

## **Functional Impairment of Reading in Patients with Dry Eye**

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1 **SUBTITLE**

- 2 Dry eye is associated with slower out-loud and silent reading. The decrement in reading speed  
3 directly correlates with the severity of dry eye disease, as measured by the Ocular Surface  
4 Disease Index and corneal staining score.

5 **ABSTRACT**

6 **Background/Aims:** To evaluate the impact of dry eye on reading performance.

7 **Methods:** Out-loud and silent reading in patients with clinically significant dry eye (n=41) and  
8 controls (n=50) was evaluated using standardized texts. Dry eye measures included tear film  
9 break-up time, Schirmer's test, and corneal epithelial staining. Symptoms were assessed by the  
10 Ocular Surface Disease Index.

11 **Results:** The dry eye group had a greater proportion of women as compared to the control group  
12 but did not differ in age, race, education level, or visual acuity ( $p \geq 0.05$  for all). Out-loud  
13 reading speed averaged 148 words per minute (wpm) in dry eye subjects and 163 wpm in  
14 controls ( $p=0.006$ ). Prolonged silent reading speed averaged 199 wpm in dry eye subjects versus  
15 226 wpm in controls ( $p=0.03$ ). In multivariable regression models, out-loud and sustained silent  
16 reading speeds were 10 wpm (95% CI= -20 to -1 wpm,  $p=0.039$ ) and 14% (95% CI = -25% to -  
17 2%,  $p=0.032$ ) slower, respectively, in dry eye subjects as compared with controls. Greater  
18 corneal staining was associated with slower out-loud (-2 wpm/1 unit increase in staining score,  
19 95% CI=-3 to -0.3 wpm) and silent (-2%, 95% CI=-4 to -0.6 wpm) reading speeds ( $p<0.02$  for  
20 both). Significant interactions were found between OSDI score and word-specific features  
21 (longer and less commonly used words) on out-loud reading speed ( $p<0.05$  for both).

22 **Conclusions:** Dry eye is associated with slower out-loud and silent reading speeds, providing  
23 direct evidence regarding the functional impact of dry eye. Reading speed represents a  
24 measurable clinical finding that correlates directly with dry eye severity.

25 **INTRODUCTION**

26 Dry eye is a common condition affecting approximately one in three individuals over the  
27 age of 50.[1–4] Although ocular discomfort may be the most bothersome symptom, visual  
28 complaints are also common. Dry eye has a substantial yet often under-appreciated impact on  
29 vision-related quality of life.[5–6] Prior research has shown that dry eye patients report difficulty  
30 in various vision-related tasks such as driving, reading, computer work, watching television, and  
31 performing work-related activities.[7–11] Arguably the most common visual complaint reported  
32 is difficulty with reading, which may affect employment or decrease work productivity.

33 In a population-based sample of elderly, we previously noted that dry eye symptoms were  
34 associated with greater perceived difficulty with reading and also the avoidance of specific  
35 reading tasks.[12] Here, we designed a clinical study to quantify reading performance through  
36 measuring actual reading speed on both a full-passage and individual word level by using several  
37 different previously validated texts.

38

39 **MATERIALS AND METHODS**

40 The study protocol was approved by the Johns Hopkins University Institutional Review  
41 Board in accordance with the Declaration of Helsinki and was Health Insurance Portability and  
42 Accountability Act compliant. Study subjects completed the study procedures between July 2009  
43 and January 2012.

44 **Study Subjects**

45 Eligible subjects had to be 50 years or older, literate by self-report, and able to  
46 communicate in English. Dry eye subjects were recruited from the Ocular Surface Diseases  
47 Clinic at Wilmer Eye Institute and had: (1) clinically significant dry eye defined as Schirmer’s

48 test result without anesthesia  $\leq 7$  mm at 5 minutes and/or bulbar conjunctival staining with  
49 lissamine green  $\geq 1$  on the Oxford scale in either eye [13], and (2) an Ocular Surface Disease  
50 Index (OSDI) score of 13 or higher. All patients were on topical treatment at the time of  
51 enrollment (including artificial tears and/or anti-inflammatories), which was not held prior to  
52 testing.

