

PRODUCT

MYOPIA AND EFFECTIVE MANAGEMENT SOLUTIONS

Myopia is becoming a real public health concern across the world.

The number of myopic people is increasing rapidly. The prevalence of high myopia is also expected to rise. Understanding myopia development and methods to slow its progression is currently one of the biggest stakes for researchers and clinicians from around the world. In this paper, a few Vision Scientists at Essilor have put together a general overview of myopia condition. In this article they review the definition of myopia, its evolution and causes. They describe available solutions for myopia management and discuss the relative efficacy for each solution. Finally, they focus on Myopilux®, the specific range of ophthalmic lenses which have been proven to effectively correct and control myopia progression in children.



Dr. Anna Yeo

B Optom (Hons); M App Sc;
PhD, Senior Vision Scientist,
Essilor Center of Innovation
& Technology Asia

Dr. Anna Yeo Chwee Hong joined Essilor R&D Asia in May 2013 as a Senior Vision Scientist after teaching optometry for 23 years at the Singapore Polytechnic. Her current research interest is adult myopia, on which she has conducted research internally at CI&T Asia and in collaboration with other teaching institutions such as Zhongshan University and Singapore and Ngee Ann Polytechnics. She is also a member of the Scientific Committee in Wenzhou-Essilor International Research Centre (WEIRC) for which she helps to review research protocols and scientific publication. Dr. Anna Yeo has been a member of the Optometry and Opticianry Board (OOB) in Singapore and the Chairperson for the Credentials Committee, OOB since 2008.



Dr. Damien Paillé

B Sc Optom; M Sc; PhD, Senior
Vision Scientist, Essilor Center of
Innovation & Technology Europe

Dr. Damien Paillé is a member of Essilor International's optical research and development team, based in Paris, France. Damien holds a degree in optometry and practiced as an optician before completing and defending a doctoral thesis in 2005 in cognitive sciences at the University of Paris VIII in collaboration with the College de France and the Renault company. He then pursued post-doctoral studies at the Laboratory for Perception and Motion Control in Virtual Environments (a joint Renault-CNRS laboratory), before joining Essilor International's research and development team in 2007. He currently works in the Vision Sciences department.



Patricia Koh

Optom; B BioMed; MPH,
Technical Manager, Essilor
Mission Division

Born and raised in Singapore, Patricia is an Optometrist with a background in Biomedical Science and a Master's in Public Health. She joined Essilor R&D Singapore in 2005, focusing on progressive myopia in children and ethnic differences such as postural behavior. In 2014, Patricia moved to Essilor Mission Division as Technical Manager to support the group's social initiatives on training and exploring base of the pyramid innovation.



Dr. Björn Drobe

B Sc Optom; M Sc; PhD, Associate
Director, Wenzhou Medical
University - Essilor International
Research Center (WEIRC)

Dr. Björn Drobe obtained a B.Sc. in Optometry, a M.Sc. in Cognitive Sciences and a Ph.D. in Vision Sciences in Paris, France. He joined the French Essilor Int. research team in 1998, working mainly on the interaction between ophthalmic lenses and the human visual system, as well as on progressive myopia in children. From 2007 to 2013, Dr. Drobe relocated to Essilor R&D Singapore for a higher involvement in myopia research. Since June 2013, he is the associate director of WEIRC (Wenzhou Medical University – Essilor International Research Center), managing an international research team on myopia in children.

KEYWORDS

Myopia, myopia control, myopia correction, high myopia risks, hyperopic defocus, accommodative lag, heredity, lifestyle, blue light, dopamine, atropine, Ortho-K, orthokeratology, prismatic bifocal lenses, multifocal contact lenses, progressive addition lenses, refractive surgery, outdoor light exposure, Myopilux

Although high rates of myopia have been reported in some Asian cities for years, recent publications have highlighted the importance of and increases in this condition throughout Asia, as well as in the US and in Europe. As a result, the number of myopic people is expected to exceed a quarter of the world's population by 2020, or 2 billion people out of a total population of 7.6 billion. The loss of quality in vision, not only affecting daily life, has also raised the biggest concern due to an expected increase in eye pathologies and blindness associated with the severity of myopia. Therefore, it is of great importance to understand myopia development and methods to slow its progression. In this paper we focus on: 1/ Myopia definition, evolution and causes, 2/ Available solutions for myopia management, 3/ Myopilux® new range of ophthalmic lenses for myopic children.

1. Myopia

1.1. A worldwide phenomenon

A recent Asian meta-analysis of 50 studies covering countries from Iran to Japan has reported an average myopia rate of ~28%¹, with strong disparities based on age and geographical region. The highest prevalence is reported among urban young people in Korea, where the rate reaches 96.5% among 19-year-old adults², whereas in Beijing, the prevalence of myopia is 74% among 17- to 18-year-olds.³

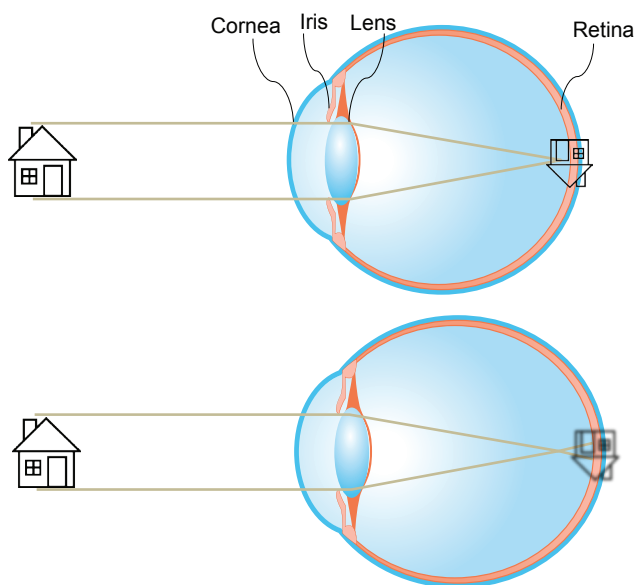


FIG. 1| Emmetropic (top) and myopic (bottom) eyes

On the other hand, the rate is as low as 5.0% among schoolchildren in rural China (5-18 y.o.)⁴ and 10.8% among the 15-year-olds in New Delhi.⁵

In the US, the literature highlights an increase in myopia, with its prevalence among 12- to 54-year-olds increasing from 25.0% between 1971-1972 to 41.6% between 1999- 2004; the highest rate is reported to be 44.0% among 25- to 34-year-olds between 1999-2004.⁶

More recently, in Europe, the prevalence of myopia has been estimated to be 30.6% among 25- to 90-year-olds, with the highest prevalence of 47.2% observed in the 25- to 29- year-old age group.⁷

1.2. What is myopia?

In most cases, myopia occurs because the eyeball is too long relative to the focusing power of the cornea and lens of the eye. This is called axial myopia.

