

Reading Speed as an Objective Measure of Improvement Following Vitrectomy for Symptomatic Vitreous Opacities

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1	Reading speed as a reliable outcome measure to assess visual
2	improvement following vitrectomy for symptomatic vitreous
3	opacities in patients with clear lenses
4	
5	Short title: Reading speed as an objective measure of improvement following vitrectomy for
6	symptomatic vitreous opacities
7	
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2	5

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33

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37

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41 MNREAD iPad app through a licensing agreement between the University of Minnesota and

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48 Abstract

49	Background and Objective: There is currently no objective measure of the visual deficits
50	experienced by patients with symptomatic vitreous opacities (SVO) that would also correlate
51	with the functional improvement they report following vitrectomy. This study aims to determine
52	whether reading speed can be used as a reliable outcome measure to assess objectively the
53	impact of both SVO and vitrectomy on patients' visual performance.
54	Study Design/Materials and Methods: 20 adult patients seeking surgery for SVO were
55	included. Measures of visual function were obtained before and after vitrectomy using the
56	ETDRS acuity chart, the NEI-VFQ and the MNREAD.
57	Results: In patients with non-opacified lenses (N=10), maximum reading speed increased
58	significantly from 138 to 159 words per minute after complete removal of SVO by vitrectomy
59	(95%CI = [14, 29], p < 0.001).
60	Conclusion: Reading speed is impaired with SVO, and improves following vitrectomy in phakic
61	and pseudophakic eyes with clear lenses. Reading speed is a valid objective measure to assess
62	the positive effect of vitrectomy for SVO on near-distance daily life activities.
63	

64 Keywords

65

66 Vitrectomy, symptomatic vitreous opacities, lens opacity, reading speed, daily-life function, functional
67 improvement, objective measurement

69 Background and Objective

Patients with symptomatic vitreous opacities (SVO) experience visual impairment from multiple 70 dense particles floating in the vitreous gel, which often cast a mobile dark shadow on the retina. 71 72 However, standard objective measures of visual function, such as Snellen visual acuity, remain often excellent in the presence of SVO^{1,2}. Nonetheless, patients with SVO report significant 73 visual improvement after their removal by vitrectomy^{3,4}. For instance, previous studies have 74 shown post-operative improvement in subjective visual quality of life^{5,6}. These results were 75 obtained with the National Eye Institute Visual Function Questionnaire (NEI-VFQ), which 76 assesses the level of difficulty experienced by individuals with chronic eye diseases during daily-77 living activities, such as driving or reading⁷. 78

79

To complement such subjective evaluations, there is a need for establishing a quantifiable 80 measure to assess objectively (1) the impairment in visual function caused by SVO and (2) the 81 improvement in visual function following vitrectomy³. First, such a clinical measure would help 82 detect patients with functional impairment from vitreous floaters. Second, it would bring 83 valuable insight to help resolve the existing controversy over vitrectomy's clinical relevance. So 84 far, intraocular straylight⁸ and contrast sensitivity⁶ have been proposed as independent objective 85 measures of visual perception with symptomatic floaters. Despite their impact on vision-related 86 quality of life, these measures do not evaluate daily life function directly. 87

88

A frequent complaint from patients with prominent opacities is interference with ease of reading.
Even if unilateral, these patients often complain of interference with binocular visual function^{9,10}.

Patients usually report moderate or extreme difficulty in reading small print⁵. In the low-vision
literature reading speed is already considered a strong objective predictor of visual ability and
vision-related quality of life for patients with ocular disorders, such as macular
degeneration^{11,12,13,14}. Here we conducted a prospective study to test whether reading
performance can also be used as reliable outcome measure to investigate the impact of SVO and
therapeutic vitrectomy on patients' visual performance.

