



Evidence-based clinical practice guidelines for the periodic eye examination in children aged 0–5 years in Canada

Joint Clinical Practice Guideline Expert Committee of the Canadian Association of Optometrists and the Canadian Ophthalmological Society

Walter T. Delperio, MD, FRCSC, (Co-chair)*¹ Barbara E. Robinson, OD, MPH, PhD, FAAO, (Co-chair)^{†,2}

Jane A. Gardiner, MD, FRCSC,^{‡,1,3} Louise Nasmith, MDCM, FCFP, FRCPC (Hon),[§]

Anne Rowan-Legg, MD, FRCPC,^{||} Benoît Tousignant, OD, MSc, MPH, FAAO^{¶,2}

ABSTRACT • RÉSUMÉ

Background: As eye disease before age 5 years is common, some form of vision screening should be performed on children before attending primary school. However, the lack of consistent national recommendations creates confusion for patients, eye care professionals, and governments alike.

Methods: The objective of this document is to provide guidance on the recommended timing, intervals, and types of ocular assessments for healthy children aged 0–5 years. A literature search yielded 403 articles. A multidisciplinary expert committee (comprising 2 optometrists, a comprehensive ophthalmologist, a pediatric ophthalmologist, a family physician, and a pediatrician) independently determined those articles deemed to be key to the clinical question. Articles that were gradable ($n = 16$) were then submitted for independent critical appraisal by an external review group, which provided a Grading of Recommendations Assessment, Development and Evaluation profile of the reviewed articles to use for assigning a grade of evidence.

Recommendations: In addition to routine screening by a primary health care professional, a comprehensive eye examination by an individual with the expertise to detect risk factors for amblyopia—such as an ophthalmologist or optometrist—is required in early childhood. The findings support the importance of early detection of amblyopia before 36 months and no later than 48 months of age via screening with at least 1 comprehensive eye examination before age 5 years.

Conclusions: Vision screening by primary health care providers during routine well-baby/child visits and scheduled vaccinations is an essential part of the detection of ocular disease. However, this early detection potential is limited, and a full oculo-visual assessment is also recommended before the child entering the school system. If amblyopia, strabismus, or other eye pathology is detected or suspected that is beyond the scope of the eye care professional examining the patient, a referral to the appropriate specialist can be made, allowing treatment to be initiated in a timely fashion.

Contexte : Étant donné que les maladies oculaires avant l'âge de 5 ans sont courantes, une certaine forme de dépistage des troubles de la vision devrait être effectuée chez les enfants avant qu'ils ne fréquentent l'école primaire. Cependant, l'absence de recommandations nationales cohérentes crée de la confusion chez les patients, les professionnels des soins oculo-visuels et les gouvernements.

Méthodes : L'objectif de ce document est de fournir des recommandations quant aux types d'examen oculaires à pratiquer chez les enfants en bonne santé de 0 à 5 ans ainsi que sur le moment et la périodicité de tels examens. Une recension des écrits a produit 403 articles. Un comité d'experts multidisciplinaire (composé de deux optométristes, d'un ophtalmologiste effectuant des examens complets de la vue, d'un ophtalmologiste pratiquant en pédiatrie, d'un médecin de famille et d'un pédiatre) a établi de façon indépendante les articles jugés essentiels à la question clinique. Les articles se prêtant à un classement [$n = 16$] ont ensuite été soumis à une évaluation critique indépendante par un groupe externe, lequel a fourni un profil « GRADE » des articles à utiliser et leur a attribué une cote.

Recommandations : En plus du dépistage de routine effectué par les professionnels de première ligne, un examen complet de la vue mené par un professionnel possédant l'expertise nécessaire à la détection des facteurs de risque de l'amblyopie (comme un ophtalmologiste ou un optométriste) est requis durant la petite enfance. Les conclusions confirment l'importance de la détection précoce de l'amblyopie avant 36 mois et au plus tard 48 mois par le dépistage assorti d'au moins un examen complet de la vue avant l'âge de 5 ans.

Conclusion : Le dépistage de la vue effectué chez les bébés et les enfants par les fournisseurs de soins de première ligne au cours des consultations de routine et des vaccinations périodiques est un élément essentiel de la détection des maladies oculaires. Toutefois, le potentiel de détection précoce est limité et un examen oculo-visuel complet est également recommandé avant que l'enfant n'entre à l'école. Si l'amblyopie, le strabisme ou une autre pathologie oculaire est détecté ou soupçonné, et que le problème dépasse le champ de compétences du professionnel qui examine le patient, celui-ci peut être dirigé vers le spécialiste approprié, ce qui permet d'amorcer le traitement en temps opportun.

© 2019 Published by Elsevier Inc. on behalf of
Canadian Ophthalmological Society.

<https://doi.org/10.1016/j.jcjo.2019.09.003>

ISSN 0008-4182



¹ Walter T. Delperio and Jane A. Gardiner are representing the Canadian Ophthalmological Society.

² Barbara E. Robinson and Benoît Tousignant is representing the Canadian Association of Optometrists.

³ Jane A. Gardiner is representing the Canadian Association of Paediatric Ophthalmology and Strabismus.

Table 1—Current Canadian recommendations for vision screening in children

Organization	COS	CAO	CFPC	CPS
Key recommendations	NA	Infants and toddlers should undergo their first eye examination between the age of 6 and 9 months; preschool children should undergo at least 1 eye examination between the ages of 2 and 5 years. ¹	Check red reflex for serious ocular diseases such as retinoblastoma and cataracts. Corneal light reflex/cover–uncover test and inquiry for strabismus. Check visual acuity at age 3 to 5 years. ^{8,9}	Check red reflex for serious ocular diseases such as retinoblastoma and cataracts. Corneal light reflex/cover–uncover test and inquiry for strabismus. Check visual acuity at age 3 to 5 years. Routine comprehensive professional eye examinations of healthy children with no risk factors have no proven benefit. ³

CAO, Canadian Association of Optometrists; CFPC, College of Family Physicians of Canada; COS, Canadian Ophthalmological Society; CPS, Canadian Paediatric Society.

