

1 **TITLE PAGE**

2 **Title:**

3 Stereopsis simulating small aperture corneal inlay and monovision conditions

4 **Authors:**

5 José J. Castro<sup>1</sup> (PhD), Carolina Ortiz<sup>1</sup> (PhD), José R. Jiménez<sup>1</sup> (PhD), Sonia Ortiz-Peregrina<sup>1</sup>  
6 (MSc), Miriam Casares-López<sup>1</sup> (MSc)

7 **Institutional affiliation:**

8 <sup>1</sup>Laboratory of Vision Sciences and Applications. Department of Optics. University of Granada  
9 (Spain)

10 **Financial support:**

11 Ministry of Economy and Competitiveness (Spain) and European Regional Development Fund  
12 (ERDF) (Grant FIS2013-42204-R) and Ministry of Education, Culture and Sport (Spain) (Grant  
13 FPU15/05571)

14 **Corresponding author:**

15 José J. Castro

16 Department of Optics. University of Granada. Avenida de Fuentenueva, s/n. 18071 Granada  
17 (SPAIN)

18 Phone number: +34 958241902

19 Fax number: +34958248533

20 E-mail address: [jjcastro@ugr.es](mailto:jjcastro@ugr.es)

21 **Statement**

22 The authors declare that there are no conflicts of interest related to this article.

23    **PRECIS**

24    Stereopsis and other binocular visual functions studied using a simulated corneal-inlay  
25    implantation under a broad range of experimental conditions show a deterioration similar to that  
26    found in other surgical techniques.

27

28    **Introduction**

29    It is well known that presbyopia affects a very high percentage of population, and therefore  
30    there is intensive research for developing or improving techniques for its compensation.  
31    Intrastromal corneal inlay with a small aperture is a surgical technique of increasing use,<sup>1-6</sup> and  
32    is usually combined with a micro-monovision.

33    An essential aspect to be analysed in all correction techniques is the binocular vision  
34    performance<sup>4,7-9</sup> especially in the case of corneal inlays since the state of both eyes is clearly  
35    different and inter-ocular differences can arise and limit binocular function.<sup>9-11</sup> Binocular vision  
36    with implanted corneal inlays is affected by induced anisocoria since the differences between  
37    the pupil sizes can be notable and affect binocular vision, depending on the observation  
38    conditions.

39    Aspects of binocular vision such as binocular summation for the contrast-sensitivity function  
40    and the night-vision disturbances have been studied in operations or simulations of corneal  
41    inlays.<sup>8,9</sup> However, stereopsis, one of the most advanced functions of our visual system, as it  
42    enables us to distinguish spatial (3D) locations of visual objects around us, has hardly been  
43    studied for corneal inlay corrections.<sup>3</sup> Furthermore, one of the drawbacks most widely  
44    mentioned concerning the monovision technique, which sometimes is used together with  
45    corneal inlays, is the reduction or loss of stereoscopic capacity. For example, Artal et al. studied  
46    stereoacuity in a simulator of adaptive optics, though limited to three observers under photopic  
47    conditions and with parallel visual axes (far distance).<sup>3</sup>

48 The aim of the present work is to analyse stereoscopic vision under a broad range of  
49 experimental conditions simulating anisocoria that could arise in the surgical correction of a  
50 corneal inlay. For this, we have studied stereoacuity in a group of subjects with healthy eyes of  
51 different pupil sizes using a partially opaque contact lens. Also, we have added the effect of  
52 monovision in order to simulate a real situation for many patients operated on for corneal inlays.  
53 Under most experimental conditions, stereoacuity was measured in low-illumination  
54 surroundings where interocular-differences in pupil sizes are larger and we can expect a higher  
55 influence on stereopsis. In addition, we measured stereoacuity for a complete range of distances:  
56 far (5.5 m), intermediate (1 m) and near distances (0.4 m) to analyse the deterioration or not of  
57 stereopsis as a function of distance, especially for distances shorter than 1.5 m, where stereopsis  
58 is decisive. These types of simulation experiments are crucial because we can analyse the  
59 potential effect of some surgical techniques under certain conditions, avoiding the surgical  
60 effects. Although our main aim is to evaluate stereopsis, we also measured other binocular  
61 functions in order to compare the results and to make a more complete analysis of binocular  
62 vision. We measured the binocular summation for binocular contrast sensitivity and for a  
63 discrimination index as a measure of night-vision disturbances. Halos and/or glare under low-  
64 illumination conditions are usually mentioned after surgical techniques.<sup>12,13</sup>

65

## 66 **Methods**

### 67 *Subjects*

68 A total of 10 subjects were studied, with ages ranging from 20 to 25 years (22.7±1.8 years).  
69 Subjects gave their informed consent in accordance with the Helsinki Declaration. Admission  
70 criteria for the experiment were: astigmatism not greater than ±1.0 D, monocular corrected  
71 distance visual acuity of 20/20 or better, and no pathological disease or condition that could  
72 limit visual performance. Distance visual acuity was measured monocularly and binocularly for  
73 all subjects at a working distance of 5.5 m with the Pola VistaVision® Visual Acuity Chart

74 System (DMD Med Tech srl. Torino, Italy). All participants received a full refractive  
75 examination and objective refraction in which the refractive error was determined by non-  
76 cycloplegic retinoscopy. Monocular and binocular subjective refraction was also performed. The  
77 distance best-corrected visual acuity (DCVA) after trial frame refraction was measured. The  
78 mean refractive error (spherical equivalent) was  $-0.49 \pm 0.71$  D. All the observers had normal  
79 stereopsis according to the Randot stereotest (40 arcsec or lower).

#### 80 *Contact lenses*

81 The effect of anisocoria generated by corneal inlays was simulated by using a partial-opaque  
82 soft contact lens (pHema 38% water content), supplied by Servilens Fit & Cover (Granada,  
83 Spain) with a clear, circular, central aperture 1.6 mm in diameter. These lenses had two base  
84 curve radii (8.4 and 8.6 mm) and a 14.5-mm lens diameter. The outer diameter of the opaque  
85 region was 11.0 mm. For an acceptable fit, the contact lens had to be well centred both  
86 vertically and horizontally and moved from 0.5 to 1 mm on a blink. The lenses were worn for at  
87 least 20 min before any measurement. The lenses were fitted on the non-dominant eye (NDE) to  
88 create the required anisocoria. Ocular dominance of all subjects was checked with the Miles  
89 test.<sup>14</sup> After biomicroscopic examination, no signs of corneal oedema or corneal staining were  
90 detected either at the baseline of the study or at the end.