97

Our main objective was to investigate whether reading performance, evaluated with the 98 standardized MNREAD test, could provide an objective measure of functional improvement in 99 patients with SVO treated with pars plana vitrectomy. To this aim, we compared pre and post-100 operative measures of (1) vision-related quality of life (subjectively obtained with the NEI VFQ) 101 and (2) reading performance (objectively obtained with the MNREAD test). Given that reading 102 performance is rapidly degraded with reduced contrast from cloudy ocular media¹⁵, these 103 104 comparisons were performed while controlling for patients' lens status (clear vs. mildly opacified). Additionally, we investigated whether a potential improvement in these subjective 105 and objective measures following vitrectomy would be correlated with pre-operative opacity 106 severity. 107

108

109

111 Patients/Materials and Methods

112 **Patients**

Patients over 21 years old were included in the present work if they (1) elected to undergo 113 vitrectomy, (2) presented symptoms consistent with examination findings of dense opacities for 114 at least 6 months, (3) had visual acuity of 20/80 (0.6 logMAR) or better in both eyes before 115 surgery and (4) did not experience a significant drop in acuity in the non-operated eye between 116 the pre- and post-surgery measurements. Phakic and pseudophakic patients were included, as 117 well as patients with or without a vitreous detachment. History of scleral buckle for retinal 118 119 detachment (RD) was acceptable if the macula was not involved. If an epiretinal membrane was noted on OCT but not clinically visible or deemed significant, patients were included in the 120 study. Patients were excluded if they had history of cognitive impairment, macula-off RD, severe 121 122 glaucoma, macular degeneration, diabetic macular edema, or other confounding ocular disorders. A total of 20 patients were recruited, tested and treated at two different sites: 11 at a private 123 retina practice in Minnesota and 9 at an academic retina practice in California. Figure 1 124 illustrates the protocol sequence. Institutional Review Board (IRB)/Ethics Committee approval 125 was obtained and written informed consent was obtained before the study from each patient 126 according to IRB guidelines. The study also complied with tenets of the Declaration of Helsinki 127 and HIPAA. 128



130 Figure 1: Protocol schematic showing the different test procedures along with the resulting outcome measures.

131 Subjective measures are represented in blue; objective measures are represented in pink.

132

133 Surgery

20 eyes of 20 patients underwent outpatient three-port 25-gauge pars plana vitrectomy using the 134 Alcon Constellation system. Inspection of the peripheral retina with indirect ophthalmoscopy and 135 136 scleral depression was performed at surgery conclusion. Leaking sclerotomies were sutured. Postoperative examinations were at 1 day, 1-2 weeks, and 4-6 weeks. Presence or absence of a 137 138 posterior vitreous detachment (PVD) was confirmed intraoperatively. Visual acuity, intraocular pressure, dilated funduscopic exam and any postoperative complications including high or low 139 140 intraocular pressure, retinal tear or detachment, and/or endophthalmitis were recorded. For each 141 patient, the non-operated eye served as control.

142

144 Subjective grading of opacity density

145 Before and after surgery, patients went through video recording of the vitreous using the infrared confocal scanning laser ophthalmoscope (SLO) combined with optical coherence tomography 146 (OCT)^{16,17}. The movie created with this technique reveals motion of the shadows projected by 147 the opacities onto the retinal surface, enabling a subjective grading of the opacity density. 148 149 Recording was performed in each eye. Patients were instructed to look to one side and then re-150 fixate, which set the vitreous opacities in motion. This step was repeated to each side several times. The pre and post-surgery videos were assessed by two experienced, masked surgeons and 151 152 given a score of 0-3, with 0 corresponding to no floaters and 3 corresponding to very dense floaters (see supplementary material for a pre-op movie graded as 2 and a post-op movie graded 153 as 0). 154

155

156 Subjective measure of vision-related quality-of-life

Before and after surgery, patients were interviewed with the NEI-VFQ-25, the 25-item version of
the VFQ test⁷. Data were scored using the standard method to calculate: 1-the near activities
VFQ score (involving reading) and 2- the composite VFQ score (encompassing many visionrelated functions). Scores ranged from 0 to 100, with higher scores representing better function.

161

162 **Objective measure of reading speed**

163 Before and after surgery, patients' reading performance was measured with the MNREAD acuity

164 chart, a standardized test designed to measure binocular and monocular reading performance¹⁸.