INTRODUCTION

Vision screening and comprehensive eye examinations are recommended throughout life as a method of uncovering treatable asymptomatic ocular disease that may otherwise go undetected.^{1–5} As eye disease before age 5 years is common, family medicine, pediatric medicine, optometry, and ophthalmology have long advised that some form of vision screening should be performed on children before attending primary school (Table 1).^{1,3–9} In addition to various recommendations from national organizations, vision screening recommendations also vary across provinces, and within provinces by county or even by school board district. The Canadian Ophthalmological Society (COS) and the Canadian Association of Optometrists (CAO) recognized that the lack of consistent national recommendations from ophthalmologists and optometrists regarding screening and comprehensive eye examination intervals was creating confusion for patients, eye care professionals, and governments alike. It was further recognized that eye care guidelines should include input from the other key health care professionals involved in primary health surveillance for children, namely, pediatricians and family physicians. COS and CAO thus invited the College of Family Physicians of Canada and the Canadian Paediatric Society to each appoint a representative to an interdisciplinary guideline expert committee to develop recommendations based on evidence and the clinical expertise and practice realities of all representatives.

Ideally, guidelines are flexible tools that are based on the best available scientific evidence and clinical information; they also reflect the consensus of professionals in the field and allow health care professionals to use their individual judgement in managing their patients.¹⁰ Guidelines are not intended to provide a “cookbook” approach to medicine or health care or to be a replacement for clinical judgement¹¹; rather, they are intended to inform patterns of practice. These guidelines should be considered in this context. Adherence to these guidelines will not necessarily produce successful outcomes in every case. Furthermore, these guidelines are not intended to define or serve as a legal standard of medical care¹² and should therefore not be used as a legal resource, as their general nature cannot provide individualized guidance for all patients in all circumstances.¹¹ Standards of medical care are specific to all the facts or circumstances involved in an individual case and can be subject to change as scientific

knowledge and technology advance, and practice patterns evolve. Indeed, health care professionals must consider the needs, preferences, values, and financial and personal circumstances of individual patients, and work within the realities of their health care setting.

The objective of this document is to provide guidance on the recommended timing, intervals, and types of ocular assessments for healthy children aged 0–5 years (e.g., not premature, without chronic systemic disease [e.g., diabetes], without hearing loss or neurodevelopmental disorders). The intended audience is any Canadian health care professional who refers or sees infants and children for an eye examination (i.e., pediatricians, family physicians, primary care providers, ophthalmologists and optometrists, nurses, and nurse practitioners). The recommended intervals of examination will also be of interest to the general public and policy makers. It is acknowledged that there are inequities in human, financial, and health care resources in different regions of the country and that these factors may affect health care professionals’ and patients’ options and decisions. To this extent, these guidelines could be used for advocacy for basic eye care for the pediatric population in underserved areas.

METHODS

These guidelines were systematically developed and based on a thorough consideration of the medical literature and clinical experience of the interdisciplinary health care professionals on the Expert Committee. Where possible, the content of this document was developed in accordance with the Canadian Medical Association *Handbook on Clinical Practice Guidelines*¹¹ and the criteria specified in the 6 domains of the *Appraisal of Guidelines Research and Evaluation II Instrument*.^{13,14} These domains cover the following dimensions of guidelines: scope and purpose, stakeholder involvement, rigor of development, clarity and presentation, applicability, and editorial independence. In addition, the guideline development checklist by Schünemann and colleagues was consulted and applied where applicable.¹⁵

The key clinical question for this guideline was, “What are the optimal times and intervals when children aged 0–5 years should undergo ocular assessment in order to promote optimal eye health?” To answer this question, searches of PubMed/Medline (1995 through April 2018) were

performed by a medical librarian, using appropriate controlled vocabulary and keywords (“amblyopia, refractive error, vision screening, strabismus” combined with variations of the term “comprehensive eye examination”). These searches were further supported by sampling searches of EMBASE, Web of Science, and the Cochrane Library. The searches were limited to children and infants 0–5 years old and published in peer-reviewed journals written in English or French. All studies were included in the search for well-conducted clinical trials and observational studies; studies of amblyopia, amblyogenic risk factors, and refractive error; studies performed in primary care and population-based settings; studies of screening tests typically available in primary care settings (e.g., visual acuity tests, red reflex, and cover test) or examination techniques used by optometrists and ophthalmologists (e.g., retinoscopy); and studies with the following outcomes: improved visual acuity, reduced amblyopia, improved school performance, and quality of life. Results were not restricted to systematic reviews, randomized controlled trials/controlled clinical trials, and observational studies. Searches were updated on a regular basis, and the bibliographies of included studies were checked for further references to relevant studies and papers. (Search strategies are available in Appendix 1. Inclusion and exclusion criteria are available in Appendix 2.)

The literature search yielded 403 articles. Committee members were asked to review article abstracts and independently indicate articles deemed to be key to the clinical question. All articles that were selected by a majority of the committee ($\geq 4/6$ members) as “key” were then reviewed by the co-chairs. Those articles that were gradable ($n = 16$) were then submitted for independent critical appraisal. Other articles that provided context and data regarding the clinical question are cited in the text of this document, but were not used to support recommendations.

Full manuscripts of the abstracts selected by the Expert Committee were examined by an external review group. This group critically appraised each article and reported back to the Expert Committee. Their evaluation included study design and purpose, directness to the study research question, methodological quality, interventions/outcomes of interest, and assessment of potential study biases. They also provided a Grading of Recommendations Assessment, Development and Evaluation (GRADE)¹⁶ profile of the reviewed articles to use for assigning a grade of evidence. The assigned grade for each study was based on criteria for assigning grade of evidence¹⁷ (Appendix 3) from the GRADE Working Group. Range for the quality of evidence is from very low to high. Upon consultation with the critical appraisers, it was agreed that all observational studies would receive the same initial ranking of low as stated in Appendix 3. Articles with high directness to the review question were used to develop the recommendations. The quality of the supporting evidence was used to determine the grade for the recommendations (Appendix 4).^{18,19} The Expert Committee met in person to review the critically appraised articles and to formulate and

grade the recommendations. According to predetermined terms of reference, consensus was required with respect to the wording and grading of each recommendation. The key evidence from the 15 articles that were critically appraised is summarized in Appendix 5. (One article, a qualitative systematic review,²⁰ is not included in Appendix 5, as it did not provide independent evidence.) The recommendations in this guideline are meant to reinforce and complement standards of practice currently recommended by CFCP and Canadian Paediatric Society (Table 1).