#### 91 *Procedures*

92 After several training sessions, all participants completed the following visual tests: visual  
93 acuity, contrast-sensitivity function, and visual-discrimination capacity (monocular and  
94 binocularly) and stereoacuity (binocularly) under natural conditions (without small-aperture  
95 contact lens).

96 To study the effect on binocular visual performance after inducing a simple anisocoria or  
97 combined with monovision, we also took measurements while observers wore the contact lens  
98 in the non-dominant eye (NDE). The measurements were performed binocularly and  
99 monocularly in the eye wearing the contact lens under the following conditions: a) no add

100 power; b) with an add power of +0.75 D, a value considered as optimum defocus in a corneal  
101 inlay modelling,<sup>3,15</sup> and c) with an add power of +1.25 D, a value used in traditional  
102 monovision.<sup>16</sup> The additions were inserted in a trial frame for each subject, as shown in other  
103 works<sup>9</sup>, and then the interpupillary distance was adjusted. These measurements were completed  
104 on different days to avoid observer fatigue. All the tests were performed at three test-distances:  
105 40 cm (near), 1 m (intermediate) and 5.5 m (far). For the CSF and the visual discrimination  
106 capacity, the observer position was fixed at the test distance using a chin and a forehead rest to  
107 minimize head movements. No mydriatics or cyclopegics were used.

#### 108 *Visual Acuity*

109 Monocular and binocular corrected distance visual acuity (CDVA) were measured using the  
110 Pola VistaVision® Visual Acuity Chart at 5.5 m. Monocular and binocular distance-corrected  
111 intermediate visual acuity (DCIVA) were measured with the Colenbrander Visual Acuity Chart  
112 NO4002 (Colenbrander, M.D., San Francisco, C.A.) at 1 m. Monocular and binocular distance-  
113 corrected near visual acuity (DCNVA) were measured with the Near Vision Chart (Promoción  
114 Optométrica, Burgos, Spain) at 40 cm.

#### 115 *Contrast-sensitivity function (CSF)*

116 Contrast sensitivity (CS) was evaluated using Gabor patches of gratings displayed on the  
117 monitor of the tablet ASUS Transformer Book T100TAF (10.1 inches). Contrast values were  
118 reported according to the Michelson contrast. The spatial frequencies tested were: 0.75, 1.5, 3.0,  
119 6.0, 12.0, and 18.0 cycles per degree (cpd), using Gabor patches of gratings with three possible  
120 orientations (vertical, left or right). The average luminance level of the monitor was 62.7 cd/m<sup>2</sup>  
121 and the test was performed in dim surroundings. The test method used was an alternate forced  
122 choice, in which, for each stimulus, the patient had to choose the grating orientation, displayed  
123 in a random order. For each spatial frequency, we determined the contrast threshold using an  
124 up-down staircase procedure with four reversals. The contrast threshold was calculated by

125 averaging the last three reversals. The grating subtended 1.3 deg for the three test distances (the  
126 CSF test was previously set for each distance).

### 127 *Night-vision disturbances*

128 The visual-discrimination capacity under low-illumination surrounding conditions was also  
129 evaluated using the test Halo, based on the freeware software Halo v1.0. This test has been  
130 successfully applied in basic and clinical research to quantify night-vision disturbances<sup>17-19</sup> In  
131 the test, the subject was shown a central high-luminance stimulus over a dark background on the  
132 monitor and, progressively, peripheral luminous stimuli were randomly shown around the  
133 central stimulus at different positions and distances from the main stimulus. We used the tablet  
134 ASUS Transformer Book T100TAF to run the Halo test. We measured the luminance of the  
135 visual stimuli using the spectroradiometer SpectraScanPR-670: 264cd/m<sup>2</sup>, 37cd/m<sup>2</sup>, and  
136 0.21cd/m<sup>2</sup> were the luminance of the main stimulus, the peripheral ones, and the monitor  
137 background, respectively. The spatial configuration was set for each test distance in such a way  
138 that the central stimulus and the peripheral ones subtended 0.46 and 0.04 deg, respectively, from  
139 the subject position.

140 The patient's task was to press the left button of the mouse whenever a peripheral stimulus  
141 could be discriminated. Once the test was finished, the software calculated the visual-  
142 disturbance index (VDI), a parameter widely studied in the literature.<sup>17-20</sup> The VDI ranges from  
143 0 to 1, in such a way that the higher the VDI, the lower the amount of peripheral stimuli  
144 discriminated, and, therefore, the worse the discrimination capacity (a greater influence of halos  
145 or night-vision disturbances).

### 146 *Stereoacuity*

147 For test distances of 1 and 5.5 m, we used the polarized stereotest implemented in the  
148 VistaVision® monitor. The stereotest evaluated stereopsis from 300 to 10 arc sec using  
149 polarized vertical lines. For each stereoacuity, a total of 5 vertical lines were displayed  
150 simultaneously on the monitor, one of them with disparity to be perceived stereoscopically. In

151 this test, the observer wore polarized glasses provided by the monitor manufacturer, and the task  
152 was to choose the line that was stereoscopically perceived.

153 For near vision (40 cm), we used the Random Dot stereo acuity test (Vision Assessment  
154 Corporation, IL) which evaluates stereoacuity from 500 to 12.5 arc sec.

### 155 *Binocular summation*

156 To compare binocular with monocular data in the CS as well as in the visual-discrimination  
157 capacity (VDI), we calculated binocular summation, a common metric used to characterize  
158 binocular visual performance.<sup>21-22</sup> Under normal binocular-vision conditions, the binocular  
159 summation of a visual function is a value higher than 1.0.

160 We reported binocular summation for the CS dividing the binocular CSF by the best monocular  
161 CSF. For each observer, binocular summation was provided as the average of the binocular  
162 summation determined for each spatial frequency.<sup>21-22</sup>

163 The binocular summation for the VDI was calculated dividing the lowest monocular VDI by the  
164 binocular VDI since as the discrimination capacity increases, the VDI decreases.<sup>9</sup>

165 Considering the binocular functions tested, the average pupil size for the eye with natural pupil,  
166 measured with a Colvard pupillometer (OASIS, Glendora, CA, USA), ranged from  $4.10 \pm 0.74$   
167 mm for the CSF and a distance of 40 cm to  $5.55 \pm 1.01$  mm for the halometer and a distance of 1  
168 m.