Test measures were obtained with the MNREAD app running on an iPad¹⁹. Viewing distance 165 was 60 cm and screen luminance was set to 200cd/m2. Patients went through six iterations of the 166 test (operated eve, non-operated eve and binocular, each condition being repeated twice), all in 167 black print on white background. MNREAD testing was performed pre-operatively and again 4-168 8 weeks after surgery. For each test performed, the four MNREAD measures were estimated 169 internally by the app¹⁸: (1) Maximum Reading Speed (MRS), (2) Critical Print Size (CPS), (3) 170 Reading Acuity (RA) and (4) reading ACCessibility index (ACC - a single-valued measure that 171 represents one's visual access to commonly encountered printed material, ranging from 0 (i.e. no 172 access to print) to 1 (i.e. average normal access) or above)²⁰. 173

174

175 Statistical analysis

Pre and post-operative scores of NEI-VFQ were compared with a Wilcoxon signed-rank test. For 176 177 each of the four MNREAD parameters, a different linear mixed-effects model including data 178 from all 20 patients was designed to compare values before and after vitrectomy for the operated eve, the non-operated eye and the binocular condition 21,22 . To control for covariate factors, the 179 following variables were also included in the models: binocular lens opacity ('clear' vs. 'mild 180 opacity'), presence of epiretinal membrane (ERM) in the operated eye ('yes' vs. 'no'), presence 181 of SVO in the non-operated eye ('yes' vs. 'no'), presence of posterior vitreous detachment 182 (PVD) in the non-operated eye ('yes' vs. 'no') and testing location ('Minnesota' vs. 183 'California'). The same random structure was chosen for all four models and included a random 184 intercept for "eyes nested within patients", assuming a different baseline performance level for 185 each patient and each eve. Post-hoc pairwise comparisons were performed using Tukey's 186

187 correction. In the Results section, mean values estimated by the models and post-hoc pairwise

188 comparisons are reported with their 95% confidence intervals (95%CI) and p-values.

189

190

191 **Results**

192 *Patients*

193 Preoperative clinical examination revealed the presence of SVO and clinical evidence of PVD in

all patients. Thirteen patients had bilateral but asymmetric opacities noted on clinical

examination, and were asymptomatic in the fellow eye. Six patients had concurrent minimally

significant epiretinal membrane. One patient had a history of scleral buckling for a macula-

sparing retinal detachment. Vitreous opacities symptoms had been present for 6 to 24 months.

198 Table 1 presents the patients' preoperative individuals characteristics.

Patient				Lens opacity	Operated eye				Non-operated eye		
ID	Location	Gender	Age	in both eyes	Pathology	SVO	Acuity	OCT-SLO Opacity grading	Pathology	SVO	Acuit y
P1	Minnesota	М	58	Clear	PVD	Yes	20/25	2	ERM	No	20/25
P2	Minnesota	М	59	Clear	PVD	Yes	20/20	1.5		Yes	20/25
P3	California	М	61	Clear	PVD	Yes	20/20	3	ERM	No	20/40
P4	Minnesota	М	62	Clear	PVD+ ERM	Yes	20/20	1.5	Scleral bucklin g + ERM	No	20/20
Р5	Minnesota	М	64	Clear	PVD+ ERM	Yes	20/15	2	PVD	Yes	20/25
P6	Minnesota	F	64	Clear	PVD	Yes	20/25	2.5		Yes	20/20
P7	Minnesota	F	64	Clear	PVD	Yes	20/20	2.5	PVD	Yes	20/25
P8	Minnesota	F	68	Clear	PVD	Yes	20/30	1	PVD	Yes	20/15
Р9	California	М	69	Clear	PVD	Yes	20/20	2.5	PVD	Yes	20/20
P10	Minnesota	F	72	Clear	PVD	Yes	20/25	2	PVD	Yes	20/25
P11	California	F	32	Mild opacity	PVD	Yes	20/25	1	PVD	Yes	20/80

