Myopia Defined
As we know myopia occurs when light from infinity converges to a point in front of the retina. This can occur from 2 main situations:
- The refractive power of the cornea/lens combination is too convergent (too much plus in power)
- The axial length of the eye too long for the refractive elements of the eye

Myopia Classified
Understanding the different classifications of myopia can allow us to consider treatments that can correct the condition and possibly slow or even prevent it. In this section, we will discuss the classification systems that have been proposed to understand myopia [1].

Rate of Progression
One classification system that has been proposed is based on the time and rate of progression of myopia, these categories include [2]:
- **Stationary myopia** occurs when there are typically low (-1.50 to -2.00 D) amounts of myopia and typically occurs prior to the age of 18 and is stable during adulthood
- **Temporarily progressive myopia** typically begins during puberty and will progress until the late 20’s at which point it tends to remain stable
- **Permanently progressive myopia** occurs at a relative rapid rate until around 30 years old, then continues to advance in “jumps” throughout life

Anatomical Causes of Myopia
Another way to think about the classification of myopia is to look at the anatomical causes of the condition. In terms of treatment or optical effects of that treatment, it can be useful to understand the underlying anatomical issues that lead to myopia. These include [3]:
- **Axial myopia** occurs when the eye is too long for its focal length
- **Refractive myopia** occurs when the refractive elements create a focal length that is too short for the length of the eye.
  - Index of refraction – causing myopia when there is a difference from physiological normal leading to a refractive change
  - Curvature – causing myopia when there is an increase in curvature (or shortening of radius) leading to a refractive change
  - Anterior chamber depth – causes myopia when there is a shortening of the anterior chamber depth that leads to myopia

Refractive Myopia
A cataract would be a classic cause of index myopia.
Keratoconus would be a classic cause of curvature myopia
Ciliary effusion secondary to topiramate would be a classic cause of anterior chamber myopia
Theory of Development
The etiology of myopia has been proposed (and have withstood scrutiny) as a way to classify myopia, current thinking includes some overlap of each theory. These theories include:

- **Environmental** – Many studies have found that an increased prevalence of myopia associated with populations who performed significant amounts of near activities [4] [5] [6]
- ** Emmetorization** – In an effort to stabilize distance vision a feedback loop that leads to axial elongation to eliminate retinal defocus (possibly hyperopic defocus) during development [7]. Additionally, this theory has been expanded to include both central and (possibly more importantly) peripheral hyperopic defocus [8] [9].

Night Myopia
Night myopia occurs in situations where there is dim illumination. The main theory is that low light levels cause stimulation of accommodation because distant objects cannot be viewed to cause relaxed accommodation. This causes the accommodative system to be suspended at a distance that is not infinite. This phenomenon occurs secondary to:

1. Increased accommodative response (~0.75 D) is thought to account for the majority of the response [10] [11]
2. Increased chromatic aberration [1]
3. Purkinje Shift [1]

Pseudomyopia
Pseudomyopia is not a true refractive error but can cause a temporary myopic shift due to a ciliary muscle spasm. Manifest refraction will typically be more minus (usually 1 D or more) than retinoscopy. Cycloplegia will reduce accommodation and eliminate the response [1].

Myopia Prevalence and Incidence
While there is a multitude of studies on the prevalence of myopia these studies are not consistent with the amount of myopia they are including as being significant enough to include in the study. For most of these studies myopia is defined as a spherical equivalent refractive error of −0.50 D but some studies will use smaller or larger amounts to define myopia [12].
Prevalence and Incidence

Children

What is the prevalence of American children between the ages of 6 months and 6 years old [13] [14]?

1. 1.2% in non-Hispanic whites
2. 3.7% in Hispanics
3. 3.98% in Asians
4. 6.6% in African Americans

What is the prevalence of myopia in Australian school-aged children of East Asian and European Caucasian ethnicity [15]?

1. East Asian
   • 12-year-olds: 42.7%
   • 17-year-olds: 59.1%
2. Caucasian
   • 12-year-olds: 8.3%
   • 17-year-olds: 17.7%

What is the annual incidence of myopia in Australian school-aged children of East Asian and European Caucasian ethnicity [15]?

