3). After performing a core vitrectomy and staining with TA (Table 4), PVD was identified in 30/106 patients (28.3%), and hyaloid attachment was noted in the remaining 76 patients (71.7%). Of the 30 eyes graded as complete PVD on OCT, 15 (50%) had pre-existing PVD at the time of surgery, and 15 eyes showed attached vitreous at the time of surgery (false-positive results). Of the 32 eyes graded as attached vitreous on OCT, 29 showed attached vitreous at the time of surgery and 3 showed pre-existing PVD at the time of surgery (false-negative results) (Figure 1). The sensitivity and the specificity of preoperative OCT assessment was 83.3% and 65.9%, respectively. The positive predictive value was 43.6% and the negative predictive value was 92.6% assuming a disease prevalence of 24% (Table 5).

		n	Percentage
Age (mean ±SD)		106	52.73 ± 12.86
Gender	Male	55	51.9
	Female	51	48.1
Eye involved	Right	59	55.7
	Left	47	44.3
Surgeon	Consultant	59	55.7
	Fellow	47	44.3

#### Table I Patient Demographics and Previous Ocular History

#### Table 2 Retinal Pathology for Surgical Indication of Vitrectomy

	n	Percentage
RRD	32	30.2
TRD	41	38.7
VH	10	9.4
VMT	3	2.8
ERM	4	3.8
FTMH	14	13.2
DLM	2	1.9

Abbreviations: RRD, rhegmatogenous retinal detachment; TRD, tractional retinal detachment; VH, vitreous haemorrhage; VMT, vitreomacular traction; ERM, epiretinal membrane; FTMH, full-thickness macular hole; DLM, Dropped Lens Material.

Table 3	Quality	of OCT	Results	Among	Study	Participants
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	Frequency	Percentage
PVD present	30	28.3
PVD absent	32	30.2
PVD could not be assessed (poor image quality)	44	41.5
Total	106	

Abbreviation: PVD, posterior vitreous detachment.

Table 4 Status of Posterior Hyaloid After TA Staining

	Frequency	Percentage
Detachment present	30	28.3
Detachment absent	76	71.7
Total	106	100

## Discussion

The main finding in this study is that preoperative OCT may not be sufficiently sensitive to detect the status of the posterior hyaloid, which seems to imply that intraoperative TA staining is recommendable in most vitreoretinal pathologies. This confirms the findings described previously by Hwang et al<sup>7</sup> and Stavrakas et al.<sup>8</sup> The new approach of

this study is that we, for the first time, quantified the ability of OCT to diagnose PVD, compared to intraoperative TA-assisted assessment, which allowed us to estimate the sensitivity and specificity of this OCT-based diagnosis, in a range of retinal conditions.

In retinal pathologies such as MH, retinal detachment, proliferative vitreoretinopathy, and VMT syndrome, a partially

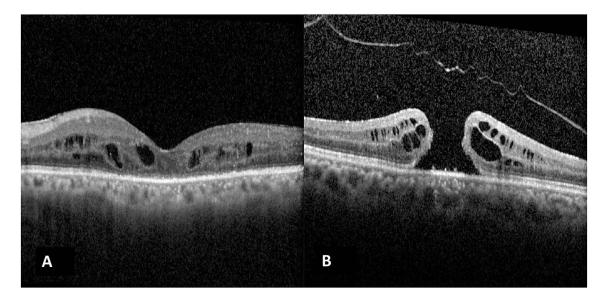


Figure I OCT image of a case labelled as Grade 4 PVD on OCT (**A**), but intraoperative TA staining determined a posterior hyaloid attachment (false-positive). Case (**B**) was labelled to have Grade 3 OCT (attached to the optic nerve head) although intraoperative staining showed that a complete PVD was present (false-negative).

separated or attached posterior hyaloid can create traction or, in patients with proliferative diabetic retinopathy, generate a framework for fibrovascular proliferation.<sup>9</sup> In this respect, the knowledge of the posterior vitreoretinal interface anatomy is the key for a successful treatment of a large variety of surgical retinal disease. In a clinical setting, posterior hyaloid status has been found not to affect the efficacy of intravitreal injection courses for diabetic macular edema or age-related macular degeneration.<sup>10–12</sup> However, in patients who have received therapy with intravitreal ocriplasmin for symptomatic vitreoretinal adhesion,<sup>13</sup> ellipsoid zone disruption has been suggested to be a preexisting finding of the disease, leading to lack of visual improvement after treatment.