169

## 170 **Results**

171 Figure 1 and Table 1 show the results for stereoacuity. We find the results for the four  
172 conditions tested (natural pupil, small-aperture contact lens, small-aperture contact lens with  
173 monovision +0.75D and small-aperture contact lens with monovision +1.25D) in each of the  
174 three distances used: far, intermediate, and near. According to the repeated-measures analysis of  
175 variance (ANOVA) we found significant differences ( $p < 0.05$ ) in the group of near and

176 intermediate distances. A *post hoc* comparison analysis, for the near and intermediate distance,  
177 indicated that the natural pupil condition differed significantly ( $p < 0.05$ ) with respect to the other  
178 three conditions, with stereoacuity being better for the natural pupil condition. For near and  
179 intermediate distance, no significant differences ( $p > 0.05$ ) appeared between the three conditions  
180 when the small-aperture contact lens was used. For the far distance, we found no statistical  
181 difference in all *post hoc* comparisons except in the case natural pupil vs. small-aperture  
182 contact-lens monovision +1.25 D.

183 Figure 2 shows the results for binocular visual acuity. According to the ANOVA analysis, we  
184 found no significant differences ( $p > 0.05$ ) for any of the conditions tested in the three distances  
185 considered. A *post hoc* comparison analysis did not indicate any significant difference, either.

186 Figures 3 and 4 show the results for the binocular CSF and the binocular summation for the  
187 CSF. For the binocular CSF, according to the ANOVA analysis, we found statistical differences  
188 ( $p < 0.05$ ) for the three distances tested: near, intermediate, and far distance. For the near and  
189 intermediate distances, we analysed *post hoc* comparisons and the natural pupil condition was  
190 significantly better than the no-add and add +1.25D conditions. *Post hoc* comparisons analysis  
191 for the far distance showed that the natural pupil condition was significantly higher than the  
192 other three conditions. For the binocular summation (Fig. 4), the ANOVA analysis gave  
193 significant differences for the intermediate and far distance ( $p < 0.05$ ). For intermediate distance,  
194 *post hoc* comparisons analyses showed statistical differences between the natural pupil and the  
195 no-add condition. For far distance, we found statistical differences between the natural pupil  
196 conditions and the other three conditions, with higher binocular summation being found for the  
197 natural pupil condition.

198 Figures 5 and 6 show the results for the visual disturbance index (VDI): binocular VDI (Fig. 5)  
199 and binocular summation (Fig. 6). The ANOVA revealed statistical differences ( $p < 0.05$ ) among  
200 conditions in the binocular VDI for the far distance and in the binocular summation for the VDI  
201 for the three distances. From the *post hoc* comparison analysis, we found that the binocular VDI  
202 was significantly lower for the natural pupil condition with respect to the other 3 conditions in



203 the case of far distance, showing that visual-discrimination capacity is better for the natural  
204 pupil conditions. In the case of binocular summation for the VDI the *post hoc* comparisons  
205 analysis showed that binocular summation was significantly higher ( $p<0.05$ ) for the natural  
206 pupil condition than for the adds +0.75D and +1.25D conditions, for near, intermediate, and far  
207 distances.

208

## 209 **Discussion**

210 The results for stereoacuity show that, although stereoscopic perception occurs for all conditions  
211 tested, there was a deterioration in stereopsis for all small-aperture contact-lens conditions with  
212 respect to natural pupil being significant mainly for near and intermediate distance. It should be  
213 taken into account that the comparison was established under experimental conditions  
214 unfavourable to stereoscopic vision. In our experiments the illumination of the setting implies a  
215 larger pupil size for the eye with the natural pupil, and induced anisocoria causes large  
216 interocular differences<sup>9-11</sup> that limit stereoscopic vision. Under conditions where the anisocoria  
217 would be more reduced, the stereoscopic deterioration would be less.

218 In the study by Fernández et al.,<sup>3</sup> where stereoacuity was tested for only 3 subjects with parallel  
219 axes (far-vision scheme) and photopic vision, the authors found that the stereoacuity with the  
220 natural pupil did not significantly differ with respect to the conditions of small-aperture and  
221 monovision. In the study by Linn et al.,<sup>23</sup> the mean stereoacuity preoperatively and  
222 postoperatively measured was not statistically significant, either, although 25% of patients had  
223 worsened stereoacuity. In our study, there were significant differences between the condition of  
224 the natural pupil and those of small-aperture contact lens with monovision for near,  
225 intermediate, and far distance. The experimental differences between the studies may explain  
226 the different results.

227 It is also important to point out that the deterioration in stereopsis found in the optical  
228 simulation of the corneal inlays also has been found in other surgical techniques for correcting

229 presbyopia<sup>24,25</sup> and myopia<sup>11</sup> in which, although post-surgical stereopsis is maintained, a  
230 significant deterioration can be appreciated.

231 Although stereopsis is the most advanced function of the binocular visual system and its  
232 analysis is key in order to assess the suitability of an emmetropization technique, it is necessary  
233 to compare the results with those from studying more binocular functions. Binocular visual  
234 acuity is usually one of the most important variables when evaluating visual performance and is  
235 widespread in clinical and optometric practice. Our results demonstrate that binocular visual  
236 acuity under the conditions that use small-aperture contact lenses is comparable to that of the  
237 natural eye. Other clinical studies have indicated that uncorrected near visual acuity with small-  
238 aperture inlays is improved without affecting uncorrected distance visual acuity.<sup>26</sup>

239 Concerning CSF, the deterioration found in our results agree with previous results<sup>8,9</sup> that  
240 reported a reduction in the CSF and binocular summation. Lin et al.<sup>8</sup> found a minor reduction in  
241 the binocular case and claimed that this reduction was similar to other surgical presbyopia  
242 correction procedures. Other surgical procedures to correct ametropia<sup>27</sup> also showed a post-  
243 surgical deterioration in the CSF and binocular summation.

244 As indicated in the Introduction,<sup>12,13</sup> night-vision disturbances often appear after surgical  
245 procedures. One way of quantifying these is the VDI used here. We found a significant  
246 deterioration for binocular VDI (far) and VDI-binocular summation (near, intermediate, and  
247 far). Our results agree with previous ones<sup>9</sup> on inlay simulations for the intermediate distance and  
248 for other surgical emmetropization techniques that result in a post-surgical increase in the night-  
249 vision disturbances.<sup>27</sup>

250 The extensive experimental analysis undertaken in this study on binocular functions for  
251 different distances and under conditions of large anisocoria allow us to generalize that there is a  
252 binocular deterioration, including stereopsis, with respect to the conditions of the natural pupil  
253 when we simulate the use of a small-aperture corneal inlay with and without monovision.