53 Control subjects were gathered from individuals followed for suspicion of glaucoma at  
54 the Glaucoma Clinic of the Wilmer Eye Institute who had (1) never been diagnosed with dry eye,  
55 and (2) had an OSDI score of 12 or less. All controls had normal visual fields in both eyes over  
56 the central 24 degrees using a size III stimulus and assessed by the Swedish interactive threshold  
57 algorithm standard testing program (HFA2, Carl Zeiss Meditec Inc., Dublin CA). Thirty-one  
58 (62%) of the control subjects were on intraocular pressure-lowering drops at the time of  
59 enrollment, which was not held prior to testing.

## 60 **Tests Performed**

61 All subjects were examined in a uniform manner using the tests performed on a single  
62 day in the following order:

### 63 *Evaluation of Vision and Covariates*

64 Sociodemographic variables were gathered using standardized forms. Visual acuity was  
65 measured binocularly with patients' habitual distance correction using the Early Treatment  
66 Diabetic Retinopathy Study vision chart, and summarized as the negative logarithm of the  
67 minimum angle of resolution(logMAR).[14,15] All subjects had at least 20/40 or better vision in  
68 both eyes.

69 Contrast sensitivity was measured using the Pelli-Robson chart under binocular  
70 conditions and converted to a log scale.[16] The presence of depressive symptoms was assessed

71 using part D of the General Health Questionnaire.[17] Cognitive ability was evaluated using the  
72 Mini Mental State Exam (MMSE).[18] After reading tests were administered, pupils were  
73 pharmacologically dilated and lens changes were graded as present or absent as described  
74 previously.[19]

### 75 *Evaluation of Reading*

76 Subjects wore their habitual reading correction for the following assessments: (a) out-  
77 loud reading speed using the Minnesota low vision reading test (MNRead)[20], (b) out-loud  
78 reading speed using a 77-word international reading speed test (IReST)[21], and (c) sustained  
79 silent reading speed using a 7,300-word validated passage read silently for 30 minutes or until  
80 the passage is finished. Greater detail regarding the administration of these three reading tests is  
81 provided elsewhere.[22]

82 Reading speed was calculated in words per minute (wpm). Maximum reading speed was  
83 calculated from MNRead times using nonlinear mixed effects models.[23] IReST passage  
84 reading speed was calculated after adjusting for reading errors. Sustained silent reading speed  
85 was calculated from the total words read and time required for reading. Details regarding these  
86 parameters are provided elsewhere.[22,23]

### 87 *Evaluation of Word-specific Reading Data*

88 Audiorecordings of the IReST passage were imported into Wave Editor Version 1.5.5  
89 (Audiofile Engineering, Minneapolis, MN) and analyzed by a masked evaluator. The start and  
90 end of each individual word was determined using the software spectrogram, and then imported  
91 into a separate database to calculate the exact duration to say each word out-loud and the  
92 following interval duration (before the start of the next word). Each word was analyzed as a  
93 word plus post-word interval unit to capture any potential interactional effect of the word-level

94 feature (i.e. word length, word frequency, and location of word in text). A detailed description  
95 of the derivation of these outcomes is described in detail elsewhere.[24]

### 96 ***Dry Eye Evaluations***

97 The Ocular Surface Disease Index (OSDI) questionnaire was administered to all subjects  
98 by a masked examiner.[10] Total scores were categorized for severity (normal=0–12, mild=13–  
99 22, moderate=23–32, severe=33–100).[25,26] A similar formula was used to compute two OSDI  
100 subscores: 1) vision-related subscore corresponding to questions 4-9 assessing the impact of dry  
101 eye on visual functioning, and 2) ocular discomfort-related subscore corresponding to questions  
102 1-3 and 10-12 evaluating symptoms relating to irritation or discomfort.[27] Subscale scores  
103 ranged from 0 to 50.