The outcomes of the current study indicate that a posterior vitreal attachment is common, with 71.7% (76/106) of patients showing this finding in association with a range of retinal baseline pathologies. The principal aim of vitrectomy is to completely remove the posterior hyaloid. Failure to achieve this step is associated with postoperative

complications. A study by Yamashita et al reinforced the theory that ERMs are produced by anomalous PVDs with vitreoschisis, leaving the topmost layer of the posterior vitreous cortex attached to the macula.<sup>14</sup> Enaida et al performed a nonrandomized study of 94 patients who underwent TA-assisted pars plana vitrectomies and 83 patients who underwent non-TA-assisted pars plana vitrectomies, reporting that 22.9% of the latter required further surgical intervention, compared to 11.7% of the TA-assisted patients.<sup>15</sup>

A possible change in PVD status from the moment of OCT acquisition until surgery has been previously proposed<sup>7</sup> and found not to be a significant variable. Most of our patients underwent surgery shortly after OCT. Some elective cases (macular holes or ERM) did have a longer period between OCT and intervention. However, due to the chronic nature of the vitreoretinal interface pathology for these patients, we did not believe that the status of the hyaloid would change. It is also

Table 5 Overall Sensitivity a	and Specificity of C	<b>D</b> ptical Coherence	Tomography versus	Triamcinolone Staining
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		Triamcinolone Staining (Gold Standard)		
		PVD Present	PVD Absent	Total
OCT findings	PVD present	15 (83.3%)	15 (34.1%)	30
	PVD absent	3 (16.7%)	29 (65.9%)	32
	Total	18 (100%)	44 (100%)	62

Notes: Sensitivity = 83.3%; specificity = 65.9%; positive predictive value = 43.56%; negative predictive value = 92.6; assuming a disease prevalence of 24%. Abbreviations: PVD, posterior vitreous detachment; OCT, optical coherence tomography.

noteworthy that if a preoperative OCT shows PVD, this is unlikely to change with time. False positives occurring by PVD development between the time of OCT and the surgery cannot be excluded.

Our current study demonstrated that preoperative OCT imaging can facilitate the diagnosis of PVD, although its reliability was low due to the presence of artifacts (media opacity and vitreoschisis). In the present study, 48.4% (30/62) of the patients had PVD based on OCT imaging. However, TA staining confirmed that only 50% (15) of these patients had a true PVD. On the other hand, 3 of the 32 patients who showed an attached hyaloid on OCT turned out to have complete PVD when stained by TA. This outcome indicates the low sensitivity and specificity of preoperative OCT (83.3% and 65.9%, respectively). Disease based specificity and sensitivity for PVD diagnosis on OCT was unable to calculated due to the largely uneven number of patients in each disease group (Table 2).

Several factors present challenges in assessing the status of the posterior hyaloid, such as media opacity (cataracts or VH) and retinal detachment, because of which an adequate scan cannot be effectively obtained to analyze the posterior hyaloid or retinal surface. In our study, 44 patients had to be excluded from OCT analysis, due to these factors. Similar to previous findings,<sup>7</sup> we also encountered difficulty when assessing extreme stages (0 or 4) of PVD. Eyes were categorized as stage 0 when the posterior hyaloid could not be clearly evaluated due to its strong adhesion to the retina, but the premacular bursa of Worst could be easily identified. Similarly, in stage 4 grading, we cannot see the posterior hyaloid due to its significant separation from the retina, but it is so far in front of the retina that the OCT cannot capture it. However, in stage 4, we would not be able to see the premacular bursa of Worst. Hwang et al compared preoperative OCT findings to operative reports describing the posterior vitreous status.7 However, triamcinolone staining was not used in 28% of their cases, leading to a wide range of possible reasons for intraoperative misdiagnosis of the posterior hyaloid, including the surgeons' level of experience, absence of TA administration, and limitations in report dictation and documentation. We decided to address all these factors by only including patients in whom TA staining had been performed and intraoperative posterior hyaloid status was documented.

