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Review Article

Visual system and motor development in children: a systematic review

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ABSTRACT.

Purpose.

The aim of this study was to review the available scientific literature on the possible relationship between the visual system and motor development in children.

Methods.

This study was performed according to the Preferred Reporting Items for Systematic Reviews (PRISMA) statement recommendations. The review protocol is available in PROSPERO (CRD42021245341). Four different databases, namely Scopus, PubMed, CINAHL and Web of Science, were assessed from April 2005 to February 2021. To determine the quality of the articles, we used the Critical Appraisal Skills Programme (CASP) Quality Appraisal Scale, and a protocol was followed to define the levels of evidence on the basis of the Centre for Evidence-Based Medicine Levels of Evidence. The search strategy included terms describing motor development in children and adolescents with visual disorders.

Results.

Among the identified studies, 23 were included in the study. All selected articles examined the relationship between the visual system and development in children. The quality of most of the studies was moderate—high, and they were between evidence levels 2 and 4.

Conclusions.

Our systematic review revealed that all included studies established a relationship between the visual system and development in children. However, the methods for measuring the visual system and motor skills lacked uniformity.

Key words: amblyopia – child development – motor skills – ocular motility disorders – visual disorders

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Introduction

Infant motor development includes the acquisition of basic skills, such as moving the head and eyes to look around, moving the arms and hands to grasp objects and moving the body to sit or go somewhere. It also includes higher-order skills, such as wielding a hammer to hit a peg and stacking boxes to reach a high object. The opportunities for motor action depend on the current state of the body: therefore, the systems involved must remain in the correct state (Adolph & Hoch 2019). In all these cases, vision plays a fundamental role in the correct acquisition of skills.

Normal visual development begins at birth and continues throughout childhood. It involves changes in visual acuity, convergence and accommodation until adequate binocular vision and stereopsis are achieved (Zimmermann et al. 2019). Binocular vision provides the visual information required to accurately perceive depth (stereopsis). In this way, the child will be able to adequately execute the movements of the upper and lower extremities (Goodale 2011; Chapman et al. 2012).

When a child suffers from visual disturbance, an alteration in motor development normally occurs. Vagge et al. (2021) reported that the presence of binocular dysfunction may be one of the factors that contribute to

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alterations in the development of a child. Other authors reported that the surgical correction of strabismus in a group of children led to the partial recovery of binocular vision and improved hand-eye coordination skills al. 2007). (Caputo et et al. (2018) stated that the proper development of the visual system is crucial for the development of correct balance. They showed that postural instability was greater in a group of children with amblyopia and strabismus than in children with normal

In addition, several studies show that children with amblyopia experience a deterioration of motor skills (Engel-Yeger 2008; Suttle et al. 2011; Webber et al. 2016; Birch et al. 2019a, 2019b). In general, these children have slower and less controlled movements than children with normal vision. Amblyopia is associated with longer saccades that decrease reading speed (Birch et al. 2019a, 2019b). Webber et al. (2016) reported that 5 weeks of treatment with visual therapy improved the fine motor skills of a group of children with amblyopia.

Other authors have established a relationship between the presence of refractive error and motor development. Atkinson et al. (2005) showed that the motor performance of hyperopic children aged between 3 and 6 years was significantly worse than that of emmetropic children of the same age. Children with hyperopia had deficits in manual dexterity, ball skills and balance.

The close relationship between visual and motor development is a topic of interest in the scientific community, although there is a lack of homogeneity, probably due to its multidisciplinary nature. Therefore, there is a need to conduct a systematic review to study the accumulated evidence.

The objective of this systematic review was to establish a relationship between visual and motor development in children. Additionally, we analysed the risk of bias assessment and publication certainty in all included studies.

Method

This systematic review was conducted according to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines (Liberati et al. 2009; Moher et al. 2009). The PRISMA statement has been designed

for systematic reviews of studies that evaluate the effects of health interventions, irrespective of the design of the included studies. However, the checklist items are applicable to reports of systematic reviews evaluating other non-health-related interventions and many items are applicable to systematic reviews with objectives other than evaluating interventions (evaluating aetiology, prevalence, prognosis *etc.*). It includes a checklist to guide the reporting of systematic reviews (Page et al. 2021).

The revision was registered in PROSPERO (Registration No. CRD42021245341). For the interpretation of all results, $p \le 0.05$ was considered statistically significant.

Search strategy

A systematic literature search was performed using the following databases: PubMed/MEDLINE (502 articles), Web of Science (27 articles), CINAHL (479 articles) and Scopus (27 articles).

The search strategy included terms describing a child's developmental stage (toddlers OR children OR Infant [Medical Subject Headings, MeSH]), terms describing motor development ("motor development" OR "motor function" OR "early motor development" OR "motor outcomes" OR "motor system" OR "motor coordination" OR "coordination skills" OR "motor skills") and terms describing visual development ("visual disorders" OR vision OR exophoria OR vergence OR exotropia OR esotropia OR heterophoria OR "ocular motility disorders" OR amblyopia OR stereoacuity OR "visual function" OR binocular OR accommodation OR accommodative OR "vertical deviation" OR "vertical disorder" OR "vertical anomalies" OR "vertical anomaly" OR "hyperdeviation" OR "strabismus" OR "eye movements" OR "visual complaints" OR "visual deterioration" OR phoria "visual development" OR stereovision OR "visual Skills" OR stereoacuity OR "Refractive errors" OR "visual acuity"). The search was updated from January to June 2021.

Inclusion criteria

Studies evaluating motor development in children with visual disorders were included. This evaluation had to be applied to children and adolescents with hyperopia, amblyopia and strabismus. The inclusion criteria were (1) studies with humans; (2) case reports; (3) case series, (4) cohort, crosssectional and case—control studies and (5) randomized clinical trials.

Exclusion criteria

Articles were excluded if: (1) they did not report data on motor or visual development; (2) the patients included were adults; (3) the patients had undergone eye surgery; (4) the patients had any motor developmental disorder or were blind; (5) the article was a letter, conference abstract, study protocol or literary review; or (6) the article was not available in English or Spanish.

If full-text reading led to the conclusion that the article did not analyse motor and visual development in typically developing healthy children and adolescents, the article was excluded.

Quality of articles, levels of evidence and data extraction

Article grading and data extraction were independently performed by two authors: MCSG and EPP. To determine the quality of the articles, two reviewers with adequate reliability (EPP and MCSG) worked independently and blindly to create a summary table (Table 1) on the basis of the Critical Appraisal Skills Programme (CASP) Quality Appraisal Scale ('CASP CHECKLISTS - CASP -Critical Appraisal Skills Programme' n.d.). CASP is a tool that analyses the quality of the articles selected in a systematic review, and it also ensures sufficient representation of the items in case-control, cohort, randomized controlled trials, as well as in cross-sectional studies. Some of the elements analysed are as follows: theoretical basis for the study; approprimethodological design; hiring information; description and representativeness of the participants; robustness of the research, including control or risk of bias; sufficiently appropriate and rigorous data analysis (including qualitative analysis, where appropriate); control of confounding factors; and clear discussion of the implications of the findings.

If the quality of the included articles was considered sufficient, a protocol was followed to define the levels of evidence on the basis of the Centre for Evidence-Based Medicine (CEBM) Levels of Evidence ('OCEBM Levels

Table 1. Quality appraisal of articles.

Author and date	Yes/total
Engel-Yeger (2008)	7/11
Webber et al. (2008)	8/11
Suttle et al. (2011)	7/11
Niechwiej-Szwedo et al. (2017)	6/11
Kelly et al. (2020)	7/11
Atkinson et al. (2005)	9/12
O'Connor et al. (2010)	8/12
Wilson and Welch (2013)	5/12
Alramis et al. (2016)	5/12
Chakraborty et al. (2017)	10/12
Fang et al. (2017)	7/12
Thompson et al. (2017)	9/12
Zipori et al. (2018)	6/12
Birch et al. (2019a)	9/12
Birch et al. (2019b)	9/12
Birch et al. (2020)	5/12
Hemptinne et al. (2020)	8/12
Niechwiej-Szwedo et al. (2020a)	9/12
Niechwiej-Szwedo et al. (2020b)	9/12
Pinero-Pinto et al. (2020)	8/12
Sá et al. (2021)	6/12
Vagge et al. (2021)	7/12
Webber et al. (2016)	7/11

Critical Appraisal Skills Programme (CASP) Quality Appraisal Scale.

of Evidence – Centre for Evidence-Based Medicine (CEBM), University of Oxford' n.d.). The CEBM levels of evidence were produced to enable the process of finding appropriate evidence and making its results explicit.

Any disagreements between the two reviewers were solved by a third unblinded reviewer (RPC). This analysis did not discard any articles. The main summary measures used in this systematic review were measures of visual development and assessment of motor development in all the included studies.

