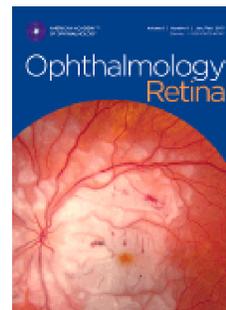


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Epiretinal Membrane Surgery After Retinal Detachment Repair: Visual Acuity Outcomes and Optical Coherence Tomography Analysis

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1 **Epiretinal Membrane Surgery After Retinal Detachment Repair: Visual Acuity Outcomes**
2 **and Optical Coherence Tomography Analysis**

3
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31

32 Acronyms:

33 CFT- central foveal thickness
34 EIFL- ectopic inner foveal layer
35 ERM- epiretinal membrane
36 ILM- inner limiting membrane
37 LogMAR- logarithm of the minimal angle resolution
38 MP- membrane peeling
39 OCT- optical coherence tomography
40 ONL -outer nuclear layer
41 PFCL- perfluorocarbon
42 PPV- pars plana vitrectomy
43 RD- retinal detachment
44 RPE-retinal pigment epithelium
45 SB- scleral buckle
46 VA- visual acuity

47 **Abstract**

48 **Purpose:** To assess visual acuity (VA) outcomes of epiretinal membrane (ERM) surgery
49 following primary rhegmatogenous retinal detachment (RD) repair.

50 **Design:** Retrospective, consecutive case series.

51 **Subjects:** Eyes undergoing pars plana vitrectomy (PPV) with membrane peel (MP) surgery for
52 ERM following primary RD repair (PPV with or without scleral buckle (SB) and gas
53 tamponade).

54 **Methods:** Retrospective chart review from 2015 to 2018. A previously described ERM grading
55 scale was utilized for OCT structural analysis.

56 **Main outcome measures:** Visual acuity (VA) and change in VA at 6 months and final follow-
57 up. Secondary outcomes included assessment of structural OCT features predictive of VA
58 outcomes.

59 **Results:** 53 eyes of 53 patients were included. VA improved significantly from logMAR $1.00 \pm$
60 0.51 (Snellen 20/200) pre-MP to 0.45 ± 0.41 (20/56) at 6 months and 0.42 ± 0.41 (20/53) at final
61 follow-up, a significant improvement ($p < 0.001$) at each timepoint. Eyes with macula on RD had
62 better 6 month [0.29 ± 0.18 (20/39) vs. 0.51 ± 0.46 (20/65), $p = 0.02$] and final VA [0.29 ± 0.14
63 (20/39) vs. 0.46 ± 0.47 (20/58), $p = 0.04$] after MP surgery, but VA improved significantly from
64 pre-MP in both macula on and macula off eyes ($p < 0.0001$, respectively). Three (5.7%) eyes were
65 graded as Stage 1, 8 (15.1%) as Stage 2, 8 (15.1%) as Stage 3, and 34 (64.2%) as Stage 4, with a
66 trend toward higher ERM stages having worse pre-MP VA ($p = 0.06$). Both MP occurring ≥ 180
67 days from RD repair and ellipsoid zone loss were associated with worse pre-MP VA [1.13 ± 0.09
68 ($20/270$) vs. 0.82 ± 0.07 (20/132), $p = 0.01$ and 1.21 ± 0.07 (20/324) vs. 0.74 ± 0.09 (20/110)
69 $p = 0.0003$, respectively]. Ellipsoid zone loss [adjusted means 0.54 ± 0.07 (20/69) vs. 0.25 ± 0.08

70 (20/36) at final visit, $p=0.006$] and RD repair with PPV/SB [0.53 ± 0.08 (20/68) vs. 0.31 ± 0.07
71 (20/41) at final visit, $p=0.03$] were significantly associated with worse VA at both 6 months and
72 final follow-up.

73 **Conclusions:** Eyes undergoing MP after RD repair have significant VA gains independent of
74 macula-status at time of RD repair. Pre-operative ellipsoid zone disruption was the OCT feature
75 best predictive of VA.

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93 **Introduction:**

94 Epiretinal membrane (ERM) formation commonly occurs after primary rhegmatogenous
95 retinal detachment (RD) repair, with a variable incidence of 6.1% to 12.8%.^{1,2} The
96 pathophysiology is thought to be due to the release of retinal pigment epithelium (RPE) cells
97 from a retinal break with subsequent proliferation on the macular surface. While some patients
98 may be asymptomatic, others may have significant reduction in visual acuity (VA) or
99 development of metamorphopsia that prompts surgical intervention.^{3,4} Small gauge pars plana
100 vitrectomy (PPV) with membrane peeling (MP) with or without inner limiting membrane (ILM)
101 peeling is the standard of care for visually significant ERM.