104 Dry eye signs was assessed by one of three masked examiners (EKA, PYR, or CAU) and  
105 in the order listed here. Tear film break-up time (TBUT) was measured with 5 microliters of  
106 anesthetic-free preservative-free 2% sodium fluorescein using the cobalt blue light of a slit lamp  
107 and a Wratten 12 yellow filter 1 minute after instilling the eye drop. Three TBUT measurements  
108 were obtained (maximum value of 10 seconds) and averaged for each eye.

109 Corneal staining was evaluated using the National Eye Institute grading system. Within 2  
110 to 3 minutes after TBUT testing, the extent of punctate epithelial erosions was graded using  
111 Wratten 12 filter paper.[28] Total corneal staining grade for each eye ranged from 0–15. Lastly,  
112 Schirmer’s test was performed without anesthesia in each eye at least 10 minutes after corneal  
113 staining assessment, read at 5 minutes, and averaged.[29]

### 114 **Statistical Methods**

115 Group differences in demographic, health, and visual features were assessed using the  
116 Student’s t-test for normally-distributed continuous variables, Wilcoxon rank sum testing for

117 non-normally distributed continuous variables, and chi-squared testing for categorical variables.  
118 The worse eye values for the TBUT, corneal staining, and Schirmer's test were used for the data  
119 analysis. Variables associated with MNRead and IReST reading speeds were evaluated using  
120 age-adjusted and multivariable linear regression models adjusting for age, sex, race, education,  
121 employment status, cognitive ability, and the presence of depressive symptoms. Sustained silent  
122 reading speeds were log-transformed and analyzed in age-adjusted and multivariable linear  
123 regression models in order to obtain normally-distributed residuals. The percent change in log  
124 sustained silent reading speeds associated with model elements was calculated as  $(10^{(\beta)}-1)*100$ .

125 Predictors of the word/post-word interval unit were evaluated using multivariate linear  
126 regression models. Covariates were included in multivariate models if they demonstrated a  
127 significant impact on word time in age-adjusted models or had been previously shown to impact  
128 reading speed.[30] Word features (i.e. word size, word frequency, location in text) were also  
129 included in multivariable models. Lastly, GEE multivariate models were used to assess  
130 interactions between dry eye severity and word features on word/post-word interval time. This  
131 interaction analysis was included to evaluate whether dry eye patients had particular difficulty  
132 with certain text features, similar to glaucoma patients.[24] All data were analyzed using STATA  
133 statistical software (STATA Release 12.1; STATA Corp., College Station, TX).

134

## 135 **RESULTS**

136 Forty-one dry eye patients and 50 controls completed study procedures and were included  
137 for analysis. One patient was excluded based on a greater than 2-fold difference between their  
138 silent and out-loud reading speeds.



139 Participant characteristics are summarized in Table 1. There was no difference between  
140 the two groups with regards to sociodemographic characteristics, cognitive ability, depressive  
141 symptoms, presence of cataracts/posterior capsular opacity, visual acuity, or contrast sensitivity.  
142 Women formed a greater proportion of the dry eye subject group as compared to the control  
143 group (90% vs. 58%,  $p=0.001$ ). Subjects with dry eye had significantly greater total (39.5 vs. 4.7,  
144  $p<0.001$ ), ocular discomfort-related (22.2 vs. 2.8,  $p<0.001$ ) and vision-related (17.3 vs. 1.8,  
145  $p<0.001$ ) OSDI scores than controls, in addition to shorter TBUTs (1.9 vs. 3.3 seconds,  $p=0.01$ )  
146 and greater corneal fluorescein staining (7.4 vs. 5.2,  $p=0.007$ ). Schirmer's test without anesthesia  
147 did not differ between the two groups ( $p=0.41$ ).

