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Successful treatment of diplopia using prism correction combined with vision therapy/orthoptics improves health-related quality of life

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Background To track improvement in diplopia symptoms with strabismus-specific health-related quality of life (HRQOL) questionnaire across a treatment consisting of prism correction followed by vision therapy/orthoptics where the former was unsuccessful.

Methods. Forty-eight participants with diplopia (mean age = 62.45) completed an Adult Strabismus-20 (AS-20) questionnaire and a Diplopia Questionnaire (DQ) before and with prism correction. Inclusion criteria before prism diplopia was "sometimes" or worse for reading and/or straight-ahead distance. The success with prism treatment was clasified as diplopia as "never" or "rarely" perceive diplopia in the items for reading and straight-ahead distance in the DQ. The failure with prism correction was determine when diplopia worsened or remained the same. In any case (success or failure), average of initial AS-20 scores werw compared with score after prism correction, taking account the AS-20 subscales (reading and general functions, and self-perception and interaction). Participants in the failure subsequently underwent a vision therapy/orthoptics programme wearing their prism correction. The analysis of the treatment success or failure was again determined using the AS-20 questionnaire (before and after vision therapy/orthoptics).

Results. Forty-three participants completed the questionnaire at the follow-up visit. Prism correction was successful for 22 patients, and failed for 21. Those participants for whom the prism correction was classified as a success showed a statistically significant improvement (P=0.01) in both reading and general functions. In the failure group, there was no significant change in AS-20 score in any of the domains (P=0.1). Following vision therapy/orthoptics treatment, 13 of the 21 participants achieved binocular vision. This improvement was transferred to a statistically significant improvement in reading and general functions (P=0.01).

Conclusions. Vision therapy/orthoptics may be an option to achieve an improvement in diplopia symptoms where prism correction has proved unsuccessful. In the 81% of the patients prism correction and adicional visual therapy is correlated with enhance in strabismus-specific HRQOL, mainly in general functions and reading.

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4 5

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19

20 Abstract

21

22 Background

- 23 To track improvement in diplopia symptoms with strabismus-specific health-related quality of life
- 24 (HRQOL) questionnaire across a treatment consisting of prism correction followed by vision
- 25 therapy/orthoptics where the former was unsuccessful.

26

- 27 **Methods.** Forty-eight participants with diplopia (mean age = 62.45) completed an Adult Strabismus-20
- 28 (AS-20) questionnaire and a Diplopia Questionnaire (DQ) before and with prism correction. Inclusion
- 29 criteria before prism diplopia was "sometimes" or worse for reading and/or straight-ahead distance. The
- 30 success with prism treatment was clasified as diplopia as "never" or "rarely" perceive diplopia in the
- 31 items for reading and straight-ahead distance in the DQ. The failure with prism correction was determine
- 32 when diplopia worsened or remained the same. In any case (success or failure), average of initial AS-20
- 33 scores werw compared with score after prism correction, taking account the AS-20 subscales (reading and
- 34 general functions, and self-perception and interaction). Participants in the failure subsequently underwent
- 35 a vision therapy/orthoptics programme wearing their prism correction. The analysis of the treatment





36	success or failure was again determined using the AS-20 questionnaire (before and after vision
37	therapy/orthoptics).
38	
39	Results. Forty-three participants completed the questionnaire at the follow-up visit. Prism correction was
40	successful for 22 patients, and failed for 21. Those participants for whom the prism correction was
11	classified as a success showed a statistically significant improvement (P=0.01) in both reading and
12	general functions. In the failure group, there was no significant change in AS-20 score in any of the
1 3	domains (P=0.1). Following vision therapy/orthoptics treatment, 13 of the 21 participants achieved
14	binocular vision. This improvement was transferred to a statistically significant improvement in reading
1 5	and general functions (P=0.01).
1 6	
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48	symptoms where prism correction has proved unsuccessful. In the 81% of the patients prism correction
19	and adicional visual therapy is correlated with enhance in strabismus-specific HRQOL, mainly in general
50	functions and reading.
51	
52	Introduction
53	Globally, the prevalence of strabismus ranges from 2 to 4%. Diplopia in adulthood is associated with
54	strabismus ² deriving from different aetiologies: decompensation of previous deviations (foria or tropia); ^{3,4}
55	paresis or paralysis of extraocular muscles, whether vascular, ^{5,6} tumour-related ⁷ or secondary to brain
56	trauma;8 autoimmune diseases, such as Graves syndrome; or problems secondary to retinal diseases, such
57	as dragged-fovea diplopia syndrome.9
58	
59	The impact of strabismus on health-related quality of life (HRQOL) in adults can be assessed using different
30	questionnaires such as the specific Adult Strabismus-20 questionnaire (AS-20),10 or the more general
31	American National Eye Institute Visual Functions Questionnaire (VFQ-25). ¹¹ Predictably, the AS-20 is
62	more sensitive than the VFQ-25 for detecting reduced HRQOL in adult strabismus. 12 A previous evaluation
63	of the psychometric properties of the AS-20 with Rasch analysis proposed the reduction of the questionnaire
64	to four distinct domains: self-perception (5 items), interactions (5 items), reading function (4 items), and
65	general function (4 items). ¹³
66	
67	The impact of diplopia on quality of life has been analysed by different authors. ¹⁴ Holmes et al. designed
86	the Diplopia Questionnaire (DQ) specifically to quantify diplopia, ¹⁵ allowing the position and distance at
69	which the patient experiences diplopia to be recorded. Researchers have proven a high correlation between