102 As visual improvement after ERM surgery may be variable, identifying pre-operative,
103 imaging-based predictors of visual improvement is helpful to guide expectations.^{5,6} This may be
104 more difficult in eyes with history of RD repair, as prior macular involvement may limit visual
105 potential. Previous predictors of visual acuity (VA) have focused on outer retinal changes, such
106 as ellipsoid zone loss.⁷ For example, Theodossiadis et al revealed that final VA was significantly
107 better in both macula-on and macula-off eyes with intact ellipsoid zone and external limiting
108 membrane layers, as compared to macula-off eyes with disrupted outer layers.⁸ More recently,
109 Govetto et al. described inner retinal features on optical coherence tomography (OCT), including
110 microcystic changes and ectopic inner foveal layers (EIFL), that may be prognostic of VA
111 improvement in idiopathic ERM.^{9,10} In their study, an OCT staging scheme based on presence
112 and morphology of an EIFL (**Figure 1**) was well-correlated with best-corrected VA both pre-
113 and post-MP in eyes with idiopathic ERM.

114 To date, limited study has been dedicated to imaging-based predictors of VA outcomes in
115 eyes with ERM after primary RD repair, particularly when examining both outer retinal features

116 and recently described inner retinal parameters. The purpose of this study was to determine if
117 pre-operative inner-retinal features, such as microcystic changes and EIFL, and outer retinal
118 changes, such as ellipsoid zone loss, may be predictive of visual outcomes in eyes undergoing
119 MP after previous primary RD repair.

120

121 Methods

122

123 The study was approved by the Institutional Review Board (IRB) at Wills Eye Hospital
124 (Philadelphia, PA). A waiver of informed consent was obtained by the IRB for this retrospective
125 study. The research adhered to the Declaration of Helsinki and was designed in compliance with
126 the Health Insurance Portability and Accountability Act regulations.

127 Subjects

128 A retrospective, consecutive review of clinical records was performed to identify subjects
129 who underwent PPV with membrane peel (MP) for ERM after PPV with or without scleral
130 buckle (SB) for primary RD in the same eye. Eyes were identified using Current Procedural
131 Terminology (CPT) codes 67041, 67042, and 67108 performed between January 1, 2015 and
132 January 1, 2018. Eyes were included if there was a minimum of 6 months follow-up after MP
133 surgery.

134 Exclusion criteria included more than one RD repair surgeries prior to ERM surgery, use
135 of silicone oil tamponade, previous pneumatic retinopexy, or eyes treated with SB only. Eyes
136 with RD after MP were excluded. Patients with uveitis or with concomitant macular pathology
137 including lamellar or full-thickness macular holes, central serous chorioretinopathy, branch or
138 central retinal vein occlusions, cystoid macular edema, diabetic macular edema, exudative age-

139 related macular degeneration, and intermediate or advanced dry age-related macular
140 degeneration were also excluded.

141 RD characteristics and details of RD repair surgery were recorded. Conventional, 3-port,
142 small gauge (23 or 25-gauge) PPV using the Alcon Constellation Vitrectomy system (Alcon,
143 Geneva, Switzerland) or the Bausch and Lomb Stellaris PC Vitrectomy system (Bausch and
144 Lomb, Bridgewater, New Jersey, United States) with or without SB was performed in all RD
145 repair cases. In regards to MP surgery, all eyes underwent small gauge (23, 25, or 27-gauge)
146 PPV. Dilute indocyanine green (ICG) assisted MP of both the ERM and ILM using 23-gauge or
147 25-gauge Greishaber ILM forceps (Alcon, Fort Worth, Texas, United States) was performed in
148 all MP surgeries. All post-operative complications noted at any point during the follow up period
149 were recorded. Data including intraocular pressure, slit-lamp biomicroscopy, and indirect
150 ophthalmoscopy at specified timepoints were recorded.

151 Visual acuity and OCT-based Imaging Parameters

152 Best-available Snellen VA with habitual correction or pinhole was collected at each of
153 the following specified timepoints: immediate pre-operative visit before RD repair, three months
154 after RD repair, immediate pre-operative visit before MP surgery, three months post-operative to
155 MP surgery, six months post-operative to MP surgery, and final visit.