The quality of the included articles was classified into 3 outcome levels of equal measure: low (yes = 0-3), moderate (yes = 4-7) and high (yes = 8-12) for observational cohorts and crosssectional studies, and low (yes = 0-3), moderate (yes = 4-7)and (yes = 8-11) for case-control studies and controlled intervention studies. On the basis of this classification, we found that all case-control studies (Engel-Yeger 2008; Webber et al. 2008; Suttle al. 2011; Niechwiej-Szwedo et al. 2017; Kelly et al. 2020) were of moderate-high quality and that the single controlled intervention study (Webber et al. 2016) was of moderate quality. Regarding the cohort and crosssectional studies, we found seven studies

(Wilson & Welch 2013; Alramis et al. 2016; Fang et al. 2017; Zipori et al. 2018; Birch et al. 2020; Sá et al. 2021; Vagge et al. 2021) with high quality, and the remaining 10 (Atkinson et al. 2005; O'Connor et al. 2010; Chakal. 2017; raborty et Thompson et al. 2017: Birch et al. 2019a, 2019b; Hemptinne et al. 2020: Niechwiei-Szwedo et al. 2020a. 2020b: Pinero-Pinto et al. 2020) had moderate quality.

On the basis of the classification of the Oxford CEBM Levels of Evidence ('OCEBM Levels of Evidence - Centre for Evidence-Based Medicine (CEBM), University Oxford' n.d.), of 12 articles (O'Connor obtained et al. 2010; Alramis et al. 2016; Webal. 2016; Chakraborty et al. 2017; Fang et al. 2017; Zipori et al. 2018; Birch et al. 2019a, 2019b; Birch al. 2020; Pinero-Pinto et et al. 2020; Sá et al. 2021; Vagge et al. 2021) from level 2, six articles (Atkinson et al. 2005; Wilson & Welch 2013; Thompson et al. 2017; Hemptinne et al. 2020; Niechwiej-Szwedo et al. 2020a, 2020b) from level 3 and five articles (Engel-Yeger 2008; Webber et al. 2008; Suttle et al. 2011; Niechwiej-Szwedo et al. 2017; Kelly et al. 2020) from level 4.

Table 1 shows the representation of the agreed-upon ratings for the CASP Quality Appraisal Scale.

Results

A total of 1035 articles were identified. After removing duplicates, 865 articles were subjected to title and abstract reading by two authors, excluding 745 articles. If there was a conflict with the selection of an article, the third author decided the outcome. The full texts of the 120 articles were read, and 97 were excluded based on the exclusion criteria. A total of 23 articles were finally included in the review.

Figure S1 shows the Preferred Reporting Items for the Systematic Reviews and Meta-Analysis Flow Chart.

Characteristics of the studies

A total of 3980 children aged 2–18 years were evaluated for visual and motor development.

The systematic review suggests that there is a relationship between the visual system and motor development. Among all the reviewed articles, 9 articles established a relationship between amblyopia and fine and gross motor (Engel-Yeger 2008; Webber et al. 2008; Suttle et al. 2011; Wilson & Welch 2013; Birch et al. 2019a, 2019b; Birch et al. 2020; Kelly et al. 2020; Sá et al. 2021), 10 articles associated binocular vision and motor development al. 2010: (O'Connor et Alramis et al. 2016: Chakraborty et al. 2017: Niechwiej-Szwedo et al. 2017; Thompson et al. 2017; Zipori et al. 2018; Hemptinne et al. 2020; Niechwiej-Szwedo et al. 2020a, 2020b; Pinero-Pinto et al. 2020; Vagge et al. 2021), 1 article associated the presence of hyperopia and the impairment of manual balance dexterity and (Atkinson et al. 2005), 1 article described how visual perception and motor coordination change in preschool children (Fang et al. 2017), and 1 article studied the state of binocular vision and the accommodative system in children with typical motor development (Niechwiej-Szwedo et al. 2020a, 2020b).

The only intervention study included in the review, Webber et al. (2016), describes the efficacy of a visual therapy treatment for 5 weeks in a group of children with amblyopia. In this research, they trained binocular vision skills by using an iPod game. Fine motor skills improved in all children, and the results were maintained over time.

Regarding the motor development assessment tests, five studies (Atkinson et al. 2005; Engel-Yeger 2008; Chakraborty et al. 2017; Kelly et al. 2020; Sá et al. 2021) used the Movement Assessment Battery for Children (MABC), and three studies (Webber et al. 2008; Webber et al. 2016; Zipori et al. 2018) used the Bruininks-Oseretsky Test of Motor Proficiency. Three studies (O'Connor et al. 2010; Suttle et al. 2011; Niechwiej-Szwedo et al. 2020a, 2020b) used tests that were specifically designed for the study, and seven studies (Wilson & Welch 2013; Alramis et al. 2016; Niechwiej-Szwedo et al. 2017; Birch et al. 2019a, 2019b; Birch et al. 2020; Hemptinne et al. 2020; Niechwiej-Szwedo et al. 2020a, 2020b) did not report the test they used or did not provide data. Regarding the remaining four studies (Fang et al. 2017; Thompson et al. 2017; Pinero-Pinto et al. 2020; Vagge et al. 2021), each study used a different test that has not been used in any other study.

Among all the articles, four articles reported data on gross motor development (Atkinson et al. 2005; Engel-Yeger 2008; Webber et al. 2008; Zipori et al. 2018), another four reported on fine motor development (O'Connor et al. 2010; Suttle et al. 2011; Webber et al. 2016; Fang et al. 2017), seven articles analysed fine and coarse motor development (Chakraborty et al. 2017; Thompson et al. 2017; Kelly et al. 2020; Niechwiej-Szwedo et al. 2020a, 2020b; Pinero-Pinto et al. 2020; Sá et al. 2021;

Vagge et al. 2021), and eight did not report data on motor development (Wilson & Welch 2013; Alramis et al. 2016; Niechwiej-Szwedo et al. 2017; Birch et al. 2019a, 2019b; Birch et al. 2020; Hemptinne et al. 2020; Niechwiej-Szwedo et al. 2020a, 2020b).

Table 2 provides the study characteristics of all included articles.

The results of each study were classified by vision outcomes (Table 3) and motor outcomes (Table 4).

Discussion

The prevalence of visual dysfunction in the paediatric population has increased significantly in recent years (Cacho-Martínez et al. 2010; Jang & Park 2015; Hussaindeen et al. 2017). An early diagnosis of visual abilities in children is necessary to prevent the appearance of possible non-strabismic binocular and accommodative alterations that could affect motor development and quality of life. There are several symptoms and

Table 2. Studies characteristics.

Autor (date)	Design	Conflict of interest	Follow-up (months)	n (subjects)	Sex (F/M)	Age
Atkinson et al. (2005)	OC	NR	24	HG 1st: 110; 2nd: 99 CG 1st: 131; 2nd:113	HG 1st: 63/47; 2nd:56/43 CG 1st: 70/61; 2nd: 62/51	1st: 3 years, 7 ± 1.6 MM 2nd: 5 years, 6 ± 1.7 MM
Engel-Yeger (2008)	CC	NR	No	AG: 22 CG: 25	AG: 11/11 CG: 13/12	AG: 5.65 ± 0.91 years CG: 5.53 ± 0.71 years
Webber et al. (2008)	CC	NR	No	AG: 82 CG: 37	AG: 45/37 CG: 18/19	AG: 8.2 ± 1.7 years CG: 8.3 ± 1.3 years
O'Connor et al. (2010)	CS	NR	No	SG: 121	SG: 91/30	18.8 years (12-28)
Suttle et al. (2011)	CC	NR	No	AG: 21 CG: 47	Not described	AG: 4–8 years CG: 4–8 years; 20–42 years
Wilson and Welch (2013)	OC	No	No	AG: 1032	AG: 493/539	3–32 years
Alramis et al. (2016)	CS	NR	No	52 Children 19 Adults	Children: 30/22 Adults: 10/9	Children: 5–13 years Adults: 18–38 years
Webber et al. (2016)	CI	NR	5 weeks 17 weeks	AG: 20 CG: 10	AG: 11/9 CG: 4/6	AG: 8.5 ± 1.3 years CG: 9.63 ± 1.6 years
Chakraborty et al. (2017)	CS	NR	No	606	287/319	4.5 years
Fang et al. (2017)	CS	No	No	151	70/81	4–6 years
Niechwiej-Szwedo et al. (2017)	CC	NR	No	SG: 19 CG: 19	SG: 10/10 CG: 10/100	SG: 8.68 ± 1.89 years CG: 8.68 ± 1.89 years
Thompson et al. (2017)	OC	No	24	375	177/198	24 months (23–25)
Zipori et al. (2018)	CS	No	No	AG: 18 SG: 16 CG: 22	AG: 10/8 SG: 6/10 CG: 12/10	AG: 8.5 ± 2.0 years SG: 10.9 ± 3.6 years CG: 10.6 ± 3.2
Birch et al. (2019a)	CS	NR	17	AG: 50 NAG: 13 CG: 18	AG: 31/19 NAG: 7/6 CG: 10/8	AG: 10.6 ± 1.3 years NAG: 10.7 ± 1.2 years CG: 10.6 ± 1.4 years
Birch et al. (2019b)	CS	Yes	28	AG: 60 NAG: 30 CG: 20	AG: 28/32 NAG: 16/14 CG: 11/9	AG: 6.3 ± 1.3 years NAG: 5.9 ± 1.3 years CG: 6.1 ± 1.1 years
Birch et al. (2020)	CS	NR	No	AG: 15 CG: 20	AG: 5/10 CG: 8/12	AG: 4.6 ± 1.0 years CG: 5.0 ± 1.0 years
Hemptinne et al. (2020)	OC	NR	No	SG: 40 CG: 18	SG: 19/21 CG: 6/12	SG: 7.25 ± 3.83 years CG: 8.33 ± 5.42 years
Kelly et al. (2020)	CC	No	No	AG: 96 NAG: 47 CG: 35	AG: 43/53 NAG: 22/25 CG: 18/17	AG: 8.2 ± 2.5 years NAG: 7.0 ± 2.6 years CG: 8.3 ± 2.8 years
Niechwiej-Szwedo et al. (2020a)	OC	No	No	57	31/26	F: 10.63 ± 1.93 years M: 10.74 ± 2.03 years
Niechwiej-Szwedo et al. (2020b)	OC	NR	No	226	110/116	F: 9.21 ± 2.58 years M: 9.70 ± 2.56 years
Pinero-Pinto et al. (2020)	CS	No	No	116	63/53	29.57 ± 3.45 months
Sá et al. (2021)	CS	No	No	NAG: 97 CAG: 37 NCAG: 31	NAG: 46/51 CAG: 22/15 NCAG: 20/11	NAG: 7.6 ± 1.2 years CAG: 7.6 ± 1.2 years NCAG: 6.9 ± 1.0 years
Vagge et al. (2021)	CS	No	No	SG: 23 CG: 24	SG: 9/14 CG: 10/14	SG: 7.5 ± 2.0 years CG: 7.2 ± 1.7 years

