ORIGINAL RESEARCH

Results of Pars Plana Vitrectomy for Different Types of Macular Holes

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Purpose: To compare different types of macular holes regarding the anatomic and functional success following pars plana vitrectomy (PPV) and internal limiting membrane (ILM) removal. **Methods:** A retrospective review of all patients with macular holes treated by PPV, ILM removal with gas tamponade from January 2014 to July 2017 in Magrabi Eye Hospital.

Results: One hundred fifty-seven eyes of 153 patients were analyzed. The eyes were classified according to the etiology of macular hole into four groups: 79 eyes with idiopathic macular hole (IMH), 51 eyes with traumatic macular hole (TMH), 16 eyes with macular hole in diabetic patients (DMH) and 11 eyes with myopic macular hole (MMH). We classified the IMH group based on the International Vitreomacular Traction Study Classification according to size into 3 subgroups; subgroup 1: $\leq 250\mu$, subgroup 2: ≥ 250 to 400 μ and subgroup 3: $\geq 400 \mu$. All types of macular hole showed statistically significant postoperative improvement in BCVA compared to the baseline except cases with MMH. Anatomic postoperative hole closure was achieved in 86.1%, 60.7%, 43.65%, an 45.46% of eyes with IMH, TMH, DMH and MMH, respectively. In eyes with IMH, closure rate in subgroup 1 was significantly higher than in subgroups 2, and 3.

Conclusion: PPV, ILM peel and C2F6 technique yielded variable anatomic and functional outcomes in different types of macular holes. Anatomic results were most favorable in IMH and least favorable in MMH. The smaller the diameter of the hole the better the results. The underlying pathogenetic mechanisms that lead to different types of macular holes are pivotal in determining the final outcome.

Keywords: ILM peeling, macular holes, pars plana vitrectomy

Introduction

Full thickness macular hole is defined as a full-thickness defect that involves all layers of the neurosensory retina from the internal limiting membrane (ILM) to the photoreceptors.¹ Macular holes may be idiopathic, traumatic, myopic with or without retinal detachment or secondary to proliferative diabetic retinopathy (PDR).^{2–4} Pars plana vitrectomy (PPV), removal of the posterior hyaloid, ILM peel and gas tamponade is the standard surgical treatment for different types of macular hole with closure rate more than 90%.^{5–7} The aim of the present study is to compare the anatomic and functional success rates of PPV, ILM peel and gas tamponade in different types of macular hole.

Patients and Methods

We conducted a retrospective review of all patients with macular holes treated by PPV, and ILM peel with gas tamponade from January 2014 to July 2017 in Magrabi

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Eye Hospital, Tanta, Egypt. The Institutional Review Board (IRB) and Research Ethics Committee of Magrabi Eye Hospitals required that all patients participating in the study signed an informed consent prior to enrollment. The consent included a statement that authorized the authors to publish the data of the patients in an anonymous manner that does not reveal their identity. The study adhered strictly to the tenets of the Declaration of Helsinki (the 2013 revision).

Inclusion Criteria

Eyes were classified according to the etiology into four groups: idiopathic macular hole (IMH), traumatic macular hole (TMH), macular hole in diabetic patient (DMH), and macular hole in myopic patient (MMH). Patients presenting with TMH were included only after the lapse of onemonth observation period to allow for spontaneous closure of the hole to happen. The study required that all participants complete a minimum follow-up period of 3 months after surgical intervention.

Exclusion Criteria

The study excluded all patients presenting with macular hole retinal detachment (MHRD), lamellar macular hole, previous surgery for macular hole, and macular holes other than the 4 specified groups.

Examinations

Preoperative evaluation included detailed history, bestcorrected visual acuity (BCVA) assessment using Snellen chart and logMAR equivalent for statistical analysis, intraocular pressure measurement (IOP), slit-lamp anterior segment examination, dilated fundus examination using indirect ophthalmoscope and slit-lamp biomicroscopy using 90D lens. Color fundus photo and optical coherence tomography (OCT) were done using Cirrus OCT (v 4.0.7; Carl Zeiss, Meditec, Inc, Dublin, CA). On OCT, the size of the macular hole was assessed by measuring the minimum linear diameter (MLD), which was defined as the distance in microns between the narrowest points of the macular hole in the mid-retina.