156 Spectral domain optical coherence tomography (SD-OCT, Heidelberg Engineering,
157 Heidelberg, Germany) was performed at each timepoint and used for both quantitative
158 measurements and qualitative evaluation. Measurements of EIFL and central foveal thickness
159 (CFT) were performed using Heidelberg Eye Explorer (Version 1.9.13). The Heidelberg caliper
160 tool was used to measure the retinal layers in accordance with the International Nomenclature for
161 Optical Coherence Tomography panel definitions.¹¹ The EIFL was identified in accordance with

162 the ERM staging system proposed by Govetto et al (**Figure 1**).^{9,10} Microcystoid changes were
163 defined as small hyporeflective cystoid spaces in the inner nuclear layer not confluent with cystic
164 spaces in other layers and without a cyst wall.¹² Ellipsoid zone disruption was defined as a non-
165 continuous hyperreflective inner segment/outer segment band, of any length, not induced by
166 shadowing. For qualitative variables, two masked graders (RS and RM) independently evaluated
167 all the OCT scans of the included ERMs. All disagreements were adjudicated by a third grader
168 (MAK).

169 The primary outcome measure was visual acuity (VA) and change in VA from prior to
170 MP to 6 month post-MP and at final follow-up. Secondary outcomes included OCT features
171 predictive of VA.

172 Statistical Analysis

173 All data were analyzed with SAS v9.4 (SAS Institute Inc., Cary, NC). Snellen VA was
174 converted to logarithm of the minimal angle of resolution (logMAR) values for statistical
175 analysis. Descriptive statistics were performed using mean and standard deviation for continuous
176 measures and proportions for categorical variables. Continuous variables were analyzed with an
177 independent two sample t-test. One-way analysis of variance (ANOVA) was used to compare the
178 differences in continuous variables between two or more groups. A Fisher exact test was used to
179 compare proportions. Univariate and multivariate generalized linear models were utilized to
180 determine the predictors of: (1) VA at pre-MP, (2) VA at 6 months after MP, (3) VA at the final
181 visit, (4) change of VA from pre-MP to 6 months post-MP and (5) change of VA from pre-MP to
182 final visit. For multivariate analyses, all predictors with $p < 0.10$ in the univariate analyses were
183 initially included in the multivariate models, and the multivariate models went through the

184 backward variable selection by only keeping the statistically significant predictors ($p < 0.05$) in
185 the final multivariate model.

186 Interobserver agreement for qualitative OCT variables was determined with Cohen kappa
187 coefficient calculation.

188

189 **Results**

190 A total of 53 eyes from 53 patients met the inclusion and exclusion criteria and were
191 included in the analysis. No patient had both eyes qualify for inclusion in the study. All eyes
192 underwent successful, single surgery RD repair and subsequent ICG-assisted removal of ERM
193 and ILM, as confirmed on post-operative SD-OCT.

194 Interobserver agreement for qualitative OCT variables, including ERM staging, presence
195 of microcystic changes, presence of EIFL, and ellipsoid zone disruption were found to be
196 excellent. The interobserver agreement and Cohen kappa coefficients are as follows for each
197 qualitative variable: ERM staging (interobserver agreement 89%, Cohen kappa coefficient of
198 > 0.81), presence of microcystic changes (interobserver agreement 90%, Cohen kappa coefficient
199 of > 0.81), ellipsoid zone disruption (interobserver agreement 84%; Cohen kappa coefficient of
200 > 0.81), and presence of EIFL (interobserver agreement 100%, Cohen kappa coefficient of
201 > 0.81).

202 Baseline characteristics, features of RD repair, and features at the time of MP are
203 described in **Table 1**. In regards to macular status at time of RD repair, 14 (26.4%) eyes had a
204 macula-on RD and 39 (73.6%) eyes had a macula-off RD. 28 (52.8%) eyes underwent PPV alone
205 and 25 (47.2%) underwent combined PPV/SB for RD repair. Prior to MP, 3 (5.7%) eyes were
206 graded as Stage 1, 8 (15.1%) eyes as Stage 2, 8 (15.1%) eyes as Stage 3, and 34 (64.2%) eyes as

207 Stage 4. An EIFL layer was present in 42 (79.3%) eyes. Sixteen (30.2%) eyes were phakic prior
208 to MP. Of the 16 phakic eyes, 1 (6.3%) was Stage 1, 3 (18.8%) were Stage 2, 2 (12.6%) were
209 Stage 3, and 10 (62.5%) were Stage 4. Eight of the 14 eyes undergoing CE/PCIOL at the time of
210 MP were Stage 4.