The presence of ERMs could also confuse the examiner and lead to misdiagnosis. Our preoperative OCT findings mimic those obtained by Kičová et al,<sup>16</sup> who compared biomicroscopy, B-scan ultrasound, and OCT for clinically detecting PVD preoperatively and indicated that OCT has the lowest reliability index (12.5% of 30 eyes) for determining the correct diagnosis among all modalities. They also sought to identify minimally invasive imaging modalities that could be used reliably and efficiently in clinical settings for the detection of PVD, reporting that B-scan ultrasound was accurate but, importantly, operator-dependent. Similar observations were published by Mugit et al and Vaz-Pereira et al.<sup>17,18</sup> who evaluated the OCT features of neovascular complexes and correlated the results of an earlier study by Chu et al.<sup>19</sup> Gupta et al found that vitreoschisis was present in 53% of patients with MH, as revealed by OCT.<sup>20</sup> Recently, Stavrakas et al<sup>8</sup> reported low sensitivity and specificity (37.5% and 31.3%) of spectral domain OCT in patients undergoing vitrectomy for diagnosis of ERM, MH and VMT. Although several studies have attempted to describe the findings in diabetic patients with ERM on time-domain OCT<sup>21</sup> and more recently on spectral-domain OCT,<sup>22</sup> our description of the intraoperative findings and poor specificity of OCT are consistent with a report by Gandorfer et al,<sup>23</sup> in which 60 of 61 eyes with diabetic macular edema (98.3%) and tangential VMT were examined with electron microscopy and showed native vitreous collagen overlying the macula. Thus, it is reasonable to conclude that the intraoperative TA-assisted technique for determining posterior hyaloid status is much more reliable than preoperative OCT imaging.

Another pathology that can lead to preoperative or intraoperative misdiagnosis is vitreoschisis, which refers to a split in the posterior vitreous cortex. Recently, combined OCT and scanning laser ophthalmoscopy (SLO) have been used to clinically identify vitreoschisis. A previous study reported that OCT/SLO detected vitreoschisis in 53% of patients with MH and in 43% of patients with macular pucker.<sup>24</sup> Additionally, Schwatz et al<sup>25</sup> retrospectively evaluated patients with late-stage proliferative diabetic retinopathy who had undergone vitrectomy over a 2-year period and found positive immunohistochemical staining for type II collagen antibody, indicating the vitreal origin of the membranes. They concluded that there was a high probability that the supplemental ERMs were the posterior leaf of torn cortical gel.

There are some limitations to this study, including the diversity of retinal pathologies and the lack of preoperative OCT data for all patients. Future studies should enroll a larger sample size with lower heterogeneity in retinal pathologies and perform additional OCT evaluations with supplementary/comparative preoperative B-scan assessments. Additionally, a new clinical imaging method for assessing posterior hyaloid status prior to TA staining should be devised with adequate validity, reliability, and reproducibility. Furthermore, studies including the preoperative assessment with wide-field OCT viewing are warranted.

## Conclusion

A key objective of vitrectomy is the complete visualization of the posterior hyaloid and vitreous cortex. The outcomes of this study indicated that preoperative OCT imaging is associated with a low sensitivity and specificity for detecting vitreoschisis and PVD. Surgeons might consider performing TA staining of the vitreous to potentially prevent complications, negative patient outcomes, and the need for reoperations.

# **Data Sharing Statement**

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

# **Ethics Approval**

The ethics review board at KKESH granted IRB approval for this study.

# **Consent for Publication**

Written informed consent for publication was not required due to the retrospective nature study of this study and the lack of identifiable patient data.

# **Consent to Participate**

Informed consent was not required due to the retrospective nature of the study.

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# Disclosure

The authors declare that they have no competing interests in this work.

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