AG = amblyopic group, CAG = corrected-amblyopic group, CC = case–control study, CG = control group, CI = controlled Intervention study, CS = cross-sectional study, F/M = female/male, F/M = hyperopic group, F/M = non-amblyopia group, F/M = non-corrected-amblyopic group, F/M = not reported, F/M = observational cohort study, F/M = strabismus group.

Table 3. Vision outcomes.

Atkinson et al. (2005) Engel-Yeger (2008) Webber et al. (2008) Stereopsis sec arc Interocular difference V (logMAR) VA better eye (logMAR) VA worse eye (logMAR) Refractive error (D) Vision assessme VA Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergenc (convergence Suttle et al. (2011) VA, binocular (Mean (SD)) VA, dom eye (IMEAN (SD)) VA, non-dom eye (logMar Mean (SD)) Interocular axis mean (SD) Interoc	Nil 800–60 ≤40	Amblyopia group (n = 82) 50 (61) 27 (33) 5 (6) 0.31 (0.06)	3 years, 7 months: +5.33 (1.48) 5 years, 6 months: +5.30 (1.49)
Webber et al. (2008) Vision assessme Stereopsis sec arc Interocular difference V (logMAR) VA better eye (logMAR) VA worse eye (logMAR) Refractive error (D) Vision assessme VA Worth 4 dot test 4 (D) base-out Motor fusion Positive vergence (convergence) Suttle et al. (2011) VA, binocular (Mean (SD)) VA, dom eye (logMar) VA, non-dom eye (logMar) Mean (SD) VA, non-dom eye (logMar) Mean (SD)	Nil 800–60 ≤40	group (n = 82) 50 (61) 27 (33) 5 (6) 0.31 (0.06)	
Stereopsis sec arc Interocular difference V (logMAR) VA better eye (logMAR) VA worse eye (logMAR) Refractive error (D) Vision assessme et al. (2010) Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergenc (convergence VA, binocular (Mean (SD) VA, dom eye (Indean (SD) VA, non-dom eye (logMar Mean (SD))	Nil 800–60 ≤40	group (n = 82) 50 (61) 27 (33) 5 (6) 0.31 (0.06)	
Interocular difference V (logMAR) VA better eye (logMAR) VA worse eye (logMAR) Refractive error (D) Vision assessme et al. (2010) Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergenc (convergence Suttle et al. (2011) Vision assessme vA, binocular (Mean (SD) VA, dom eye (IMean (SD) VA, non-dom eye (logMar Mean (SD)	800–60 ≤40	50 (61) 27 (33) 5 (6) 0.31 (0.06)	
Interocular difference V (logMAR) VA better eye (logMAR) VA worse eye (logMAR) Refractive error (D) Vision assessme et al. (2010) Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergenc (convergence Suttle et al. (2011) Vision assessme vA, binocular (Mean (SD) VA, dom eye (IMean (SD) VA, non-dom eye (logMar Mean (SD)	800–60 ≤40	27 (33) 5 (6) 0.31 (0.06)	
Interocular difference V (logMAR) VA better eye (logMAR) VA worse eye (logMAR) Refractive error (D) Vision assessme et al. (2010) Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergenc (convergence Suttle et al. (2011) Vision assessme vA, binocular (Mean (SD) VA, dom eye (IMean (SD) VA, non-dom eye (logMar Mean (SD)	≤40	5 (6) 0.31 (0.06)	
difference V (logMAR) VA better eye (logMAR) VA worse eye (logMAR) Refractive error (D) Vision assessme VA Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergenc (convergence Suttle et al. (2011) Vision assessme VA Vision assessme VA, binocular (Mean (SD) VA, dom eye (Indean (SD) VA, non-dom eye (logMar Mean (SD)	4	,	
VA better eye (logMAR) VA worse eye (logMAR) Refractive error (D) Vision assessme VA Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergenc (convergence Suttle et al. (2011) Vision assessme VA Vision assessme VA, binocular (Mean (SD) VA, dom eye (Index (SD) VA, non-dom eye (logMar Mean (SD)		0.10 (0.01)	
VA worse eye (logMAR) Refractive error (D) Vision assessme VA Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergenc (convergence Suttle et al. (2011) Vision assessme VA Vision assessme VA, binocular (Mean (SD) VA, dom eye (IMEAN (SD) VA, non-dom eye (logMar Mean (SD)		0.10 (0.01)	
O'Connor et al. (2010) Wision assessme VA Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergenc (convergence) Suttle et al. (2011) Vision assessme VA, binocular (Mean (SD)) VA, dom eye (I Mean (SD)) VA, non-dom eye (logMar Mean (SD))		0.38 (0.05)	
et al. (2010) Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergence (convergence) Suttle Vision assessme VA, binocular (Mean (SD)) VA, dom eye (I Mean (SD)) VA, non-dom eye (logMar Mean (SD))		2.30 (0.25)	
Worth 4 dot tes 4 (D) base-out Motor fusion Positive vergence (convergence) Vision assessme et al. (2011) VA, binocular (Mean (SD) VA, dom eye (I Mean (SD) VA, non-dom eye (logMar Mean (SD)	ıt	Strabismus group	
4 (D) base-out Motor fusion Positive vergene (convergence) Suttle et al. (2011) Vision assessme VA, binocular (Mean (SD)) VA, dom eye (IMean (SD)) VA, non-dom eye (logMan Mean (SD))		of the poorer seeing eye ranged from 0.2 to 1.6 logMAR of the fellow eyes from 0.24 to 0.30 logMAR	
Motor fusion Positive vergence (convergence Vision assessme et al. (2011) VA, binocular (Mean (SD)) VA, dom eye (I Mean (SD)) VA, non-dom eye (logMan Mean (SD))	t	Fusion $(n = 99)$ (a response of four lights) Suppression $(n = 16)$ (a response	
Positive vergence (convergence) Suttle et al. (2011) Vision assessme VA, binocular (Mean (SD)) VA, dom eye (I Mean (SD)) VA, non-dom eye (logMan Mean (SD))	est	of two or three lights) Bifoveal (n = 94) (Vergence movement of eye not	
(convergence Suttle et al. (2011) VA, binocular (Mean (SD) VA, dom eye (IMean (SD)) VA, non-dom eye (logMan Mean (SD))		under prism following conjugate version movements by both eyes) Central suppression (<i>n</i> = 6) (no vergence movement) Prism bar Normal (<i>n</i> = 101) Reduced (<i>n</i> = 7)	Risley prism Normal $(n = 84)$ Reduced $(n = 17)$
(convergence Suttle et al. (2011) VA, binocular (Mean (SD) VA, dom eye (IMean (SD)) VA, non-dom eye (logMan Mean (SD))		Nil $(n = 13)$	Nil $(n = 14)$
(convergence Suttle et al. (2011) VA, binocular (Mean (SD) VA, dom eye (IMean (SD)) VA, non-dom eye (logMan Mean (SD))	e	Normal $(n = 97)$	Normal $(n = 61)$
et al. (2011) VA, binocular (Mean (SD) VA, dom eye (IMean (SD)) VA, non-dom eye (logMan Mean (SD))		Reduced $(n = 9)$	Reduced $(n = 41)$
et al. (2011) VA, binocular (Mean (SD) VA, dom eye (IMean (SD)) VA, non-dom eye (logMan Mean (SD))		Nil $(n = 15)$	Nil (n = 14)
Mean (SD) VA, dom eye (I Mean (SD) VA, non-dom eye (logMar Mean (SD)		Group normal child	Amblyopic children
VA, dom eye (I Mean (SD) VA, non-dom eye (logMar Mean (SD)	logMar)	Early = $0.01 (0.05)$	_
Mean (SD) VA, non-dom eye (logMar Mean (SD)		Middle = $-0.02 (0.07)$ Late = $-0.06 (0.06)$	
Mean (SD) VA, non-dom eye (logMar Mean (SD)	ogMar)	Early = $0.06 (0.06)$	_
VA, non-dom eye (logMar Mean (SD)	/giviai)	Middle = $0.04 (0.08)$	
eye (logMar Mean (SD)		Late = $0.01 (0.08)$	
Mean (SD)		Early = $0.08 (0.06)$	_
` /)	Middle = $0.02 (0.07)$	
Interocular		Late = $0.03 (0.08)$	
		Early = $0.05 (0.04)$	Mild amblyopia
difference (IOD Mean (SD)	ı	Middle = 0.04 (0.04) Late = 0.04 (0.03)	(IOD = 0.11-0.3) Moderate-to-severe
C4	assad	$E_{0} = 45 (12)$	amblyopia (IOD ≥0.31
Stereo acuity cr (arc sec)	Joseu	Early = $45 (13)$ Middle = $44 (24)$	_
Mean (SD)		Late = $51 (15)$	
Stereo acuity u	icrossed	Early = $57 (21)$	_
(arc sec)		Middle = $63 (28)$	
Mean (SD)		Late = $50 (19)$	
Stereo acuity (a Mean (SD)		_	Mild amblyopia and Moderate-to-severe amblyopia