211 *Time-course of ERM development*

212 ERM formation was first diagnosed on OCT at a mean of 91 ± 64 days [range, 15 - 289
213 days] after RD repair. The ERM was determined to be visually significant, defined as the visit at
214 which the surgeon and patient elected to proceed with MP surgery, at a mean of 206 ± 190 days
215 (range, 45 -1151 days) after RD repair. There was an interval of mean 101 ± 115 days (range, 0
216 - 458 days) between first diagnosis of ERM on OCT and the visit at which the ERM was
217 determined to be visually significant.

218 Twenty-three (43.4%) eyes underwent MP surgery ≤ 180 days from the time of RD repair.
219 Of these eyes, 17 were macula off (73.9%), 19 were graded as Stage 4 (82.6%), 19 had
220 microcystoid changes (82.6%), and 17 had ellipsoid zone disruption on pre-operative OCT
221 (73.9%). There was no difference in macular status at time of RD repair ($p>0.99$) or presence of
222 pre-operative microcystoid changes ($p=0.13$) between eyes undergoing MP surgery ≤ 180 days
223 and those undergoing MP surgery >180 days. Eyes with Stage 4 ERM were significantly more
224 likely to have MP surgery ≤ 180 days from RD repair compared to eyes with Stage 1, 2, or 3
225 ERMs, collectively ($p=0.021$).

226 *Visual Acuity Outcomes Over Time and by Macula Status at RD Repair*

227 Table 2 summarizes VA and change in VA outcomes at each timepoint for all eyes and
228 stratified by macula status. Following ERM removal, VA significantly improved from the pre-
229 MP VA at each subsequent time point. Compared to macula off eyes, VA was significantly

230 better in macula-on eyes prior to RD repair, at 6 months post-MP, and at final follow-up. At all
231 timepoints, change in VA from pre-RD repair was greater in eyes with macula-off RD compared
232 to macula-on RD.

233 Visual acuity outcomes by ERM Stage

234 Prior to the ERM removal, mean VA tended to be worse with higher ERM stages [Stage
235 1, 0.62 ± 0.36 (20/83)]; [Stage 2, 0.72 ± 0.43 (20/105)]; [Stage 3, 0.84 ± 0.52 (20/138)]; and
236 [Stage 4, 1.13 ± 0.50 (20/270), $p=0.06$]. There was no significant association between ERM
237 stage and VA at three months, six months, or at final visit after MP. There was no significant
238 association of ERM stage with change in VA from the pre-RD or pre-MP timepoint.

239 OCT structural thickness

240 OCT-based thickness measurements were assessed following MP surgery. Mean CFT
241 improved from $565 \pm 168 \mu\text{m}$ pre-MP to $359 \pm 79 \mu\text{m}$ at three months post-MP, an improvement
242 of $206 \pm 154 \mu\text{m}$ ($p<0.0001$). Mean \pm SD outer nuclear layer (ONL) thickness improved from
243 $132 \pm 96 \mu\text{m}$ pre-MP to $101 \pm 56 \mu\text{m}$ at three months post-MP, an improvement of $33 \pm 82 \mu\text{m}$
244 ($p=0.01$). In the 42 eyes with preoperative EIFL, the mean \pm SD EIFL thickness improved from
245 $376 \pm 143 \mu\text{m}$ pre-MP to $174 \pm 86 \mu\text{m}$ at three months post-MP, an improvement of 202 ± 142
246 μm ($p<0.0001$). Among 42 eyes (79.2%) with EIFL pre-MP, 15 (35.7%) had full resolution of
247 EIFL at three months post-MP. Among 37 eyes (69.8%) with microcystic changes pre-MP, 11
248 (29.7%) had full resolution of microcystic changes at three months post-MP. Of 31 eyes (58.5%)
249 with ellipsoid zone disruption pre-MP, 22 (71.0%) did not have ellipsoid zone disruption at 3
250 months post-MP.