Surgical Procedures

A single surgical technique was used for all patients. A single surgeon (HG) performed all surgeries at the same center. The surgical technique consisted of 3-port PPV using transconjunctival cannulated 20 or 23 gauge. Lensectomy was done if the lens was cataractous or if lens

touch occurred intraoperatively. Core vitrectomy was performed and followed by induction of posterior vitreous detachment (PVD). PVD continued anteriorly as safely as possible. ILM was peeled in all cases using Eckardt end-gripping forceps. ILM peel was extended over at least an area of 2 disc diameters centered onto the macular hole. Triamcinolone acetonide (TA) was used to assist visualization during peeling of ILM. The signs of ILM removal were whitening of the retina with or without petechial hemorrhage, and dull appearance of denuded retina. Brilliant Blue stain, 0.2 mL (Brilliant Peel; Geuder, Heidelberg, Germany) was used if the ILM could not be identified by TA. The surgeon did not attempt to perform an inverted ILM flap technique in any case. Laser barrage 360° was done routinely, followed by air-fluid exchange and gas injection (16% C2F6; Hexafluoroethane). We used C2F6 in all macular holes included regardless of the diameter because we wanted to standardize the treatment approach in all patients and explore the efficacy of this standard approach across different categories of holes. Patients were instructed to maintain face-down position for 1 week after surgery. All patients were examined on the 1st postoperative day. Postoperative follow-up was scheduled at 1 week, 3 weeks, 6 weeks and 8-weekly thereafter. Fundus examination, color fundus photography and OCT were done whenever possible.

Statistical Analysis

Results were collected, tabulated and statistically analyzed by an IBM compatible personal computer with SPSS statistical package version 23 (SPSS Inc. Released 2015. IBM SPSS statistics for windows, version 23.0, Armonk, NY: IBM Corp.). Kruskal–Wallis test was used for comparison of quantitative variables between more than two groups of not-normal distributed data with Tamhane's test as post hoc test. Wilcoxon test was used to compare different readings of not-normally distributed data in the same group. Chi-square test (χ 2) was used to study association between qualitative variables. Whenever any of the expected cells were <5, Fischer's Exact test with Yates correction was used.

Results (Tables 1-4, Figures 1-3)

The study reviewed the medical records of 201 patients with macular hole. Thirty-one patients with MMH associated with retinal detachment were excluded. Another 17 patients who did not complete the required minimum follow-up period were excluded. Table I Demographic, Preoperative BCVA, Postoperative BCVA and Functional Outcomes of the Different Groups

Macular Hole Type	IMH n = 79	TMH n = 51	DMH n = 16	MMH n = 11	p value	Kruskal- Wallis	Fisher's Exact Test
†Mean age (years)	63 ± 7.6	27 ± 15.2	54.3 ± 7.5	42 ± 13	<0.001	76.5	-
Preoperative BCVA Log	1.1±0.31 (20/	1.2±0.3 (20/	1.2±0.4 (20/	1.1±0.2 (20/	0.19	4.7	-
MAR(Snellen)	250±20/40)	300±20/40)	300±20/50)	250±20/30)			
††Postoperative BCVA Log	0.7±0.35 (20/	0.9±0.3 (20/	1.1±0.3 (20/	0.9±0.3 (20/	<0.001	13.3	-
MAR(Snellen)	100±20/35)	160±20/40)	250±20/40)	160±20/40)			
Macular hole closure	86.1%	61%	44%	45.4%	<0.001	-	16.7