No

7 (64%)

9 (64%)

17 (68%)

394 (91%) 313 (93%)

33 (100%)

4.3(0.03)

amblyopia 13 (52%)

Recovered

1 (4%) 1 (9%)

0 (0%)

1 (4%)

6 (2%) 2 (1%)

0 (0%)

3.5(0.47)

amblyopia

Wilson and Welch (2013)	TNO best score (arc sec)	Amblyopia	Possible amblyopia
()	Absent	6 (24%)	5 (20%)
	480	0 (0%)	3 (27%)
	240	2 (14%)	3 (22%)
		` /	
	120	1 (4%)	6 (24%)
	60	5 (1%)	27 (6%)
	30	1 (<1%)	19 (6%)
	15	0 (0%)	0 (0%)
	Mean (SD)	2.1(0.05)	3.7(0.19)
Alramis	Vision assessment	Group child	
et al. (2016)	VA	Monocular RE	
		Monocular LE ≥	_
		Binocular ≥0.1 lo	og MAR
	Stereoacuity	Young = 20–70	
	range (arc sec)	Middle = 20-70	
		Older = 20-30	
Webber	Vision assessment	Treated amblyop	
et al. (2016)	VA worst eye	Baseline = 0.46 (` '
	(logMAR)		eatment) = $0.37 (0.16)$
	Mean (±SD)	17 weeks (12 we	
			$d) = 0.36 \ (0.18)$
	VA both eyes	Baseline = -0.03	3 (0.09)
	(logMAR)	5 weeks (post-tre	eatment) = $-0.06 (0.09)$
	Mean (±SD)	17 weeks (12 we	eks After
		Treatment Ceas	sed) = -0.03 (0.06)
	BF score	Baseline $= 3.44$ ((1.27)
	(log stereoacuity)	5 weeks (Post-tre	eatment) = $2.88 (1.05)$
	Mean (±SD)	17 weeks (12 we	eks After
		Treatment Ceas	sed) = 2.74 (1.06)
Chakraborty	Vision assessment	Group child	
et al. (2017)	VA; N (%)	$\leq 0.3 \log MAR =$	578 (95.37)
		$>0.3 \log MAR =$	28 (4.62)
	Stereoacuity; N (%)	<100 = 476 (78.5	54)
	(arc sec)	>100-800 = 84 (13.86)
		>800 = 46 (7.59)	
Fang et al.	Vision assessment	Group child	
(2017)	Visual motor integration	4 years = 107.58	(10.98); Range = 83–141
	Mean (±SD)	5 years = 112.38	(10.99); Range = 95–138
		6 years = 105.80	(10.46); Range = 90–138
	Visual perception	4 years = 111.53	(16.10); Range = $67-144$
	Mean (±SD)	5 years = 117.70	(11.45); Range = $92-146$
		6 years = 115.07	(8.94); Range = 92–146
Niechwiej-Szwedo	Vision assessment	Group child	
et al. (2017)	VA (logMAR) distance/	-0.01 ± 0.02 (-	0.10 to
, ,	near $(n = 19)$	$0.00)/0.01 \pm$	0.02 (0.00-0.10)
	$(Mean \pm SD)$	Snellen range: 20	
	Stereopsis (sec of arc)	$28 \pm 10 \ (20-50)$	
	(n = 19)		
	(Mean \pm SD) [range]		
	Positive fusional	$27 \pm 12 (12-45)$	$/17 \pm 8 \ (8-30)$
	vergence (BO)		
	Near – break/recovery		
	(n = 17)		
	(Mean \pm SD) [range]		
	Negative fusional	$12 \pm 5 (6-20)/9$	$\pm 5 (2-20)$
	vergence (BI)		
	Near – break/recovery		
	(n = 17)		
	(Mean ± SD) [range]		
	Binocular accommodation	$8 \pm 3 \ (3-11.5)$	
	facility (cpm) $(n = 15)$. /	
	$(Mean \pm SD)$ [range]		
	Monocular	$8 \pm 3 (3-13)/8 \pm$	± 3 (3–13)
	accommodation	. //-	. ,
	facility		

	RE and LE (cpm) (n = 16) (Mean \pm SD) [range] Amplitude of accommodation RE and LE (D) $(n = 18)$ (Mean \pm SD) [range]		$12 \pm 2 (8-16)/12 \pm 2 (8-16)$
Thompson et al. (2017)	Vision assessment Binocular visual acuity (LogMAR) mean (SD) [range] Stereoacuity (sec of arc) mean (SD) [range] Vision impairment score, n (%)	Group child 0.06 (0.15) [-0.20, 1.00] 366 (196) [200, 1200] n = 290 (77%) (normal) n = 69 (18%) (internal or external ocular health problem or strabismus or abnormal motility or absence of stereopsis or binocular visual acuity worse than 0.5 logMAR) n = 16 (4%) (two or more	-10)
	Refractive error score, n (%)	visual dysfunctions) $n = 177 (90.5\%)$ (normal) $n = 17 (9\%)$ (Hyperopia (mean sphere [M] $\geq +4.00$ diopter [D])) or myopia (M ≤ -1.00 D) or astigmatism (cylinder [C] ≤ -1.50 D in any meridian) or anisometropia (difference in M between eyes of ≥ 3.00 D in either the most positive or negative meridian) $n = 1 (0.5\%)$ (two or more refractive errors)	
Zipori et al. (2018)	Vision assessment	Unilateral amblyopia	Strabismus without amblyopia
	VA (logMAR)	BCVA between (0.3 logMAR) and (1.3 logMAR) in the amblyopic eye VA of (0.1 logMAR) or better in the non- amblyopic eye	Normal VA in both eyes
	Strabismus	–	Esotropia $(n = 9)$ Exotropia $(n = 7)$
	Stereoacuity	$3000 \ (n=2)$	3000 $(n = 10)$
T	(arcsecond)	$140 \ (n=1)$	$20-60 \ (n=8)$
Birch et al. (2019a)	Vision assessment VA (logMAR)	Group child With amblyopia	Without amblyopia
ct al. (2019a)	VA (logNIAR) ≤0.1	0 (0%)	0 (0%)
	0.2-0.3	30 (60%)	0 (0%)
	0.4-0.5	12 (24%)	0 (0%)
	0.6–0.7 >0.7	4 (8%) 4 (8%)	0 (0%) 0 (0%)
	Mean (SD); (range)	0.41 (0.33); (0.2–1.9)	-0.01 (0.03); (-0.1 to 0.1)
	Fellow eye visual acuity (logMAR)	With amblyopia	Without amblyopia
	-0.1 0.0	34 (68%) 13 (26%)	7 (50%) 5 (43%)
	0.1	3 (6%)	1 (7%)
	Mean (SD); (range)	-0.06 (0.06); (-0.1 to 0.1)	0.00 (0.02); (-0.1 to 0.1)
	Stereoacuity (log arcsec).	With amblyopia	Without amblyopia
	Median (range) (Nil stereoacuity was	4.00 (1.6-nil)	2.30 (1.6-nil)
	assigned a value of 4.0 log		
	arc per second)	C 131	
	Vision assessment	Group child	