251 Predictors of VA

252 Factors associated with VA at the pre-MP timepoint are summarized in **Table 3**. In
253 multivariate analysis, MP ≥ 180 days from RD repair and ellipsoid zone disruption retained
254 significance for pre-MP VA. In univariate analysis, presence of EIFL and CFT $>550 \mu\text{m}$ at the
255 pre-MP visit timepoint were both associated with a significantly worse VA; however, these
256 variables became non-significant in multivariate analysis.

257 Multivariate analysis of factors associated with VA and change in VA from pre-MP at 6
258 months and final follow-up is summarized in **Table 4**. Ellipsoid zone disruption and RD repair
259 with PPV/SB were significantly associated with worse VA at both 6 months and final follow-up.
260 In multivariate analysis of factors associated with VA change from pre-MP, undergoing MP
261 ≥ 180 days from RD repair was significantly associated with greater VA improvement at both 6
262 months and final follow-up.

263 A complete summary of univariate and multivariate analysis associated with VA and
264 change in VA from pre-MP at 6 months is available in Table 5 (available at
265 <https://www.opthalmologyretina.org/>). A complete summary of univariate and multivariate
266 analysis of factors associated with VA and change in VA from pre-MP at final follow-up is
267 available in Table 6 (available at <https://www.opthalmologyretina.org/>).

268

269 **Discussion**

270 Our retrospective, consecutive case series of eyes undergoing ERM surgery after retinal
271 detachment repair explored visual outcomes and the functional significance of OCT parameters
272 on predicting visual outcomes. In this series, we found an overall improvement in vision after
273 MP surgery at all timepoints, with similar improvement in eyes with history of macula-on and
274 macula-off RD. Furthermore, we analyzed outcomes using the ERM grading system established

275 by Govetto et al.^{9,10} ERM stage trended with pre-MP VA ($p=0.06$) and presence of an EIFL was
276 associated with worse pre-MP VA ($p=0.03$) on univariate analysis. The OCT feature that best
277 predicted 6 month and final visual outcomes in eyes with ERM post-RD repair was presence of
278 ellipsoid zone disruption.

279 In our study, mean VA improved by 4.5 lines at 6 months and the final visit. These visual
280 outcomes were comparable to the 5.6 Snellen line improvement noted by Katira et al. and the 4
281 line gain noted by Council et al, which also evaluated outcomes of MP surgery after prior RD
282 repair.^{2,13} Interestingly, no significant difference in VA existed between macula on and macula
283 off eyes at the pre-MP timepoint. Similarly, there was no significant difference in the degree of
284 VA improvement after MP between macula-on and macula-off eyes. This suggests that ERM
285 formation has a clinically meaningful effect on VA independent of macula status once ERM
286 development has occurred. This finding is similar to that of Council et al, which found no
287 significance of macula status regarding change in VA from the pre-MP timepoint.¹³

288 In previous studies, ERM occurred in 12.1% to 35% of post-RD repair eyes.^{14,15} Ishida et
289 al. found the majority of ERMs (76.9%) were diagnosed within the first 3 months postoperative
290 to RD repair¹⁴, while Katira et al found that of eyes requiring surgery for ERM after RD repair
291 the mean time to MP was 5.4 months.² Our study similarly reveals a brief period from RD repair
292 to first diagnosis of ERM and diagnosis of visually significant ERM (3 and 6 months,
293 respectively). Furthermore, our study suggests that ERM formation soon after RD repair is
294 characterized by a more severe disorganization of macular anatomy compared to idiopathic
295 ERM. In Govetto et al., only 12.6% of idiopathic ERMs were found to be Stage 4, whereas
296 64.2% of post-RD eyes in our study were found to be Stage 4.¹ Undergoing MP surgery \square 180
297 days from RD repair was significantly associated with Stage 4 ERM characteristics ($p=0.021$).

298 This may help explain the association between surgical timing, pre-MP visual acuity, and
299 observed VA improvement after MP in this study. Eyes undergoing ERM surgery ≤ 180 days
300 from RD repair had worse pre-operative VA ($p=0.01$) and greater visual acuity improvement
301 post MP ($p=0.002$) compared to eyes undergoing MP in >180 days. This greater, relative VA
302 improvement may represent a ceiling effect as eyes undergoing surgery >180 days from RD
303 repair had better pre-MP VA and, thus, less VA to be gained. However, the significant VA gain
304 in eyes undergoing ERM surgery ≤ 180 days from RD repair is encouraging, reflecting a benefit
305 to MP surgery even in eyes with early, more severe ERM characteristics.