Figure 1 shows an emmetropic eye and a myopic eye. In an emmetropic eye, light rays from far objects are focused on the retina resulting in a clear image. In a myopic eye, light rays from far objects are focused in front of the retina resulting in a blurred image.

In practice, without any correction, a myope experiences blurry vision when looking at far-away objects. The higher the level of myopia, the shorter the distance of clear vision from the eye. Typically, a -2.00 D myope will see clearly at approximately 50 cm, whereas a -5.00 D myope will see clearly only at approximately 20 cm.

1.3. From myopia to high myopia and longer term risks

Myopia is a progressive phenomenon in which onset and strongest progression are mainly reported during childhood.⁸ On average, myopia progression rates are -0.55 D per year among Caucasian children, and at a higher rate of -0.82 D per year among Asian children.⁹ With such a rapid progression during childhood, the risk to become highly myopic in adulthood is high (currently, high myopia is defined as below -6.00 D). In Taiwan, the prevalence of high myopia has increased from 10.9% in 1983 to 21.0% in 2000 among 18-year-old students.¹⁰ In Singapore, the prevalence of high myopia increased from 13.1% between 1996-1997 to 14.7% between 2009-2010 among 17-29-year-old men.¹¹ In Europe, a 5.9% prevalence of high myopia prevalence is reported among 15- to 19-year-olds, according to data collected in 2013.⁷

Although myopia may not have any eye health impact, being highly myopic may have a great impact on ocular health. It has been shown that a -8.00 D myope has 10 times more risk for the development of retinal pathologies than a -4.00 D myope (Fig. 2).^{12,13} High myopia has also been reported to be a risk factor for other ocular pathologies, including glaucoma, choroidal neovascularization, and myopic macular degeneration.¹⁴ Regarding cataract, there are divergent studies on its link with high myopia.¹⁵ Overall, high myopia is a leading cause of visual impairment worldwide.^{16,17}

Therefore, it is of great importance to understand myopia development and to find ways to slow the progression of myopia during childhood.

1.4. Myopia, a multi-factorial refractive error

Myopia development during childhood (onset and progression) is due to multiple factors, which are commonly split into two groups: heredity and lifestyle, often referred to as nature and nurture.

Regarding heredity, it has been shown that children with two myopic parents are on average two to three times more likely to be myopic than children with non-myopic parents.¹⁸ More specifically, genetic studies have identified numerous candidate genes and loci that may contribute to myopia development.¹⁹

Regarding lifestyle, near-vision-demanding tasks and limited time spent outdoors are known to influence myopia development.

Intense near vision activities performed by children have been associated with myopia development in many studies.²⁰⁻²⁴ When looking at a near object, the accommodative response of a myopic child is lower than the proximity of the object, resulting in a slightly defocused image (Fig. 3); light rays from near objects are focused behind the retina. This phenomenon is called the accommodative lag. It has been found to be higher in myopes than in emmetropes.²⁵⁻²⁷

The accommodative lag increases with proximity (Fig. 4) and creates a stimulus for the eye to elongate, leading to myopia progression.^{26,28} The risk of developing myopia increases as the working distance is shorter and the amount of near work is greater.

A large amount of near work combined with a lack of outdoor activities are also highly associated with higher myopia prevalence in children.²⁹⁻³¹ It is still unclear how outdoor activities impact myopia, and several hypotheses have been raised. Recent studies have suggested the existence of interactions between light conditions and myopia development. As light intensities are much higher outdoors than indoors³², pupils are more constricted outdoors. This would result in a greater depth of field and less image blur, resulting in less myopia progression.³¹