70	the functional results of the AS-20 and measurements provided by the DQ in patients with diplopia in
71	primary gaze and reading.
72	
73	Regarding the effects of different strabismus treatments on HRQOL, a number of studies propose surgery
74	as an option associated with significant and enduring functional and psychological improvement in patients
75	with strabismus. 16,17,18 Moreover, all the evidence seems to indicate that these improvements in HRQOL
76	are greater in diplopia patients than those without diplopia. ¹⁹
77	Hatt et al. ²⁰ studied the effects of prism correction in participants with binocular diplopia, achieving a 68%
78	success rate measured with the DQ in participants with diplopia rarely or never perceived in primary gaze
79	and reading. Improvement was also recorded in the reading and general functions subcategories of the AS-
80	20, but did not extend to either the self-perception or interactions categories. ²⁰
81	
82	Vision therapy/orthoptics) has proven its effectiveness in improving vergence ranges in patients with
83	esotropia ²¹ and convergence insufficiency; ²² to the best of our knowledge, however, the effects on HRQOL
84	of this treatment have never been evaluated. The aim of the present study is to assess the impact on the
85	HRQOL of adults with diplopia of a protocol that involves prism correction and vision therapy/orthoptics
86	as required.
87	
88	Materials & Methods
89	Design
90	This is a prospective case series study. The participants were adults (>18 years old) with acquired dip on a,
91	and all were patients at the Ikusgune Optometric Center and Begira Ophthalmologic Clinic (Basque
92	Country, Spain). Patients with severe ar plopia (BCVA <0.2), monocular diplopia, nystagmus, and/or
93	mental or cognitive impairments that would rule out the use of a HRQOL assessment, were excluded.
94	The study followed the tenets of the Declaration of Helsinki and was approved by the Basque Country
95	Ethics Committee of Clinical Research (PI2021059). The participants signed a consent form after receiving
96	a verbal and written explanation of the study.
97	
98	Evaluation protocol
99	HRQOL assessments
100	The DQ provides a self-evaluation of diplopia severity on a five-point scale (never, rarely, sometimes,
101	often, always) in seven gaze positions (reading, straight-ahead distance, right gaze, left gaze, up gaze, down
102	gaze, and any other gaze position). According to the authors' criteria, 15 where participants rate diplopia
103	perceived in reading and straight-ahead distance gaze positions as "sometimes", "often" or "always", this



104 should be included in the analysis. Prism correction success was defined as diplopia rated on the DO as 105 "never" or "rarely" for reading and straight-ahead distance. The validated version of the AS-20 106 questionnaire was also used.¹³ Each of the four dometric was scored independently and finally consolidated 107 into a unique 0 to 100 score (worst to best HRQOL) to facilitate interpretation. Both the AS-20 108 questionnaire and the DQ were taken before and after prism correction, and at the end of the vision 109 therapy/orthoptics where prism correction was rated unsuccessful.

110 111

Clinical evaluation

112 The participants were evaluated by an experienced optometrist. Refractive error was corrected, and best 113 corrected visual acuity (BCVA) was obtained using the HOTV visual acuity chart with crowding bars 114 (Smart4Vision, Spain). Binocular vision was analysed using the Worth Four Dot test at a distance of four 115 metres with scotopic illumination, and the Random Dot Preschool Stereoacuity test (Stereo Optical Company Inc., United States) at near distance. Deviation angle was measured using two different 116 117 procedures. The first of these was a cover test with an accommodative stimulus, at near and far distances, 118 placing the prism over the strabismic eye, using stimuli based on characters two lines below the participant's 119 BCVA. Where the strabismus combined horizontal and vertical deviations, the primary deviation was 120 corrected first (e.g., the vertical deviation in 4th cranial nerve palsy). The second procedure involved the use of a synoptophore. Subjective and objective deviations were evaluated using a traditional synoptophore 122 (Oculus, Germany), and a modern version based on virtual reality glasses with an eye tracker (Visionary VR, 123 Visionary Tool S.L., Spain). Deviations in cyclotorsion were measured with both synoptophore devices in

The clinical evaluation included an ocular health exam (using tropicamide for pupil dilation),

biomicroscopy, an indirect ophthalmoscope, and optical coherence tomography (i to model and

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Treatment protocol

manufacturer).