306 Our study builds upon a recent series of publications utilizing a novel, OCT based
307 grading scale centered on the presence or absence of an EIFL to describe outcomes in eyes with
308 ERM.^{9,10} Govetto et al found that the EIFL-based staging system could predict VA in eyes with
309 idiopathic ERM, with a progressive decline in vision from Stage 1 [0.02 ± 0.6 (20/21)] to Stage 4
310 [0.61 ± 0.26 (20/81)] ($p < 0.001$). The presence of EIFL was significantly associated with lower
311 BCVA, ($p=0.001$) suggesting not only that the inner foveal microanatomy is particularly
312 susceptible to disruption in ERM formation but also that such disruption may profoundly affect
313 function.⁹ In a subsequent study, the group evaluated the use of the staging system to predict pre-
314 and post-operative VA in eyes with idiopathic ERM undergoing MP surgery. ERM stage
315 ($p < 0.001$), presence of an EIFL layer ($p < 0.001$), and EIFL thickness ($r=0.58$, $p < 0.001$) were
316 negatively associated with pre-operative VA.¹⁰ Moreover, lower ERM stage was significantly
317 and positively associated with VA at all post-operative timepoints through 12 months [Stage 2
318 0.06 ± 0.08 (20/23) vs. Stage 4 0.31 ± 0.26 (20/41), $p < 0.001$].¹⁰

319 In our study, EIFL presence was associated with pre-MP VA in univariate analysis
320 ($p=0.03$), but EIFL presence and thickness were not associated with pre-operative VA in

321 multivariate analysis. In addition, pre-operative EIFL presence and thickness were not significant
322 predictors of mean VA or change in VA at any timepoint on multivariate analysis. Unlike
323 Govetto et al., where Stage 3 and 4 eyes had the greatest change in VA after surgery ($p < 0.001$),
324 our study found similar changes in VA amongst all stages and ERM stage was not associated
325 with post-operative VA at any postoperative timepoint. The lack of association between EIFL
326 presence or thickness with VA in our study may be a result of smaller sample size especially in
327 the Stage 1-3 groups, or may indicate that eyes with prior RD may have additional ultrastructural
328 changes, in addition to EIFL thickness, which are of more predictive value compared to eyes
329 with idiopathic ERM.

330 Our work emphasizes the importance of outer retinal layer disruption as a key
331 prognosticator of function. Ellipsoid zone disruption pre-operatively was the single OCT feature
332 in our study significantly associated with pre-MP and post-MP mean VA. The significance of the
333 ellipsoid zone in determining visual acuity in post-RRD repair eyes has been previously noted by
334 prior authors.^{8,16} Wakabayashi et al. evaluated OCT microstructural changes in eyes undergoing
335 primary RD repair and found that only macula-off eyes had disruption of the ellipsoid zone
336 ($p < 0.001$) and that post-operative ellipsoid zone changes were associated with post-operative VA
337 ($r = 0.805$, $p < 0.001$).¹⁶ Theodossiadis et al. found that ellipsoid zone loss pre-operatively was a
338 significant predictor of final VA at 6 months after MP.⁸ From these studies and from our own,
339 the disruption of the ellipsoid zone may indicate structural and functional damage to
340 photoreceptors that contribute to visual prognosis post-RD and post-MP repair.

341 Limitations of this study are inherent in its retrospective nature. Use of Snellen visual
342 acuity with habitual correction or with pinhole and without refraction may have underestimated
343 VA outcomes. Our sample is weighted to a higher proportion of patients with Stage 4 ERMs,

344 which may lead to an underestimation of visual potential in post-RD, post-MP eyes. Moreover,
345 eyes in this sample were chosen for MP surgery based on surgeon discretion, and selection bias
346 against eyes with poorer visual potential post RD may be present. While lens status was not
347 significantly associated with VA or VA change at final follow-up, the fact that the majority of
348 phakic eyes had Stage 4 ERMs could have diminished the significance of VA differences by
349 ERM stage. Possible factors contributing to the severity of ERM or visual impairment, including
350 the intensity or area encompassed by laser or cryotherapy treatment during RD repair, were not
351 quantified in the present study. The strengths of our study include characterization of OCT
352 markers using SD-OCT in all cases, standardization of surgical technique across surgeons, and a
353 relatively high case number of eyes undergoing ERM peel post RD repair compared to prior
354 studies of a similar population.