254 Although it should be taken into account that no binocular function was found that deteriorated

255 in a generalized way for all the distances tested, depending on the visual function analysed, in  
256 some cases the significant deterioration corresponds to one distance and in other cases to  
257 another. As indicated above, this deterioration is similar to that reported in other studies using  
258 different surgical emmetropization techniques and presbyopia corrections.<sup>8,9,27</sup> It should be  
259 pointed out that the experiment conducted here simulated the optical conditions of the technique  
260 of emmetropization with and without monovision and that the surgical variables are expected to  
261 influence binocular performance. Most experimental conditions in this work were performed  
262 under illumination conditions that generate high anisocoria. Under other illumination  
263 conditions, the interocular differences in pupil size could be reduced and therefore, better visual  
264 performance could be expected. On the other hand, a corneal inlay has an outer diameter of 3.8  
265 mm and in our simulation the opaque region of the contact lens has a diameter of 11.0 mm.  
266 Under low-illumination conditions, a presbyopic patient implanted with a corneal inlay would in  
267 most cases reach a pupil size of greater than 3.8 mm. In this situation, light passes through the  
268 small aperture but also through the peripheral portion of the cornea, resulting in an opaque  
269 annulus in front of the pupil, thereby deteriorating visual performance. In addition, mesopic and  
270 scotopic pupils of the young participants could be larger compared with presbyopic patients,  
271 although, in near vision, myosis due to the accommodation in young eyes could partially  
272 compensate for this. In our work, we found pupil sizes under mesopic conditions similar to  
273 those reported by Tomita et al.<sup>28</sup> after KAMRA inlay implantation, in which the influence of  
274 pupil size on visual acuity was evaluated. These authors found no significant differences in  
275 UNVA (uncorrected near visual acuity) or in CNVA (corrected near visual acuity) under  
276 mesopic lighting conditions between the small (< 6 mm) and large (> 6 mm) pupil groups. The  
277 range of pupil sizes evaluated in the work of Tomita et al. was 6.00 to 8.32 mm. On average,  
278 these pupil sizes are larger compared with the range analysed in our study (from 3.00 to 7.00  
279 mm). Other authors have studied the effect of pupil size on visual performance in a large group  
280 of patients implanted with a small-aperture corneal inlay.<sup>29,30</sup> The results showed no impact on  
281 visual acuity and visual symptoms in photopic pupil sizes and minimal influence in mesopic  
282 pupils. Regarding larger pupils under scotopic conditions, subjects implanted with a small-

283 aperture corneal inlay showed slightly worse visual performance, although not statistically or  
284 clinically significant. These findings support the contention that the effect on visual  
285 performance of a pupil size higher than the corneal-inlay would be negligible compared with the  
286 eye wearing the small-aperture contact lens used here and the same pupil size. Apart from  
287 differences associated with the dimensions of the opaque annulus, the optical effects of the  
288 contact lens would be expected to be similar to those of the corneal inlay, since the stromal  
289 depth of the flap/pocket in which the inlay is placed measures only 0.20 mm.<sup>31</sup> Another factor to  
290 consider in the use of the small-aperture contact lens is the movement of the lens after a blink,  
291 which decentres the contact lens, worsening visual performance. In this sense, the task of  
292 discriminating or detecting visual stimulus in the tests of the present study were much longer in  
293 time than the re-centring of the contact lens after a blink, allowing the patient to perform the  
294 visual tests in an effective way, without the influence of the contact-lens movement. However,  
295 apart from the re-centring, the contact lens centration after fitting cannot be completely  
296 guaranteed compared with an intrastromal corneal inlay which has been surgically aligned.

297 An aspect to be highlighted in the present study is that the participants were young patients,  
298 whereas patients implanted with a small-aperture corneal inlay are pre-presbyopic or  
299 presbyopic. Higher interocular differences would be expected in presbyopic patients for near  
300 distance (0.4 m), due to smaller pupil sizes and reduced accommodation, deteriorating visual  
301 performance. Tomita and Waring analysed the influence of age on visual outcomes of  
302 emmetropized patients implanted with corneal-inlays,<sup>5</sup> for ages ranging from 40 to 65 years,  
303 finding on average the same UDVA (uncorrected distance visual acuity) of 20/20 for the  
304 different age groups studied, and a mean UNVA of 20/30 or 20/40, with no significant  
305 differences, although younger patients generally had better UDVA and UNVA. Regarding the  
306 CDVA (corrected distance visual acuity) and the CNVA, no significant differences were found  
307 between age groups. Nor were there significant differences in subjective symptoms between the  
308 age groups. Patients from the youngest age group of that study were able to accommodate to  
309 some degree and older patients had reduced accommodative amplitudes, as opposed to young

310 participants of our study, who had a full accommodative amplitude. In this sense, we expected  
311 slight differences in our results in stereopsis, halo perception or CSF (more deteriorated) both in  
312 normal anisocoria as well as anisocoria combined with monovision when patients had been pre-  
313 presbyopic or presbyopic. However, it should also be considered that most of the visual tests  
314 were performed under the most unfavourable lighting conditions, i.e. in a dim ambience, in  
315 order to reach the highest interocular differences and check visual functions under these  
316 conditions, and therefore under normal conditions, such as those in daily visual tasks, better  
317 results would be expected for visual performance.

318 With respect to monovision, we have checked three different conditions in participants wearing  
319 the small-aperture contact lenses: a) no add power; b) an add power of +0.75 D, and an add  
320 power of +1.25 D. These add-power values are clinically informative to analyse since the add  
321 power of +0.75D causes a defocus that is considered optimum.<sup>3</sup> Furthermore, using optical  
322 modelling of a corneal inlay in real eyes, some authors have demonstrated that the best residual  
323 defocus is within the range of -0.75 to -1.00D,<sup>15</sup> as opposed to traditional monovision, from -  
324 1.00 to -2.00D, with optimum results for -1.25 and -1.50D.<sup>32</sup> In our work the optimum add  
325 power was +0.75D, as this gave us the best results of visual function for near and intermediate  
326 distances, in agreement with other findings.<sup>3,9</sup> Whether traditional or micro-monovision, the  
327 final add power depends not only on achieving the best optical quality or visual function, but  
328 also on the need according to the patient's lifestyle or occupation. Thus it bears analysing the  
329 visual function for two different values of add power combined with the small-aperture contact  
330 lens, as previously done.<sup>9</sup> It would also have been informative to study the stereopsis of the  
331 participants under traditional monovision conditions (without contact lens, and with an add  
332 power of +1.25 or +1.50D), but because of the time required for each test, we focused the  
333 present work on analysing stereoscopic vision under a broad range of experimental conditions  
334 simulating anisocoria. Although patients implanted with a corneal inlay show continuous  
335 functional vision over a range of 3.5D,<sup>33</sup> we should mention the limitation of our patients, that  
336 is, that they were younger and had a full accommodative amplitude compared with presbyopes.

