INTRODUCTION

Based on classical studies on adult eyes performed by Tron\(^1\) and Stenstrom,\(^2\) it has been generally accepted that among the ocular components of the eye, axial length (AL) shows the strongest correlation with refractive error, whereas corneal power and lens power show much weaker correlations. In the mentioned studies, the distribution of the optical components was normal, contrasting with the tight, leptokurtic distribution of refraction (i.e., a more acute peak around the mean and flatter tails). Since random association of normally distributed optical components would not produce the observed distributions of refraction, these early studies led to the concept of an active process of...
emmetropization, where the optical components of refraction appear to develop in concert.

In particular, active matching of AL to corneal power has been implicated since the ratio of AL to corneal radius (CR) of curvature (AL/CR) shows a more leptokurtic distribution and a stronger correlation with refraction than does AL by itself. In contrast, the role of changes in lens power in determining the refractive state of the eye at rest has received little emphasis, although François and Goes in 1977 reported that lens power may be involved after they studied the ocular components of 100 emmetropic eyes with the newly developed ultrasonic biometry.

Studies on refractive development in children have confirmed the importance of AL and the AL/CR ratio, and in addition have considered changes in lens power. Sorsby et al., had showed that from the age of three, the lens lost power with age as the eye increased in AL. They also found that changes in corneal power had little impact on refractive development after the age of three. They also noted in cross-sectional data on subjects aged 20-60 that eyes in the emmetropic range had a normal distribution of ALs and that there lens power was lower in longer eyes. Their findings therefore supported an important role for changes in lens power in the final determination of refractive status. The importance of axial elongation and the contribution of reductions in lens thickness and lens power during the development of refraction during childhood has subsequently been confirmed by Larsen and Goes and more recently by Mutti et al.

Gordon and Donzis analyzed ocular components in 148 eyes, from newborn to 36-year-old adults, and found an age-related increase in AL with reductions in corneal and lens powers, with the most dramatic changes occurring in the 1st year or two of life. Subsequent studies have confirmed the rapid development of kurtosis in the distribution of refraction postnatally. Thus, the overall picture of refractive development during childhood is that of rapid matching of AL to the power of the refractive surfaces, followed by continuing axial elongation which can be balanced, at least in some cases, by reductions in lens power. As a result, in young adults, hyperopic eyes have, on average, slightly higher lens power than emmetropic or myopic eyes, and in the population as a whole (if mainly composed of emmetropic eyes), shorter eyes have, on average, higher lens power than longer eyes.

The lens thins and flattens until around the age of 10. From this age onward, lens curvatures steepen (and the lens becomes axially thicker), which might be expected to produce a more powerful lens and a myopic shift in refraction with ageing. However, quite substantial hyperopic shifts in refraction are commonly observed in adults, even with cycloplegic refraction. The conflict between these two observations has become known as the lens paradox, which could be explained by loss of effective refractive index with age. Recent studies on ocular components in children and adults have shown that the calculated effective index of the lens decreases with increasing age, consistent with this hypothesis.

The final refractive state of the older adult eye thus depends on a complex set of changes in the ocular components which continue throughout childhood and adult life. Although there is now a number of population-based studies on refraction and ocular biometry in older adults, the Reykjavik Eye Study was the first to report lens power. It found a negative correlation between lens power and refractive error; that is, in contrast to the situation at the end of childhood, the lens has less power in hyperopic eyes and vice versa, in myopic eyes. But they also reported a negative correlation between lens power and AL, such that shorter eyes (which would normally be expected to be more hyperopic) would have higher lens power. The authors recognized this paradox but did not explore it further.

The present analysis from another population-based study, the Shahroud Eye Cohort Study (ShECS), was limited to subjects aged 55-64 years to assess continuing hyperopic shifts in refraction, and excluded subjects with nuclear cataracts, which are well-known to be associated with myopic shifts.

**METHODS**

The ShECS population consists of residents in Shahroud aged 40-64. A total of, 5,190 subjects of both genders participated in this study. The detailed methodology has been previously published. Multistage sampling was applied to the 40 to 64-year-old population of Shahroud, a city in the North of Iran. The study was approved by the ethics committee of Shahroud University of Medical Sciences and adhered to the tenets of the Declaration of Helsinki for research in human subjects.

For analysis of the ocular components of refraction, only subjects aged 55-64 years with nuclear opacity no more than grade 0 and 1 (i.e. without clinically significant cataract) were included to maximize the inclusion of subjects with the age-related hyperopic shifts occurring between ages 40-60, before myopic shifts due to nuclear cataracts develop.

Corneal power was calculated using the CR and assuming a refractive index of 1.3315 as proposed by Olsen. Crystalline lens power calculation was based on measurements of distance cycloplegic refraction, corneal power, anterior chamber depth, lens thickness, and AL using the formula proposed by Bennett. The A and B constants in this formula were calculated using Gullstrand’s reduced eye model.
As described by Grosvenor and Scott,[3] the emmetropic eye has an AL/CR near 3:1. In children, hyperopic eyes tend to have lower AL/CR ratio due to a short AL or a flatter cornea. The AL/CR ratio is apparently set by the end of childhood since CR does not change much after the first few years of life, while AL stabilizes in early adulthood. In an attempt to distinguish subjects who would have been hyperopic from childhood from those who were initially emmetropic or possibly mildly myopic and became hyperopic later in life due to hyperopic shifts, a cut-off point for AL/CR ratio was established at 2.92. Which represents subtraction of one standard deviation (SD) from a mean (±SD) value of 3.04 ± 0.11.