- 130 Treatment lasted from 1 to 4 months, with two well-differentiated phases (Figure 1).
- 131 Phase I. Prism correct 📶
- 132 Fresnel prisms were prescribed at the first visit. Where the deviation was lower than 6 prism diopters (D),
- 133 the prism was located over the participant's dominant eye. Where the deviation was higher than 6 D, the
- 134 value was divided between two prisms, with the prism used in front of the dominant eye having twice the
- 135 power of that used over the non-dominant eye.

addition to a Double Maddox rod test.²³

- After one months at the follow-up visit the participant was required to complete both AS-20 and DQ 136
- 137 questionnaires again. Where the participant rated the prism correction as successful (diplopia rated "never"





or "rarely" on the DQ for reading and straight-ahead distance), new glasses with the prism correction were prescr. Where the prism correction was rated as unsuccessful (diplopia rated "sometimes" or "always"), the participant was included in phase II.

Phase II. Vision therapy/orth op ics

Vision therapy/orthoptics included exercises sessions at the centre and at home. The exercise sessions at the centre lasted for 2 months, with visits scheduled every 15 days (4 sessions in total), and used traditional orthoptics materials and instruments such as a synoptophore, analyphs and vectograms, and a Brock string with and without prisms. Gamified training exercises using the previously cited virtual reality synoptophore were also used at the clinic (VisionaryVR, VisionaryTool S.L., Spain). Exercises at home were prescribed at the same time, also with a 2-month duration, consisting of 20 minutes per day, 5 days per week, of game play involving computerised vergence exercises with analyphs (Figure 2). Two similar programs were used: Vision Builder Version 2.7 for Windows (Haraldseth Software, Norway) and VisionaryTool (VisionaryTool S.L., Spain) (Figure 2).

Data analysis

Following Holmes et al.'s criteria, 15 prism treatment success occurs when diplopia is rated "never" or "rarely" in both reading and straight-ahead distance gaze positions. The mean with standard deviation of the AS-20 and DQ scores were calculated for the four domains before and after prism correction, and after vision therapy/orthoptics treatment where prism correction was rated a failure. The participants completed the questionnaires during the first visit, before prism correction or intervention of any kind. After prism correction, data were taken at the follow-up visit (after 1 month), irrespective of whether the prism power was charged. AS-20 questionnaire and DQ scores before/after prism correction and before/after vision therapy/orthoptics were compared using a Wilcoxon signed rank test, for the whole group and in sub-groups of success/failure, and according to the prism value. In addition, a chi-square test was performed to determine whether the success number was significant. Baseline differences between patients were also analysed, considering success or failure in both prism correction and vision therapy/orthoptics. Cronbach's alpha coefficient was used to evaluate the reliability of the scales used, despite the questionnaires having been previously validated for this purpose.

Results

A total of 48 adults (21 women and 27 men) with diplopia were recruited for the study, with a mean age of 62.45 ± 16.00 years old (within the age range 26 to 86 years). The baseline variables can be consulted in annexed 1 and 2. Eleven of the participants had received previous treatment to correct the diplopia: five of



172	these had been treated with botulinum toxin, three had undergone strabismus surgery, and another three had
173	been treated with botulinum toxin followed by strabismus surgery.
174	
175	The spheric refraction equivalent was -1.12 ± 4.29 D (-22.00 to 4.25 D range) for the right eye and -0.94 \pm
176	4.28 D (-22.00 to 3.50 D range) for the left. Presbyopia was compensated in 37 subjects with a mean
177	addition of 1.90 ± 1.10 D (1.50 to 3.00 D range). The best corrected visual acuity media in decimal acuity
178	was 0.92 ± 0.16 (0.40 to 1.00 range) for the right eye and 0.92 ± 0.15 for the left (0.40 to 1.00 range). A
179	Worth Four Dot test recorded results of diplopia in far vision in all participants except two, who achieved
180	sensorial fusion (convergence insufficiency). Of the 48 participants, 32 (66.67%) patients were diagnosed
181	as stereo blind in near vision; mean values were 914.60 ± 559.92 " (arc seconds) within a 40" to 1300"
182	range.
183	
184	Thirty-four of the 48 participants showed paretic deviation, with paresis of the 4th (17) and 6th (17) cranial
185	nerves; eight had a decompensated deviation (overaction of the superior oblique muscle or exotropia
186	following esotropia surgery); four exhibited restrictive symptoms (scleral buckling for retinal detachment,
187	myopic myopathy or thyroid surgery); and two had convergence insufficiency.
188	
189	The left-right strabismus distribution was 22 participants (45.83%) with strabismus of the right eye, and 26
190	(54.17%) with strabismus of the left. Twenty participants (41.67%) had a pure horizontal deviation, (with
191	no additional vertical deviation), 17 (35.42%) had an isolated vertical deviation, and 11 (22.91%) presented
192	a mixed deviation.
193	The mean value of the horizontal deviation was 5.40 ± 6.58 PD (4 to 30 PD range) at far, and 4.26 ± 8.57
194	PD (4 to 30 PD range) at near distance. The vertical deviation mean value was 3.75 ± 4.55 PD (4 to 20 PD
195	range) at far, and 3.85 ± 4.66 PD (2 PD to 20 PD range) at near distance. Excyclotropia deviation was 1.55
196	\pm 2.76 degrees (2 to 10 degree range).
197	
198	Initial visit and pre-prism correction results
199	The DQ results obtained prior to prism prescription (rated from 0 to 100, worst to best) indicated a mean
200	of 64.01 ± 24.46 (22.50 to 100 rate range). Forty-five participants showed diplopia at far and reading
201	distance; three experienced diplopia when reading only. Of these, perceived diplopia was reported when
202	looking to the right (30 participants), to the left (34), upwards (26), downwards (30), and in intermedial
203	positions (29).
204	