The final guideline document was approved by the relevant governing bodies of the COS, the Canadian Association of Paediatric Ophthalmology and Strabismus, and the CAO.

NATURAL HISTORY OF REFRACTIVE ERROR, AMBLYOPIA, AND STRABISMUS

Visual impairment can affect 1%–7% of children, depending on the definition.^{21,22} Some studies report an even higher prevalence of vision disorders in childhood.²³ The most frequent and easily corrected ocular problems, by far, are refractive errors.^{24–26} Refractive errors resulting in anisometropia are the most common risk factor for amblyopia.

Refractive Error

Refractive error is a defect in the ability of the eye to focus on an image accurately. Uncorrected refractive errors may be responsible for as much as 69% of childhood visual impairment.²⁷ Refractive errors result in decreased vision because the image of regard is not focused on the retina. If the axial length of the eye is too short, hyperopia results, whereas if the axial length of the eye is too long, myopia results. If the refracting power of the eye is different in one meridian compared with another, astigmatism results. Depending on the degree of refractive error and the age of the child, uncorrected refractive errors could be potentially amblyogenic.⁶

Amblyopia

Amblyopia is defined as decreased vision, not correctable by glasses, in an otherwise healthy eye. Prevalence in childhood is estimated to range from 1% to 3%, depending on the definition, and is the leading cause of monocular vision loss between age 20 and 70 years.^{28–33} Risk factors include prematurity, neurological disorders, genetic syndromes, and positive family history.^{6,22} A diagnosis of amblyopia is made when there is a 2-line difference in best-corrected vision between eyes. Bilateral amblyopia is considered vision worse than 20/40 in the better seeing eye at age ≥ 4 years of age, or worse than 20/50 at ≤ 3 years of age.^{6,33} The opposite eye can have subtle deficits.^{34,35} Amblyopia is caused by visual deprivation in the amblyopic eye in the ocular developmental period—generally accepted to be until the age of 10 years, although some studies suggest later—leading to structural abnormalities in the brain.³⁶ (Newer studies have challenged

the concept of a complete loss of plasticity in the adult brain,^{37,38} suggesting that vision can be improved after the end of the conventional critical ocular development period^{39–43}; however, early intervention may still result in better vision outcomes.⁴⁴) Amblyopia can result from anisometropia, strabismus, or from deprivation due to an obstruction of the visual axis (e.g., media opacity, obstruction from lid). The remainder of the eye and visual pathway is normal. Approximately 40% of amblyopes have anisometropia, 40% have strabismus, and 20% have a combined mechanism. A small proportion have obstruction of the visual axis.⁴⁵ Anisometropia is an unequal refractive error in each eye, resulting in relative optical defocus. Hyperopic (far-sighted) anisometropia is particularly amblyogenic. High isometropia (equal, but high refractive error) can be amblyogenic in bilateral cases.⁴⁶ A recent pooled report from 2 of the largest population-based pediatric eye disease studies provides estimates of risk of amblyopia with various levels of refractive error and types of strabismus.²¹

Amblyopia is the second most treatable ocular disease (after refractive error) if detected and treated early.^{44,47–50} Overall the benefits of screening and treatment where disease is uncovered outweigh any harm and cost.^{20,51–53} Indeed, treating amblyopia has been shown to be one of the most cost-effective medical procedures in the world.^{54,55} Untreated or insufficiently treated amblyopia may result in life-long impairment in visual function and quality of life. Treatment fails in more than 20% of cases,⁴⁵ and amblyopia can recur after treatment in as many as 25% of cases.^{56,57} Earlier diagnosis may mean more successful treatment.

Strabismus

Strabismus results when the eyes are not aligned. It deprives the visual cortex of simultaneous input from corresponding retinal areas, leading to rivalry and suppression of the input from the nondominant eye. The result is amblyopia

in as many as 50% of cases of strabismus.^{58,59} Treatment of strabismic amblyopia consists of penalizing the “good” eye, although there are newer behavioural therapies—including dichoptic training and perceptual learning^{60,61}—that hold some promise.

VISION SURVEILLANCE IN PRIMARY CARE

Family physicians, general pediatricians, and other primary care practitioners in Canada use the Rourke Baby Record⁸ or the ABCDaire⁹ (in Quebec) to guide their routine health surveillance and examinations of infants and children. Both are based on best evidence and consensus by experts. Specific elements of the eye examination are found in Table 1. The recommendations in this guideline document are meant to reinforce and complement these standards of practice.

SCREENING

A summary of Canadian vision screening recommendations is shown in Tables 1 and 2,^{1–9,62} and highlights the fact that Canadian health care professionals are confronted with inconsistent recommendations. A 2013 survey assessed adherence of family physicians and pediatricians in Ontario to the vision screening guidelines for children as recommended by the CPS and the Rourke Baby Record. From a total of 3000 mailed surveys, 719 completed surveys were included in the analysis (23.5% response rate). Vision screening at every well-child visit was reported to be performed by 65% of family physicians/general practitioners and 52% of general pediatricians. Red reflex was reported to be checked by 94% of physicians in children <3 years of age, but only by 25% for children >3 years of age. Thirty-seven percent of all physicians reported never performing a visual acuity test in any age group.⁶³

In the context of eye care, screening consists of the summary assessment of visual function and ocular anatomy. Screening is

Table 2—Current published vision screening guidelines*

Guideline	0–3 Months	3–6 Months	6–9 Months	3 Years	2–5 Years	6–19 Years	20–39 Years	40–64 Years	56–65 Years	>65 Years
AAP 2003 ⁶²				Screen		Screen	NA			
AAPOS 2012 ^{6,7}	Screen					Every 1 to 2 years	NA			
CADTH 2007 ⁴	Preschool vision screening programs varied from province to province, from public health nurse to full optometric exam; none shown to be superior.				Screen	NA				
CAO 2012 ¹			Eye exam		Eye exam	Annually	Every 2 to 3 years	Every 2 years	Every 2 years	Annually
COS 2007 ²	NA						At least every 10 years	At least every 5 years	At least every 3 years	At least every 2 years
CPS 2009 ³	Screen		Screen		Screen	Screen	NA			
USPSTF 2017 ⁵		Screen			Screen	NA				

*Please see original documents for full details. This table is intended to highlight the differences in scope and recommendations of various guidelines, and is not intended to summarize completely each document.