Notes: †Post-hoc analysis of mean age in years in different types of macular hole: IMH vs TMH (p < 0.001), IMH vs DMH (p=0.003), IMH vs MMH (p=0.06), DMH vs TMH (p < 0.001), TMH vs MMH (p=0.05), and DMH vs MMH (p=0.14). ††Post-hoc analysis of postoperative visual acuity in different types of macular hole: IMH vs TMH (p=0.01), IMH vs DMH (p=0.02), IMH vs MMH (p=0.16), DMH vs TMH (p=0.35), TMH vs MMH (p=0.16), DMH vs MMH (p=0.16), DMH vs TMH (p=0.35), TMH vs MMH (p=1.0), and DMH vs MMH (p=0.6).

Abbreviations: DMH, diabetic macular hole; IMH, idiopathic macular hole; MMH, myopic macular hole; n, number; TMH, traumatic macular hole; BCVA, best-corrected visual acuity.

 Table 2 Comparison Between Pre- and Postoperative BCVA of Different Groups of Macular Holes

Macular Hole Type	IMH n = 79	TMH n = 51	DMH n = 16	MMH n = 11
Preoperative BCVA Log MAR (Snellen)	1.1 ± 0.31(20/250±20/40)	1.2±0.3 (20/300±20/40)	1.2±0.4 (20/300±20/50)	1.1±0.2 (20/250±20/30)
Postoperative BCVA Log MAR (Snellen)	0.7±0.35 (20/100±20/35)	0.9±0.3 (20/160±20/40)	1.1±0.3 (20/250±20/40)	0.9±0.3 (20/160±20/40)
Wilcoxon test	5.9	4.5	2.3	1.7
p value	<0.001	<0.001	0.02	0.08

Abbreviations: DMH, diabetic macular hole; IMH, idiopathic macular hole; MMH, myopic macular hole; n, number; TMH, traumatic macular hole; BCVA, best-corrected visual acuity.

Macular Hole Size (µ)	<250 (n = 36)	250–400 (n = 33)	>400 (n = 10)
Preoperative VA Log MAR (Snellen)	0.9 ± 0.3 (20/160±20/40)	1.2 ± 0.2 (20/300±20/30)	1.4 ± 0.04 (20/500±20/22)
Postoperative VA Log MAR (Snellen)	0.4 ± 0.3 (20/50±20/40)	0.9 ± 0.3 (20/160±20/40)	0.9±0.35 (20/160±20/45)
Wilcoxon test	3.9	3.5	2.2
p value	<0.001	<0.001	0.05
[#] Macular hole closure (%)	97.2	79	70
Fisher's exact test	7.3	7.3	7.3
p value	0.02	0.02	0.02

Table 3 Anatomic and Functional Outcomes of Subgroups of Idiopathic Macular Hole

Notes: [#]Pairwise comparison between different sizes of IMH in terms of closure rate: $<250\mu$ vs $250-400\mu$ (p=0.04), 250μ vs $>400\mu$ (p=0.03), $250-400\mu$ vs $>400\mu$ (p=0.09). Abbreviations: μ , micron; n, number; VA, visual acuity.

Macular Hole Type	IMH n = 79	TMH n = 51	DMH n = 16	MMH n = 11
Closed Log MAR (Snellen)	0.60±0.3 (20/80±20/40)	0.76±0.3 (20/100±20/40)	1±0.3 (20/200±20/40)	0.90±0.14 (20/160±20/25)
Open Log MAR (Snellen)	0.95±0.41 (20/180±20/50)	1.2±0.2 (20/315±20/30)	1.2±0.35 (20/315±20/45)	1.04±0.15 (20/220±20/28)
p value	0.008	<0.001	0.142	0.286

Abbreviations: DMH, diabetic macular hole; IMH, idiopathic macular hole; MMH, myopic macular hole; n, number; TMH, traumatic macular hole; BCVA, best-corrected visual acuity.