D:I-	VA (IMAD)	W/:4h Ahl		W/:4L4 A Ll: -
Birch et al. (2019b)	VA (logMAR) ≤0.1	With Amblyopia		Without Amblyopia 30 (100%)
et al. (2019 b)	0.2–0.3	22 (37%)		0 (0%)
	0.4–0.5	21 (35%)		0 (0%)
	0.6-0.7	5 (8%)		0 (0%)
	>0.7	12 (20%)		0 (0%)
	Mean (SD); (Range)	0.49 (0.27; [0.2–1.4]		0.04 (0.07); [-0.1 to 0.2]
	Fellow eye visual acuity (logMAR)	With amblyopia		Without amblyopia
	-0.1 (20/16)	12 (20%)		5 (17%)
	0.0 (20/20) 0.1 (20/25)	27 (45%) 21 (35%)		18 (60%) 7 (23%)
	Mean (SD) [range]	0.01 (0.07); [-0.1 to 0.1]		0.01 (0.06); [-0.1 to 0.1
	Snellen equivalent	20/20; [20/16–20/25]		20/20; [20/16–20/25]
	Stereoacuity (log arcsec).	3.75 (0.58); [1.8 to nil]		3.33 (0.83); [1.8 to nil]
	Median (range)			
	(Nil stereoacuity			
	was assigned			
	a value of 4.0 log			
Birch et al. (2020)	arc per second) Vision assessment	Amblyopia group		
Birch et al. (2020)	VA (logMAR)	Ambiyopia group		
	Amblyopic eye BCVA	0.4-2.0; (20/50-2000)		
	Mean (SD) [range]	, (., ,		
	Fellow eye visual acuity	-0.1 to 0.2; (20/15-30)		
	Mean (SD) [range]			
	VA in each eye	_		
TT	Stereoacuity (arcsecond)	Nil		
Hemptinne et al. (2020)	Vision assessment Strabismus type	Strabismus group Infantile esotropia ($n = 13$)	3)	
et al. (2020)	Strabishius type	secondary esotropia (<i>n</i> = 13		
		acquired esotropia (n		
	Binocularity degree	Absent $(n = 21)$, partial $(n = 21)$,	
		normal $(n = 6)$		
	Deviation angle,	0Δ -12 Δ ($n = 29$), 12Δ -20 Δ	Δ	
	corrected	$(n=5), \ge 20\Delta \ (n=6)$		T.F.
	Distance VA (logMAR)	RE		LE Nihit (n = 1)
		Nihil $(n = 2)$ Hand motions $(n = 1)$		Nihil $(n = 1)$ Hand motions $(n = 1)$
		$0.0 \ (n = 22)$		0.0 (n = 25)
		0.1 (n = 7)		0.1(n = 1)
		0.2 (n = 4)		0.2 (n = 6)
		0.3 (n = 3)		0.3 (n = 1)
		0.4 (n = 0)		$0.4 \ (n=3)$
17 11	*****	0.5 (n = 1)		0.5 (n = 2)
Kelly et al. (2020)	Visión assessment BCVA (logMAR)	Amblyopic 0.4 ± 0.3		Non-amblyopic 0.1 ± 0.1
et al. (2020)	(amblyopic eye)	0.4 ± 0.3 0.1–1.9		-0.1 ± 0.1
	Mean ± SD; range	0.1 1.5		0.1 to 0.5
	BCVA (logMAR)	0.0 ± 0.1		0.1 ± 0.1
	(fellow eye)	-0.1 to 0.2		-0.1 to 0.3
	Mean \pm SD; range			
	Stereoacuity (arcsecond)	3.4 ± 0.8		3.1 ± 1.0
	Mean ± SD; range	$(1.8-4)$ 0.4 ± 0.4		$(1.6-4)$ 0.3 ± 0.6
	Extent of suppression (log deg) Mean \pm SD; range	(-0.2 to 1.2)		(-0.2 to 1.2)
	Depth of suppression (CBI)	4.8 ± 3.6		3.0 ± 2.9
	Mean \pm SD; range	(0.2–11.0)		(0.2–11.0)
Niechwiej-Szwedo	Vision assessment		ly developing	
et al. (2020a)			dren. Mean \pm SD (range)	
	Visual acuity (logMAR)		± 0.07 (-0.10-0.18)/	
	distance/near		\pm 0.03 (0.00–0.18) range: 20/15–20/25	
	Stereopsis (secof arc)	Shellen 24 ± 7		
	Phoria (PD) distance/near		$0 (8-45)/18 \pm 11 (2-45)$	
	Negative fusional vergence (BI,		$(4-25)/11 \pm 4 (4-20)$	
	divergence) near - break/			
	recovery (PD)		(5.00)	
	Vergence facility (cpm)	14 ± 4	(5–24)	

	Binocular accommodative	$8 \pm 3 \ (0.5 - 14.5)$	
	facility (cpm)		
	Amplitude of accommodation– RE and LE (D)	$11 \pm 2 (7-16)/11 \pm 2 (6-16)$	
	Accuracy of accommodation	$1.06 \pm 0.40 \; (0.25 - 2.00) /$	
	(MEM)–RE and LE (D)	$21.06 \pm 0.40 (0.25 - 2.00)$	
Niechwiej-Szwedo et al. (2020b)	Vision assessment	Typically developing children. Mean \pm SD (range)	
` ,	Binocular visual acuity	$0.00 \pm 0.11 (-0.2 \text{ to } 0)$	
	(logMAR) distance	, ,	
	Monocular visual acuity (logMAR) distance	$0.04 \pm 0.12 \; (-0.1 \; \text{to} \; 0)$	
	Interocular visual acuity difference (logMAR)	$0.06 \pm (0.10) (0.0 – 0.6)$	
	stereoacuity (sec of arc)	$45 \pm 27 \ (20-200)$	
Pinero-Pinto	Visual development parameter	Gross motor quotient < 100	Gross motor quotient > 100
et al. (2020)	Visual acuity (CardiffTest—	$0.18 \pm 0.10/0.18 \pm 0.10$	$0.18 \pm 0.10/0.18 \pm 0.09$
ct al. (2020)	LogMAR) (RE)/(LE)	$0.10 \pm 0.10/0.10 \pm 0.10$	0.16 ± 0.10/0.16 ± 0.09
	Visual acuity (Broken	$0.36 \pm 0.05/0.36 \pm 0.04$	$0.37\pm0.04/0.37\pm0.04$
	Wheels—LogMAR) (RE)/(LE)	,	
	Retinoscopy refraction	$+1.27 \pm 0.91/+1.35 \pm 0.92$	$+1.35 \pm 0.73/+1.49 \pm 0.74$
	(Diopters D) (RE)/(LE)		
	Spherical equivalent refraction	4 (100/) /7 (17 50/) /20 (72 50/)	1 (1 20/) (20 (26 90/) (47
	Kappa angle	4 (10%)/7 (17.5%)/29 (72.5%)	1 (1.3%)/28 (36.8%)/47
	(negative/0/positive) (RE)	4 (100/) (0 (200/) (20 (700/)	(61.8%)
	Kappa angle	4 (10%)/8 (20%)/28 (70%)	2 (2.26%)/26 (34.3%)/48
	(negative/0/positive) (LE)	2 (7 70/)/7 (17 05)/20 (74 40/)	(63.2%)
	Hirshberg reflex (temporal/	3 (7.7%)/7 (17.95)/29 (74.4%)	1 (1.3%)/30 (39.5%)/45
	centred/nasal) (RE) Hirshberg reflex (temporal/	3 (7.9%)/8 (21.1%)/27 (71.1%)	(59.2%) 2 (2.6%)/30 (39.5%)/44
	centred/nasal) (LE)	3 (7.970)/8 (21.170)/27 (71.170)	(57.9%)
	Krismky test (normal/deviated)	70 (87.5%)/10 (12.5%)	36 (100%)/0 (0%)
	Near point of convergence	2.46 ± 4.07	1.00 ± 2.02
	(centimetre, cm)	2.40 ± 4.07	1.00 ± 2.02
	Base-out 6ΔPrism Test	27 (33.8%)/53 (66.3%)	16 (44.4%)/20 (55.6%)
	(prism diopters, Δ)	27 (33.870)/33 (00.370)	10 (44.470)/20 (33.070)
	(negative/positive)		
	Base-In 6ΔPrism Test	60 (75%)/20 (25%)	26 (72.2%)/10 (27.8%)
	(prism, diopters, Δ	00 (73 /0)/20 (23 /0)	20 (72.270)/10 (27.070)
	(negative/positive)		
	Stereopsis lang test	303.89 ± 143.67	282.35 ± 131.35
	(second arc)	505105 ± 1.6107	202100 ± 101100
	(200", 400" y 600")		
	Bruckner test	71 (88.8%)/9 (11.3%)	36 (100%)/0 (0%)
	(normal/deviated)		(), . ()
	Fixation test	64 (80%)/16 (20%)	26 (72.2%)/10 (27.8%)
	(passed/not passed)		
	Reflection and head	46 (57.5%)/34 (42.5%)	21 (58.3%)/15 (41.0%)
	(saccades movements)		
	Head (saccades movements)	12(15%)/33(41.3%)	3(8.3%)/21(58.3%)
	(motionless/slight/	24(30%)/11(13.85)	6(16.7%)/6(16.7)
	medium/strong)		
Sá et al. (2021)	Amblyopic group		
	Visual acuity values <0.7 on the		
	Snellen scale in one or both		
	eyes or difference in vision between		
	the eyes greater than two lines		
	on that scale.		
Vagge et al. (2021)	Vision assessment	Strabismus group	
	Stereopsis	Normal $(n = 11)$; absent $(n = 12)$	
	Amblyopia	n = 9	
	Non-amblyopia	n = 14	

BCVA = best-corrected visual acuity, BCVA = monocular best-corrected visual acuity, BF = binocular function, BI = base-in, BO = base-out, CBI = Contrast Balance Index, CPM = cycles per minute, D = diopters, Dom Eye = dominant eye, LE = left eye, Log deg = log degrees (Extent of suppression scotoma), MEM = monocular estimate method, Non-dom eye = nondominant eye, PD = prism diopter; Positive fusional vergence (BO, convergence) near – break/recovery (PD), RE = right eye, SD = standard deviation, Δ = prism diopter.