AAP, American Academy of Pediatrics; AAPOS, American Association for Pediatric Ophthalmology and Strabismus; CADTH, Canadian Agency for Drugs and Technology in Health; CAO, Canadian Association of Optometrists; COS, Canadian Ophthalmological Society; CPS, Canadian Paediatric Society; NA, these guidelines did not address these age groups; USPSTF, United States Preventive Services Task Force.

not meant to be diagnostic; suspicious or positive results are referred to the appropriate professional for diagnosis and treatment.⁶⁴ Basic vision screening performed at the well-baby visit by a family physician or pediatrician can identify treatable vision issues at an early stage.³ In low-risk, asymptomatic children, vision screening in preschool-aged children is aimed at early disease detection for more prevalent conditions such as amblyopia, strabismus, and uncorrected refractive error, as well as for rarer conditions or diseases such as retinoblastoma and congenital cataracts.^{3,62,65} These screening procedures may be performed by pediatricians, family physicians, or other primary care providers during well-baby/child visits.^{64,66} Screening for amblyopia involves screening for risk factors, as they can be diagnosed before amblyopia itself (i.e., before formal vision can be checked). Screening involves checking for refractive error, strabismus, and obstructions to the visual axis. Procedures used to screen for strabismus and amblyopia may include visual acuity testing and the cover/uncover test, while procedures for screening for retinoblastoma and congenital cataracts may include red reflex and fundus testing.^{3,62} Techniques for these procedures can be found in the literature.^{3,62,65,67} Patients with a positive screening test should be referred to an eye care professional for further evaluation.

Simple screening (i.e., a family history of vision problems and any baby concerns, as well as red reflex test and observation of eye movement, lids, and pupils) is quick and easy to perform, but lacks sensitivity and specificity.^{20,51} More rigorous testing is time-consuming and resource-intensive, but provides better sensitivity and specificity.^{20,51–53} Uncorrected refractive error is the most common finding and simplest to correct, but is not easily identified by simple screening.^{24–26} Amblyopia, which affects 2%–4% of the population, can be suspected with basic screening, but requires a full assessment before proceeding with treatment.^{47,53} Although not conclusive, it would appear that the earlier the treatment is initiated, the better the final outcome.^{44,47–50}

COMPREHENSIVE EYE EXAMINATIONS

Comprehensive eye examinations performed by an optometrist or ophthalmologist allow for a fuller assessment that addresses both amblyogenic and nonamblyogenic, yet treatable, ocular disease.^{1,3} This includes, but is not limited to, refractive errors, subtle strabismus, lid and lacrimal disease, and retinal pathology.^{1,2,20,51–53,66} These examinations are meant to be diagnostic and lead to the management of eye conditions, including, but not limited to, amblyopia, strabismus, and uncorrected refractive error.^{1,2,6,66} The main components of a comprehensive eye examination consist of the assessment of refractive status, visual acuity, strabismus/binocular vision/ocular motility, and ocular anatomy (external and internal).^{1,2,6,62,66}

Examination techniques for strabismus and amblyopia in the pediatric population (age 0–5 years) may include fixation assessment; visual acuity; cover/uncover tests; red reflex

(Bruckner method); corneal light reflexes (Hirschberg reflex); sensory fusion (red filter/Worth 4-dot, etc.); stereopsis testing; and ocular motility testing. Examination techniques for refraction assessment may include retinoscopy (static or dynamic); manifest (subjective) refraction; and autorefractometry (not generally used in this age group). Cycloplegic agents (eye drops that inhibit accommodation temporarily) should be used in conjunction with these techniques. Supplemental examination techniques include pupillary testing, visual field testing, intraocular pressure, colour vision testing, funduscopy, and slit lamp or external ocular health assessment.^{1,6,62,66}

REVIEW OF THE EVIDENCE AND RATIONALE FOR RECOMMENDATIONS

The primary goal of the literature review was to find studies that assessed the impact, if any, of vision screening on the prevalence of amblyopia in childhood. Of the 15 key articles included in the summary of findings (Appendix 5), 11 provided evidence related to our study question and recommendations. Performance (quality or efficacy) of screening tests by family physicians, pediatricians, orthoptists, optometrists, or ophthalmologists was not explored specifically.

No masked randomized clinical trial has evaluated the effectiveness of vision screening in children aged 0–5 years; however, prospective cohort studies have provided consistent and strong evidence that vision screening from 8 to 48 months reduces the prevalence of amblyopia at 7–8 years. Four studies from 3 countries (Israel, England, and the Netherlands) were directly related to our main research question and were rated as moderate in overall quality. The Israeli study⁴⁴ was a prospective cohort trial in Haifa that included 808 children who were screened between 1 and 2.5 years of age with a follow-up examination completed at 8 years of age (screening included Hirschberg test [i.e., corneal light reflex test], cover test, and retinoscopy without cycloplegia). The children were screened by members of the Bnai-Zion Ophthalmology Department for amblyopia and amblyogenic risk factors. Children ($n = 782$) in a comparable population, but without early screening, were also examined at 8 years of age. Amblyopia was 2.6 times more likely to be present in the cohort that was not screened (2.6% vs 1.0%, respectively, $p = 0.0098$). Children who were not screened also had more severe amblyopia (1.7% vs 0.1% in screened children, $p = 0.00026$).