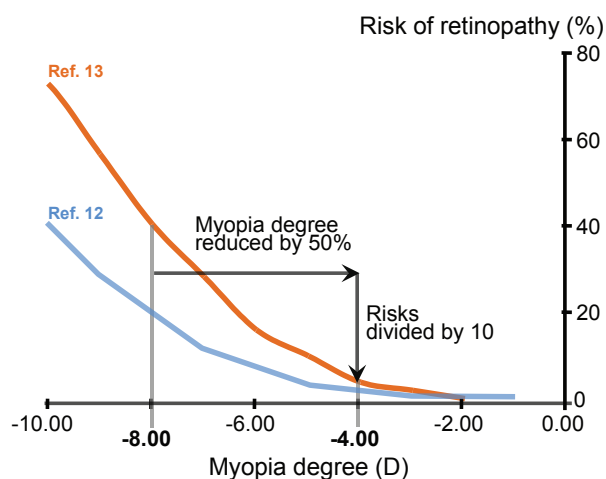


FIG. 2 | Risks of developing retinopathy as a function of myopia degree

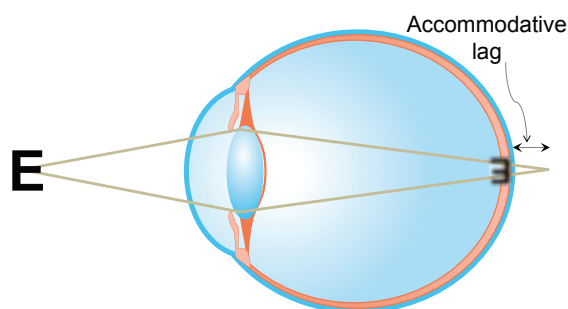


FIG. 3 | The accommodative lag in near vision tasks

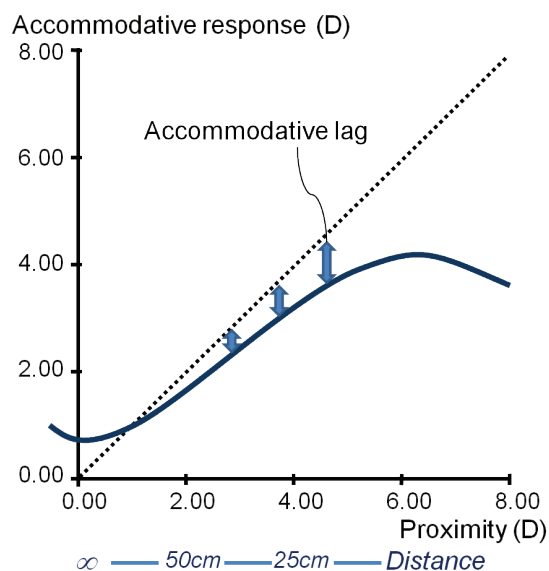


FIG. 4| Influence of proximity on accommodative response

Another hypothesis is the release of dopamine from the retina, which would act as an inhibitor for eye growth, and which is known to be stimulated by blue light in the range of 460-500 nm. With higher amounts of light outdoors, dopamine secretion would prevent the eye from elongating.³³

In practice, modern lifestyle in cities, associated with limited outdoor activities and intense near vision tasks,

favors myopia development. In particular, higher educational levels and hand-held digital device use tend to favor indoor work while exerting a higher demand on our eyes. For instance, research has shown that when using handheld video games, children adopt closer working distances, which in turn may favor myopia onset and progression.³⁴

2. Solutions for myopia management

There are currently several options available to manage myopia. They can be classified according to their ability to correct and slow myopia progression during childhood as shown in Figure 5.

2.1. Solutions that correct myopia but do not control its progression

Single vision lenses are the most common non-invasive solutions for myopia correction. Contrary to common belief, under-correction of myopia does not prevent it from progressing. One study showed that undercorrection of 0.75 D led to a 30% more myopic prescription after 2 years, which was statistically significant.³⁵ Another study showed that undercorrection of 0.50 D led to a 21% more myopic prescription after 1.5 years.³⁶ Other studies also showed that over-correction is not recommended for myopia control either.^{37,38} As a consequence, to correct myopia and to avoid the risk of more rapid myopia progression, full correction should always be chosen based on regular eye examinations.

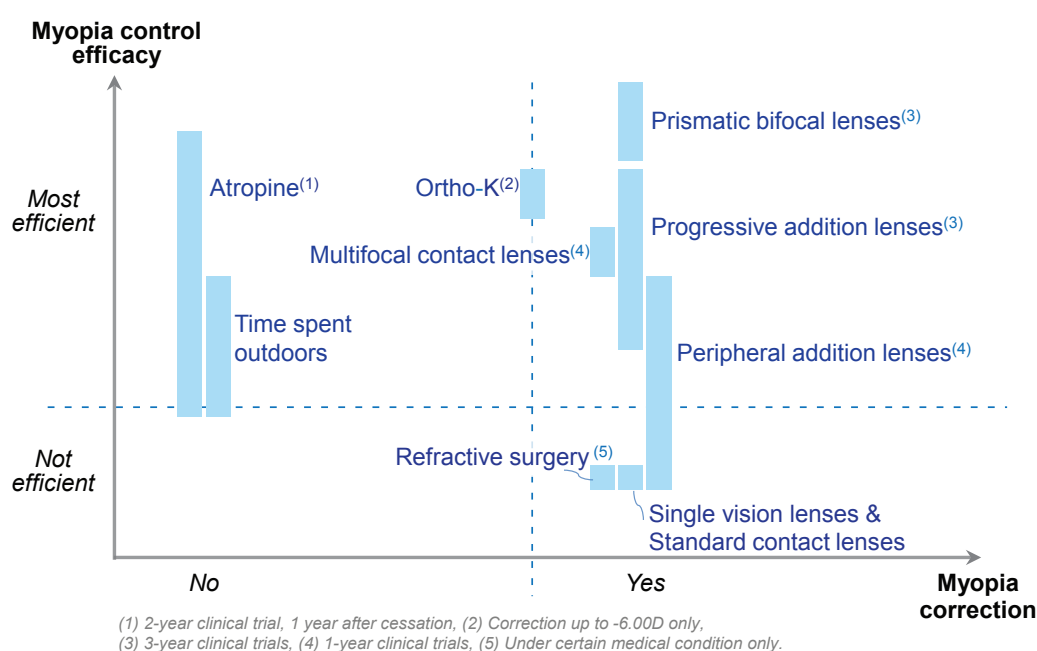


FIG. 5| Solutions for myopia management, classified according to their ability to correct myopia progression

Contact lenses have long been used to correct myopia. However, the clinical efficacy of wearing standard soft contact lenses in myopia control has not been demonstrated.³⁹

As an alternative, refractive surgery, such as LASIK, offers a proven solution for correcting myopia in adulthood. However, the method is invasive and does not control myopia or limit the risks of developing ocular pathologies linked to high myopia. Indeed, refractive surgery modifies the shape of the cornea at the front part of the eye, but it does not change the axial length of the eyeball.

2.2. Solutions that control myopia progression but do not correct it

The least invasive method for myopia control is undoubtedly to increase the time spent outdoors. A meta-analysis performed on the association between time spent outdoors and the risk of developing myopia in children has indicated that spending one hour outdoors per week during childhood reduces the risk of developing myopia by 2%: in other words, a child spending 10 hours more per week outdoors than another child has 20% less chance to become a myope later on.⁴⁰

Atropine eye drops are also used in some countries in clinical practice to slow down myopia progression. Initially it had been suggested that paralyzing accommodation would result in less myopization, but later studies showed alternative mechanisms and sites of action for atropine at either the retina or the sclera.⁴¹ Atropine has thus been studied in several clinical trials. One of them compared several dosages of atropine.⁴² The high dosages (above 0.1%) were efficient during treatment but were associated with a myopic rebound after the cessation of treatment.

The lowest dosage (0.01%) showed a moderate myopia slowing effect that was more sustained after cessation of the treatment. Unfortunately, this study did not include a control group to be able to quantify the effects. Moreover, in addition to its short-term side effects (photophobia due to pupil dilatation, and reduced accommodation power), atropine's long-term side effects have not been documented in children to date.