Baseline Characteristics of the Study Population

One hundred fifty-seven eyes of 153 patients were analyzed. The eyes were classified according to the etiology of the macular hole into four groups: 79 eyes with IMH, 51 eyes with TMH, 16 eyes with DMH and 11 eyes with MMH. We further classified the IMH group based on the International Vitreomacular Traction Study Classification according to size into 3 subgroups. Subgroup 1, included eyes with hole size $\leq 250 \mu$; n = 36. Subgroup 2, included eyes with hole size ranging from $\geq 250-400\mu$; n = 33. Subgroup 3, included eyes with hole size $\geq 400 \mu$; n = 10.

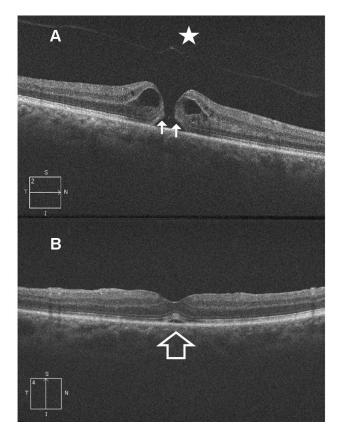


Figure I (**A**) High-definition 5-line raster OCT image of the right eye of a 66-yearold male with idiopathic FTMH. Note that the posterior hyaloid is fully detached along with an operculum (star). The hole edges are thickened with cystic spaces in a typical pregnant draw-bridge appearance. Both ends of the ELM and IS/OS junction layers are drawn upwards (2 white arrows). MLD of the hole was 216µ. Preoperative BCVA was 6/24. (**B**) Postoperative OCT image of the same patient taken at I-month follow-up visit. Note U-type hole closure with restoration of ELM and IS/OS layers. There is a residual shallow sub-foveal neurosensory detachment (white arrow), Postoperative BCVA was 6/9.

The mean preoperative BCVA was distributed among the 4 groups as follows; In the IMH group; 1.10 logMAR \pm 0.31, in the TMH group; 1.2 logMAR \pm 0.3, in the DMH; 1.21 logMAR \pm 0.4 and in the MMH group; 1.1 \pm 0.2. Eyes with IMH size $\leq 250\mu$ had significantly better preoperative BCVA (0.9 logMAR \pm 0.3) than eyes with hole size $\geq 250-400\mu$ (1.2 logMAR \pm 0.2) and eyes with hole size $\geq 400\mu$ (1.4 logMAR \pm 0.04). Eyes with hole size $\geq 400\mu$ had better BCVA than eyes with hole size $\geq 400\mu$.

Postoperative Visual Outcome

Mean postoperative BCVA was 0.7 logMAR \pm 0.35, 0.9 logMAR \pm 0.3, 1.1 logMAR \pm 0.3, and 0.9 logMAR \pm 0.3 in IMH, TMH, DMH, and MMH groups, respectively. All macular holes showed statistically significant postoperative improvement in BCVA compared to baseline except

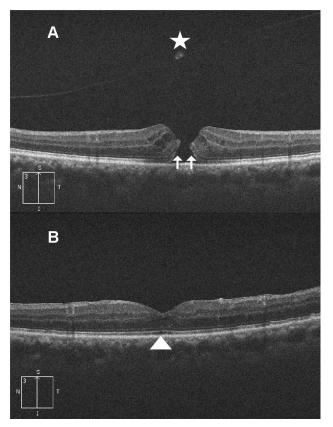


Figure 2 (**A**) High-definition 5-line raster OCT image of the left eye of a 75-yearold female with idiopathic FTMH. Note the fully detached posterior hyaloid and the hole operculum (star). The hole shows thickened edges with cystic spaces, and upward lift of the ELM and IS/OS junction layers (2 white arrows). MLD of the hole was 200µ. Preoperative BCVA was 6/36. (**B**) Postoperative OCT image of the same patient taken at 3-month follow-up visit. Note U-type hole closure with restoration of ELM and IS/OS layers. Note the residual defect in the IS/OS junction layer (white arrow head). Postoperative BCVA improved to 6/9.

cases with MMH (p< 0.05 in IMH, TMH, DMH and 0.08 in MMH).