148 In unadjusted analyses, dry eye subjects demonstrated slower reading speeds than  
149 controls for the IReST passage (148 vs. 163 wpm,  $p=0.006$ ) and sustained silent reading (199 vs.  
150 226 wpm,  $p=0.03$ ) but did not demonstrate slower maximum reading speeds in the MNRead test  
151 (180 vs. 186 wpm,  $p=0.22$ )(Table 2). No significant differences were noted in other MNRead  
152 parameters including reading acuity and critical print size ( $p>0.05$  for both)(Table 2).

153 In multivariable models, dry eye was associated with significantly reduced IReST  
154 passage reading speed (-10 wpm, 95% CI=-20 to -1 wpm,  $p=0.04$ ) and sustained silent reading  
155 speed (14% slower, 95% CI=-25 to -1%,  $p=0.03$ ), but not with a slower maximum MNRead  
156 speed (Table 3). In separate multivariable models, reduction in the MNRead, IReST, and  
157 sustained silent reading speeds correlated with total OSDI scores ( $p\leq 0.05$  for all). Ocular-  
158 discomfort-related and vision-related subscores were associated with slower IReST and sustained  
159 silent reading ( $p\leq 0.05$  for both), but not for the MNRead passage. As compared to those with  
160 normal OSDI scores, those with severe scores had significantly slower IReST (-18 wpm, 95%

161 CI=-31 to -7,  $p=0.003$ ) and sustained silent reading (26% slower, 95% CI=-38 to -13%,  
162  $p<0.001$ ).

163 Additional multivariable models were run to determine the association between ocular  
164 surface measures and reading speed (Table 3). Worse-eye TBUT was not significantly associated  
165 with reading speed for all three tests. Corneal staining was associated with changes in IReST (-2  
166 wpm/1 unit change in staining score, 95% CI=-3 to -0.3,  $p=0.015$ ) and sustained silent reading  
167 speeds (-2%/1 unit change in staining score, 95% CI=-4 to -0.6,  $p=0.009$ ), but not with  
168 maximum reading speed calculated from the MNRead test ( $p=0.93$ ). African American race and  
169 lower MMSE score were significantly associated with reduced reading speed for at least one  
170 reading test.

171 Multivariate GEE models (using the exchangeable correlation structure) assessing the  
172 time required to read individual word/post-word interval durations demonstrated that higher  
173 OSDI (+1.1 ms/1 point increase in OSDI; 95%CI = 0.6 to 1.5;  $p<0.001$ ) and corneal staining  
174 scores (+3.0 ms/1 point increase in corneal staining; 95%CI = 0.1 to 5.8;  $p=0.045$ ), but not  
175 TBUT or Schirmer's ( $p>0.05$  for both), were associated with longer word/post-word interval  
176 complex durations. Greater word/post-word interval durations were also associated with  
177 increased word size, word frequency, and word location (end of line versus any other location)  
178 ( $p<0.05$  for all).

179 Interactions between dry eye severity and text features on word/post-word interval  
180 durations were also analyzed in separate multivariate GEE models for each dry eye metric. Each  
181 interaction model included the metric of dry eye severity, word feature of interest, interaction  
182 term (dry eye metric x word feature), and all relevant non-visual metrics. Significant interactions

183 were noted between greater OSDI score and both word length (p=0.002) and word frequency  
184 (p=0.02), but not with any other dry eye measures or features (p>0.05 for all).(Table 4)

185

## 186 **DISCUSSION**

187 In this clinic-based patient population, dry eye was associated with reduced reading  
188 speeds using a variety of reading tests. This decrement correlated directly with the severity of  
189 symptoms as measured with OSDI. Individuals with severe dry eye symptoms (OSDI score>33)  
190 had substantial reductions in sustained silent reading (26% decrease in wpm). These findings  
191 suggest that dry eye symptoms impair reading performance, and likely interfere with daily  
192 activities for which reading is critical.