1. East Asian
   • 12-year-olds: 6.9%
   • 17-year-olds: 7.3%
2. Caucasian
   • 12-year-olds: 1.3%
   • 17-year-olds: 2.9%

Is there a difference in myopia prevalence between rural and urban areas?

1. Children between the ages of 5-15 in rural northern China showed a prevalence of 16.2% [16].
2. Children between the ages of 5-15 in rural Nepal showed a prevalence of 1.2% [17]
3. Children between the ages of 5-15 in large urban Chinese cities showed a prevalence of 38.1% (Guangzhou) [18] and 36.7% (Hong Kong) [19].

Adults

How does the prevalence of myopia change in adult populations [20]?

1. Adults aged 43–54: 42.9%
2. Adults aged 55–64: 25.1%
3. Adults aged 65–74: 14.8%
4. Adults aged 75+: 14.4%
Myopia Risk Factors
While we still do not fully understand the underlying mechanism of the development or progression of myopia, most of the evidence suggests that myopia results from a combination of genetic and environmental factors. These risk factors will be discussed below.

Family History

What is the impact of parental refractive error on the risk of myopia development in children [21] [22]?

1. If both parents are emmetropic the myopia prevalence for their offspring is 6.3-7.6%
2. If one parent is myopic the myopia prevalence for their offspring is 14.9-18.2%
3. If both parents are myopic the myopia prevalence for their offspring is 32.9-43.6%

Is myopia progression secondary to near work (environmental risk factor) an inherited trait [21]?

1. Researchers did not find that children with myopic parents can inherit a susceptibility to myopia progression due to near work

Socioeconomic Status

What socioeconomic factors increase the prevalence of myopia [23] [24] [25]?

1. Higher levels of education
2. Better housing
3. Higher income
4. Occupations associated with near work

Near Work

What impact do near activities (reading, writing, device use) have on the prevalence of myopia?

1. In one study, children who read more than 2 books per week were 3X more likely to develop myopia (-3.00 D or more) [26].
2. This same study also found that children who read for more than 2 hours per day were no more likely to develop myopia than children who read less [26].
3. Another study found that the association between parental myopia and near work were not significant. This indicates that there is no increased risk associated with near work with an increase in number of parents with myopia. To say this another way, in this study while there was an increased risk of myopia with increasing number of parents with myopia, increasing near work did not increase the risk of myopia [21].
4. The Sydney Adolescent Vascular and Eye Study (SAVES) found that children who became myopic performed 2 more hours of near work per week (19.4 vs. 17.6 hours; P=0.02) than patients who did not become myopic. This effect is less strong the older the patient is [15].

Family Trends
Prevalence in myopia tends to increase in children as the number of parents with myopia increases.

Near Trends
Prevalence of myopia increases with increasing near work. This may be secondary to an increase in axial length during near work or a peripheral hyperopic defocus.
What is the potential mechanism that leads to the relationship between near activities and myopia?

1. In early onset and progressing myopes there is a greater temporary increase in axial length while doing near work than in emmetropic patients [27].
2. Patients with a high lag of accommodation during near work will have a hyperopic retinal defocus and that may lead to myopia progression [28].

Outdoor Activity

What impact do outdoor activities have on the prevalence of myopia?

1. More hyperopic refractions and lower myopia prevalence was found in 12-year-olds with higher levels of outdoor activity (sports or leisure) [29].
   - Children with high levels of near work AND low levels of outdoor activity had the least hyperopic mean refraction
   - Children with low levels of near work AND high levels of outdoor activity had the most hyperopic mean refraction
   - The odds of developing myopia were the lowest in children reporting the highest levels of outdoor activity
   - No difference between males and females

2. In younger (6-year-old) patients there was no association between refraction and outdoor activity [29].

3. Another study to evaluate outdoor activity in 7-12 year old children risk of myopia progression and onset found [30]:
   - **Design**
     i. 571 students: 333 interventional, 238 control
     ii. Recess time: 80 minutes/day
     iii. Outdoor P.E.: 2 hours/wk
     iv. Patients on atropine therapy prior to and throughout the study:
        1. 29% of intervention
        2. 20% of control
   - **Results:**
     i. Myopia Onset After 1 year:
        1. 8.41% - Interventional
        2. 17.61% – Control
        3. NNT - 10
     ii. Myopia Progression After 1 year:
        1. -0.25D/year – Interventional
        2. -0.38D/year – Control

Outdoor Trends

Prevalence of myopia decreases with increasing outdoor activities. The “light-dopamine” theory was proposed as a possible mechanism for this effect.
How much time should children spend outdoors to make an impact on myopia?