P12	California	М	52	Mild opacity	PVD+ ERM	Yes	20/25	2.5	Vitreous Syneresis	No	20/20
P13	California	М	54	Mild opacity	PVD+ ERM	Yes	20/40	3	NPDR	No	20/20
P14	California	F	54	Mild opacity	PVD	Yes	20/25	2.5	PVD	Yes	20/80
P15	Minnesota	М	63	Mild opacity	PVD+ ERM	Yes	20/40	2.5	ERM	Yes	20/25
P16	California	М	63	Mild opacity	PVD+ ERM	Yes	20/80	2	ERM	No	20/25
P17	Minnesota	М	64	Mild opacity	PVD	Yes	20/20	2.5	Vitreous Syneresis	Yes	20/20
P18	California	F	65	Mild opacity	PVD	Yes	20/30	2.5	PVD	Yes	20/25
P19	California	F	67	Mild opacity	PVD	Yes	20/30	2.5	ERM	No	20/25
P20	Minnesota	М	68	Mild opacity	PVD	Yes	20/20	1.5	PVD	Yes	20/25

<u>Table 1: Patients' individual characteristics prior to surgery.</u> SVO stands for symptomatic vitreous
 opacities; ERM stands for epiretinal membrane. PVD stands for posterior vitreous detachment;

202 NPDR stands for non-proliferative diabetic retinopathy; Visual acuity is given in Snellen notation.

203

204 Surgery

205 No complications were seen. No cataract progression was observed in phakic patients during the

short period of follow-up (6 weeks). Complete removal of the central vitreous opacities was

documented by examination and video SLO in all 20 cases. Prior to surgery, OCT-SLO grading

of opacity was on average 2.2, ranging from 0 to 3 (Table 1). After vitrectomy, opacity grading

score was 0 for all 20 patients.

210

211 Visual function

In the operated eye, mean visual acuity was 0.11±0.16 logMAR before surgery and 0.09±0.16

logMAR after surgery. The difference between pre- and post-op visual acuity was not significant

(p = 0.36). Both NEI-VFQ scores improved significantly after vitrectomy, but this improvement

was dependent on the lens opacity (Figure 2). Among patients with clear lenses (N=10), the

216	average near activities sub-score went from 47.5 to 74.2. This significant increase of 26.7 points
217	(95%CI = [16.2, 37.1], p < 0.001) corresponds to an overall 56.2% improvement (Figure 2A-
218	left). For patients with opacified lenses however (N=10), vitrectomy did not improve the near
219	activities sub-score. For patients with clear lenses, the average pre-op composite score was 64.6
220	and increased by 19.8 points (95%CI = $[13.9, 25.7]$, p < 0.001) after vitrectomy, representing a
221	30.6% improvement (Figure 2B-left). The improvement was somewhat smaller for patients with
222	opacified lenses, whose score went from 71.4 to 85.8, representing a significant gain of 20.2%
223	(14.4 points, 95% CI = [-6.3, 23.4], p = 0.003). The overall improvement for both subgroups on
224	the composite score was 26.3%. There was no correlation between the opacity grading score
225	prior surgery and the amount of NEI-VFQ score improvement following surgery (Pearson's
226	correlation coefficients was -0.36 and -0.39 for the near activities sub-score and the composite
227	score respectively).



Figure 2: Pre and post-operative NEI-VFQ scores grouped by lens opacity status. Points show the mean estimates for the near activity sub-score (A) and the overall composite score (B), both before and after surgery, as estimated by the mixed effects models, for patients with clear lenses in blue

(N=10) and patients with mildly opacified lenses in orange (N=10). Error bars represent the 95%
 confidence intervals.

234

235 *Reading performance*

236 <u>Maximum Reading Speed (MRS)</u>

First, we included data from all 20 patients in the mixed-effects model without any distinction on 237 their lens opacity status. MRS before surgery was on average 137 words/minute (wpm) for the 238 operated eve (95%CI = [125, 149]). It was significantly higher by 13 wpm in the non-operated 239 eve (95%CI = [5, 22], p = 0.003) and by 15 wpm in the binocular condition (95%CI = [7, 24], p 240 < 0.001). After surgery, MRS in the operated eye increased significantly to 146 wpm (i.e. a 9 241 wpm increase; 95%CI = [3, 15], p = 0.007). Post-operatively, MRS did not change significantly 242 in the non-operated eye (1 wpm increase; 95%CI = [-12, 14]; p = 0.8) or in the binocular 243 condition (3 wpm increase; 95%CI = [-9, 17]; p = 0.23). 244 Second, we included an interaction between the "surgery" and "lens opacity" factors in the 245 model. For patients with clear lenses only (N=10), MRS prior surgery was on average 138 246 247 words/minute (wpm) for the operated eye (95%CI = [120, 156]; Figure 3-left panel). It was

significantly higher by 13 wpm in the non-operated eye (95%CI = [6, 20], p < 0.001) and by 14

wpm in the binocular condition (95%CI = [7, 22], p < 0.001). After surgery, MRS in the

operated eye increased significantly to 159 wpm (i.e. a 21 wpm increase; 95%CI = [14, 29], p <