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205 Globally averages AS-20 scores before to prism correction were 51.39 ± 22.52 points in the general function 206 domain, 53.50 ± 31.80 points in the reading function domain, and 89.30 ± 15.74 points and 86.31 ± 21.24 207 in the self-perception and interaction domains, respectively (Figure 3). The mean prism correction values 208 were 7.73 ± 5.80 PD (4 to 30 PD range) at far, and 6.92 ± 7.05 PD (0 to 30 PD range) at near distance. 209 210 **Post-prism treatment overall results** 211 Five of the 48 participants did not complete the DQ at the end of the prism correction period (three dropped 212 out of the study, one died, and one declined to wear the prism for work reasons but did attend the vision 213 therapy protocol). Based on a reso i definitions of success and failure based on participants' responses to 214 the DQ, 22 of 43 participants (51%) were classified as prism correction successes and 21 (49%) as prism 215 correction failes, (P > 0.1, according to chi-square analysis). 216 217 In all participants, mean AS-20 scores were statistically significant improved after prism correction 218 compared to scores obtained prior to prism correction in both general function and reading function 219 domains, improving from 51.39 ± 22.52 to 68.48 ± 24.12 , and from 53.50 ± 31.80 to 68.66 ± 32.55 , 220 respectively (P < 0.01 in both). No significant changes in scores (P > 0.1 in both) were recorded in the self-221 perception and interaction domains (Figure 3). 222 Where the prism treatment was rated successful, new glasses with the prism correction were prescribed. At 223 far distance, 5 subjects did not require a prism, while the mean prism power for the remaining subjects was 224 6.95 ± 6.13 PD, within a 2 to 30 PD range. At reading distance, 11 subjects did not require a prism, while 225 the remainder were prescribed a mean prism power of 6.40 ± 7.66 PD, within a 3 to 30 PD range. 226 Before prism treatment, binocular vision tested with the Worth Four Dot test indicated 46 participants with di ia and 32 with null stereoacuity. Following prism treatment, Worth Four Dot test results f is 18 227 228 participants (37.5%) with null stereoacuity (mean scores 673.49 ± 567.88 "; P = 0.01), and 18 patients with 229 diplopia (mean scores 4.42 ± 0.50 ; P = 0.01). 230 231 Post-prism results: success or failure The DQ results pre- and post-prism treatment indicated no significant differences in either the success or 232 233 failure group (P > 0.05). 234 235 For the 22 participants for whom the prism treatment was a success, mean AS-20 scores significantly

For the 22 participants for whom the prism treatment was a success, mean AS-20 scores significantly improved in the reading function domain, from 65.99 ± 32.39 to 86.45 ± 23.24 points after prism correction (P = 0.01); and in the general function domain, from 55.68 ± 22.96 to 83.32 ± 17.37 after prism correction



- 238 (P < 0.01). No significant differences were found in either the self-perception (P = 0.17) or interaction
- domains (P = 0.50), as shown in Figure 3.
- 240 In the 21 participants in whom the prism treatment was not effective, the mean AS-20 scores showed no
- significant improvement in any domain (P = 1.00). These participants were older, had a greater angle of
- deviation, and worse binocular vision (Table 1); they also obtained a higher score on the DQ and more
- 243 symptoms on the AS-20 questionnaire (Table 2).

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Post-vision therapy/orthoptics overall results

- 246 Phase II (Vision therapy/orthoptics) commenced with 22 participants: 21 from the prism treatment failure
- 247 group plus 1 who declined to wear the prism correction; one participant dropped out. The treatment
- succeeded in 13 participants (62%) and failed in 8 (38%). The number of participants that experienced an
- 249 improvement was not significant (P = 0.27, using a chi-square test). The DQ results pre- and post-vision
- 250 therapy exhibit significant e differences (P = 0.05).