355 Eyes undergoing MP after RRD repair did exhibit substantial visual gains post MP,
356 regardless of ERM stage and macular status at time of RD repair. While presence of an EIFL has
357 been previously associated with pre-operative and post-operative visual acuity in idiopathic
358 ERMs,^{10,12} ellipsoid zone disruption was the OCT biomarker most associated with pre-MP, 6
359 month, and final VA in eyes with prior history of RD repair. Larger, prospective studies are
360 needed to further evaluate the utility of inner and outer retinal OCT alterations on predicting
361 function in eyes with secondary ERM after RD repair.

Figure 1: ERM staging system as based on optical coherence tomography ectopic inner foveal layer (EIFL) presence and disruption of inner foveal layers.^{9,10}

A- Stage 1 is defined as the presence of ERM with minimal disruption in inner foveal contour.

B-Stage 2 is defined as the presence of ERM with loss of inner foveal contour but no EIFL. C-

Stage 3 is defined as the presence of an EIFL but clear distinction between all retinal layers. D-

Stage 4 is defined as the presence of an EIFL but loss of distinction between retinal layers.

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Table 1: Baseline characteristics of study subjects and study eyes undergoing membrane peel after primary rhegmatogenous retinal detachment repair.

Age (years)

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CE/IOL=cataract extraction/intraocular lens insertion. ERM=epiretinal membrane. EIFL=ectopic inner foveal layer. MP= membrane peel. PFC=perfluorocarbon liquid. PCIOL=posterior chamber intraocular lens. PVR=proliferative vitreoretinopathy. RD=retinal detachment. SB=scleral buckle. SD= standard deviation.

Table 2: Visual acuity and change in visual acuity at each time point as stratified by macula status.

Timepoint	All eyes (N=53)	Macula status Off (N=39)
	Mean \pm SD in logMAR (Snellen)	Mean \pm SD in logMAR (Snellen)

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Table 3: Analysis of variables associated with mean visual acuity at the pre-membrane peel surgery visit.

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* From the generalized linear model with a specific predictor in the model.