251 0.001). In the non-operated eye, MRS did not change post-operatively, with a non-significant

increase of 3 wpm (95%CI = [-7, 39], p = 0.43). In the binocular condition, the limited increase

of 8 wpm following vitrectomy barely reached significance (95%CI = [-0.38, 45], p = 0.04).

For patients with mildly opacified lenses (N=10), there was no significant difference in MRS
before and after surgery in any of the three conditions tested (operated eye, un-operated eye and
binocular; Figure 3-right panel).

For all 20 patients there was no correlation between the opacity grading score in the operated eye prior surgery and the amount of MRS improvement following surgery (Pearson's correlation coefficients was -0.13).



260

Figure 3: Effect of pre/post-surgery condition on MRS for the operated eye (top – triangles), the

262 <u>non-operated eye (center - circles) and the binocular condition (bottom – squares) grouped by lens</u>

263 <u>opacity: clear (left – blue) vs. mildly opacified (right - orange).</u> Solid lines connect the estimates for

each sub-group as given by the mixed-effects model. Errors bars (black) represent their standard
 errors. Dashed lines connect the MRS values for each patient, numbered from P1 to P20.
 266

267 <u>Reading Accessibility Index (ACC)</u>

268	As for MRS,	we first inc	luded data	from all 20	patients in t	he mixed	l-effects model,	without any
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distinction on their lens opacity status. Before surgery, ACC was on average 0.61 for the

operated eye (95%CI = [0.55, 0.68]). It was significantly higher by 0.09 wpm in the non-

operated eye (95%CI = [0.04, 0.15], p = 0.002) and by 0.11 in the binocular condition (95%CI

[0.05, 0.17], p < 0.001). After surgery, ACC in the operated eye increased significantly to 0.67

273 (i.e. a 0.06 increase; 95%CI = [0.01, 0.10], p = 0.01). Post-operatively, ACC did not change

significantly in the non-operated eye (0.002 increase; 95%CI = [-0.09, 0.09]; p = 0.95) or in the

binocular condition (0.03 increase; 95%CI = [-0.06, 0.12]; p = 0.22).

- 276 Second, we included an interaction between the "surgery" and "lens opacity" factors in the
- 277 model. For patients with clear lenses only (N=10), ACC was 0.65 in the operated eye before
- surgery (95%CI = [0.56, 0.74], p < 0.001; Figure 4 left). It was marginally but significantly

better for the non-operated eye, with a value of 0.72 (0.07 difference; 95%CI = [0.01, 0.14], p =

0.02) and significantly better in the binocular condition, with a value of 0.75 (0.1 difference;

281 95%CI = [0.04, 0.17], p = 0.002). Following surgery, ACC was significantly increased by 0.1 in

the operated eye (95%CI = [0.05, 0.16], p < 0.001), reaching a value of 0.75. In the non-operated

- eye, ACC remained unchanged after surgery (0.01 difference; 95%CI = [-0.09, 0.28], p = 0.5).
- In the binocular condition, ACC increased by 0.05 after vitrectomy but this change did not reach
- significance (95%CI = [-0.03, 0.34], p = 0.06).

For patients with mildly opacified lenses (N=10), there was no significant difference in ACC
before and after surgery in any of the three conditions tested (operated eye, un-operated eye and
binocular; Figure 4 – right panel).
For all 20 patients, there was no correlation between the opacity grading score prior surgery in

the operated eye and the ACC increase following surgery (Pearson's correlation coefficient was -

291 0.41).