193 Previous studies have demonstrated the functional impact of dry eye on various everyday  
194 tasks, such as reading.[5-10,31-32] We previously demonstrated self-reported difficulty with  
195 reading in an elderly population-based cohort.[12] In that study, dry eye did not significantly  
196 affect reading speed, although dry eye subjects reported reading difficulty and avoidance of  
197 newspaper reading. This discrepancy can be attributed to the fact that subjects from our prior  
198 study were derived from a population-based sample who are likely to have less severe disease,  
199 compared to the patients in the current study who were cared for at a tertiary dry eye center.  
200 Additionally, in our previous study, reading speed was only measured using short out-loud text  
201 passages. Finally, limited objective measures were available to categorize the severity of the dry  
202 eye in our prior work.

203 Only two other studies to our knowledge have evaluated reading speed in dry eye. One  
204 study used the Wilkins Rate of Reading test, which consists of simple words without context that  
205 are read aloud and takes less than 2 minutes to complete.[33, 34] Dry eye subjects exhibited

206 slower reading speeds ( $134.9 \pm 4.95$  wpm) than controls ( $158.3 \pm 8.40$  wpm,  $p=0.046$ ), but were  
207 not undergoing treatment at the time of evaluation which may have resulted in a larger difference  
208 in reading speed than we observed. Another recent small-scale case-control study reported  
209 slower reading rates in dry eye patients as well, but its association with subjective or objective  
210 measures of dry eye disease was not studied.[35] Our study improves on the methodology of  
211 prior studies by using reading tests that more closely mimic reading scenarios which patients  
212 encounter in their day-to-day lives.

213 An interesting finding in our study was that the impact of dry eye on reading speed  
214 differed based on the type of reading test employed. Of the two out-loud reading tests, the  
215 magnitude of the associations found between dry eye measures and IReST reading speed was  
216 greater as compared to MNRead maximum reading speed. One possible reason for this  
217 difference is that dry eye exerts its impact on reading speed through visual disturbances that were  
218 not identified in the current study (our groups had similar distance/reading acuity and contrast  
219 sensitivities). MNRead reading speeds are modeled as the maximum reading speed observed for  
220 the sentences presented at different text sizes, and perhaps larger text size can overcome the  
221 visual disturbances associated with dry eye. We found a greater impact of dry eye on sustained  
222 silent reading speed. In multivariable models, dry eye was associated with 14% slower sustained  
223 silent reading (20 wpm decrement at the mean reading speed,  $p=0.03$ ), while the reduction with  
224 IReST testing was 15 wpm ( $p=0.04$ ). Our findings therefore support the validity and utility of  
225 sustained silent reading speed as an important measure to evaluate patients with dry eye  
226 disease.[22] Finally, our interaction analysis showed that dry eye patients do not appear to have  
227 particular difficulty with word-specific features, in contrast to what has been demonstrated in the  
228 glaucoma population.[24] These results suggest that dry eye disease likely affects reading in a

229 more diffuse manner, as opposed to a distinct process which manifests with particular text  
230 features (i.e. peripheral visual constriction in glaucoma patients leading to particular difficulty  
231 during line transitions). For example, decreased ocular optical qualities due to dry eye disease  
232 (i.e. those captured by dynamic aberrometry) may represent the mechanism of decreased reading  
233 speed.[36] Therefore, visual rehabilitation may be more difficult to specifically tailor to the dry  
234 eye population as compared to other ocular conditions.

235         The limitations of our study include the fact that a great majority of the participants were  
236 on topical therapy (artificial tears, anti-inflammatories, or intraocular pressure-lowering drops),  
237 which was not held prior to enrollment. It is possible that the overall reading disability measured  
238 here is understated, given that dry eye patients were getting appropriate therapy that was not held  
239 prior to testing. The participants represented a convenience sample; therefore, perhaps less  
240 symptomatic dry eye patients were less likely to participate, biasing our findings in a positive  
241 direction. Also, patients with best-corrected visual acuity (BCVA) better than 20/40 were  
242 included, but may have had other pathology influencing reading speed. However there was no  
243 statistical difference in the BCVA between the two groups, and the associations were observed to  
244 exist independent of BCVA. Additionally, we included glaucoma suspects as controls and not  
245 individuals without any signs of dry eye. We considered the possibility that using this control  
246 group could bias our findings towards the null hypothesis if reading speed was affected by eye  
247 drop therapy. However, in our sensitivity analyses we found no difference in reading speed on  
248 any of the tests between controls using eye drops to those who did not. In addition, controls who  
249 attend essentially the same clinic as cases are more likely to be similar on unmeasured factors.  
250 Recruitment of entirely normal controls (i.e. spouses or friends accompanying patients to clinic)  
251 would likely exclude individuals who are less likely to venture outside the home due to poorer