2.3. Solutions that correct myopia and control myopia progression

Ophthalmic lenses with near vision addition have been shown to be efficient in both correcting and slowing myopia progression and will be detailed in part 3. These lenses have dedicated additional optical power in the near vision zone that compensates for accommodative lag in the myopic eye while the upper part of the lens allows full myopia correction for far vision (Fig. 6). These lenses can either be prismatic bifocal lenses or progressive addition lenses with an addition value and a design adapted to children's physiology. As of today, an addition value of 2.00 D has been shown to be the most efficient compared to lower addition values for myopia control,⁴³ with up to 62% reduction in myopia evolution for prismatic bifocal lenses.⁴⁴

Other ophthalmic lens designs, such as peripheral addition lenses, have also been studied. The elongated shape of myopic eyes results in a defocused image in the periphery even with a perfect central focus (Fig. 7).⁴⁵ It has been shown that this can cause elongation of the eyeball.⁴⁶ Peripheral addition lenses are thus intended to compensate for the peripheral hyperopic defocus and include two visual zones: the central zone of the lens allows full myopia correction and the peripheral zone of

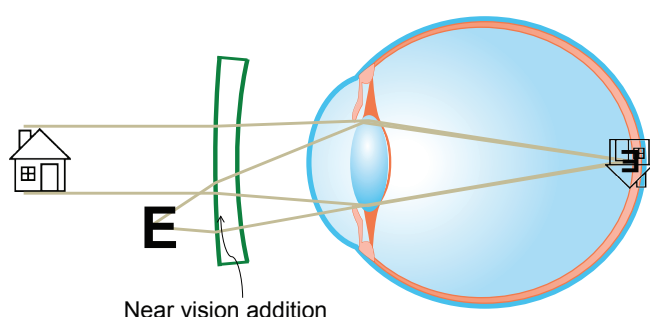


FIG. 6| Near vision addition lenses

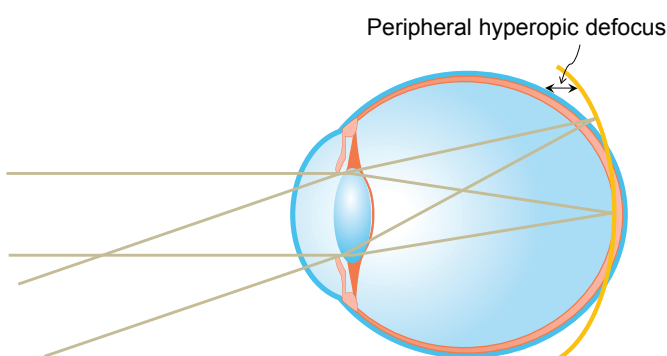


FIG. 7| Peripheral hyperopic defocus

the lens presents a power addition for correcting the hyperopic defocus. In the main study conducted on this concept, no statistically significant differences were observed with the new designs compared to single vision lenses. However, for the subgroup of younger children with at least one myopic parent, myopia progression was reduced by ~30%.⁴⁷ Nonetheless, it was only a one-year study. Moreover, a two-year clinical trial showed that peripheral addition lenses do not enhance the therapeutic efficacy in slowing myopia progression versus near vision addition lenses only.⁴⁸

As an alternative, in recent years, various multifocal contact lenses have been designed to retard the progression of myopia. Two one-year studies have shown a reduction of ~35% in myopia progression with multifocal soft contact lenses.^{49,50} Although these studies showed promising results, there are no available results beyond the first year, thus no evaluation of rebound risks upon the cessation of wearing multifocal soft contact lenses. Several new clinical trials are currently in progress.

Another option is Orthokeratology (Ortho-K), also known as corneal reshaping. The patient wears rigid contact lenses overnight, with a specific reversed geometry; this flattens

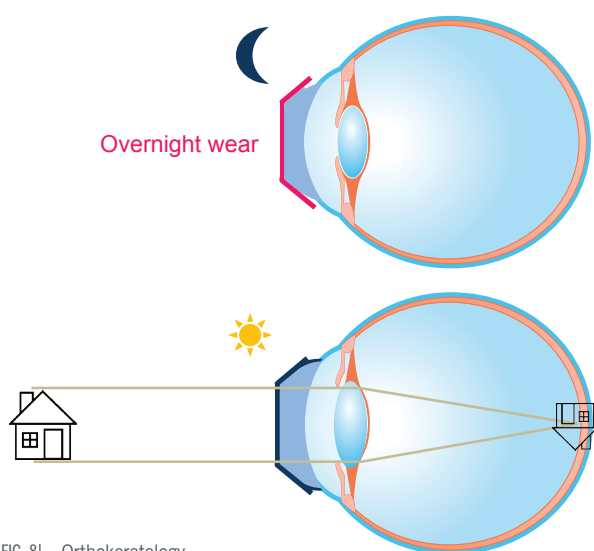


FIG. 8| Orthokeratology

the cornea temporarily to push the focal point back to the retina (Fig. 8). With a proper fitting protocol, Ortho-K can correct myopia up to -6.00 D during daytime. Several recent meta-analyses also showed that Ortho-K slows down myopia progression by approximately 40% with careful education and regular monitoring to ensure safety.⁵¹⁻⁵³ Nonetheless, the long-term efficacy (including a possible rebound effect) as well as the long-term side effects have not been assessed yet and should be evaluated through further large-scale studies.

3. Focus on Myopilux® lenses

Myopilux® is an all-in-one non-invasive range of near vision addition ophthalmic lenses for both myopia correction and myopia control throughout childhood.

3.1. More than 10 years of research

Resulting from more than 10 years of exploratory research by Essilor International myopia experts, Myopilux® lenses are based on a deep understanding of myopic children's natural posture and physiology to ensure good ergonomics and comfortable vision and provide a non-invasive solution for myopia control.

Regarding children's posture, two studies were conducted in China and Singapore. Children were asked to perform their usual reading and writing tasks while their posture was recorded in real time.^{54,55} The results highlighted that when performing near vision activities, children adopt a closer working distance than adults, leading to higher convergence between far and near vision tasks, and that children also prefer to use head over eye declination. These findings were taken into consideration when designing the lateral and vertical positioning of the visual zones in Myopilux® lenses.

Regarding children's physiology, the Myopilux® range has been defined by taking into account children's near phoria: esophoria (tendency to "over convergence"), and exophoria (tendency to "under convergence") (Fig. 9).⁵⁶

When wearing near vision addition lenses, as accommodation drives convergence, the reduction in accommodation will result in less convergence in the eyes, meaning an exophoric shift.⁵⁷

For esophoric profiles, near vision addition lenses will be comfortable because the exophoric shift induced by the addition will partially compensate for their natural esophoria.