290



Figure 4: Effect of pre/post-surgery condition on ACC for the operated eye (top – triangles), the
 non-operated eye (center - circles) and the binocular condition (bottom – squares) grouped by lens
 opacity: clear (left – blue) vs. mildly opacified (right - orange). Solid lines connect the estimates for
 each sub-group as given by the mixed effects model. Errors bars (black) represent their standard
 errors. Dashed lines connect the MRS values for each patient, numbered from P1 to P20.

299 Critical Print size (CPS) and Reading Acuity (RA)

300 For both CPS and RA, we found no significant difference between the operated eye and the non-

301 operated eye or the binocular condition before surgery. None of these measures changed

302 significantly after surgery in the tested eyes.

303

304 Correlation between reading performance change and daily life visual function improvement

Lastly, we inspected the correlation between the improvement in reading performance and the

improvement in NEI-VFQ near activities sub-score in the operated eye of all 20 patients. We

307 found no correlation between the percentage of improvement in MRS and the increase in NEI-

VFQ near activities sub-score (r = 0.4, 95%CI = [-0.12, 0.75], p = 0.12). On the other hand, the

improvement in ACC was significantly correlated with the near activities sub-score (r = 0.74,

310 95%CI = [0.39, 0.90], p = 0.001; Figure 5).



Figure 5: Post-operative NEI-VFQ near activity sub-score improvement as a function of post operative reading accessibility index improvement for all 20 patients.

311

315 **Discussion**

316

The symptomatic relief experienced by patients following vitrectomy has been demonstrated 317 before by the use of the NEI-VFQ subjective test^{5,6}. Our study confirmed the literature results, 318 with a significant overall improvement of 26% on the test composite score. This value is in line 319 with previously reported improvement results, ranging from 19% to 29%, in patients treated for 320 symptomatic floaters^{6,23}. The present analysis also revealed a significant interaction between the 321 impact of surgery on the VFQ scores and the opacity status of the patient's lenses. For near 322 distance activities, vitrectomy only improved patients' score if their lenses were clear, whereas 323 the overall composite score (which includes both near- and far-distance activities) improved even 324

if the lenses were mildly opacified. To our knowledge, this result was never reported before and
suggests that the removal of SVO may have a significant impact on near-distance daily life
activities, but only in the absence of cataract or lens opacification. Because near distance
activities rely on fine central vision, for which performance is rapidly degraded past a critical
contrast threshold²⁴, SVO removal may not be sufficient to help improve performance if contrast
sensitivity is still reduced from lens opacification.

331

Our second result is the poor MRS achieved in all 10 patients with SVO and clear lenses (138 332 333 wpm on average in the operated eye prior surgery) compared to normal values. According to Calabrèse et al., 2016, normal readers between 58 to 68 years old should reach a MRS comprised 334 between 183.2 and 189.2 wpm when reading with one or both eyes^{25,26}. This 35% decrease 335 suggests that reading speed may be considered as an objective measure of functional impairment 336 in the presence of SVO. However, this finding should be interpreted with caution, given that 337 other confounding clinical factors (e.g. cognitive or visual) may have also contributed to 338 reducing reading speed. 339

340

Our third outcome is the significant change in MRS, measured after vitrectomy in patients with clear lenses, with a 15% improvement in the operated eye. For these patients, the non-operated eye served as control and showed no improvement post-surgery, confirming that the improvement measured in the fellow eye was not due to a practice effect. More importantly, this improvement did not occur in eyes with mildly opacified lenses, either from cataract (phakic eyes) or posterior capsule opacification (pseudophakic eyes). Taken all together, these results suggest that reading speed may be a valid objective measure to quantify the positive impact of 348 vitrectomy on visual function, but only if contrast sensitivity is not still altered by lens opacification. There is evidence that a main effect of vitrectomy is to restore normal contrast 349 sensitivity function for individuals with clear lenses^{6,27}. We hypothesize that for patients with 350 mildly opacified lenses, who experienced no post-surgery improvement in MRS, reduced 351 contrast sensitivity from cloudy ocular media created a bottleneck for any potential increase in 352 reading speed. We noted that binocular MRS was not improved post-surgery. Since, our 353 population was not restricted to patients with non-pathological fellow eyes, we did not expect to 354 see monocular vitrectomy having a significant impact on binocular performance. 355