Two studies in England from the Avon Longitudinal Study of Parents and Children, also rated as moderate in quality, explored early screening and the prevalence of amblyopia.^{48,49} A randomized trial nested in a prospective cohort compared children who received multiple orthoptic screenings from 8 to 37 months of age (intensive group, $n = 2029$) to children who received only one screening at 37 months of age (control group, $n = 1490$). Amblyopia was less prevalent at 7.5 years of age in the intensive

group (1.45% vs 2.66% in the control group, $p = 0.06$). A major concern regarding this study is that only 54% of the initial intensive group and 55% of the control group were assessed at the 7.5-year examination.⁴⁹ The second study, an observational trial nested in the prospective cohort, examined the impact of orthoptic screening offered at age 4–5 years versus no screening.⁴⁸ The prevalence of amblyopia at 7.5 years was 45% lower in children who received preschool screening than in those who did not (1.1% vs 2.0%, respectively; $p = 0.052$). The power of this study was too low to show statistically significant results regarding the prevalence of amblyopia when the data were adjusted for several potentially confounding factors related to amblyopia.

Studies from the Netherlands also explored early and multiple vision screenings in a single birth cohort. Children born in Rotterdam between September 1996 and May 1997 were followed to the age of 7 years. One study, evaluated as moderate in overall quality, examined the effect of multiple screenings from age 1 to 72 months (2964 of the original Rotterdam Amblyopia Screening Effectiveness Study cohort) on the prevalence of amblyopia at 7 years. There was a 3.4% prevalence of amblyopia at 7 years and a dose–response effect with children who attended more screenings having lower rates of amblyopia.⁴⁷ Another report on this same cohort stated that preschool screening from age 3 years contributed most to amblyopia detection.⁵¹ The authors also reported that refractive error was the most common cause of amblyopia. Neither of these studies included a control group of children who were not screened.

The above studies, from different countries and employing a variety of screening tests at different ages, all reported similar findings: lower prevalence of amblyopia by age 7–8 years in screened versus unscreened children and with multiple screenings versus single screenings.

Ascertaining evidence from published studies regarding the best age at which to screen children was more difficult. Additional studies that explored the importance of age at the time of screening were found, but all were evaluated as low in overall study quality. One article from the Netherlands did not show a difference in rate of referral to ophthalmologists or in prevalence of amblyopia between a screened versus an unscreened cohort at the age of 6–9 months.⁶⁸ This finding may have reflected a problem with the tests or screeners used, rather than the age of children. Referrals in both groups were based primarily on observed strabismus, and 25%–50% of the screeners were found to have inadequately performed the screening tests.

The impact of early referral for treatment on visual acuity outcomes and prevalence of amblyopia was explored in 2 other prospective studies that were evaluated as low in overall study quality. A study of children referred during a screening program in Alaska found that children referred for treatment before 2 years of age had a greater chance of achieving a visual acuity of 6/12; however, this study's results were vulnerable to bias because <25% of potential participants were included in the final outcome assessment.⁵⁰ Atkinson and colleagues reported on 2 Cambridge Infant Screening Programs with a focus on children

with hyperopia.⁵³ The first program screened 3166 children (born from 1981 to 1983) at 7–8 months of age with a follow-up between 1 and 3 years of age and visual acuity testing at 4 years of age. The second program screened 5142 children (born from 1992 to 1994) at 8 months of age and then administered up to 11 follow-up visits by 7 years of age. Both programs reported a decreased prevalence of amblyopia in hyperopic children with early spectacle wear when evaluated at 4 years of age and 7 years of age, compared with hyperopic children who did not wear spectacles.⁵³

Of the 3 cross-sectional studies that were evaluated, all received low or very low overall study quality ratings due to concerns about potential selection bias.^{31,26,69} The studies evaluated as low in overall study quality were from the United States. One study from Tennessee with 5548 children aged 1–6 years found a very high prevalence of amblyopia in children with anisometropia (454/724 or 62.7%).⁶⁹ Donahue also reported that by age 3 years, nearly two-thirds of children with >1.0 diopter anisometropia had amblyopia, and the prevalence of amblyopia increased with age among anisotropic children. The Vision in Preschoolers Study found a high prevalence of unilateral amblyopia (296/3869 or 7.7%) in children aged 3–5 years in Head Start programs.²⁶ In this group, the increased risk of amblyopia was independently associated with the presence of strabismus and significant refractive errors (e.g., myopia, hyperopia, astigmatism, and anisometropia).

A cross-sectional study from Australia was rated as very low in overall study quality, as a large number of children were excluded from the study due to low visual acuity testability.³¹ This study, part of the Sydney Paediatric Eye Study, recruited 2461 children between the age of 6 and 72 months, but results were reported for only 1422 of them. The prevalence of amblyopia was found to be 27/1422 (1.9%) and was significantly associated with hyperopia, astigmatism, anisometropia, and strabismus.³¹

In summary, there is very strong evidence from well-conducted prospective studies that cohorts of children screened at an early age will have a lower prevalence and severity of amblyopia at age 7–8 years, compared with unscreened cohorts.^{44,47–49,51} The age at which the screenings in these studies took place varied, but it is known from other studies that the earlier the detection and treatment of potential risk factors for amblyopia, the better the visual outcomes.^{50,53} Risk factors that must be detected include refractive errors, anisometropia, and strabismus.^{26,31,69}

Based on this review, the Expert Committee concluded that, in addition to routine screening by a primary health care professional, a comprehensive eye examination by an individual with the expertise to detect risk factors for amblyopia—such as an ophthalmologist or optometrist—is recommended in early childhood. Overall, the findings support the importance of early detection of amblyopia before 36 months and no later than 48 months of age via screening with at least 1 comprehensive eye examination before age 5 years.