251

- Mean AS-20 scores significantly improved in the reading function domain, from 51.71 ± 29.19 to $72.78 \pm$
- 253 27.94 points after undergoing the vision therapy/orthoptics programme (P = 0.00); and in the general
- function domain, from 52.95 ± 20.07 to 67.54 ± 22.40 points after vision therapy/orthoptics (P < 0.01). No
- significant differences were recorded in either self-perception (P = 0.31) or interaction domains (P = 0.21),
- as shown in Figure 4.

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- 258 Prior to vision therapy/orthoptics treatment, binocular vision tests showed 18 participants with dir and
- 259 null stereoacuity. After vision therapy/orthoptics, only 5 participants exhibited with diplopia with the Worth
- Four Dot test (mean scores: 4.12 ± 0.32 , P < 0.01), and all of them also exhibited null stereoacuity (mean
- 261 scores: 427.90 ± 432.87 "; P < 0.01).

262

- After vision therapy/orthoptics, the prism power needed to achieve binocular vision was significantly lower
- at far (mean scores: 3.07 ± 5.80 PD; P < 0.01) and near distance (mean scores: 2.93 ± 5.66 PD, P < 0.01).

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Post-vision therapy/orthoptics results: success or failure

- For the 13 patients successfully treated with vision therapy/orthoptics, mean AS-20 scores significantly
- 268 improved after undergoing the vision therapy/orthoptics programme in the general function and reading
- function domains. General function scores improved from 54.74 ± 21.45 points post-prism correction to 73.21
- \pm 20.36 points subsequent to undergoing the vision therapy/orthoptics programme (P = 0.02). Reading
- function domain scores improved from 57.79 ± 29.05 points post-prism to 85.09 ± 19.97 points post-vision



therapy/orthoptics (P = 0.01). No significant difference was found between pre- and post-vision therapy/orthoptics scores in the self-perception and interaction domains (P = 0.77 and P = 0.35, respectively), as shown in Figure 4.

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For the 8 patients for whom vision therapy treatment was unsuccessful, mean AS-20 scores improved significantly in the reading function domain, from 41.83 ± 28.42 points post-prism correction to 52.77 ± 28.43 points post-vision therapy/orthoptics (P = 0.04). In the general function, self-perception and interaction domains, no difference was found between pre- and post-vision therapy/orthoptics scores (P = 1.00) for both domains, as shown in Figure 4.

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Summary

- 283 The proposed treatment prism correction followed by vision therapy/orthoptics as required was
- successful in $\frac{1}{2}$ articipants, or 81% of the study sample (P < 0.01, using a chi-square test). In addition,
- 285 the participants showed significant improvement in their binocular vision tested with the Worth Four Dot
- 286 test, from 4.95 ± 0.20 to 4.12 ± 0.32 (P < 0.01); and in stereoacuity, from 914.60 ± 559.92 " to $427.90 \pm$
- 287 432.87" (P < 0.01).

288 289

Discussion

In this study, prism correction alone solved the diplopia in 51% of patients. Where the prism correction was instruction, the orthoptic/vision therapy prescribed resolved the diplopia in 62% of patients. Overall, the proposed treatment of prism correction followed by vision therapy/orthoptics as required was successful in 81% of patients.

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Participants successfully treated with prisms improved in both the general function and reading function domains. These results are congruent with those obtained by Hatt et al.,²⁰ underlining the importance of prisms in the treatment of diplopia. Interestingly, those participants that failed to obtain stable fusion at far distance or when reading, according to DQ results, showed no significant improvement in either general or reading function domains. This result obtained in our study was also pointed out by Hatt et al.²⁰

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Those participants that continued with the vision therapy/orthoptics treatment also improved in both the general and reading function domains. Again, as with the prism correction, participants that showed no improvement on the DQ recorded no improvement on the AS-20 either. The lack of AS-20 improvement in participants classified as prism or vision therapy/orthoptics treatment failures suggests that the improvement recorded for the successfully treated participants was not attributable to a placebo effect.





Very few studies have studied the prism correction effect in adult subjects with diplopia.^{25–27} Tamhankar and Ying²⁸ is a retrospective study in subjects with 4th cranial nerve palsy, classifing the prism treatment results into three categories (totally satisfied, mostly satisfied and dissatisfied). The analysis performed in this study, using a specific DQ and analysing the transference with the AS-20 PQOL questionnaire, represents a step forward.

To the best of the author's knowledge, no previous study has analysed the impact of vision therapy/orthoptics treatment on the HRQOL of subjects with diplopia. Previous studies have evaluated how an improvement in vergence response transfers to diministrate reading skills in subjects with convergence insufficiency.²⁹ Therefore, it makes sense that improvements in reading function should be obtained alongside improved binocular vision in diplopia subjects.