1. A meta-analysis of 7 studies found that the odds of developing myopia is reduced by 2% per additional hour of time spent outdoors per week [31].

What is the potential mechanism that leads to the inverse relationship between outdoor activities and myopia?

1. One theory is that the increased light intensity during outdoor activities leads to an increase in dopamine release. It is hypothesized that dopamine reduces axial elongation leading to lower prevalence of myopia [29] [32].

Treatment and Management of Myopia Progression

Undercorrection

During development, the process of emmetropization must occur. During this process, retinal blur or image defocus ought to provide a feedback mechanism that guides the growth of the eye to eliminate refractive error [33]. In order for emmetropization to work perfectly, this feedback mechanism must be able to discern between the blur associated with myopia and blur associated with hyperopia.

One possible theory on why emmetropization occurs is that the visual system interprets retinal defocus as either myopic or hyperopic, thus undercorrecting myopia should halt or slow the progression of the condition.

Does undercorrecting myopia help slow the progression of myopia [34]?

1. 94 myopic (-0.50 or greater with less than -2.00 cyl) children between the ages of 9 and 14 were divided into 2 groups and followed for 2 years:
   a. Fully corrected
   b. Undercorrected to 20/40 (~ +0.75 D added to BSCVA)

2. After 2 years of monitoring every 6 months the study found:
   a. Myopia progression
      i. Fully corrected: −0.77 D
      ii. Undercorrected: −1.00 D (statistically significant)
   b. Axial length increase
      i. Fully corrected: 0.68 mm
      ii. Undercorrected: 0.60 mm (statistically significant)
Bifocal and Progressive Addition Lenses (PALs)
Utilizing a bifocal or PAL in slowing myopia progression may work by eliminating or reducing the accommodative demand or lag associated with near tasks [35] [36].

How do bifocal glasses impact myopia progression in patients with near esophoria [37]?
1. In a single masked randomized trial, 82 myopic children with near esophoria were randomized into 2 groups (single-vision lenses or bifocals with a +1.50 add) and followed for 2.5 years.
2. After 30 months of treatment the study found the following average myopia progression:
   a. Single-vision: $-1.24$ D
   b. Bifocals +1.50 add: $-0.99$ D (statistically significant)

How does the addition of BI prism with executive bifocals impact myopia progression [38]?
1. 135 children (age 8-13 years) with progressive myopia (-0.50 D in the preceding year) were randomized into 3 groups (single-vision lenses, +1.50-D executive bifocals or +1.50-D executive bifocals with 3-Δ base-in prism in the near segment of each lens)
2. After 36 months of treatment the study found the following:
   a. Myopia progression:
      i. Single-vision: $-2.06$ D
      ii. Executive bifocals: $-1.25$ D
      iii. Executive bifocals and BI prism: $-1.01$ D
   c. Average axial length increase:
      i. Single-vision: 0.82 mm
      ii. Executive bifocals: 0.57 mm
      iii. Executive bifocals and BI prism: 0.54 mm

How do PALs impact myopia progression in patients with near esophoria and high accommodative lag [39]?
1. In a double-masked multicenter randomized trial, 118 myopic (spherical equivalent of -0.75 to -2.50) children with near esophoria (≥2 PD) and a high accommodative lag (accommodative response less than 2.50 D for a 3.00-D demand) were randomized into 2 groups either single-vision lenses or PALs with a +2.00 add and followed every 6 months for 3 years.
2. After 36 months of treatment the study found the following average myopia progression:
   a. Single-vision: $-1.15$ D
   b. PALs +2.00-D add: $-0.87$ D (statistically significant)
c. Additionally, patients with large fusional vergence ranges had a greater treatment effect.