However, for exophoric profiles, near vision addition lenses lead to discomfort as they add exophoric shift and require a higher fusional vergence demand. Nonetheless, it has been shown that near base-in prisms can reduce the exophoria induced by near vision addition lenses. More precisely, a 3D base-in prism combined with a +2.00 D near addition on each lens brings visual comfort to the child, with a phoria at its initial state.⁵⁸ It results in an efficient usage of these near vision addition eyeglasses.

3.2. An innovative range of ophthalmic lenses

Based on the above long-term exploration, as well as on sophisticated lens surface calculation methods, high performance production means and efficient methods for controlling lens manufacturing processes, the Myopilux® range of lenses is protected by six Essilor patents and is available in three product versions: Myopilux® Lite, Myopilux® Plus, and Myopilux® Max.

Myopilux® Lite:

Myopilux® Lite lenses are recommended for esophoric children with progressive myopia. Its design includes a progressive optical design, with a recommended addition of +2.00 D for better efficacy in myopia control (Fig. 10).

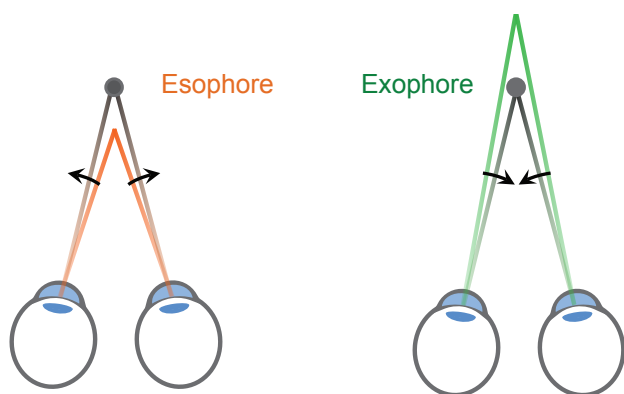


FIG. 9| Near phoria.

The lens is adapted to children's posture; its inset is higher and its progression length is shorter than those for adults. This is to fit to children's closer working distance and preferred usage of head over eye declination (Fig. 11).

Myopilux® Plus:

Myopilux® Plus lenses should be chosen by parents looking for an advanced solution for their esophoric children with progressive myopia. In addition to Myopilux® Lite lenses, it is tailored to each child's specific visual ergonomics and benefits from Wave Technology point-by-point calculation. It ensures tailored lateral positioning of the whole visual zones for enhanced visual comfort and it provides the child with better visual resolution (Fig. 10).

Myopilux® Max:

Myopilux® Max lenses are highly recommended for children whose myopia progression is more than -1.00 D per year. Its design includes a prismatic bifocal made of two wide and aberration-free optical zones separated by a segment line (Fig. 10):

- The upper part of the lens offers the visual correction adapted to the prescription.
- The lower part is dedicated to near vision with an addition of +2.00 D and 3D base-in prism.
- The wide visual zones as well as the short segment height have been designed specifically for children.

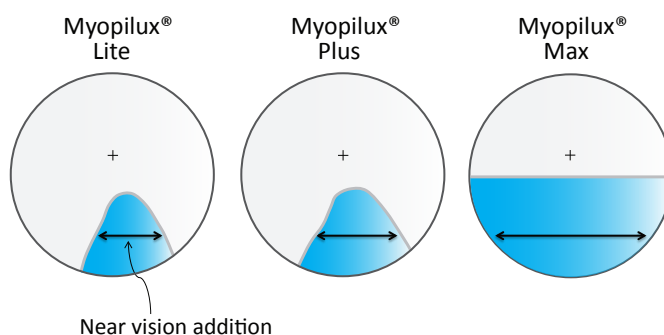


FIG. 10| Near vision zone for Myopilux® Lite (left), Myopilux® Plus (center) and Myopilux® Max (right)

3.3. Validation through clinical trials on 600 children

Myopilux® lenses' concept has been validated through two major clinical trials with approximately 600 children, with third party ethics committees approval.

The concept at the heart of Myopilux® Lite and Myopilux® Plus lenses was tested in the Correction of Myopia Evaluation Trial (COMET) study. The purpose was to evaluate the effect of progressive addition lenses (PALs) compared with single vision lenses (SVLs) on the progression of juvenile-onset myopia.⁵⁹ A total of 469 children were recruited in this study. The children were randomly assigned to either wearing single vision lenses or PALs with +2.00 D addition. The children were monitored for three years with six monthly follow-up visits. The primary outcome measure was progression of myopia, which was determined by auto-refraction after cycloplegia. The retention rate was extremely high with only 1% dropout rate. At the end of three years, the overall PALs group had a statistically significant reduction of 14% in myopia progression compared with single vision lenses (SVLs) that served as a control. However, a better effect of the PALs was observed in esophoric children with high lags of accommodation, whereas there was a statistically significant reduction of 37.2% in myopia progression compared to the SVLs group.⁶⁰

The concept of the Myopilux® Max lens was tested in a 3-year clinical trial. The objective of this study was to determine whether bifocal and prismatic bifocal spectacles control myopia progression in children with high rates of myopia progression compared to SVLs. A total of 135 children aged seven to 13 years old were recruited and randomly assigned to wear SVLs, bifocal and prismatic bifocal lenses. The children were monitored for three years with visits every six months. The primary outcome was cycloplegic auto-refraction and the secondary outcome was axial length growth.

The two-year and three-year results were published in the Archives of Ophthalmology in 2010 and in the Journal of the American Medical Association Ophthalmology in 2014.

At year two, the progression of myopia in children wearing prismatic bifocal lenses was reduced by 55% compared to children wearing SVLs.⁴⁴ This difference was highly significant. The best results were seen in the exophoric group of children; those in the prismatic bifocal group had a reduction of 62% in myopia progression compared with those wearing SVLs.

At year three, children in the prismatic bifocal group had their myopia progression reduced by 51% (Fig. 12).⁶¹ Moreover, contrary to other myopia control spectacle lenses, prismatic bifocals were efficient in slowing myopia progression for all children in different age groups, near phoria types, lag of accommodation or number of myopic parents.