Baseline clinical differences between the success and failure groups were also analysed. Participants with several diplopia, higher HRQOL symptomatology, worse binocular vision and higher deviation angle showed the poorest results after prism treatment. This result was not observed in Hatt et al., ²⁰ perhaps due to the fact that the present study included participants for whom surgery did not resolve the issue, with baseline deviations of > 10 PD. Since the higher the prism power, the greater the adaptation properties in the present study were more likely to fail prism treatment in isolation. Following orthoptic/vision therapy, those participants needed lower-power prisms, facilitating adaptation and raising the likelihood of the final success of the treatment.

Where prism correction was insufficient, vision therapy/orthoptics resolved the problem in 62% of participants. There was a clear transference towards improved HRQOL, as in the case of prism success. Most of these participants had a history of strabismic surgery and/or botulinum toxin, hence prism treatment and orthoptic/vision therapy was the only remaining therapeutic option.

One participant declined prism treatment for work reasons, but opted to try vision therapy/orthoptics treatment, with a successful result. In future studies, it would be interesting to compare results obtained with only orthoptic/vision therapy versus prism treatment, particularly where refraction is not tolerated (e.g., severe anisometropia).³⁰

Previous studies have demonstrated the effectiveness of vision therapy/orthoptics in convergence insufficiency,³¹ but not, to the author's knowledge, in the case of diplopia. The present study sample included two participants with convergence insufficiency, but only one improved; importantly, the participant that showed no improvement had another condition that may have affected his prognostic:



340	Parkinson's disease. The prevalence of Parkinson's in adults with diplopia is high (18.1%). ³² Another
341	participant with Parkinson's did achieve successful fusion using the prism. The success of the proposed
342	treatment in Parkinson's sufferers is therefore deserving of future study.
343	
344	The present study sample included participants with systemic diseases that increase the risk of depression.
345	One of these participants, recently diagnosed with CREST syndrome, dropped out of the study. Hatt et al.
346	(IOVS 2013;54: ARVO E-Abstract 5987) also describes how depression reduces self-perception of
347	HRQOL in subjects with diplopia.
348	
349	The use of new technologies and gamification strategies may contribute to increase motivation and
350	compliance. Where new technologies are available both at the treatment centre and at home, with remote
351	compliance and performance tracking, adherence to the treatment is increased, as previous studies have
352	shown in cases of convergence insufficiency. ²²
353	
354	This was a prospective study with 48 participants, making it reliable, but the authors are aware of a number
355	of limitations. Firstly, the inclusion of a control group was opposed by the ethical committee in defense of
356	research participants' interests. Nevertheless, placebo effects are unlikely to have occurred, since those
357	participants who perceived diplopia after treatment did not report any improvement in HRQOL self-
358	perception.
359	
360	Secondly, the sample was heterogeneous. Most of the participants showed 4th and 6th cranial nerve palsy,
361	while others exhibited different types of strabismus, such as convergence insufficiency in the geriatric
362	population or childhood-onset esotropia. Childhood-onset esotropia usually exhibits high deviation angles,
363	even sensory adaptations, that make treatment and prognosis challenging, and this may have had a statistical
364	effect on this study.
365	Finally, an assessment of the stability of the improvements – six months after the end of the treatment, for
366	example – would have been desirable.
367 368	Conclusions
369	Effective prism correction of diplopia is correlated with enhance in function-related HRQOL. Where prism
370	correction fails, successful vision therapy/orthoptics treatment also improves function-related HRQOL.
371	Prism correction together with vision therapy/orthoptics as needed offers a valuable non-surgical treatment
372	option that could be particularly helpful in strabismus with small- and medium-angle.
373	Acknowledgements
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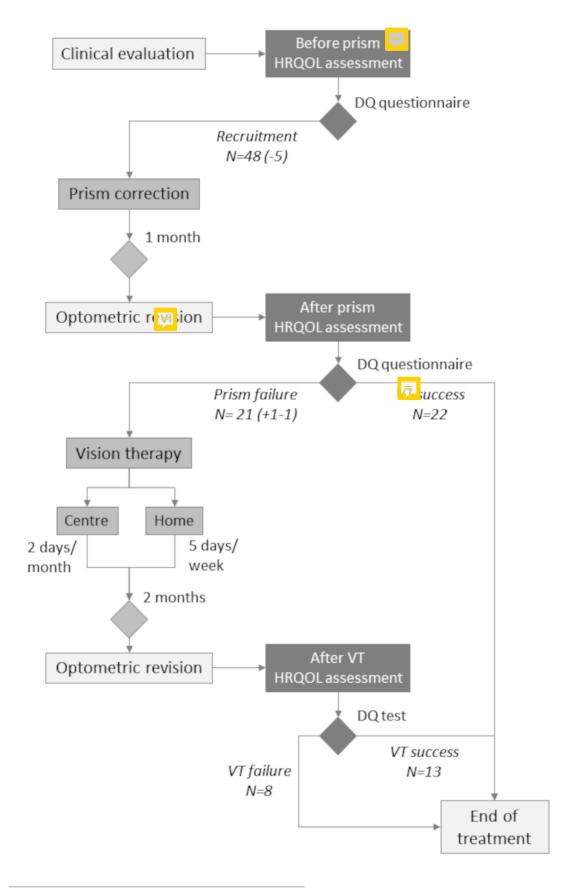
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Flowchart of Methods section