252 general health, mood, or cognitive ability, thus producing a “supranormal” group of controls.  
253 Also our data collection did not include blink frequency, which could affect reading time and dry  
254 eye measurements. Our findings pertain to a specific set of office-based environmental testing  
255 conditions, and the effect of dry eye on reading may differ under other conditions such as higher  
256 or lower humidity or air drafts or different lighting conditions. Future studies may consider  
257 using dynamic aberrometry of the tear film in the future, which could be utilized as a  
258 standardized surrogate marker and potentially facilitate multicenter clinical trials.[37]

259 In summary, our findings provide direct evidence for the impact of dry eye on reading  
260 performance. Our results show that reading speed could be utilized as a tool to directly measure  
261 functional impairment from dry eye.

**Table 1.** Characteristics of subjects with physician-diagnosed dry eye versus controls participating in reading evaluations.

	<b>Control* (n=50)</b>	<b>Dry Eye (n=41)</b>	<b><i>p</i> value</b>
<b>Demographics</b>			
Mean age, years (SD)	67.4 (8.5)	65.7 (10.3)	0.42
African-American, n (%)	9 (18.0)	3 (7.3)	0.13
Female, n (%)	<b>29 (58.0)</b>	<b>37 (90.2)</b>	<b>0.001</b>
Education, years (SD)	15.6 (2.1)	15.1 (1.9)	0.25
Employed, n (%)	24 (48.0)	16 (39.0)	0.39
<b>Vision</b>			
Better-eye acuity, logMAR, mean (SD)	-0.01 (0.11)	0.02 (0.10)	0.18
Binocular log CS, mean (SD)	1.93 (0.12)	1.88 (0.16)	0.11
Cataract/PCO, either eye, n (%)	4 (8.0)	6 (14.6)	0.31
<b>Health</b>			
MMSE Score, mean (SD)	27.6 (1.5)	26.8 (2.4)	0.06
Depressive symptoms, n (%)	3 (6.0)	6 (14.6)	0.17
<b>Dry Eye Measures (SD)</b>			
Mean OSDI Total score	<b>4.7 (3.8)</b>	<b>39.5 (21.1)</b>	<b>&lt;0.001</b>
Mean OSDI Discomfort subscore	<b>2.8 (2.8)</b>	<b>22.2 (12.2)</b>	<b>&lt;0.001</b>
Mean OSDI Vision subscore	<b>1.8 (2.1)</b>	<b>17.3 (1.9)</b>	<b>&lt;0.001</b>
Mean TBUT in worse eye	<b>3.3 (3.0)</b>	<b>1.9 (2.0)</b>	<b>0.01</b>
Mean corneal staining score in worse eye	<b>5.2 (3.8)</b>	<b>7.4 (3.6)</b>	<b>0.007</b>
Mean Schirmer's test in worse eye	10.2 (9.2)	8.6 (9.2)	0.41

\*Control patients included were glaucoma suspects without any history or symptoms of dry eye disease

**CS:** Contrast sensitivity; **logMAR:** Logarithm of the minimum angle of resolution; **MMSE:** Mini Mental State Exam; **OSDI:** Ocular Surface Disease Index; **PCO:** Posterior capsular opacity (in pseudophakic subjects); **SD:** Standard deviation; **TBUT:** Tear film break up time

**Table 2.** Comparison of reading parameters in subjects with physician-diagnosed dry eye versus controls: Unadjusted values.