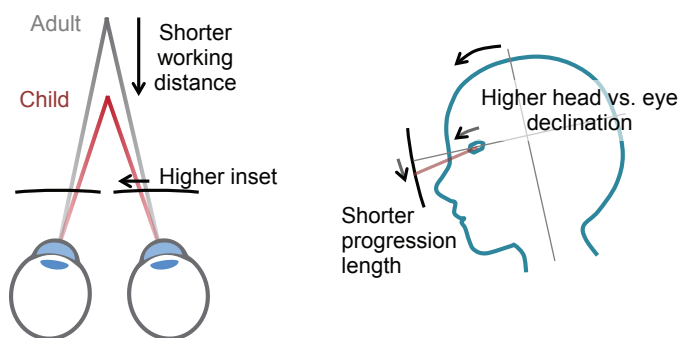


FIG. 11| Child posture

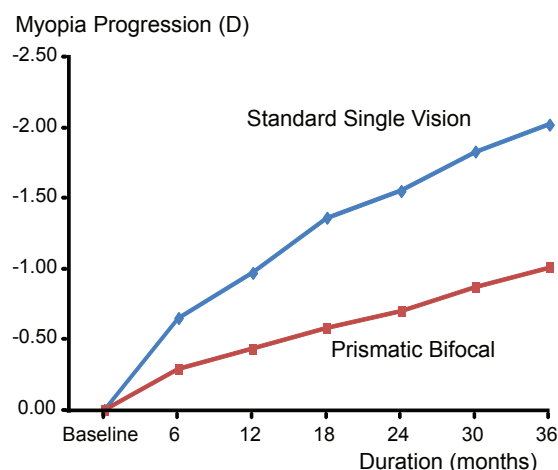


FIG. 12| Myopia progression of children wearing bifocal prismatic addition lenses vs. single vision lenses over three years.

Conclusion

Based on the current scientific state of the art and the scope of clinician's practice, a number of options for myopia correction and myopia control are worthy of consideration. As far as non-invasive solutions are concerned, ophthalmic lenses such as Myopilux®* can be prescribed for effective myopia correction and control.

In terms of protocol, the ideal recommendation would be:

- 1/ Practice eye examinations at least annually
- 2/ Update child corrections when needed
- 3/ In case of ophthalmic lens prescription, choose near vision addition lenses with a design adapted to children needs (see chapter 3.2 for Myopilux® designs)
- 4/ Encourage outdoor activities. •

*Myopilux®: a non-invasive range of near vision addition ophthalmic lenses designed by Essilor for both myopia correction and myopia control. The availability of Myopilux lenses can vary depending on country and should be checked locally by contacting an Essilor representative.



KEY TAKEAWAYS

- Myopia is a progressive phenomenon in which onset and strongest progression are mainly reported during childhood.
- Myopia development during childhood (onset and progression) is due to multiple factors, which are commonly split into two groups: heredity and lifestyle, often referred to as nature and nurture.
- Regarding heredity, it has been shown that children with two myopic parents are on average two to three times more likely to be myopic than children with non-myopic parents.
- Regarding lifestyle, near-vision-demanding tasks and limited time spent outdoors are known to influence myopia development.
- There are currently several options available to manage myopia and they can be classified according to their ability to correct and slow myopia progression during childhood:
 - Solutions that correct myopia but do not control its progression are: single vision ophthalmic lenses, regular contact lenses, refractive surgery
 - Solutions that control myopia progression but do not correct it are: time spent outdoors, atropine eye drops
 - Solutions that correct myopia and control myopia progression are: ophthalmic lenses with near vision addition (such as Myopilux® offer), various multifocal contact lenses and Orthokeratology (Ortho-K).
- Myopilux® is an all-in-one non-invasive range of near vision addition ophthalmic lenses (prismatic bifocal and progressive designs) for both myopia correction and myopia control throughout childhood,
- Resulting from more than 10 years of exploratory research by Essilor International myopia experts, Myopilux® lenses are based on a deep understanding of myopic children's natural posture and physiology to ensure good ergonomics and comfortable vision.