Table 4. Motor outcomes.

Author et al. (year	Motor measures	i						
Atkinson et al. (2005) Movement Assessment Battery for Children (MABC-2) Mean (SD) Engel-Yeger (2008) Balance and ball skills sub-tests from the Movement Assessment Battery for Children (MABC-2) Mean ± SD Webber et al. (2008) Bruininks-Oseretsky Test of Motor Proficiency (BOTMP)		Children I	Parameters Catch bean bag (bags caught) Roll ball (balls rolled into goal) Walk (steps walked) Jump score Balance (PL) (seconds balanced) Balance (NPL) (seconds balanced) Coin (PH) (seconds taken) Coin (NPH) (seconds taken) Beads: (6 beads) (seconds taken) Bicycle trail (errors made) Static balance Dynamic balance Mean dynamic balance score Total mean ball skills score Amblyopic group Visual motor control Cutting circle Drawing crooked path Drawing straight path Drawing circle Copying triangle Copying diamond Copying pencils Sum item 7		Hyperopic group Age 31/2 years 6 (5.9) 4 (3.9) 5 (6.4) 0 (1.5) 4 (4.3) 3 (4.5) 26 (26.3) 28 (30.5) 46 (49.2)		8 6 13 0 11 7 19 22	Age 51/2 years 8 (8.2) 6 (6.4) 13 (12.1) 0 (0.6) 11 (9.5) 7 (7.9) 19 (19.8) 22 (22.3) 59 (63.0)
		l skills m the Sassessment l Children			6 (7.0) Standing on one leg Jumping over cord Walking heels raised Catch			0 (0.6) Amblyopic group 1.9 ± 1.99 0.85 ± 1.76 1.45 ± 1.81 1.15 ± 1.09 1.4 ± 1.05 1.9 ± 1.86
		tsky Test Officiency			Variable Victor Victor		s 9 (1–10) ds 3 (1–7) eads 2 (1–4) pegs 4 (2–7) ertical lines 5 (0–8) eles 4 (1–8) ts 5 (1–9)	
			0		Bead task time (s) Water ta Mean ± SD Mean ±)
	Worth 4 dot response	Worth 4 do	Suppression Bifoveal	16.8 ± 1.6 15.2 ± 1.3 16.8 ± 1.5	Large 49.2 ± 5.2 58.4 ± 7.0 48.9 ± 5.2	Small 57.2 ± 7.9 72.3 ± 9.9 56.5 ± 7.4	Error (ml) 1.1 ± 1.0 1.56 ± 1.09 1.1 ± 0.9	Time (s) 43.4 ± 10.7 46.1 ± 11.7 43.5 ± 10.7
	Prism fusion range total amplitude	Prism bar Risley	CS Normal Reduced Nil Normal	15.3 ± 2.0 16.7 ± 1.6 17.0 ± 1.3 15.0 ± 1.4 16.6 ± 1.6	52.5 ± 3.8 49.4 ± 5.3 51.7 ± 9.9 58.5 ± 5.7 49.3 ± 5.4	66.7 ± 7.1 57.1 ± 7.6 66.4 ± 11.0 72.2 ± 11.0 58.1 ± 7.7	1.0 ± 0.6 1.1 ± 1.0 1.4 ± 0.5 1.5 ± 1.2 1.2 ± 0.9	48.0 ± 11.0 43.2 ± 10.0 44.1 ± 11.0 47.2 ± 12.1 42.3 ± 9.3
	Prism fusion-adjusted positive vergence measure	Prism bar Risley	Reduced Nil Normal Reduced Nil Normal Reduced Nil	17.2 ± 1.6 15.1 ± 1.4 16.7 ± 1.6 17.3 ± 1.5 15.1 ± 1.4 16.7 ± 1.6 16.7 ± 1.6 15.1 ± 1.4	49.8 ± 7.0 58.1 ± 5.7 49.2 ± 5.2 49.8 ± 59.1 59.1 ± 6.7 49.3 ± 5.7 49.5 ± 5.8 58.1 ± 5.7	55.6 ± 9.8 72.2 ± 10.5 57.0 ± 7.6 62.1 ± 10.6 72.0 ± 10.2 58.3 ± 8.4 56.8 ± 7.6 72.2 ± 10.5	0.9 ± 1.0 1.5 ± 1.2 1.1 ± 1.0 1.0 ± 0.7 1.5 ± 1.1 1.4 ± 1.0 0.8 ± 0.7 1.5 ± 1.2	45.6 ± 14.3 45.8 ± 12.8 42.7 ± 10.2 51.7 ± 11.4 45.3 ± 12.3 41.0 ± 9.4 45.8 ± 11.2 45.8 ± 12.8
Suttle et al. (2011) Parameter Mean ± SD Movement time Reaching		SD	Bin 105	ablyopic group socular 66 ± 66 0 ± 25		1	Dom eye 122 ± 45 649 ± 28	Non-don eye 1118 \pm 52 537 \pm 25
	Reach du Time to p Low velo Grasping	eration, ms beak dec, ms city phase, ms	844 511 326	3 ± 52 4 ± 21 5 ± 47		8 5 3	377 ± 43 312 ± 26 355 ± 38	889 ± 46 512 ± 22 364 ± 42
		aperture, ms contact, mm		± 2 ± 1			75 ± 2 54 ± 1	75 ± 2 57 ± 2

Wilson and	Time to peak grip, ms Grip closure time, ms Grip application time, ms Reach-grasp coupling Peak dec-to-peak grip, ms At object contact, ms NR	237 174 66	3 ± 38 7 ± 20 4 ± 16 ± 25 ± 8			603 ± 26 251 ± 21 174 ± 16 88 ± 19 75 ± 5	$654 \pm 34 244 \pm 28 185 \pm 23$ $95 \pm 21 85 \pm 9$
Welch (2013) Alramis et al. (2016) Webber et al. (2016)	ND Bruininks-oseretsky test of motor proficiency Mean (SD)		IS ne motor skills (Fl IS age-standardize	,	ed score	Amblyopic gro 35.80 (4.53) 14.10 (3.37)	ир
Chakraborty et al. (2017)	Movement Assessment Batt for Children (MABC-2) Mean (SD)	Gr	ne motor control poss motor control tal motor control	percen	tile $(n = 587)$	36.3 (27.78) 43.8 (22.12) 37.8 (26.89)	
Fang et al. (2017)	Beery developmental test package (motor coordination task-MC task)		ng the VMI 34–6-year-old	p sl c	or coordination redicting the VMI kills of 4-year-old hildren 0.40 (p < 0.001)	assoc VMI old c	oordination iated with the skills of 5-year- nildren (p < 0.1)
Niechwiej-Szwedo et al. (2017)	ND	2 0.27 (0.001)	2	oo (p 0.001)	5 0.20	(P 0.1)
Thompson et al. (2017)	Bayley Scales of Infant Development (Bayley III) Mean ± SD	Subtest Composite Fine motor Gross motor	subtest	-0.0	ciation with tereoacuity 4 (-0.06, -0.02) 5 (-0.07, -0.03) 2 (-0.04, 0.00)	-0.65 (- -0.94 (-	on with accuity -1.08, -0.21) -1.41, -0.47) -0.54, 0.30)
Zipori et al. (2018)	The Bruininks- Oseretsky Test of Motor Proficiency (Balance subtest)	Amblyopia 9.0 ± 3.1		Stral a 8.6 = Esot	bismus with mblyopia group ± 2.4 ropia	Exotropi	a
Birch et al. (2019a) Birch et al. (2019b) Birch et al. (2020) Hemptinne et al. (2020)	Mean ± SD NR NR NR NR			8.0 =	Ŀ 1.7	9.9 ± 3.7	
Kelly et al. (2020)	Movement Assessment Batte for Children, Second Edi Mean ± SD	•	Total motor Manual dexteri Aiming & catch Balance	-	Amblyopic 7.4 ± 2.7 7.6 ± 2.7 8.8 ± 3.4 7.8 ± 3.1	Non-amblyopic 8.2 ± 2.9 8.5 ± 3.0 9.3 ± 3.2 8.3 ± 3.1	
Niechwiej-Szwedo et al. (2020a)	Bead threading measures Mean \pm SD		Total movemen Peak velocity (r Reach duration Grasp duration Placement dura Reach-to-bead duration (m Reach-to-needle interval dur	m/s) (ms) (ms) tion (maccelerates) decelerates) e accele	(ms) ation interval ation interval ration	3.5 ± 3.1 1551 ± 302 0.886 ± 0.137 411 ± 48 173 ± 78 559 ± 190 182 ± 32 236 ± 34 212 ± 32	
Niaghuiai Sawada	NID		Reach-to-needle	e decele	ration	193 ± 28	
Niechwiej-Szwedo et al. (2020b) Pinero-Pinto et al. (2020)	NR Peabody Developmental Mo Scale-Second Version (Pl Mean ± SD (range)		Static percentile Locomotion per Handling percentile Coordination per Gross motor per Overall motor que Fine motor que Fine motor que Overall motor que	rcentile ntile e ercentile ercentile centile percenti	e e e	72.04 ± 19.90 15.87 ± 11.08 43.43 ± 21.20 73.53 ± 24.16 37.79 ± 18.76 42.40 ± 21.00 56.68 ± 24.33 49.71 ± 22.32 96.81 ± 9.15 104.52 ± 14.90 98.66 ± 14.55	(2.00–50.00) (5.00–95.00) (5.00–99.00) (2.00–84.00) (8.00–95.00) (12.00–99.00) (4.00–96.00) 79.00–124.00) (14.00–151.00)