337 Our results could have been worse if the participants had been presbyopes, and the effect of the  
338 monovision could have been different in the two types of patients for the near distance.  
339 However, Tomita and Waring<sup>5</sup> found no significant differences in visual function between  
340 different age groups, one of which was formed by pre-presbyopic patients who had some  
341 accommodative amplitude, comparable, in some cases to the young participants of our study.  
342 An option to minimize the effect of accommodation had been to use cycloplegic, but we  
343 performed the visual tests under natural pupil conditions, with no dilation, in order to avoid the  
344 shift of the pupil centre, which affects ocular parameters, such as aberrations.<sup>34</sup> Furthermore,  
345 repercussions not only in ocular parameters but also in visual function have been reported in  
346 decentred small pupils,<sup>35</sup> and thus the use of cycloplegic could deteriorate visual function and  
347 optical quality, especially in the eye wearing the small-aperture contact lens.

348 Therefore, our simulation using small-aperture contact lenses, based on the same optical  
349 principle as the small-aperture corneal inlay (to increase the depth-of-focus by a pin-hole) is  
350 quite acceptable to analyse stereoscopic vision under a broad range of experimental conditions  
351 simulating anisocoria, although, given the potential limitations of this work, such as the  
352 decentration of the contact lens with respect to a surgically aligned corneal inlay, additional  
353 studies of small-aperture corneal-inlay simulations should be undertaken with presbyopic  
354 patients, older than those studied here and simulating more realistic conditions of the corneal  
355 inlay (a contact lens like the one used in this work but with an outer diameter of close to 3.8 mm  
356 and taking into consideration the effect of contact-lens decentration on visual functions).

357 Additionally, new conditions should also be taken into account as well as new designs of  
358 corneal inlay, such as diffractive corneal inlays<sup>36</sup> so as to broaden and complement existing  
359 studies on the visual performance in patients with corneal inlays.

360 We conclude that after a complete binocular analysis, including stereopsis (stereoacuity),  
361 contrast-sensitivity, visual acuity, and visual disturbance index at three different distances: near,  
362 intermediate, and far, we have shown that optical simulations of small corneal-inlay aperture  
363 (with and without monovision) show a deterioration of the binocular visual performance,

364 although this deterioration can be acceptable for patients subjected to this surgical technique  
365 since the binocular deterioration found was similar to other surgical procedures of  
366 emmetropization.

367

### 368 **References**

- 369 1. Lindstrom RL, MacRae SM, Pepose JS, Hoopes, PC. Corneal inlays for presbyopia  
370 correction. *Curr Opin Ophthalmol*. 2013;24(4):281-287.
- 371 2. Keates RH, Martines E, Tennen DG, Reich C. Small-diameter corneal inlay in presbyopic  
372 or pseudophakic patients. *J Cataract Refr Surg*. 1995;21(5):519-521.
- 373 3. Fernández EJ, Schwarz C, Prieto PM, Manzanera S, Artal P. Impact on stereo-acuity of  
374 two presbyopia correction approaches: monovision and small aperture inlay. *Biomed Opt*  
375 *Express*. 2013;4(6):822-830.
- 376 4. Taberero J, Schwarz C, Fernández EJ, Artal P. Binocular visual simulation of a corneal  
377 inlay to increase depth of focus. *Invest Ophth Vis Sci*. 2011;52(8):5273-5277.
- 378 5. Tomita M, Waring GO. One-year results of simultaneous laser in situ keratomileusis and  
379 small-aperture corneal inlay implantation for hyperopic presbyopia: Comparison by age. *J*  
380 *Cataract Refr Surg*. 2015;41(1):152-161.
- 381 6. Dexl K, Jell G, Strohmaier G, Seyeddain O, Riha W, Rueckl T, Bachernegg A, Grabner G.  
382 Long-term outcomes after monocular corneal inlay implantation for the surgical  
383 compensation of presbyopia. *J Cataract Refr Surg*. 2015;41(3):566-575.
- 384 7. Schwarz C, Manzanera S, Prieto PM, Fernández EJ, Artal P. Comparison of binocular  
385 through-focus visual acuity with monovision and a small aperture inlay. *Biomed Opt*  
386 *Express*. 2014;5(10):3355-3366.
- 387 8. Lin L, van de Pol C, Vilupuru S, Pepose JS. Contrast sensitivity in patients with  
388 emmetropic presbyopia before and after small-aperture inlay implantation. *J Refract Surg*.  
389 2016;32(6):386-393.

- 390 9. Castro JJ, Soler M, Ortiz C, Jiménez JR, Anera RG. Binocular summation and visual  
391 function with induced anisocoria and monovision. *Biomed Opt Express*. 2016;7(10):4250-  
392 4262.
- 393 10. Jiménez JR, Castro JJ, Jiménez R, Hita E. Interocular differences in higher-order  
394 aberrations on binocular visual performance. *Optom Vis Sci*. 2008;85(3):174-179.
- 395 11. Jiménez JR, Castro JJ, Hita E, Anera RG. Upper disparity limit after LASIK. *J Opt Soc Am*  
396 *A*. 2008;25(6):1227-1231.
- 397 12. Fan-Paul NI, Li J, Miller JS, Florakis GJ. Night vision disturbances after corneal refractive  
398 surgery. *Surv Ophthalmol*. 2002;47(6):533-546.
- 399 13. Klyce SD. Night vision disturbances after refractive surgery: haloes are not just for angels.  
400 *Brit J Ophthalmol*. 2007;91(8):992-993.
- 401 14. Miles W. Ocular dominance demonstrated by unconscious sighting. *J Exp Psychol*.  
402 1929;12(2):113-126.
- 403 15. Taberero J, Artal P. Optical modeling of a corneal inlay in real eyes to increase depth of  
404 focus: Optimum centration and residual defocus. *J Cataract Refr Surg*. 2012;38(2):270-  
405 277.
- 406 16. Vandermeer G, Legras R, Gicquel JJ, Pisella PJ. Quality of vision with traditional  
407 monovision versus modified monovision. *Acta Ophthalmol*. 2013;91(s252):S085.
- 408 17. Castro JJ, Jiménez JR, Ortiz C, Alarcón A, Anera RG. New testing software for  
409 quantifying discrimination capacity in subjects with ocular pathologies. *J Biomed Opt*.  
410 2011; 16(1):015001.
- 411 18. Ortiz C, Castro JJ, Alarcón A, Soler M, Anera RG. Quantifying age-related differences in  
412 visual-discrimination capacity: Drivers with and without visual impairment. *Appl Ergon*.  
413 2013;44(4):523-531.
- 414 19. Castro JJ, Ortiz C, Pozo AM, Anera RG, Soler M. A visual test based on a freeware  
415 software for quantifying and displaying night-vision disturbances: study in subjects after  
416 alcohol consumption. *Theor Biol Med Model*. 2014;2014(11):S1.