RECOMMENDATIONS

- Routine age-appropriate screening as recommended by Rourke and ABCDaire (red reflex test, cover/uncover test, and visual acuity) of infants and children by a primary health care provider or pediatrician should continue.^{8,9}
- If an infant or child is identified with an abnormality, they should be referred to the appropriate eye care professional.
- In addition to age-appropriate screening, children aged 0–5 years should undergo ocular assessment by an individual with the expertise to detect risk factors for amblyopia. [1B^{44,47–49}]
 - Ideally, the ocular assessment should occur by age 3 years. [1B^{44,47–49}]
 - The ocular assessment should include refraction and ocular motility evaluation. [1B^{44,47–49}]

LIMITATIONS

The main limitation to the implementation of this guideline may be access and the increased resources required to sustain such a screening process. Further efforts should thus focus on advocating that children have access to oculo-visual assessments that detect treatable eye conditions.

CONCLUSIONS

Vision screening performed by primary health care providers during routine well-baby/child visits and scheduled vaccinations have been—and will continue to be—an essential part of the detection of ocular disease. Obtaining an appropriate history while performing an assessment of the red reflex and examination of the external adnexa provides an opportunity for the early detection of not only amblyogenic pathology but also other potentially vision-threatening (e.g., cataracts, glaucoma) and life-threatening diseases (e.g., retinoblastoma). However, this early detection potential is limited and a full oculo-visual assessment is also recommended prior to the child entering the school system. Although comprehensive eye examinations are possible from birth by certain eye care professionals by adapting techniques, by age 3 years it is expected that the child may be able to cooperate in a complete oculo-visual assessment. This would include visual acuity testing, ocular motility evaluation, slit lamp examination, dilated fundus examination, and cycloplegic refraction. If amblyopia, strabismus, or other eye pathology is detected or suspected that is beyond the scope of the eye care professional examining the patient, a referral to the appropriate specialist can be made, allowing treatment to be initiated in a timely fashion.

SUPPLEMENTARY MATERIALS

Supplementary material associated with this article can be found in the online version at doi:10.1016/j.jcjo.2019.09.003.

REFERENCES

1. Robinson BE, Mairs K, Glennly C, Stolee P. An evidence-based guideline for the frequency of optometric eye examinations. *Prim Health Care*. 2012;2:121.

2. Clinical Practice Guideline Expert Committee. Canadian Ophthalmological Society evidence-based clinical practice guidelines for the periodic eye examination in adults in Canada. *Can J Ophthalmol*. 2007;42:39–45.
3. Amit M. Canadian Paediatric Society. Community Paediatrics Committee. Vision screening in infants, children and youth. *Paediatr Child Health*. 2009;14:246–51.
4. Dunfield L, Keating T. *Preschool Vision Screening* [Technology Report number 73]. Ottawa, ON: Canadian Agency for Drugs and Technologies in Health; 2007.
5. U.S. Preventive Services Task Force, Grossman DC, Curry SJ, Owens DK, et al. Vision screening in children aged 6 months to 5 years: U.S. Preventive Services Task Force recommendation statement. *JAMA*. 2017;318:836–44.
6. American Academy of Ophthalmology Pediatric Ophthalmology/Strabismus Panel. *Pediatric Eye Evaluations Preferred Practice Pattern: I. Vision Screening in the Primary Care and Community Setting; II. Comprehensive Ophthalmic Examination*. San Francisco, CA: American Academy of Ophthalmology; 2012.
7. American Association for Pediatric Ophthalmology and Strabismus. Vision screening recommendations. https://aapos.org/client_data/files/2014/1076_aapos_visscreen.pdf [accessed 27 August 2019].
8. Rourke Baby Record. Evidence-based infant/child health maintenance guide. www.rourkebabyrecord.ca; 2017 [accessed 27 August 2019].
9. ABCDaire. Recommandations concernant le dépistage des troubles de la vue chez les nourrissons et les enfants. <https://enseignement.chusj.org/ENSEIGNEMENT/files/e7/e77f612e-86b5-4e5d-a5d4-950459006a81.pdf> [accessed 27 August 2019].
10. Jacobson PD. Transforming clinical practice guidelines into legislative mandates: proceed with abundant caution. *JAMA*. 2008;299:208–10.
11. Davis D, Goldman J, Palda VA. *Handbook on Clinical Practice Guidelines*. Ottawa, ON: Canadian Medical Association; 2007.
12. Canadian Medical Protective Association. Clinical practice guidelines: what is their role in legal proceedings? *CMPA Perspect*. 2011;(September):3–5.
13. AGREE Enterprise. Appraisal of guidelines research and evaluation II (AGREE II) instrument. <http://www.agreetrust.org> [accessed 27 August 2019].
14. Brouwers M, Kho ME, Browman GP, et al. AGREE Next Steps Consortium. AGREE II: advancing guideline development, reporting and evaluation in healthcare. *CMAJ*. 2010;182:E839–42.
15. Schünemann HJ, Wiercioch W, Etzeandia I, et al. Guidelines 2.0: systematic development of a comprehensive checklist for a successful guideline enterprise. *CMAJ*. 2014;186:E123–42.
16. Guyatt G, Oxman AD, Akl EA, et al. GRADE guidelines: 1. Introduction—GRADE evidence profiles and summary of findings tables. *J Clin Epidemiol*. 2011;64:383–94.
17. GRADE Working Group. <http://www.gradeworkinggroup.org/>; 2017 [accessed 27 August 2019].
18. Guyatt GH, Oxman AD, Vist GE, et al. GRADE Working Group. GRADE: an emerging consensus on rating quality evidence and strength of recommendations. *BMJ*. 2008;336:924–6.
19. Fletcher RH, Fletcher SW, Fletcher GS. *Clinical Epidemiology: The Essentials*. 5th ed. Baltimore: Lippincott Williams & Wilkins; 2014:227.
20. Solebo AL, Cumberland PM, Rahi JS. Whole-population vision screening in children aged 4–5 years to detect amblyopia. *Lancet*. 2015;385:2308–19.
21. Tarczy-Hornoch K, Varma R, Cotter SA, et al. Joint Writing Committee for the Multi-Ethnic Pediatric Eye Disease Study and the Baltimore Pediatric Eye Disease Study Groups. Risk factors for decreased visual acuity in preschool children: the Multi-Ethnic Pediatric Eye Disease and Baltimore Pediatric Eye Disease Studies. *Ophthalmology*. 2011;118:2262–73.
22. Pai AS, Wang JJ, Samarawickrama C, et al. Prevalence and risk factors for visual impairment in preschool children in the Sydney Paediatric Eye Disease Study. *Ophthalmology*. 2001;188:495–500.
23. Drover JR, Kean PG, Courage ML, Adams RJ. Prevalence of amblyopia and other vision disorders in young Newfoundland and Labrador children. *Can J Ophthalmol*. 2008;43:89–94.
24. Irving EL, Harris JD, Machan CM, et al. Value of routine eye examinations in asymptomatic patients. *Optom Vis Sci*. 2016;93:660–6.