REFERENCES

- 1 Pan CW, Dirani M, Cheng CY, Wong TY, Saw SM. The age-specific prevalence of myopia in Asia: a meta-analysis. *Optom Vis Sci*. 2015 Mar;92(3):258-66.
- 2 Jung SK, Lee JH, Kakizaki H, Jee D. Prevalence of myopia and its association with body stature and educational level in 19-year-old male conscripts in Seoul, South Korea. *Invest Ophthalmol Vis Sci*. 2012 Aug 15;53(9):5579-83.
- 3 You QS, Wu LJ, Duan JL, Luo YX, Liu LJ, Li X, Gao Q, Wang W, Xu L, Jonas JB, Guo XH. Prevalence of myopia in school children in greater Beijing: the Beijing Childhood Eye Study. *Acta Ophthalmol*. 2014 Aug;92(5):e398-406.
- 4 Li Z, Xu K, Wu S, Lv J, Jin D, Song Z, Wang Z, Liu P. Population-based survey of refractive error among school-aged children in rural northern China: the Heilongjiang eye study. *Clin Experiment Ophthalmol*. 2014 May-Jun;42(4):379-84.
- 5 Murthy GV, Gupta SK, Ellwein LB, Muñoz SR, Pokharel GP, Sanga L, Bachani D. Refractive error in children in an urban population in New Delhi. *Invest Ophthalmol Vis Sci*. 2002 Mar;43(3):623-31.
- 6 Vitale S, Sperduto RD, Ferris FL 3rd. Increased prevalence of myopia in the United States between 1971-1972 and 1999-2004. *Arch Ophthalmol*. 2009 Dec;127(12):1632-9.
- 7 Williams KM, Bertelsen G, Cumberland P, et al.; European Eye Epidemiology (E3) Consortium. Increasing Prevalence of Myopia in Europe and the Impact of Education. *Ophthalmology*. 2015 Jul;122(7):1489-97.
- 8 Goss DA, Rainey BB. Relation of childhood myopia progression rates to time of year. *J Am Optom Assoc*. 1998 Apr;69(4):262-6.
- 9 Donovan L, Sankaridurg P, Ho A, Naduvilath T, Smith EL 3rd, Holden BA. Myopia progression rates in urban children wearing single-vision spectacles. *Optom Vis Sci*. 2012 Jan;89(1):27-32.
- 10 Lin LL, Shih YF, Hsiao CK, Chen CJ. Prevalence of myopia in Taiwanese schoolchildren: 1983 to 2000. *Ann Acad Med Singapore*. 2004 Jan;33(1):27-33.
- 11 Koh V, Yang A, Saw SM, Chan YH, Lin ST, Tan MM, Tey F, Nah G, Ikram MK. Differences in prevalence of refractive errors in young Asian males in Singapore during between 1996-1997 and 2009-2010. *Ophthalmic Epidemiol*. 2014 Aug;21(4):247-55.
- 12 Vongphanit J, Mitchell P, Wang JJ. Prevalence and progression of myopic retinopathy in an older population. *Ophthalmology*. 2002 Apr;109(4):704-11.
- 13 Liu HH, Xu L, Wang YX, Wang S, You QS, Jonas JB. Prevalence and progression of myopic retinopathy in Chinese adults: the Beijing Eye Study. *Ophthalmology*. 2010 Sep;117(9):1763-8.
- 14 Morgan IG1, Ohno-Matsui K, Saw SM. Myopia. *Lancet*. 2012 May 5;379(9827):1739-48.
- 15 Pan CW1, Cheng CY, Saw SM, Wang JJ, Wong TY. Myopia and age-related cataract: a systematic review and meta-analysis. *Am J Ophthalmol*. 2013 Nov;156(5):1021-1033.
- 16 Iwase A, Araie M, Tomidokoro A, Yamamoto T, Shimizu H, Kitazawa Y; Tajimi Study Group. Prevalence and causes of low vision and blindness in a Japanese adult population: the Tajimi Study. *Ophthalmology*. 2006 Aug;113(8):1354-62.
- 17 Wu L, Sun X, Zhou X, Weng C. Causes and 3-year-incidence of blindness in Jing-An District, Shanghai, China 2001-2009. *BMC Ophthalmol*. 2011 May 5;11:10.
- 18 Zhang X, Qu X, Zhou X. Association between parental myopia and the risk of myopia in a child. *Exp Ther Med*. 2015 Jun;9(6):2420-2428.
- 19 Simpson CL, Wojciechowski R, Oexle K, et al. Genome-wide meta-analysis of myopia and hyperopia provides evidence for replication of 11 loci. *PLoS One*. 2014 Sep 18;9(9):e107110.
- 20 Saw SM, Wu HM, Seet B, Wong TY, Yap E, Chia KS, Stone RA, Lee L. Academic achievement, close up work parameters, and myopia in Singapore military conscripts. *Br J Ophthalmol*. 2001 Jul;85(7):855-60.
- 21 Saw SM, Hong RZ, Zhang MZ, Fu ZF, Ye M, Tan D, Chew SJ. Near-work activity and myopia in rural and urban schoolchildren in China. *J Pediatr Ophthalmol Strabismus*. 2001 May-Jun;38(3):149-55.
- 22 Vera-Díaz FA, Strang NC, Winn B. Nearwork induced transient myopia during myopia progression. *Curr Eye Res*. 2002 Apr;24(4):289-95.
- 23 Yi JH, Li RR. Influence of near-work and outdoor activities on myopia progression in school children. *Zhongguo Dang Dai Er Ke Za Zhi*. 2011 Jan;13(1):32-5. Chinese.
- 24 Saw SM, Chua WH, Hong CY, Wu HM, Chan WY, Chia KS, Stone RA, Tan D. Nearwork in early-onset myopia. *Invest Ophthalmol Vis Sci*. 2002 Feb;43(2):332-9.
- 25 Abbott ML, Schmid KL, Strang NC. Differences in the accommodation stimulus response curves of adult myopes and emmetropes. *Ophthalmic Physiol Opt*. 1998 Jan;18(1):13-20.
- 26 Gwiazda JE, Thorn F, Bauer J, Held R. Myopic children show insufficient accommodative response to blur. *Invest Ophthalmol Vis Sci*. 1993 Mar;34(3):690-4.
- 27 Yeo AC, Kang KK, Tang W. Accommodative stimulus response curve of emmetropes and myopes. *Ann Acad Med Singapore*. 2006 Dec;35(12):868-74.
- 28 Harb E, Thorn F, Troilo D. Characteristics of accommodative behavior during sustained reading in emmetropes and myopes. *Vision Res*. 2006 Aug;46(16):2581-92.
- 29 Lu B, Congdon N, Liu X, Choi K, Lam DS, Zhang M, Zheng M, Zhou Z, Li L, Liu X, Sharma A, Song Y. Associations between near work, outdoor activity, and myopia among adolescent students in rural China: the Xichang Pediatric Refractive Error Study report no. 2. *Arch Ophthalmol*. 2009 Jun;127(6):769-75.
- 30 Hepsten IF, Evereklioglu C, Bayramlar H. The effect of reading and near-work on the development of myopia in emmetropic boys: a prospective, controlled, three-year follow-up study. *Vision Res*. 2001 Sep;41(19):2511-20.
- 31 Rose KA, Morgan IG, Ip J, Kifley A, Huynh S, Smith W, Mitchell P. Outdoor activity reduces the prevalence of myopia in children. *Ophthalmology*. 2008 Aug;115(8):1279-85.