Postoperative Closure Outcome

Macular hole closure was achieved in 86.1%, 61%, 44%, and 45.4% of eyes with IMH, TMH, DMH, and MMH groups, respectively. The rate of hole closure was significantly higher in IMH than other types of holes. Subgroup analysis of the IMH group revealed macular hole closure rate of 97.2%, 79%, and 70% in subgroups 1, 2, and 3, respectively. Macular hole closure rate was significantly higher in subgroup 1 than in subgroups 2, and 3. The postoperative VA stratified according to macular hole status in different groups of macular holes is shown in Table 4.

Discussion

Different pathologies lead to different types of macular hole. IMH may occur due to the anteroposterior and

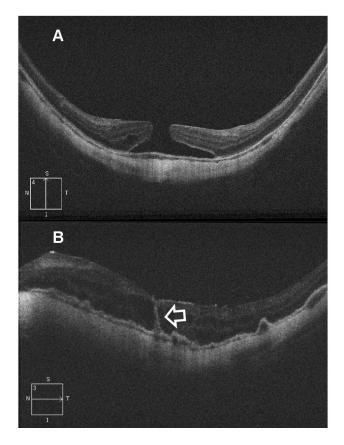


Figure 3 (**A**) High-definition 5-line raster OCT image of the left eye of a 46-yearold female with myopic FTMH. Note the OCT features of myopia as posterior bowing of the sclera, thinning of the choriocapillaris and dome-shaped macula. MLD of the hole was 328μ . Preoperative BCVA was 5/60. (**B**) Postoperative OCT image of the same patient taken at 1-month follow-up visit. The hole has closed though with interrupted ellipsoid zone. Note the hyporeflective intra-retinal cystic spaces. The hyperreflective vertical line traversing the neurosensory retina from the fovea to the RPE represents a track line which is a marker of previous injury to ELM layer and damage to the photoreceptors around the line (white arrow). The RPE is thickened and shows multiple pigment epithelial detachments. Postoperative BCVA remained 5/60.

tangential vitreomacular traction exerted by the posterior vitreous cortex at the fovea.^{8,9} DMH may occur due to tangential traction of the premacular fibrosis superimposed on long standing cystoid macular edema, rupture of the cyst or traction on the already weakened ischemic fovea.^{3,10} The pathogenesis of TMH is related to the avulsion force applied to the fovea by the vitreous due to anterior-posterior compression or persistent vitreoretinal adhesion.^{11,12} In MMH, tangential traction with posterior staphyloma may be a predisposing factor.¹³ In the current study we compared the anatomic and functional outcomes in patients with IMH, TMH, DMH and MMH that underwent PPV and ILM peeling with gas injection. Patients with IMH had significantly higher age than all other types of macular hole. There was a significant improvement in postoperative BCVA of all types of macular holes as