- 417 20. Castro JJ, Pozo AM, Rubiño M, Anera RG, Jiménez del Barco L. Retinal-image quality  
418 and night-vision performance after alcohol consumption. *J Ophthalmol.*  
419 2014;2014:704823.
- 420 21. Castro JJ, Jiménez JR, Hita E, Ortiz C. Influence of interocular differences in the Strehl  
421 ratio on binocular summation. *Ophthal Physiol Opt.* 2009;29(3):370-374.
- 422 22. Pardhan S, Elliott DB. Clinical measurements of binocular summation and inhibition in  
423 patients with cataract. *Clin Vision Sci.* 1991;6(5):355-359.
- 424 23. Linn SH, Skanchy DF, Quist TS, Desautels JD, Moshirfar M. Stereoacuity after small  
425 aperture corneal inlay implantation. *Clin Ophthalmol.* 2017;24(11): 233-235.
- 426 24. Levinger E, Trivizki O, Pokroy R, Levartovsky S, Sholohov G, Levinger S. Monovision  
427 surgery in myopic presbyopes: visual function and satisfaction. *Optom Vis Sci.*  
428 2013;90(10):1092-1097.
- 429 25. Durrie DS. The effect of different monovision contact lens powers on the visual function of  
430 emmetropic presbyopic patients (An American Ophthalmological Society Thesis). *Trans*  
431 *Am Ophthalmol Soc.* 2006;104:366-401.
- 432 26. Yilmaz OF, Alagoz N, Pekel G, Azman E, Aksoy EF, Cakir H, Bozkurt E, Demirok A.  
433 Intracorneal inlay to correct presbyopia: long-term results. *J Cataract Refract Surg.*  
434 2011;37(7):1275–1281.
- 435 27. Jiménez JR, Villa C, Anera RG, Gutiérrez R, and Jiménez del Barco I. Binocular visual  
436 performance after LASIK. *J Refract Surg.* 2006;22(7):679-688.
- 437 28. Tomita M, Kanamori K, Waring GO, Huseynova T. Retrospective evaluation of the  
438 influence of pupil size on visual acuity after KAMRA inlay implantation. *J Refract Surg.*  
439 2014;30(7):448-453.
- 440 29. Vilupuru S, Ling L. Effect of pupil size on visual performance of presbyopes with small-  
441 aperture corneal inlay. *Invest Ophth Vis Sci.* 2016;57(12):4872.
- 442 30. Schallhorn S. Effect of pupil size on visual performance with small-aperture corneal inlay.  
443 Abstract from the XXXIV Congress of the ESCRS (European Society of Cataract &  
444 Refractive Surgeons) held September 10-14, 2016, in Copenhagen, Denmark.

- 445 31. Atchison DA, Blazaki S, Suheimat M, Plainis S, Charman WN. Do small-aperture  
446 presbyopic corrections influence the visual field? *Ophthal Physiol Opt.* 2016;36(1):51-59.
- 447 32. Hayashi K, Yoshida M, Manabe S, Hayashi H. Optimal amount of anisometropia for  
448 pseudophakic monovision. *J Refract Surg.* 2011;27(5):332-338.
- 449 33. Vilupuru S, Lin L, Pepose JS. Comparison of contrast sensitivity and through focus in  
450 small-aperture inlay, accommodating intraocular lens, or multifocal intraocular lens  
451 subjects. *Am J Ophthalmol.* 2015;160(1):150-162.
- 452 34. Yang Y, Wu F. Technical note: comparison of the wavefront aberrations between natural  
453 and pharmacological pupil dilations. *Ophthal Physiol Opt.* 2007;27(2):220-223.
- 454 35. Artal P, Marcos S, Iglesias I, Green DG. Optical modulation transfer and contrast  
455 sensitivity with decentered small pupils in the human eye. *Vision Res.* 1996;36(22):3575-  
456 3586.
- 457 36. Furlan WD, García-Delpech S, Udaondo P, Remón L, Ferrando V, Monsoriu JA.  
458 Diffractive corneal inlay for presbyopia. *J Biophotonics.* 2017;10(9):1110-1114.
- 459
- 460
- 461
- 462
- 463
- 464
- 465
- 466
- 467
- 468

469

470

471

472

473

474

475 **Tables**

476 Table 1. Average stereoacuity (arcsec) for the distances and conditions tested. \*Significantly  
 477 different from the other three conditions ( $p < 0.05$ ); §significantly different from small-  
 478 aperture contact-lens monovision +1.25D ( $p < 0.05$ ).

Distance (m)	Natural pupil	Natural pupil DE/1.6-mm small-aperture contact lens NDE		
		No Add	Add +0.75D	Add +1.25D
0.4	22.55 ± 9.24*	48.90 ± 25.37	38.40 ± 13.00	34.40 ± 13.26
1.0	58.00 ± 23.94*	152.00 ± 124.44	152.00 ± 110.84	168.00 ± 112.43
5.5	46.00 ± 30.62§	65.00 ± 32.40	72.00 ± 35.21	101.00 ± 69.35