25. Huang J, Maguire MG, Ciner E, et al. Vision in Preschoolers Study Group. Risk factors for astigmatism in the Vision in Preschoolers Study. *Optom Vis Sci.* 2014;91:514–21.
26. Pascual M, Huang J, Maguire MG, et al. Vision In Preschoolers Study Group. Risk factors for amblyopia in the Vision In Preschoolers study. *Ophthalmology.* 2014;121:622–9.
27. Varma R, Tarczy-Hornoch K, Jiang X. Visual impairment in preschool children in the United States: demographic and geographic variations from 2015 to 2060. *JAMA Ophthalmol.* 2017;135:610–6.
28. Hendler K, Mehravaran S, Lu X, Brown SI, Mondino BJ, Coleman AL. Refractive errors and amblyopia in the UCLA Preschool Vision Program: first year results. *Am J Ophthalmol.* 2016;172:80–6.
29. Attebo K, Mitchell P, Cumming R, Smith W, Jolly N, Sparkes R. Prevalence and causes of amblyopia in an adult population. *Ophthalmology.* 1998;105:154–9.
30. Friedman DS, Repka MX, Katz J, et al. Prevalence of amblyopia and strabismus in white and African American children aged 6 through 71 months. The Baltimore Pediatric Eye Disease Study. *Ophthalmology.* 2009;116:2128–34.
31. Pai AS, Rose KA, Leone JF, et al. Amblyopia prevalence and risk factors in Australian preschool children. *Ophthalmology.* 2012;119:138–44.
32. McKean-Cowdin R, Varma R, Cotter SA, et al. Multi-Ethnic Pediatric Eye Disease Study and the Baltimore Pediatric Eye Disease Study Groups. Risk factors for astigmatism in preschool children: the Multi-Ethnic Pediatric Eye Disease and Baltimore Pediatric Eye Disease studies. *Ophthalmology.* 2011;118:1974–81.
33. Wallace DK, Repka MX, Lee KA, et al. American Academy of Pediatric Ophthalmology/Strabismus Preferred Practice Pattern Pediatric Ophthalmology Panel. Amblyopia Preferred Practice Pattern. *Ophthalmology.* 2018;125:105–42.
34. González EG, Wong AMF, Niechwiej-Szwedo E, Tarita-Nistor L, Steinbach MJ. Eye position stability in amblyopia and in normal binocular vision. *Invest Ophthalmol Vis Sci.* 2012;53:5386–94.
35. Meier K, Giaschi D. Unilateral amblyopia affects two eyes: fellow eye deficits in amblyopia. *Invest Ophthalmol Vis Sci.* 2017;58:1779–800.
36. Hubel DH, Wiesel TN. Receptive fields, binocular interaction and functional architecture in the cat's visual cortex. *J Physiol.* 1962;106:106–54.
37. Assaf AA. The sensitive period: transfer of fixation after occlusion for strabismic amblyopia. *Br J Ophthalmol.* 1982;66:64–70.
38. Wiesel TN, Hubel DH. Single-cell responses in striate cortex of kittens deprived of vision in one eye. *J Neurophysiol.* 1963;26:1003–17.
39. Taylor V, Bossi M, Bunce C, Greenwood JA, Dahlmann-Noor A. Binocular versus standard occlusion or blurring treatment for unilateral amblyopia in children aged three to eight years. *Cochrane Database Syst Rev.* 2015;8:CD011347.
40. Sato M, Stryker MP. Distinctive features of adult ocular dominance plasticity. *J Neurosci.* 2008;28:10278–86.
41. Evans BJ, Yu CS, Massa E, Mathews JE. Randomised controlled trial of intermittent photic stimulation for treating amblyopia in older children and adults. *Ophthalmic Physiol Opt.* 2011;31:56–68.
42. Levi DM. Perceptual learning in adults with amblyopia: a reevaluation of critical periods in human vision. *Dev Psychobiol.* 2005;46:222–32.
43. Levi DM, Li RW. Improving the performance of the amblyopic visual system. *Philos Trans R Soc Lond B Biol Sci.* 2009;364:399–407.
44. Eibschitz-Tsimhoni M, Friedman T, Naor J, Eibschitz N, Friedman Z. Early screening for amblyogenic risk factors lowers the prevalence and severity of amblyopia. *J AAPOS.* 2000;4:194–9.
45. Pediatric Eye Disease Investigator Group. A randomized trial of atropine vs. patching for treatment of moderate amblyopia in children. *Arch Ophthalmol.* 2002;120:268–78.
46. Barrett BT, Bradley A, Candy TR. The relationship between anisometropia and amblyopia. *Prog Retin Eye Res.* 2013;36:120–58.
47. De Koning HJ, Groenewoud JH, Lantau VK, et al. Effectiveness of screening for amblyopia and other eye disorders in a prospective birth cohort study. *J Med Screen.* 2013;20:66–72.
48. Williams C, Northstone K, Harrad RA, Sparrow JM, Harvey I, ALSPAC Study Team. Amblyopia treatment outcome after preschool screening v school entry screening: observational data for a prospective cohort study. *Br J Ophthalmol.* 2003;87:988–93.
49. Williams C, Northstone K, Harrad RA, Sparrow JM, Harvey I, ALSPAC Study Team. Amblyopia treatment outcomes after screening before or at age 3 years: follow up from randomised trial. *BMJ.* 2002;324:1549.
50. Kirk VG, Clausen MM, Armitage MD, Arnold RW. Preverbal photo-screening for amblyogenic factors and outcomes in amblyopia treatment: early objective screening and visual acuities. *Arch Ophthalmol.* 2008;126:489–92.
51. Groenewoud JH, Tjiam AM, Lantau VK, et al. Rotterdam Amblyopia Screening Effectiveness Study: detection and causes of amblyopia in a large birth cohort. *Invest Ophthalmol Vis Sci.* 2010;51:3476–84.
52. Vision in Preschoolers Study Group. Does assessing eye alignment along with refractive error or visual acuity increase sensitivity for detection of strabismus in preschool vision screening? *Invest Ophthalmol Vis Sci.* 2007;48:3115–25.
53. Atkinson J, Braddick O, Nardini M, Anker S. Infant hyperopia: detection, distribution, changes and correlates – outcomes from the Cambridge Infant Screening Programs. *Optom Vis Sci.* 2007;84:84–96.
54. Membreno JH, Brown MM, Brown GC, Sharma S, Beauchamp GR. A cost-utility analysis of therapy for amblyopia. *Ophthalmology.* 2002;109:2265–71.
55. König HH, Barry JC. Cost-effectiveness of treatment for amblyopia: an analysis based on a probabilistic Markov Model. *Br J Ophthalmol.* 2004;88:606–12.
56. Holmes JM, Beck RW, Kraker RT, et al. Pediatric Eye Disease Investigator Group. Risk of amblyopia recurrence after cessation of treatment. *J AAPOS.* 2004;8:420–8.
57. Holmes JM, Melia M, Bradfield YS, Cruz OA, Forbes B, Pediatric Eye Disease Investigator Group. Factors associated with recurrence of amblyopia on cessation of patching. *Ophthalmology.* 2007;114:1427–32.
58. Von Noorden GK, Dowling JE. Experimental amblyopia in monkeys. II. Behavioral studies in strabismic amblyopia. *Arch Ophthalmol.* 1970;84:215–20.
59. Baker FH, Grigg P, von Noorden GK. Effects of visual deprivation and strabismus on the response of neurons in the visual cortex of the monkey, including studies of striate and prestriate cortex in the normal animal. *Brain Res.* 1974;66:185–208.
60. Hess RF, Thompson B. Amblyopia and the binocular approach to its therapy. *Vis Res.* 2015;114:4–16.
61. Birch EE, Li SL, Jost RM, et al. Binocular iPad treatment for amblyopia in preschool children. *J AAPOS.* 2015;19:6–11.
62. American Academy of Pediatrics. Committee on Practice and Ambulatory Medicine. Section on Ophthalmology. American Association of Certified Orthoptists. American Association for Pediatric Ophthalmology and Strabismus. American Academy of Ophthalmology. Eye examination in infants, children and young adults by pediatricians. *Pediatrics.* 2003;111:902–7.
63. Le TD, Raashid RA, Colpa L, Noble J, Ali A, Wong A. Paediatric vision screening in the primary care setting in Ontario. *Paediatr Child Health.* 2018;23:e33–9.
64. Lennerstrand G, Jakobsson P, Kvarnström G. Screening for ocular dysfunction in children: approaching a common program. *Acta Ophthalmol Scand Suppl.* 1995;214:26–38.
65. Schmidt P, Maguire M, Dobson V, et al. Vision in Preschoolers Study Group. Comparison of preschool vision screening tests as administered by licensed eye care professionals in the Vision in Preschoolers Study. *Ophthalmology.* 2004;111:637–50.
66. American Optometric Association. Comprehensive pediatric eye and vision examination: evidence-based clinical practice guideline. 2017:1–67.
67. Goldbloom R. Pediatric Clinical Skills. 4th ed. Philadelphia, PA: Saunders; 2010.
68. Sloot F, Sami A, Karaman H, et al. Effect of omission of population-based eye screening at age 6–9 months in the Netherlands. *Acta Ophthalmol.* 2015;93:318–21.
69. Donahue SP. Relationship between anisometropia, patient age, and the development of amblyopia. *Am J Ophthalmol.* 2006;142:132–40.