Sá et al. (2021)	Motor competence assessment (MA Mean (SD) Physical activity questionnaire (I Mean (SD)	Shifting Jumping Standin, Shuttle Ball thre Ball kick MC con Locomo Manipu MCA to	owing velocity (km/h) king velocity (km/h) nponents stability ttor lative	Non-amblyopia group 82.18 (22.33) 50.31 (24.96) 68.46 (23.13) 50.54 (24.21) 64.73 (26.77) 51.94 (29.67) 66.24 (19.36) 59.50 (20.18) 58.33 (22.47) 66.24 (19.36) 2.5 (0.3)	Corrected amblyopia 69.32 ± 31.94 42.70 ± 24.75 57.92 ± 27.61 39.59 ± 27.29 56.51 ± 30.02 51.19 (26.91) 56.01 (23.5) 48.75 (24.40) 53.85 (21.40) 56.01 (23.57) 2.5 (0.3)	Non-Corrected amblyopia 67.29 ± 25.63 36.13 ± 27.68 61.97 ± 23.57 31.26 ± 23.31 50.71 ± 28.90 46.26 (31.64) 51.71 (23.20) 46.61 (20.01) 48.48 (25.19) 51.71 (23.20) 2.4 (0.3)
Vagge et al. (2021)	Developmental Coordination Disorder Questionnaire (DCDQ) 2007 Mean ± SD	Strabismus group (n = 23) Total DCDQ score 58.7 ± 11.3 Control during movement 24.2 ± 6.5 Fine motor 13.3 ± 4.4 General coordination 22.3 ± 3.4	Normal stereopsis group (n = 11) Total DCDQ score 67.3 ± 4.8 Control during movement 28.8 ± 1.8 Fine motor 14.3 ± 4.2 General coordination 24.2 ± 2.2	Absent stereopsis $(n = 12)$ Total DCDQ score 50.8 ± 9.5 Control during movement 19.9 ± 6.3 Fine motor 12.3 ± 4.6 General coordination 20.7 ± 3.6	Amblyopia ($n = 9$) Total DCDQ score 55.4 ± 5.9 Control during movement 22.9 ± 3.3 Fine motor 11.3 ± 3.8 General coordination 21.2 ± 2.8	No amblyopia $(n = 14)$ Total DCDQ score 60.7 ± 13.5 Control during movement 25.0 ± 7.9 Fine motor 14.5 ± 4.9 General coordination 23.1 ± 3.8

CS = central suppression, NR = not reported, VMI = visual motor integration.

signs that define the visual state of a subject (Cacho-Martínez et al. 2014; Cacho-Martínez et al. 2015). However, in the paediatric population, there is a lack of consensus on the diagnostic criteria, mainly in preschool-age children, in whom visual and cognitive abilities are still developing.

The presence of visual disturbances negatively affects the development of motor skills. Children follow a developmental pattern that is highly dependent on the subject's ability to focus, as well as on eye movements and refraction. The studies included in this review established a relationship between amblyopia, binocular vision, accommodative alterations and refractive status with gross and fine motor development in children.

Amblyopia and development

Amblyopia is a decrease in visual acuity without any organic lesion to justify it. The involvement is generally unilateral and occurs as a consequence of a lack of adequate visual stimulation during the critical period of visual development. Amblyopia may affect both eyes, if both have suffered a long period of visual deprivation

(DeSantis 2014), and it is related to the presence of an asymmetric refractive error that has not been detected or treated during childhood.

There is a close relationship between visual acuity deficit and motor delay. Several studies revealed the importance of vision in relation to balance and coordination (Atkinson et al 2005; Chakraborty et al. 2017; Fang et al. 2017; Thompson et al. 2017; Zipori et al. 2018; Hemptinne et al. 2020; Sá et al. 2021), although no study found significant results in this association (Wilson & Welch 2013).

Most of the studies indicated that fine motor skills may be affected if there is any alteration of vision, particularly in the case of amblyopia and strabismus, and can be improved in cases of correct binocular vision (O'Connor et al. 2010; Suttle et al. 2011; Alramis et al. 2016; Webber et al. 2016; Niechwiei-Szwedo et al. 2017; Webber 2018; Kelly et al. 2019; Niechwiej-Szwedo et al. 2020a, 2020b; Vagge et al. 2021). Gross and fine motor skills have been shown to be reduced in children with amblyopia (Engel-Yeger 2008; Webber et al. 2008). Manual dexterity tasks require more time for execution and planning. Reading speed and hand-eye coordination are also

affected (Suttle et al. 2011; Birch et al. 2019a, 2019b; Birch et al. 2020).

The deficiencies in motor performance were greater in manual dexterity tasks, which require speed and precision. Children with amblyopia are slower in planning and executing reaching movements and have a less precise grip than children without amblyopia.

In addition, children with amblyopia present postural instability, which is a consequence of poor static balance (Kelly et al. 2020; Sá et al. 2021). Children with amblyopia are more cautious when walking, take shorter steps and slow down as a result of poor visual processing (Buckley et al. 2010).

Several studies have identified that lower self-perception of peer acceptance and physical competence identity is associated with worse motor skills, which may be related to the wideranging effects of impaired visual development in children with amblyopia in their daily lives (Birch et al. 2019a, 2019b; Birch et al. 2020). One study also revealed that amblyopia can negatively affect children's motor skills, as expressed by objective measures in daily life, whereas self-perception is less affected (Engel-Yeger 2008).

Binocular vision and development

Binocular vision is the ability to integrate two images into one. This requires both eyes to be perfectly aligned on the fixation point. Normal binocular vision positively influences the optimal development of fine motor skills and tasks related to reading. Niechwiej-Szwedo et al. (2017) assessed motor performance in a group of children with reading difficulties by using two tasks: threading beads and pegboard. Children with reading problems had difficulty in the task of threading beads but not with the pegboard. This group performed poorly on a single task that relied heavily on binocular information.

Chakraborty et al. (2017) and Thompson et al. (2017) evaluated binocular vision (visual acuity, stereopsis, alignment of visual axes, ocular motility and self-refraction) and showed that it is strongly related to motor function by using the MABC-2 scales, Peabody Developmental Motor Scale-2nd Version and Bayley Scales of Infant Development, all of which are valid and reliable for measuring infant motor development. In addition, Pinero-Pinto et al. (2020) performed a comprehensive study of binocular vision in a group of typically developing children. They confirmed that children with slower motor development had greater exophoria and a closer convergence point further away, which hindered fusion and binocular vision. In addition, other authors have highlighted the influence of age and affirm that the role of vision in the performance of fine motor skills depends on both the task and age (Alramis et al. 2016; Niechwiej-Szwedo et al. 2020a, 2020b).

Strabismus is an anomaly of binocular vision consisting of the loss of parallelism of both eyes. The lack of binocularity and stereopsis in children with strabismus is associated with the significant impairment of motor skills, particularly for static balance and capture tasks (Hemptinne et al. 2020; Vagge et al. 2021). Furthermore, when normal binocular vision is interrupted in childhood due to strabismus and/or amblyopia, vision and posture are affected, and balance is reduced (Zipori et al. 2018).

Hyperopia and development

Toddlers typically have uncorrected hyperopes (Mayer et al. 2001). Uncorrected hyperopia presents a greater accommodative demand that causes a closure of the visual axes (endophoria) (Leone et al. 2010). A total of 20% of children with high hyperopia (>3.5 D) develop convergent strabismus (Anker et al. 2004; Babinsky and Candy 2013). Atkinson et al. (2005) compared motor skills in hyperopic and emmetropic children by using the MABC-2 as a motor development measurement tool. Hyperopic children performed worse on at least one test in each category (manual dexterity, balance and ball skills). This implies an impairment in fine motor skills in hyperopic children.

This review has several limitations, particularly with regard to the difficulty of extracting data via varied methodologies and different visual systems and motor development assessment tools. Furthermore, there could be some inherent bias due to the professional interests of the authors that are unknown to us. To the best of our knowledge, this is the first review to analyse the relationship between motor development and the visual system. Therefore, this review provides valuable information for the evaluation and treatment of children by professionals from different disciplines in relation to paediatrics.

Conclusions

All included studies confirmed a relationship between the visual system and development in children, although they also demonstrated a lack of uniformity in the methods of visual system measurement and developmental assessment.