356

ACC showed the same pattern as MRS, suggesting that this measure, which is potentially 357 quicker to obtain (in terms of testing and calculation time), could be a good alternative in clinical 358 359 settings where time is often limited. More interestingly, the improvement in ACC induced by vitrectomy was significantly correlated with the improvement in near distance activities score 360 measured with the NEI-VFQ. This result alone suggests that improved reading performance 361 following vitrectomy will also have a positive impact on the overall patients' quality of life. The 362 simple objective assessment of ACC post-operatively may therefore provide some insight to the 363 patient and his/her care team about his/her overall quality of life improvement. 364

365

Surprisingly, neither CPS nor RA were sensitive to the presence of dense floaters. Even more, we found no effect of vitrectomy on any of these measures. In their study of 110 treated eyes, Nie et al., 2013 reported that 71% of their patients had difficulty in reading small print, which markedly improved after surgery⁵. Based on their results, we had hypothesized that RA (i.e. the smallest print one can read) would improve following vitrectomy. However, our results do not support this hypothesis and suggest that these reading measures may not be valid to quantify theimpact of floaters on daily visual function.

373

We had expected patients with the eyes having the most prominent vitreous opacities to exhibit 374 the greatest improvement in both NEI-VFQ scores and reading performance. This was not the 375 case. In clinical practice, patients with a wide range of vitreous debris are seen, and often 376 individuals with very substantial opacities can be essentially asymptomatic (as in asteroid 377 hvalosis)²⁸. Our result, as well as the wide variability in dysfunction among patients with similar 378 vitreous opacities, suggests that the location and motion characteristics of the opacities may be 379 more significant drivers than the level of opacity itself in the decision to seek symptomatic relief 380 with surgery. However, the ability to show the degree of vitreous opacification using the video 381 SLO was found to be helpful for educational purposes, both pre- and post-operatively. First, to 382 show family members dynamically what the patients were seeing. Second, to help persuading 383 patients with significant complaints but mild opacities on SLO testing that surgery would not be 384 prudent. Finally, to document the absence of the opacities post-surgery. 385

386

Our work presents some limitations. The main one is the restricted number of patients. In the future, our results should be replicated with larger sets of patients to confirm our findings. Another limitation is that, given the nature of the MNREAD, the current study only measured fluent reading for short sentences. Therefore, it remains to be determined whether speed is also improved (and to what extent) for spot reading (i.e. for isolated words, such as tag labels) and sustained reading (i.e. for long texts).

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398

399 Ethical Approval

400 This study was conducted in accordance with the Declaration of Helsinki. Ethical approval for

401 this study was obtained from the Institutional Review Board (IRB) at the University of South

402 Carolina. The collection and evaluation of all protected patient health information was performed

403 in a Health Insurance Portability and Accountability Act (HIPAA)-compliant manner.

404

405 Statement of Informed Consent

406 Written informed consent was obtained before the study from each patient according to IRB

407 guidelines, including permission for publication of all videos included herein.

408

409

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474

475 **Figure legends**

476 <u>Figure 1:</u> Protocol schematic showing the different test procedures along with the resulting
477 outcome measures. Subjective measures are represented in blue; objective measures are
478 represented in pink.

Figure 2: Pre and post-operative NEI-VFQ scores grouped by lens opacity status. Points show
the mean estimates for the near activity sub-score (A) and the overall composite score (B), both
before and after surgery, as estimated by the mixed effects models, for patients with clear lenses
in blue (N=10) and patients with mildly opacified lenses in orange (N=10). Error bars represent
the 95% confidence intervals.

485

Figure 3: Effect of pre/post-surgery condition on MRS for the operated eye (top – triangles), the
non-operated eye (center - circles) and the binocular condition (bottom – squares) grouped by
lens opacity: clear (left – blue) vs. mildly opacified (right - orange). Solid lines connect the
estimates for each sub-group as given by the mixed-effects model. Errors bars (black) represent
their standard errors. Dashed lines connect the MRS values for each patient, numbered from P1
to P20.