compared to baseline except in MMH group (P <0.05 in IMH, TMH, DMH and 0.08 in MMH). In IMH group, we detected that eyes in subgroups 1, and 2 had significant visual improvement (P < 0.001). On the other hand, we did not detect similar significant improvement in visual acuity in eyes in subgroup 3 (P=0.05). In addition, visual acuity improvement was related to the size of the hole in the sense that the smaller the hole diameter the better the visual outcome. These results are in accordance with those of Huang et al¹¹ who reported that larger IMHs (basal and apical area) were associated with worse visual acuity. Similarly, previous studies reported that postoperative visual acuity improvement was more frequently demonstrated in patients with small hole sizes <350 μ m.^{14–19} Classifying patients with IMH by the size of the hole into <400 µm group or >400 µm might be clinically significant for counselling the patients about possible improvement after surgery.²⁰ This was evident in present study as the closure rate was 97.2%, 78%, 70% in IMH subgroups 1, 2, and 3, respectively. In addition, we found that the closure rate of small IMH <250µ was statistically significant than the other 2 groups (P= 0.02). Liu et al^{20} found that the rate of Type 1 closure for patients with smaller preoperative MHs ($\leq 250\mu$) was 100%; whereas rate of closure of holes with size 250 μ and \leq 400 μ , was 97%). These figures were significantly better than the closure rate of IMHs \geq 400µ (89%). In DMH group, there was statistically significant improvement of the mean logMAR postoperative BCVA compared to baseline (from 1.21 ± 0.39 to 1.09 ± 0.34) (P= 0.02). The closure rate was 44% of eyes. Ghoraba³ reported 4 cases with full thickness macular holes without retinal detachment in diabetic patients with vitreous hemorrhage that was discovered intraoperatively. The author did not peel the ILM in any case. Visual acuity improved, though did not exceed 20/200. Macular hole closure rate was 50% and he suggested that diabetic ischemic changes at the macula, prolonged macular detachment, toxicity of sub-macular blood and severe cystoid macular edema might explain the poor visual prognosis in DMH patients. In the TMH, there was highly significant improvement in the mean visual acuity (p <0.001) compared to the diabetic and myopic types (p=0.02and 0.08 respectively). The closure rate in TMH was 61% versus 44% and 45.4% for diabetic and myopic types, respectively. This relatively good recovery of TMH cases might be due to early diagnosis compared to the other 2 types and to better healing in younger patients.^{21–23} The postoperative BCVA was significantly better in cases with closed macular holes in idiopathic and traumatic groups (p 0.008 and <0.001 respectively); however, in PDR and myopic macular holes there was no significant difference in the BCVA between closed or opened postoperative holes. (p=0.142 and 0.286 respectively).

Previous studies reported significant improvement in visual acuity, and in closure rates that ranged from 42% to 82%.^{4,24,25} In our series, the visual outcome of MMH patients was poorer than that reported by other studies. Moreover, our closure rate was 45.46%, which was clearly less than that reported by other authors and that exceeded 60%.^{26,27} Several reasons may explain these results. Firstly, these studies relied only on clinical assessment to determine hole closure and not on OCT finding. Secondly, all myopic eyes included in our study had extensive retinal pigment epithelial atrophy, which limited visual recovery. Thirdly, all myopic eyes were associated with posterior staphyloma, which rendered ILM peeling difficult and required several attempts to perform it and that eventually led to damage of the inner retinal tissues. Finally, all studies that reported the outcome of MMH were limited by small number of cases.

Conclusion

PPV, ILM peel and C2F6 technique yielded variable anatomic and functional outcomes in different types of macular holes. Anatomic results were most favorable in IMH and least favorable in MMH. The smaller the diameter of the hole the better the results. The underlying pathogenetic mechanisms that lead to different types of macular holes are pivotal in determining the final outcome.

Summary

Macular hole is a full-thickness defect that involves all layers of the neurosensory retina. Macular hole leads to central visual loss. The aim of our study is to evaluate the surgical outcome of different types of macular hole; Idiopathic macular hole, traumatic macular hole, myopic macular hole and diabetic macular hole. We did vitrectomy, internal limiting membrane peeling and gas injection to allow the closure of the hole and restore the central vision. We detected different closure rates in the different types of macular holes. The difference in the results was related to the pathology of each type. We found that Idiopathic macular holes had the best anatomic results; whereas myopic macular holes had the least favorable anatomic outcome. The postoperative visual acuity was significantly better in cases with closed macular holes in idiopathic and traumatic groups but not in diabetic and myopic macular holes. The smaller the hole diameter the better the results.

Data Sharing Statement

The data of patients used to support the findings of this study are restricted by the (Magrabi research ethics board). Data are available for researchers who meet the criteria for access to confidential data upon request from Professor Hammouda Ghoraba, Head of vVtreoretinal Department, Magrabi Eye Centers. Email: hghoraba@email.com.

Disclosure

This work was self-funded by the authors. The authors report no conflicts of interest for this work.

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