Flowchart of proposed treatment of prism correction followed by orthoptic/vision therapy. The number of volunteers at each step includes the number of abandons or new inclusions in parentheses.

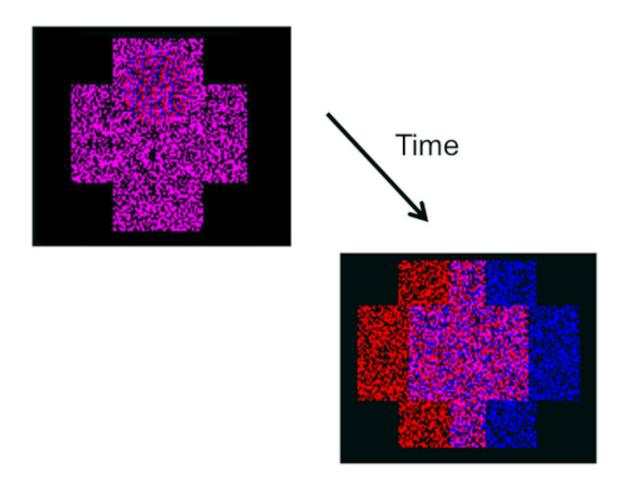






graphical example of the gamified visual therapy.

Logical process of the game . The participant's task is to locate a ball situated at one of the four extremes of a cross (top, bottom, right or left). The program automatically adjusts the vergence difficulty during the training session. Where the participant responds correctly, the software will split the image into two anaglyph crosses which the participant must then merge using their own vergence system. Should the participant fail to respond correctly, the two crosses are merged by the program.

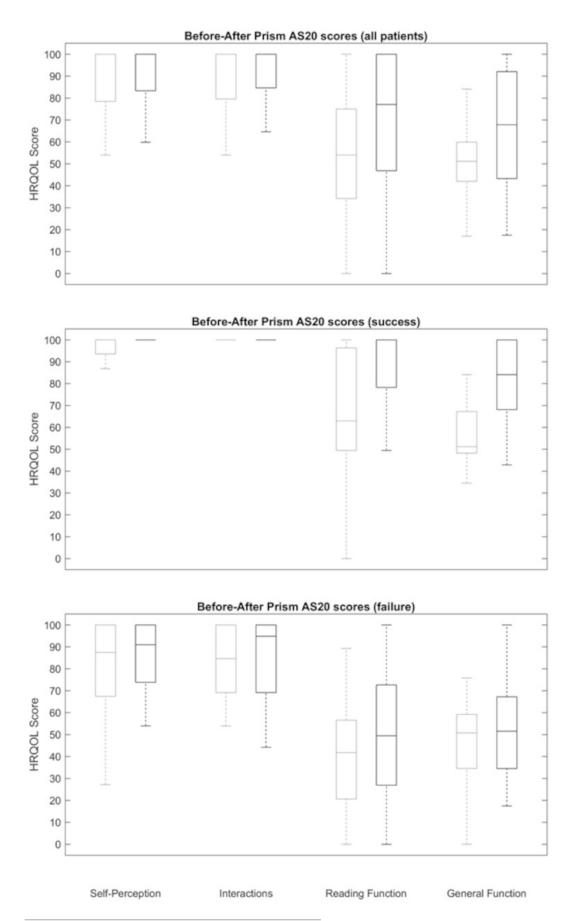




Outcomes after prisma correction

Box and whisker plots of Adult Strabismus-20 Health-Related Quality of Life scores in participants with diplopia treated with prism. Clear line boxes show pre-prism scores and shaded line boxes show scores in prism correction. The centre line represents the median, lower and upper quartiles; the whiskers represent the extremes. Top (A): All participants; Centre (B): Successfully treated participants only; Bottom (C): Participants who failed prism treatment.





Outcomes after visual therapy treatment

Box and whisker plots illustrate Adult Strabismus-20 Health-Related Quality of Life scores in participants with diplopia treated with orthoptic/vision therapy. Clear line boxes show pre-orthoptic/vision therapy scores and shaded line boxes show scores after orthoptic/vision therapy. The centre line represents the median, lower and upper quartiles; the whiskers represent the extremes. Top (A): All patients; Centre (B): Successfully treated participants only; Bottom (C): Participants who failed prism treatment.