**Soft Multifocal Contact Lenses**
Clinically we typically utilize soft bifocal contact lenses for presbyopic patients as an alternative to monovision to help reduce the need for reading glasses. Center distance soft multifocal lenses also create peripheral myopic defocus which may slow myopic progression by reducing the stimulus (peripheral hyperopic defocus) for axial elongation [40].

**What effect do center distance soft multifocal contact lenses have on myopia progression and axial elongation?**

1. 40 myopic (-1.00 D to -6.00 D spherical component and less than 1.00 D cyl) children (8-11 years old) fitted with soft multifocal contact lenses with a **+2.00 D add (Proclear Multifocal "D"; CooperVision)**. These patients were then age- and gender-matched to control participants from a previous study who were fitted with single-vision contact lenses (1 Day Acuvue; Vistakon). They were monitored annually for 2 years.

2. After 24 months of treatment the study found the following:
   a. Average SE myopia progression:
      i. Single-vision: -1.03 D
      ii. Multifocal: -0.51 D (statistically significant)
   b. Average axial elongation:
      i. Single-vision: 0.41 mm
      ii. Multifocal: 0.29 mm (statistically significant)

**What effect do center distance soft multifocal contact lenses have on myopia progression and axial elongation in patients with an eso fixation disparity at near?**

1. In a prospective, randomized, clinical trial, 86 children (8 to 18 years) with progressive myopia (-0.50 D or more since previous examination) and an eso fixation disparity at near were randomized to wear single-vision soft contact lenses (Acuvue 2) or bifocal soft contact lenses (Acuvue Bifocal) with enough add power to neutralize the associated phoria and followed for at 6 months and 1 year.

2. After 12 months of treatment the study found:
   a. Average SE myopia progression:
      i. Single-vision: -0.79 ± 0.43 D
      ii. Bifocals: -0.22 ± 0.34 D (statistically significant)
   b. Average axial elongation:
      i. Single-vision: 0.24 ± 0.17 mm
      ii. Bifocals: 0.05 ± 0.14 mm (statistically significant)

**Orthokeratology**
Orthokeratology (ortho-k) is utilized to temporarily reduce the need for daytime vision correction by flattening the central cornea. Similar to multifocal soft contact lenses, it is thought that the underlying mechanism for slowing myopia progression is by creating peripheral myopic defocus.
which may slow myopic progression by reducing the stimulus (peripheral hyperopic defocus) for axial elongation [43].

**What effect does orthokeratology have on axial elongation [44]?**

1. In a single-masked randomized clinical trial 102 myopic (between 0.50 and 4.00 diopters (D) and astigmatism not more than 1.25D) children (6 to 10 years) were randomized to ortho-k lenses or single-vision glasses and monitored every 6 months for 2 years.
2. After 24 months of treatment the study found:
   a. Average axial elongation:
      i. Single-vision spectacles: 0.63 ± 0.26 mm
      ii. Ortho-K: 0.36 ± 0.24 mm (statistically significant)

**Topical Pharmaceutical Agents**

Anti-muscarinic topical pharmaceutical medications used to dilate the pupil or paralyze accommodation have also been used in an attempt to control myopia progression. The theory of how these medications slow the progression of myopia is unclear. It has been hypothesized that the mechanisms may include a reduction in accommodation or through inhibition of the muscarinic receptors in the ciliary body and iris but the evidence for both of these hypothesis are conflicting.

The medications that have been evaluated for myopia control include atropine and pirenzepine. One advantage of pirenzepine over higher doses of atropine is that pirenzepine has action only on M1 anti-muscarinic receptors, which are not as prevalent in the iris or ciliary body, so pirenzepine does not dilate the pupil or reduce accommodation as much as atropine. For the purposes of this course we will focus on atropine because it is a medication that has a long track record utilization in eye care and lower doses reduce the side effect profile as well.