- 32 Dharanani R, Lee CF, Theng ZX, Drury BV, Ngo C, Sandar M, Wong TY, Finkelstein EA, Saw SM. Comparison of measurements of time outdoors and light levels as risk factors for myopia in young Singapore children. *Eye (Lond)*. 2012 Jul;26(7):911-8.
- 33 McCarthy CS, Megaw P, Devasas M, Morgan IG. Dopaminergic agents affect the ability of brief periods of normal vision to prevent form deprivation myopia. *Exp Eye Res*. 2007 Jan;84(1):100-7.
- 34 Bao J, Drobe B, Wang Y, Chen K, Seow EJ, Lu F. Influence of Near Tasks on Posture in Myopic Chinese Schoolchildren. *Optom Vis Sci*. 2015 Aug;92(8):908-15.
- 35 Chung K, Mohidin N, O'Leary DJ. Undercorrection of myopia enhances rather than inhibits myopia progression. *Vision Res*. 2002 Oct;42(22):2555-9.
- 36 Adler D, Millodot M. The possible effect of undercorrection on myopic progression in children. *Clin Exp Optom*. 2006 Sep;89(5):315-21.
- 37 Goss DA. Overcorrection as a means of slowing myopic progression. *Am J Optom Physiol Opt*. 1984 Feb;61(2):85-93.
- 38 Kushner BJ. Does overcorrecting minus lens therapy for intermittent exotropia cause myopia? *Arch Ophthalmol*. 1999 May;117(5):638-42.
- 39 Walline JJ, Jones LA, Sinnott L, Manny RE, Gaume A, Rah MJ, Chitkara M, Lyons S; ACHIEVE Study Group. A randomized trial of the effect of soft contact lenses on myopia progression in children. *Invest Ophthalmol Vis Sci*. 2008 Nov;49(11):4702-6.
- 40 Sherwin JC, Reacher MH, Keogh RH, Khawaja AP, Mackey DA, Foster PJ. The association between time spent outdoors and myopia in children and adolescents: a systematic review and meta-analysis. *Ophthalmology*. 2012 Oct;119(10):2141-51.
- 41 McBrien NA, Moghaddam HO, Reeder AP. Atropine reduces experimental myopia and eye enlargement via a nonaccommodative mechanism. *Invest Ophthalmol Vis Sci*. 1993;34:205-15.
- 42 Chia A, Chua WH, Wen L, Fong A, Goon YY, Tan D. Atropine for the treatment of childhood myopia: changes after stopping atropine 0.01%, 0.1% and 0.5%. *Am J Ophthalmol*. 2014 Feb;157(2):451-457.
- 43 Leung JT, Brown B. Progression of myopia in Hong Kong Chinese schoolchildren is slowed by wearing progressive lenses. *Optom Vis Sci*. 1999 Jun;76(6):346-54.
- 44 Cheng D, Schmid KL, Woo GC, Drobe B. Randomized trial of effect of bifocal and prismatic bifocal spectacles on myopic progression: two-year results. *Arch Ophthalmol*. 2010 Jan;128(1):12-9.
- 45 Mutti DO, Sholtz RI, Friedman NE, Zadnik K. Peripheral refraction and ocular shape in children. *Invest Ophthalmol Vis Sci*. 2000 Apr;41(5):1022-30.
- 46 Smith EL 3rd, Hung LF, Huang J. Relative peripheral hyperopic defocus alters central refractive development in infant monkeys. *Vision Res*. 2009 Sep;49(19):2386-92.
- 47 Sankaridurg P, Donovan L, Varnas S, Ho A, Chen X, Martinez A, Fisher S, Lin Z, Smith EL 3rd, Ge J, Holden B. Spectacle lenses designed to reduce progression of myopia: 12-month results. *Optom Vis Sci*. 2010 Sep;87(9):631-41.
- 48 Hasebe S, Jun J, Varnas SR. Myopia control with positively aspherized progressive addition lenses: a 2-year, multicenter, randomized, controlled trial. *Invest Ophthalmol Vis Sci*. 2014 Sep 30;55(11):7177-88.
- 49 Sankaridurg P, Holden B, Smith E 3rd, Naduvilath T, Chen X, de la Jara PL, Martinez A, Kwan J, Ho A, Frick K, Ge J. Decrease in rate of myopia progression with a contact lens designed to reduce relative peripheral hyperopia: one-year results. *Invest Ophthalmol Vis Sci*. 2011 Dec 9;52(13):9362-7.
- 50 Anstice NS, Phillips JR. Effect of dual-focus soft contact lens wear on axial myopia progression in children. *Ophthalmology*. 2011 Jun;118(6):1152-61.
- 51 Si JK, Tang K, Bi HS, Guo DD, Guo JG, Wang XR. Orthokeratology for myopia control: a meta-analysis. *Optom Vis Sci*. 2015 Mar;92(3):252-7.
- 52 Sun Y, Xu F, Zhang T, Liu M, Wang D, Chen Y, Liu Q. Correction: Orthokeratology to Control Myopia Progression: A Meta-Analysis. *PLoS One*. 2015 Jun 11;10(6):e0130646.
- 53 Wen D, Huang J, Chen H, Bao F, Savini G, Calossi A, Chen H, Li X, Wang Q. Efficacy and Acceptability of Orthokeratology for Slowing Myopic Progression in Children: A Systematic Review and Meta-Analysis. *Journal of Ophthalmology*, vol. 2015, Article ID 360806, 12 pages, 2015.
- 54 Drobe B, Seow EJ, Bao J, Wang Y, Lu F. Near vision posture in myopic Chinese children. *ARVO Poster*, 2011.
- 55 Seow EJ, Drobe B, Tang FL. Influence of Language and Task on Working Distance in Singaporean Chinese Bilinguals. *ARVO Poster*, 2007.
- 56 Millodot M, 2009, Dictionary of Optometry and Visual Science, 7th edition, Butterworth-Heinemann.
- 57 Jiang BC, Tea YC, O'Donnell D. Changes in accommodative and vergence responses when viewing through near addition lenses. *Optometry*. 2007 Mar;78(3):129-34.
- 58 Cheng D, Schmid KL, Woo GC. The effect of positive-lens addition and base-in prism on accommodation accuracy and near horizontal phoria in Chinese myopic children. *Ophthalmic Physiol Opt*. 2008 May;28(3):225-37.
- 59 Gwiazda JE, Hyman L, Hussein M, Everett D, Norton TT, Kurtz D, Leske MC, Manny R, Marsh-Tootie W, Scheiman M. A randomized clinical trial of progressive addition lenses versus single vision lenses on the progression of myopia in children. *Invest Ophthalmol Vis Sci*. 2003 Apr;44(4):1492-500.
- 60 Gwiazda JE, Hyman L, Norton TT, et al.; COMET Group. Accommodation and related risk factors associated with myopia progression and their interaction with treatment in COMET children. *Invest Ophthalmol Vis Sci*. 2004 Jul;45(7):2143-51.
- 61 Cheng D, Woo GC, Drobe B, Schmid KL. Effect of bifocal and prismatic bifocal spectacles on myopia progression in children: three-year results of a randomized clinical trial. *JAMA Ophthalmol*. 2014 Mar;132(3):258-64.