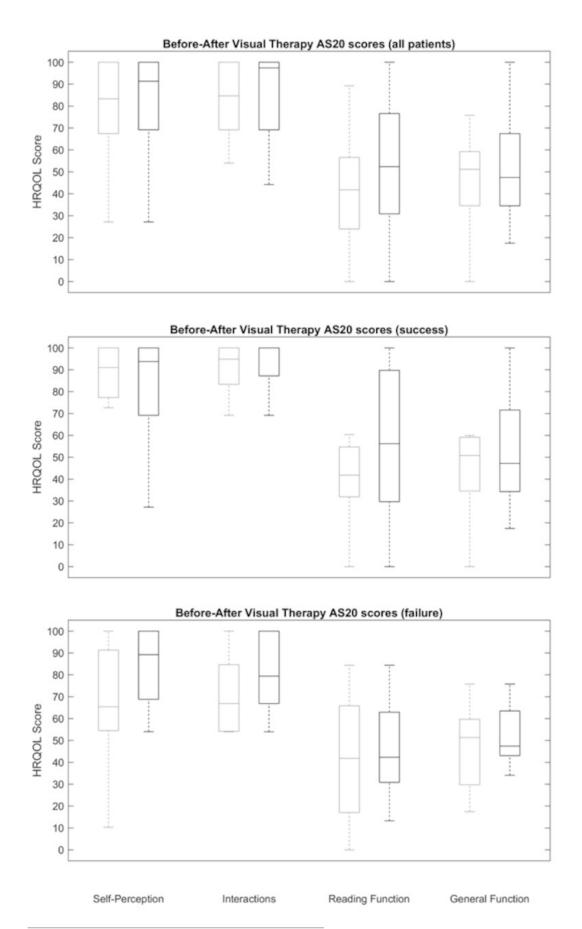




Table 1(on next page)

Results after prism treatment

Comparative analysis between success and failure groups. Statistical analysis performed using Mann-Whitney U test. Abbreviations: RE = right eye; LE = left eye; RPST = random-dot preschool stereo- acuity test.

Table 1. Comparative analysis between success and failure groups. Statistical analysis performed using Mann–Whitney U test. Abbreviations: RE = right eye; LE = left eye; RPST = random-dot preschool stereo- acuity test.

	Success group	Failure group N = 21	P
	N = 22	N = 21	
Age	70.09 ± 13.15	53.85 ± 15.39	<0.01
Refraction RE	-1.50 ± 4.88	-0.98 ± 4.09	0.51
Refraction LE	-1.22 ± 4.88	-1.01 ± 4.02	0.69
Visual acuity RE	0.88 ± 0.81	0.96 ± 0.8	0.41
Visual acuity LE	0.96 ± 0.81	±0.91 ± 0.169	0.57
Worth test	5	4.90 ± 0.30	0.14
RPST	607 ± 69	1219.5 ± 278.60	<0.01
Far strabismus horizontal deviation	3.65 ± 3.43	8.05 ± 8.61	0.16
Far strabismus vertical deviation	2.64 ± 3.82	5.05 ± 5.39	0.08
Near strabismus horizontal deviation	0.91 ± 2.52	8.57 ± 9.25	<0.01
Near strabismus vertical deviation	2.77 ± 3.93	5.43 ± 5.37	0.06
Torsional strabismus deviation	0.55 ± 1.26	2.52 ± 3.59	0.06
Far prismatic correction	5.85 ± 2.49	10.48 ± 7.59	0.03
Near prismatic correction	3.55 ± 4.04	11.33 ± 7.83	<0.01



Table 2(on next page)

symptomatology analysis

Comparative symptoms analysis of the two groups (success and failure) prior to prism correction, using the Adult Strabismus-20 questionnaire and the Diplopia Questionnaire. Statistical analysis performed with the Mann-Whitney U test.





Table 2: Comparative symptoms analysis of the two groups (success and failure) prior to prism correction, using the Adult Strabismus-20 questionnaire and the Diplopia Questionnaire. Statistical analysis performed with the Mann–Whitney U test.

Adult Strabismus-20 questionnaire	Success group	Failure group	P
Self-perception (0–100)	93.23 ± 13.74	79.07 ± 25.33	0.01
Interaction (0–100)	95.12 ± 11.82	83.21 ± 17.24	<0.01
Reading function (0–100)	65.99 ± 32.39	40.24 ± 25.90	<0.01
General function (0–100)	55.68 ± 22.96	46.89 ± 21.69	0.18
Diplopia Questionnaire	54.29 ± 26.09	72.33 ± 19.04	0.01