	<b>Control* (n=50)</b>	<b>Dry Eye (n=41)</b>	<b>p value</b>
<b>Out-loud reading, MNRead acuity card (SD)</b>			
Mean maximum reading speed, wpm	186 (21)	180 (25)	0.22
Mean critical print size	0.14 (0.16)	0.21 (0.21)	0.08
Mean reading acuity, logMAR	-0.05 (0.11)	-0.01 (0.15)	0.09
<b>Out loud reading, IReST passage</b>			
Mean reading speed (SD) wpm	<b>163 (22)</b>	<b>148 (27)</b>	<b>0.006</b>
<b>Sustained silent reading passage (SD)</b>			
Median reading speed, wpm	<b>226 (59)</b>	<b>199 (82)</b>	<b>0.03</b>

\*Control patients included were glaucoma suspects without any history or symptoms of dry eye disease

**IReST:** International Reading Speed Text; **MNRead card:** The Minnesota low vision reading test; **SD:** Standard deviation; **wpm:** words per minute



**Table 3.** Associations between MNRead, IReST, and sustained silent reading speeds with dry eye status, vision, demographic, and health variables in subjects with physician diagnosed dry eye versus controls: Multivariable analyses

Variable	Interval	Outloud (MNRead) Reading Speed Change in wpm (95% CI)	Outloud (IReST) Reading Speed Change in wpm (95% CI)	Sustained Silent Reading Speed % Change (95% CI)
<b>Vision Parameters</b>				
Dry eye (OSDI $\geq 13$ )	vs. control	-1 (-11 to 9)	<b>-10 (-20 to -1)</b>	<b>-14% (-25 to -1)</b>
OSDI Discomfort score	5 units lower	-2 (-4 to 0.1)	<b>-4 (-3 to -1)</b>	<b>-4% (-7 to -2)</b>
OSDI Vision score	5 units lower	-2 (-4 to 0.2)	<b>-3 (-5 to -1)</b>	<b>-5% (-8 to -2)</b>
OSDI Total score	5 units lower	<b>-1 (-2 to -0.002)</b>	<b>-2 (-3 to -1)</b>	<b>-3% (-4 to -1)</b>
OSDI Total Score Severity				
Mild(13 to 22)	vs. normal (0 to 12)	9 (-6 to 23)	-2 (-16 to 13)	1% (-19 to 21)
Moderate(23 to 32)		-6 (-20 to 9)	-6 (-21 to 9)	-5% (-23 to 17)
Severe (33 to 100)		-5 (-17 to 7)	<b>-18 (-31 to -7)</b>	<b>-26% (-38 to -13)</b>
TBUT in worse eye (seconds)	1 unit lower	-2 (-3 to 0.1)	-0.3 (-2 to 1)	0.3% (-2 to 3)
Corneal staining in worse eye	1 unit worse	-0.1 (-1 to 1)	<b>-2 (-3 to -0.3)</b>	<b>-2% (-4 to -0.6)</b>
Schirmer's in worse eye	1 mm greater	-0.04 (-1 to 0.4)	0.02 (-1 to 1)	-0.3 (-1.0 to 0.5)
<b>Non-Visual Parameters*</b>				
Age	5 years older	-0.2 (-3 to 3)	-0.2 (-2 to 2)	-1% (-5 to 3)
Male	vs. female	7 (-4 to 17)	1 (-10 to 12)	1% (-13 to 18)
Black	vs. non-black	-8 (-22 to 5)	-11 (-25 to 3)	<b>-28 (-41 to -13)</b>
Education	4 years less	-5 (-15 to 4)	-2 (-12 to 8)	-10% (-21 to 4)
Employed	vs. not employed	10 (-2 to 21)	9 (-2 to 20)	9% (-7 to 29)
MMSE score	5 points lower	<b>-13 (-25 to -1)</b>	<b>-24 (-37 to -13)</b>	-14% (-31 to 6)
Depressive symptoms	Present	-2 (-17 to 13)	-5 (-21 to 10)	-17% (-33 to 2)

**Bolded values represent statistical significance (p<0.05)**

\*The values for non-visual parameters taken from a single model including dry eye covariate and all nonvisual variables are shown. All other visual parameter values were derived from a separate multivariable model including the non-visual variables shown.