492

Figure 4: Effect of pre/post-surgery condition on ACC for the operated eye (top – triangles), the
non-operated eye (center - circles) and the binocular condition (bottom – squares) grouped by
lens opacity: clear (left – blue) vs. mildly opacified (right - orange). Solid lines connect the
estimates for each sub-group as given by the mixed effects model. Errors bars (black) represent
their standard errors. Dashed lines connect the MRS values for each patient, numbered from P1
to P20.

499

500 Figure 5: Post-operative NEI-VFQ near activity sub-score improvement as a function of post501 operative reading accessibility index improvement for all 20 patients.

503 **Tables**

Patient	Patient Lens opacity							Non-operated eye			
ID	Location	Gender	Age	in both eyes	Pathology	SVO	Acuity	OCT-SLO Opacity grading	Pathology	SVO	Acuit y
P1	Minnesota	М	58	Clear	PVD	Yes	20/25	2	ERM	No	20/25
P2	Minnesota	М	59	Clear	PVD	Yes	20/20	1.5		Yes	20/25
P3	California	М	61	Clear	PVD	Yes	20/20	3	ERM	No	20/40
P4	Minnesota	М	62	Clear	PVD+ ERM	Yes	20/20	1.5	Scleral bucklin g + ERM	No	20/20
P5	Minnesota	М	64	Clear	PVD+ ERM	Yes	20/15	2	PVD	Yes	20/25
P6	Minnesota	F	64	Clear	PVD	Yes	20/25	2.5		Yes	20/20
P7	Minnesota	F	64	Clear	PVD	Yes	20/20	2.5	PVD	Yes	20/25
P8	Minnesota	F	68	Clear	PVD	Yes	20/30	1	PVD	Yes	20/15
Р9	California	М	69	Clear	PVD	Yes	20/20	2.5	PVD	Yes	20/20
P10	Minnesota	F	72	Clear	PVD	Yes	20/25	2	PVD	Yes	20/25
P11	California	F	32	Mild opacity	PVD	Yes	20/25	1	PVD	Yes	20/80
P12	California	М	52	Mild opacity	PVD+ ERM	Yes	20/25	2.5	Vitreous Syneresis	No	20/20
P13	California	М	54	Mild opacity	PVD+ ERM	Yes	20/40	3	NPDR	No	20/20
P14	California	F	54	Mild opacity	PVD	Yes	20/25	2.5	PVD	Yes	20/80
P15	Minnesota	М	63	Mild opacity	PVD+ ERM	Yes	20/40	2.5	ERM	Yes	20/25
P16	California	М	63	Mild opacity	PVD+ ERM	Yes	20/80	2	ERM	No	20/25
P17	Minnesota	М	64	Mild opacity	PVD	Yes	20/20	2.5	Vitreous Syneresis	Yes	20/20
P18	California	F	65	Mild opacity	PVD	Yes	20/30	2.5	PVD	Yes	20/25
P19	California	F	67	Mild opacity	PVD	Yes	20/30	2.5	ERM	No	20/25
P20	Minnesota	М	68	Mild opacity	PVD	Yes	20/20	1.5	PVD	Yes	20/25

504

505 <u>Table 1: Patients' individual characteristics prior to surgery.</u> SVO stands for symptomatic

vitreous opacities; ERM stands for epiretinal membrane. PVD stands for posterior vitreous

507 detachment; NPDR stands for non-proliferative diabetic retinopathy; Visual acuity is given in

508 Snellen notation.

510 Supplementary material

511	Movie 1. Video 1, preoperative video SLO. The video is live streaming of scanning laser
512	ophthalmoscopic images from the Heidelberg OCT machine. This is recorded in "Movie Max"
513	mode, (in avi) then converted to .mov. A patient with prominent vitreous opacities months post
514	scleral buckling was instructed to look left and re-fixate, then look right and re-fixate. Shadows
515	from mobile vitreous opacities were projected on the stabilized retinal surface and thus imaged
516	with the infrared camera.

517

518 Movie 2. Video 2, postoperative video SLO. The video is recorded in the same manner 519 (with saccades) as Video 1, and of the same eye one week post vitrectomy. Absence of 520 shadowing from vitreous opacities is noted.