479

480

481

482

483

484

485

486

487

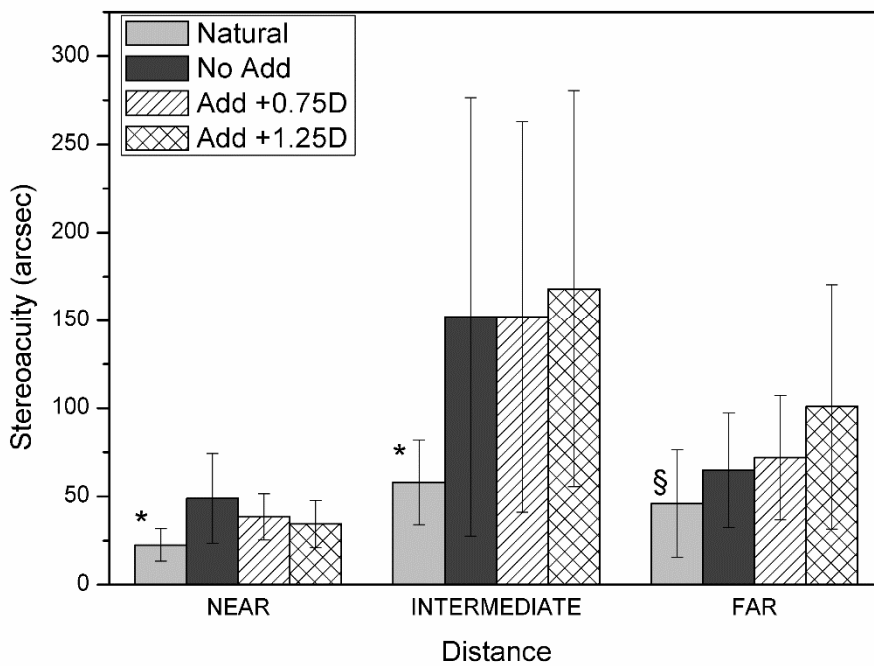
488

489

490

491

492 **Figures**



493

494 Figure 1. Average stereoacuity (arcsec) for natural conditions and wearing the small-aperture

495 contact lens in the NDE (with no add power and with add powers of +0.75 and +1.25D). The

496 results are shown for the three distance conditions tested: near (0.4 m), intermediate (1.0 m),

497 and far (5.5 m) vision. \*Significantly different from other three conditions ( $p < 0.05$ );

498 §significantly different from small-aperture contact-lens monovision +1.25D ( $p < 0.05$ ).

499

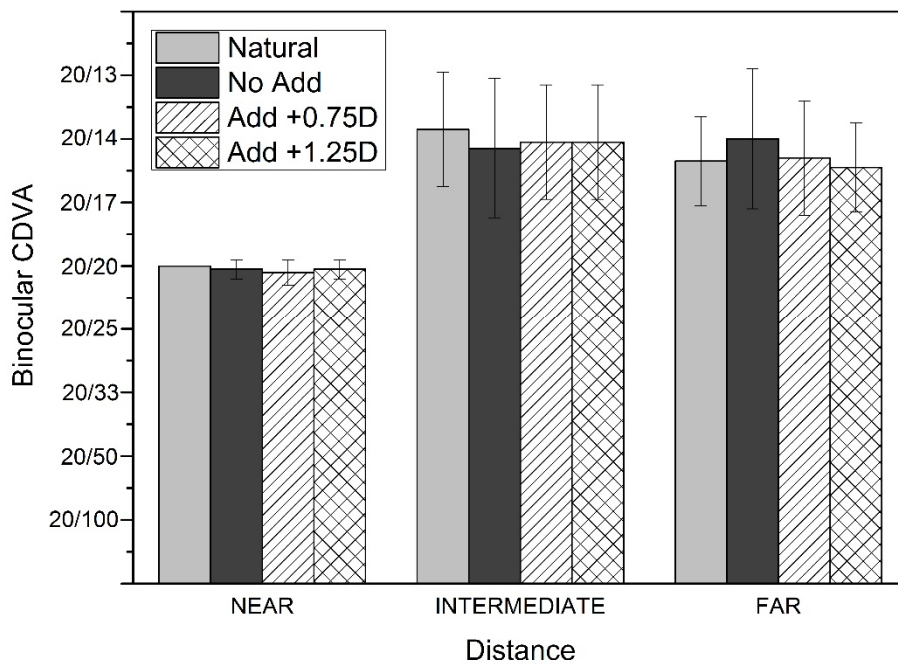
500

501

502

503

504



505

506 Figure 2. Average binocular CDVA for the distances and conditions tested. No significant

507 differences for any of the conditions at the three distances (near, intermediate, far).

508

509

510

511

512

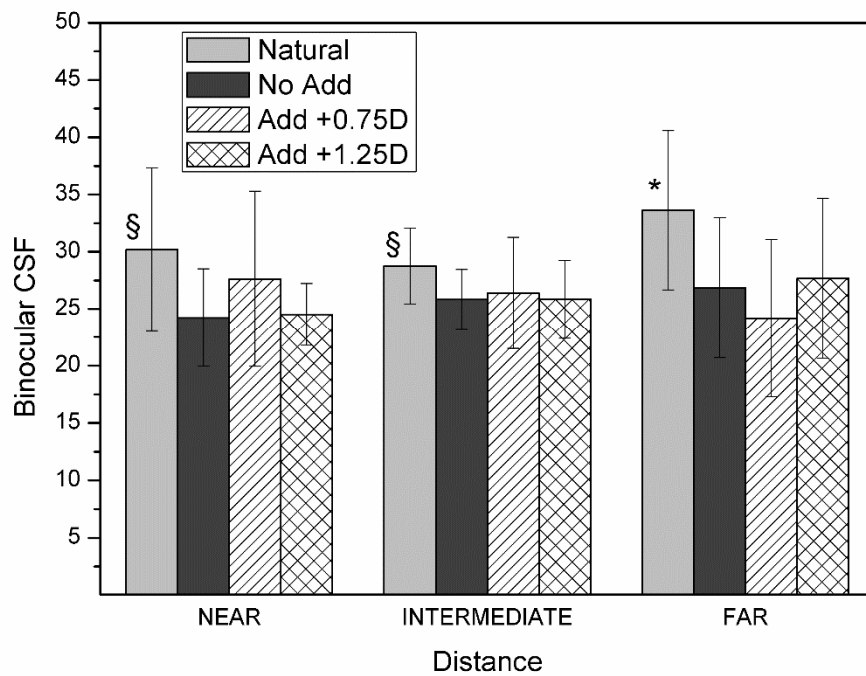
513

514

515

516

517



518

519 Figure 3. Average binocular CSF for all the conditions and distances tested. \*Significantly  
520 different from other three conditions ( $p < 0.05$ ); §significantly different from no add and  
521 monovision +1.25D conditions ( $p < 0.05$ ).