** From the generalized linear model with all statistically significant predictors in the final model.

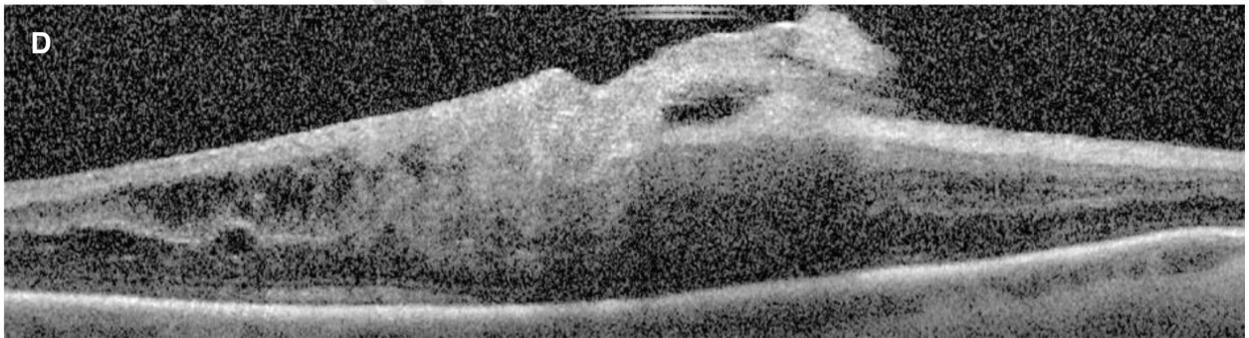
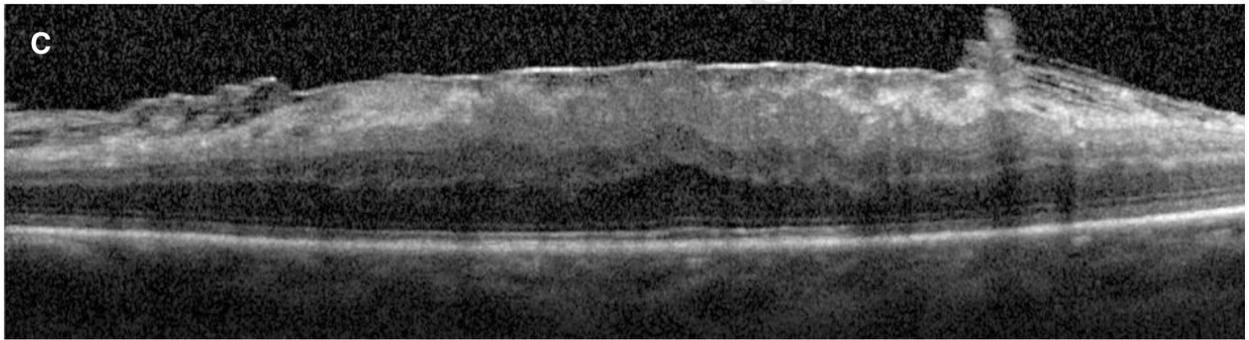
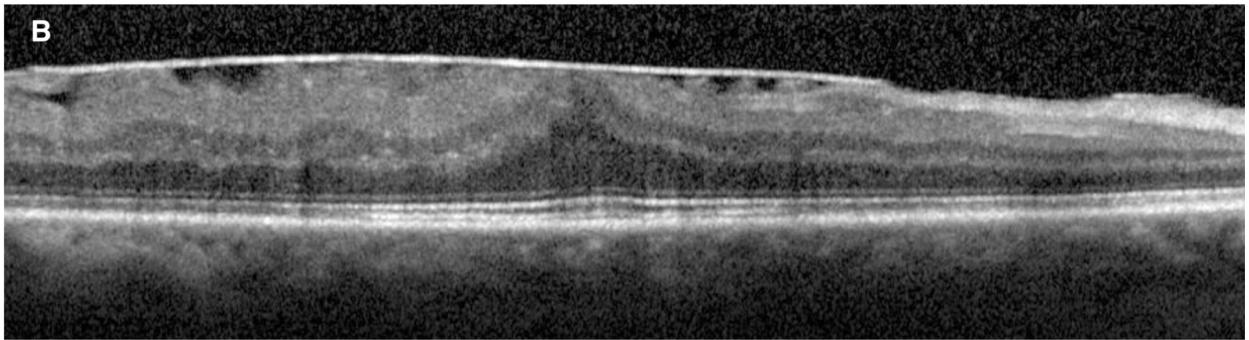
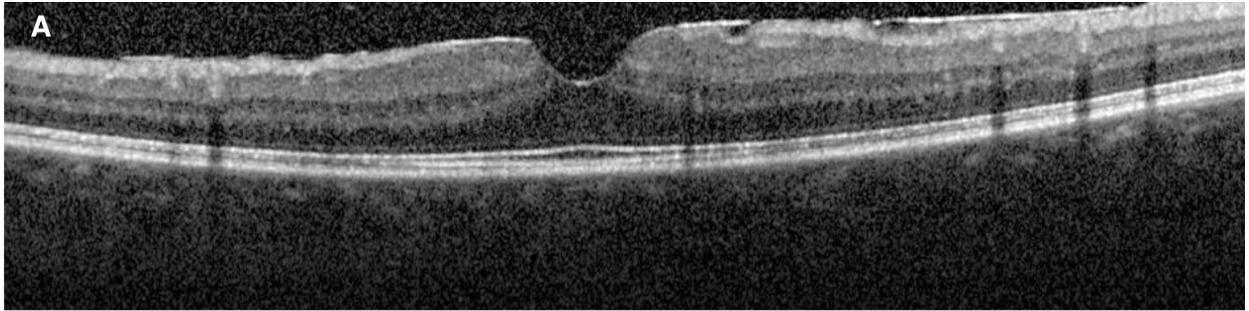
CFT=central foveal thickness. EIFL=ectopic inner foveal layer. MP= membrane peel. Pre-op= pre-operative. PPV=pars plana vitrectomy. RD= retinal detachment. SB=scleral buckle. VA=visual acuity.

Table 4: Factors associated with visual acuity and change in visual acuity from pre-MP at 6 months and final follow-up in multivariate analysis.

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* From a generalized linear model with all statistically significant predictors in the final model.

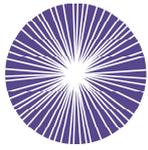
CFT=central foveal thickness. EIFL=ectopic inner foveal layer. MP= membrane peel. Pre-op= pre-operative. PPV=pars plana vitrectomy. RD= retinal detachment. SB=scleral buckle. VA=visual acuity.



Precis:

Eyes undergoing epiretinal membrane surgery after prior retinal detachment (RD) repair have significant visual acuity gains independent of macula-status at time of RD repair. Pre-operative ellipsoid layer disruption was the OCT feature best predictive of final visual acuity.

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