**What effect does atropine have on myopia progression [45]?**

1. In ATOM1 a parallel-group, placebo-controlled, randomized, double-masked study of 400 myopic (SE -1.00 to -6.00 (D) and -1.50 DC or less) children (6-12 years) and were randomized to receive either 1% atropine or saline drops 1 time per day in one eye and followed for 2 years.
2. After 24 months of treatment the study found myopia progression of:
   a. Placebo: -1.20 ± 0.69 D
   b. 1% atropine: -0.28 ± 0.92 D (statistically significant)

**After stopping atropine is there a rebound of myopia progression [46]?**

1. Utilizing the same cohort as above, this study followed them for an additional year after stopping 1% atropine to determine the extent, if any, rebound myopic progression occurred.
2. 12 months after stopping 1% atropine treatment this study found:
   a. Average myopia progression in final 12 months:
      i. Placebo: -0.38+/-.039 D
      ii. 1% atropine: -1.14+/-.080 D
   b. Average myopia progression in total 36 months (24 months of treatment, 12 months without treatment):
i. Placebo:  -5.22±/-1.38 D  
ii. 1% atropine:  -4.29±/-1.67 D

Can we minimize side effects and still reduce myopia progression with lower doses of atropine [47]?

3. In the ATOM2 study patients and were randomized to receive either 0.5%, 0.1% or 0.01% atropine since there was already proven benefit of atropine the control arm was utilized from ATOM1 since it would not be ethical to prevent patients from receiving treatment that was shown to be effective in the prior study. Medication was administered 1 time per day in each eye and patients were followed for 2 years.

4. After 24 months of treatment the study found myopia progression of:
   a. 0.5% atropine:  -0.30 D  
   b. 0.1% atropine:  -0.38 D  
   c. 0.01% atropine:  -0.49 D

5. During the study, photochromic glasses with a reading add were prescribed if patients complained of glare or difficulty reading. The side effect profile was less with lower doses of atropine and while the 0.01% atropine group had the greatest rate of myopia progression, accommodative amplitudes, visual acuities and pupil sizes were all less impacted than those in the other treatment groups. Further, the study found that the number of children complaining of near blur and requiring a near add was:
   a. 0.5% atropine:  70%  
   b. 0.1% atropine:  61%  
   c. 0.01% atropine:  6%

EyeCode Clinical Pearls

1. Studies suggest that the visual system of myopes have an abnormal mechanism for detecting the direction of retinal image blur leading to axial elongation and myopia progression in the presence of myopic retinal defocus.

2. Undercorrection of myopia should not be utilized as a strategy for slowing myopia progression.

3. While myopia progression is slowed with bifocals spectacles and progressive addition lenses in a statistically significant way, I would argue that reducing myopia by 0.25 D over 2.5 years (for FT) and 0.28 D over 3 years is not clinically significant.

4. Executive bifocal spectacles can slow myopia progression in children by approximately 40% over 3 years. It is unclear whether the benefit seen in this study is due to an executive BF (compared to studies using a FT) or if the reason there was a more significant slowing of myopia progression because the study used patients who were already progressing in their myopia as an inclusion factor.

5. There may be some additional benefit of utilizing BI prism at near to slow myopia progression more than executive BFs alone in patients with low accommodative lags (less than 1.01 D).

6. Center distance soft multifocal contact lenses can reduce both myopia progression (by 50%) and axial elongation (by 29%) over 2-years.
7. Center distance bifocal contacts showed approximately a 70% reduction in myopia progression and an 80% reduction in axial elongation in children with an eso fixation disparity at near after 1 year of treatment. While rarely done in clinical practice, factoring in near fixation disparities and prescribing add powers to neutralize the associated phoria may slow myopia progression more than utilizing a generalized add power.

8. Ortho-K contact lenses can reduce axial elongation (by 43%) over 2-years.

9. The rate of microbial keratitis with overnight ortho-k lenses is similar to that of overnight soft contact lens wear [48], so it is prudent to ensure the benefits outweigh the risks prior to initiating overnight wear with any contact lenses. Clinically, it may be wise to ensure that patients can safely and responsibly wear daytime contact lenses prior to initiating overnight wear options.

10. When using 1% atropine for myopia control expect a more rapid progression in myopia after stopping treatment.

11. There is a significant benefit of reducing myopia progression with 1% atropine even though the effect is not as great once atropine is stopped.

12. To minimize side effects, consider utilizing 0.01% atropine for myopia control.

Bibliography


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