522

523

524

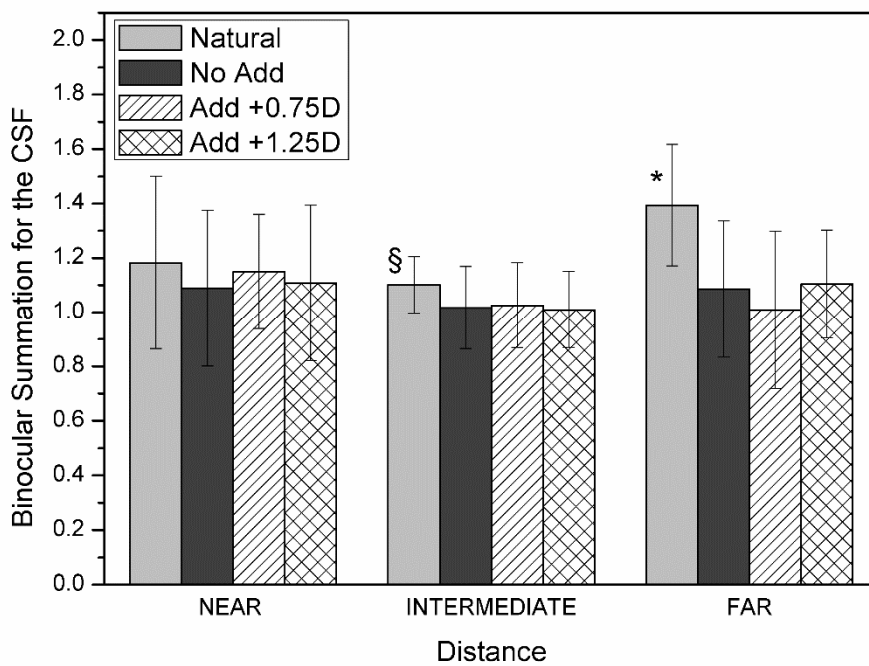
525

526

527

528

529



530

531 Figure 4. Average binocular summation for the CSF under all the conditions and distances

532 tested. \*Significantly different from other three conditions ( $p < 0.05$ ); §significantly

533 different from no add condition ( $p < 0.05$ ).

534

535

536

537

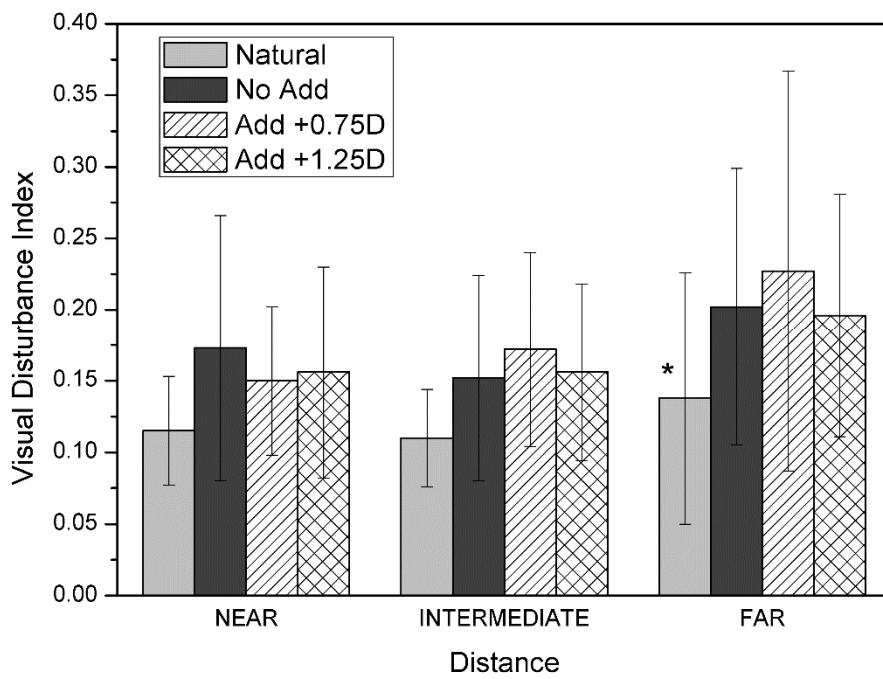
538

539

540

541

542



543

544 Figure 5. Average binocular visual disturbance index for all the conditions and distances tested.

545

\*Significantly different from other three conditions ( $p < 0.05$ ).

546

547

548

549

550



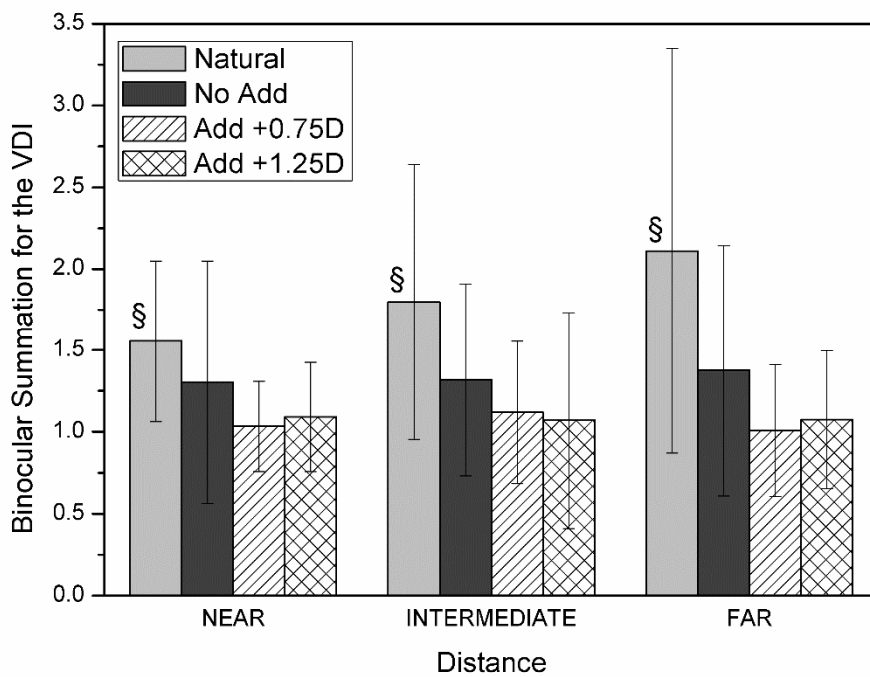
551

552

553

554

555



556

557 Figure 6. Average binocular summation for the visual disturbance index under all the  
558 conditions and distances tested. §Significantly different from small-aperture contact lens  
559 monovision +0.75D and +1.25D conditions ( $p < 0.05$ ).

560