Footnotes and Disclosure:

Expert Committee members disclosed any and all potential conflicts/dualities of interest for the previous 2 years (regardless of whether these relationships had any bearing on the guideline topic). J.A.G. participated in a clinical trial supported by Vertex. The other authors have no affiliation (financial or otherwise) with a commercial organization.

This article is being simultaneously published in the *Canadian Journal of Optometry*. The Expert Committee acknowledges the assistance of Cynthia Lank, BSc, project manager and executive editor, and Mona Frantzke, BSc, MLS, medical librarian. In addition, the Expert Committee acknowledges the independent critical appraisal of the evidence by the following: Ya-Ping Jin, MD, PhD, Department of Ophthalmology and Vision Sciences, University of Toronto; Andrei-Alexandru Szigiato, Hon BSc, MD, research fellow, Glaucoma and Advanced Anterior Segment Surgery,

Department of Ophthalmology and Vision Sciences, University of Toronto; Alex Lai Chi Tam, MSc, Department of Ophthalmology and Vision Sciences, University of Toronto; Sophia Liu, BMSc, Faculty of Medicine, University of Toronto.

This project was jointly funded by the Canadian Ophthalmological Society and the Canadian Association of Optometrists. No industry funding was sought or provided.

*Assistant Professor, Department of Ophthalmology, University of Ottawa, Ottawa, Ont.; †Professor Emeritus, School of Optometry and Vision Science, University of Waterloo, Waterloo, Ont.; ‡Clinical Professor, Department of Ophthalmology and Vision Science, University of British Columbia, British Columbia Children's Hospital, Vancouver, B.C.; §Professor, Department of Family Practice, The University of British Columbia, Vancouver, B.C.; || Consultant Paediatrician, Children's Hospital of Eastern Ontario and Renfrew Victoria Hospital; Assistant Professor, Department of Pediatrics, University of Ottawa, Ottawa, Ont.; ¶Assistant Professor, School of Optometry, Université de Montréal, Montréal, Que.