References

- Adolph KE & Hoch JE (2019, January 4): Motor development: embodied, embedded, enculturated, and enabling. Annu Rev Psychol 70: 141–164.
- Alramis F, Roy E, Christian L & Niechwiej-Szwedo E (2016): Contribution of binocular vision to the performance of complex manipulation tasks in 5–13 years old visuallynormal children. Hum Mov Sci 46: 52–62.
- Anker S, Atkinson J, Braddick O, Nardini M & Ehrlich D (2004): Non-cycloplegic refractive screening can identify infants whose visual outcome at 4 years is improved by spectacle correction. Strabismus 12: 227–245.
- Atkinson J, Nardini M, Anker S, Braddick O, Hughes C & Rae S (2005): Refractive errors in infancy predict reduced performance on the movement assessment battery for

- children at 3 1/2 and 5 1/2 years. Dev Med Child Neurol **47**: 243–251.
- Babinsky E & Candy TR (2013): Why do only some hyperopes become strabismic. Invest Ophthalmol Vis Sci **54**: 4941–4955.
- Birch EE, Castañeda YS, Cheng-Patel CS, Morale SE, Kelly KR, Beauchamp CL & Webber A (2019a): Self-perception of school-aged children with amblyopia and its association with reading speed and motor skills. JAMA Ophthalmol 137: 167–174.
- Birch EE, Castañeda YS, Cheng-Patel CS, Morale SE, Kelly KR, Beauchamp CL & Webber A (2019b): Self-perception in children aged 3 to 7 years with amblyopia and its association with deficits in vision and fine motor skills. JAMA Ophthalmol 137: 499–506.
- Birch EE, Castañeda YS, Cheng-Patel CS, Morale SE, Kelly KR & Wang SX (2020): Self-perception in preschool children with deprivation amblyopia and its association with deficits in vision and fine motor skills. JAMA Ophthalmol 138: 1307–1310.
- Buckley JG, Panesar GK, MacLellan MJ, Pacey IE & Barrett BT (2010): Changes to control of adaptive gait in individuals with long-standing reduced stereoacuity. Investig Ophthalmol Vis Sci 51: 2487–2495.
- Cacho-Martínez P, Cantó-Cerdán M, Carbonell-Bonete S & García-Muñoz Á (2015): Characterization of visual symptomatology associated with refractive, accommodative, and binocular anomalies. J Ophthalmol 2015: 895803.
- Cacho-Martínez P, García-Muñoz Á & Ruiz-Cantero MT (2010): Do we really know the prevalence of accomodative and nonstrabismic binocular dysfunctions? J Optom 3: 185–197.
- Cacho-Martínez P, García-Muñoz Á & Ruiz-Cantero MT (2014): Is there any evidence for the validity of diagnostic criteria used for accommodative and nonstrabismic binocular dysfunctions? J Optom 7: 2–21.
- Caputo R, Tinelli F, Bancale A, Campa L, Frosini R, Guzzetta A, Mercuri E & Cioni G (2007): Motor coordination in children with congenital strabismus: effects of late surgery. Eur J Paediatr Neurol 11: 285–291.
- CASP CHECKLISTS CASP Critical Appraisal Skills Programme (n.d.).
- Chakraborty A, Anstice NS, Jacobs RJ et al. (2017): Global motion perception is related to motor function in 4.5-year-old children born at risk of abnormal development. Vision Res 135: 16–25.
- Chapman GJ, Scally A & Buckley JG (2012): Importance of binocular vision in foot placement accuracy when stepping onto a floor-based target during gait initiation. Exp Brain Res 216: 71–80.
- DeSantis D (2014): Amblyopia. Pediatr Clin **61**: 505–518.
- Engel-Yeger B (2008): Evaluation of gross motor abilities and self perception in children with amblyopia. Disabil Rehabil **30**: 243–248.
- Fang Y, Wang J, Zhang Y & Qin J (2017): The relationship of motor coordination, visual

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- perception, and executive function to the development of 4–6-year-old chinese preschoolers' visual motor integration skills. Biomed Res Int **2017**: 1–8.
- Goodale MA (2011): Transforming vision into action. Vision Res **51**: 1567–1587.
- Hemptinne C, Aerts F, Pellissier T, Ramirez Ruiz C, Alves Cardoso V, Vanderveken C & Yüksel D (2020): Motor skills in children with strabismus. J AAPOS 24: 76.e1–6.
- Hussaindeen JR, Rakshit A, Singh NK, George R, Swaminathan M, Kapur S, Scheiman M & Ramani KK (2017): Prevalence of non-strabismic anomalies of binocular vision in Tamil Nadu: report 2 of BAND study. Clin Exp Optom 100: 642–648.
- Jang JU & Park IJ (2015): Prevalence of general binocular dysfunctions among rural schoolchildren in South Korea. Taiwan J Ophthalmol 5: 177–181.
- Kelly KR, Morale SE, Beauchamp CL, Dao LM, Luu BA & Birch EE (2020): Factors associated with impaired motor skills in strabismic and anisometropic children. Invest Ophthalmol Vis Sci 61: 43.
- Kelly KR, Morale SE, Wang SX, Stager DRJ & Birch EE (2019): Impaired fine motor skills in children following extraction of a dense congenital or infantile unilateral cataract. J AAPOS Off Publ Am Assoc Pediatr Ophthalmol Strabismus 23: 330.e1–6.
- Leone JF, Cornell E, Morgan IG, Mitchell P, Kifley A, Wang JJ & Rose KA (2010): Prevalence of heterophoria and associations with refractive error, heterotropia and ethnicity in Australian school children. Br J Ophthalmol 94: 542–546.
- Liberati A, Altman DG, Tetzlaff J et al. (2009, July): The PRISMA statement for reporting systematic reviews and meta-analyses of studies that evaluate health care interventions: explanation and elaboration. PLoS Med 6: e1000100.
- Mayer DL, Hansen RM, Moore BD, Kim S & Fulton AB (2001): Cycloplegic refractions in healthy children aged 1 through 48 months. Arch Ophthalmol 119: 1625–1628.
- Moher D, Liberati A, Tetzlaff J, et al. (2009, July): Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. PLoS Med. 6: e1000097.
- Niechwiej-Szwedo E, Alramis F & Christian LW (2017): Association between fine motor

- skills and binocular visual function in children with reading difficulties. Hum Mov Sci **56**: 1–10.
- Niechwiej-Szwedo E, Meier K, Christian L, Nouredanesh M, Tung J, Bryden P & Giaschi D (2020a): Concurrent maturation of visuomotor skills and motion perception in typically-developing children and adolescents. Dev Psychobiol 62: 353–367.
- Niechwiej-Szwedo E, Thai G & Christian L (2020b): Contribution of stereopsis, vergence, and accommodative function to the performance of a precision grasping and placement task in typically developing children age 8–14 years. Hum Mov Sci 72: 102652.
- O'Connor AR, Birch EE, Anderson S & Draper H (2010): Relationship between binocular vision, visual acuity, and fine motor skills. Optom Vis Sci 87: 942–947.
- OCEBM Levels of Evidence Centre for Evidence-Based Medicine (CEBM), University of Oxford (n.d.).
- Page MJ, Moher D, Bossuyt PM et al. (2021): PRISMA 2020 explanation and elaboration: updated guidance and exemplars for reporting systematic reviews. BMJ 372: n160.
- Pinero-Pinto E, Pérez-Cabezas V, De-Hita-Cantalejo C, Ruiz-Molinero C, Gutiérrez-Sánchez E, Jiménez-Rejano J-J, Sánchez-González J-M & Sánchez-González MC (2020): Vision development differences between slow and fast motor development in typical developing toddlers: a cross-sectional study. Int J Environ Res Public Health 17: 3597.
- Sá CSC, Luz C, Pombo A, Rodrigues LP & Cordovil R (2021): Motor competence in children with and without ambliopia. Percept Mot Skills 128: 746–765.
- Suttle CM, Melmoth DR, Finlay AL, Sloper JJ & Grant S (2011): Eye-hand coordination skills in children with and without amblyopia. Invest Ophthalmol Vis Sci **52**: 1851–1864.
- Thompson B, McKinlay CJD, Chakraborty A et al. (2017): Global motion perception is associated with motor function in 2-year-old children. Neurosci Lett **658**: 177–181.
- Vagge A, Pellegrini M, Iester M, Musolino M, Giannaccare G, Ansaldo R & Traverso CE

- (2021): Motor skills in children affected by strabismus. Eye **35**: 544–547.
- Webber AL (2018): The functional impact of amblyopia. Clin Exp Optom **101**: 443–450.
- Webber AL, Wood JM, Gole GA & Brown B (2008): The effect of amblyopia on fine motor skills in children. Invest Ophthalmol Vis Sci 49: 594–603.
- Webber AL, Wood JM & Thompson B (2016): Fine motor skills of children with amblyopia improve following binocular treatment. Invest Ophthalmol Vis Sci 57: 4713–4720.
- Wilson GA & Welch D (2013): Does amblyopia have a functional impact? Findings from the Dunedin Multidisciplinary Health and Development Study. Clin Experiment Ophthalmol 41: 127–134.
- Zimmermann A, de Carvalho KMM, Atihe C, Zimmermann SMV & VLM R (2019): Visual development in children aged 0 to 6 years. Arq Bras Oftalmol 82: 173–175.
- Zipori AB, Colpa L, Wong AMF, Cushing SL & Gordon KA (2018): Postural stability and visual impairment: assessing balance in children with strabismus and amblyopia. PLoS One 13: e0205857.

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Supporting Information

Additional Supporting Information may be found in the online version of this article:

Figure S1. Flow chart.