**IReST:** International Reading Speed Text; **OSDI:** Ocular Surface Disease Index; **SD:** Standard deviation; **TBUT:** Tear film break-up time; **wpm:** Words per minute

1 **Table 4:** Significant Interactions between Dry Eye Severity and Word Features on Word/Post-Word Interval Complex Duration, Multivariable  
 2 Analysis\*

<u>Variable</u>	<u>Interval</u>	<u>OSDI</u> Word/Post-Word Interval Complex (ms) β (95% CI)	<u>Tear Break Up Time</u> Word/Post-Word Interval Complex (ms) β (95% CI)	<u>Corneal Staining</u> Word/Post-Word Interval Complex (ms) β (95% CI)	<u>Schirmer's Test</u> Word/Post-Word Interval Complex (ms) β (95% CI)
<b>Dry Eye &amp; Word Size</b>					
Dry eye metric*	1 unit increase	0.2 (-0.5 to 0.9)	2 (-4 to 7)	0.1 (-4 to 4)	0.3 (-1.2 to 2)
Word Size	1 letter longer	<b>23 (19 to 28)</b>	<b>29 (24 to 34)</b>	<b>24 (18 to 29)</b>	<b>28 (23 to 33)</b>
Dry eye metric • Word Size <sup>†</sup>		<b>0.2 (0.1 to 0.3)</b>	-0.6 (-1 to 0.3)	0.7 (-0.1 to 1.4)	-0.1 (-0.3 to 0.2)
<b>Dry Eye &amp; Word Frequency<sup>‡</sup></b>					
Dry eye metric*	1 unit increase	<b>2 (1 to 3)</b>	-2 (-7 to 4)	<b>5 (0.2 to 10)</b>	-0.2 (-2 to 2)
Word Frequency	10 fold less common	<b>44 (41 to 48)</b>	<b>46 (43 to 49)</b>	<b>45 (40 to 49)</b>	<b>46 (43 to 50)</b>
Dry eye metric • Word Frequency <sup>†</sup>		<b>0.1 (0.02 to 0.2)</b>	0.1 (-0.6 to 0.8)	0.3 (-0.3 to 0.9)	0.02 (-0.2 to 0.2)
<b>Dry Eye &amp; Last Word of Line</b>					
Dry eye metric*	1 unit increase	<b>1 (0.5 to 1.5)</b>	-1 (-4 to 3)	3 (-0.4 to 5)	-0.3 (-1 to 1)
Last Word of Line	vs. not last word	<b>30 (7 to 53)</b>	<b>44 (18 to 71)</b>	21 (-9 to 51)	<b>44 (17 to 70)</b>
Dry eye metric • Last Word of Line <sup>†</sup>		0.4 (-0.4 to 1)	-2 (-7 to 4)	3 (-1 to 8)	-0.5 (-2 to 1)

3 **Bolded values represent outcomes with p<0.05.** Positive values indicate slower reading (longer word/post-word interval complex reading times) for words that  
 4 were longer, less frequently used, or found at the end of a line of text for the respective dry eye metric. Negative values represent faster reading (shorter  
 5 word/post-word interval complex reading time).

6 \* Four dry eye metrics used: OSDI (unit= 1 point), Tear Film Breakup Time (unit=1 second), Corneal Staining (unit= 1 point), and Schirmer's Test (unit= 1  
 7 millimeter).

8 † The impact of each interaction derived from a separate model including the dry eye metric, the word feature of interest, the interaction term (dry eye metric x  
 9 word feature), and all relevant non-visual metrics (age, gender, race, education, mini-mental state exam, word size, word frequency).

10 ‡ Represented by negative log of word frequency per million words used in common English language

11 **CI-** Confidence interval; **mm-** millimeter

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