As the ocular components had normal distributions, Pearson correlations were calculated (r values are presented). One-way analysis of variances (ANOVA) with post-hoc Scheffe tests was employed for mean values of ocular parameters grouped by refractive error categories. The definitions of refractive error categories [Table 1]. P values < 0.05 were considered as significant.

RESULTS

A total of 1,006 subjects aged 55 to 64 years with nuclear opacity grade 0 and 1, including 496 (49.63%) men were studied. The distribution of data was quite symmetrical for spherical equivalent and principal ocular components of refraction [Figures 1-4] As presented in Table 2, only refractive error and AL/CR had high kurtosis.

Table 1 shows the number of subjects in each refractive group and their definitions, with the mean spherical equivalent, corneal power, AL, lens thickness, and calculated lens power. As expected, myopic eyes were longer, hyperopic eyes were shorter, and interestingly there was no difference in corneal power between emmetropes and hyperopes.

Table 3 shows correlation coefficients between the optical components that contribute to refraction. The correlation between AL and refraction was strongly negative, as expected. The correlation between AL and lens power was also strongly negative. In other words,
longer eyes were more myopic and had lower lens powers. Correlations of refraction and AL with lens thickness were much weaker, but lens power and lens thickness were moderately correlated; in other words, thicker lenses had more power.

Some important relationships between variables are illustrated graphically in Figures 5 and 6. Figure 5a shows that longer eyes had lower calculated lens powers and shorter eyes had higher calculated lens powers. In an apparent contradiction, Figure 5b shows that more myopic eyes tended to have lower lens powers, but the correlation and regression coefficients were low and not statistically significant. Figure 6 shows that lens power and lens thickness were positively correlated, with thicker lenses having higher lens power, despite the documented loss of lens power as it thickens during adult life.

When examined as refractive groups, there were significant differences in AL between the three refractive groups [Table 1]. Compared to emmetropes and hyperopes, myopes showed lower lens thickness [Table 1] while hyperopes had the lowest lens power although this difference was not significant [Table 1 and Figure 7].

It has been shown that hyperopic subjects in early adulthood have low AL/CR.[3] AL/CR ratios are likely to become stable in adult life since neither AL nor CR changes significantly. Given the hyperopic shift that take place in adult life, hyperopic subjects at the age of 55-64 consist of two groups: The first are subjects who were hyperopic at the end of childhood with low AL/CR ratios, while the second group consists of individuals who were emmetropic at the end of childhood, with medium AL/CR ratios, but developed hyperopic shifts during adult life. Hyperopic subjects were thus analyzed using an AL/CR cut-off of 2.92 as explained earlier. As evident form Table 4, 75 out of 269 hyperopes (28%) had AL/CR ratio ≤ 2.92. These subjects had, on average, less powerful corneas, shorter AL, and interestingly, higher lens power than emmetropes. In contrast, the group of hyperopes with more emmetropic AL/CR ratios had AL and corneal powers similar to emmetropic subjects, but lower lens powers, suggesting that they may have been emmetropes who had lost lens power during adult life.

**DISCUSSION**

The principal findings of this study confirm previously reported correlations between refractive components of the eye in an adult population. Only subjects aged 55-64 with low amounts of nuclear opacity in the ShECS population were considered for this analysis. This group was chosen to search for the impact of hyperopic shifts and avoid cataract-related myopic shifts. The main ocular component correlated with refractive error was AL. The correlation between lens power and refractive error was very low and negative; hyperopic eyes had lower lens powers.

Studying ocular refractive components in the Reykjavik Eye Study, Olsen et al.,[24] reported relationship between the different ocular parameters in an adult population, showing a significant negative correlation ($r=-0.23$) between spherical equivalent refraction and calculated lens power.
lens power treated as continuous variables (myopic eyes had higher lens power and vice versa, hyperopic eyes had lower lens power). In the same study, the negative correlation between lens power and AL in the whole sample demonstrated that eyes with longer ALs also had lower lens powers (and vice versa), with a significant greater correlation of −0.44.

The present data from ShECS confirms the negative relationship between lens power and AL but differs from the results of the Reykjavik Eye Study in finding little correlation between lens power and refraction as originally described by Tron in younger subjects.[1] Nevertheless, the same paradox exists, as longer eyes would be expected to be more myopic with lower lens power in childhood. The difference may be due to the fact that the Reykjavik Eye Study analysis did not exclude cataract subject, and the myopic shifts related to nuclear cataract may have produced a group of older myopic subjects with high lens power that would make the correlation between refraction and lens power even more negative as was found in the Central India Eye and Medical Study where cataract subjects were included.[28]

The situation in adults, therefore, appears to be different from that at the end of childhood,[13,29-32] where hyperopic eyes have greater lens power, emmetropic eyes have less lens power, and myopic eyes have the least. The relationships at the end of childhood result from children starting at birth with hyperopic eyes of varying sizes. The matching of AL to the refractive surfaces over the first 1-2 years of life leads to a tight and kurtotic distribution of refractive error, which is centered around mild hyperopia, such that shorter eyes tend to have higher lens powers for a given level of ametropia, thus establishing the basic association between hyperopia, higher lens power, and shorter AL. Subsequent axial elongation during childhood tends to result in myopic shifts, but these are partly compensated for by reductions in lens power, which only strengthen the association between hyperopia, high lens power, and shorter AL (or, in the same manner, of myopia, low lens power, and longer AL).

Figure 5. (a) Negative correlation between lens power and axial length ($P < 0.001$) (b) negative correlation between lens power and refraction ($P = 0.08$).

Figure 6. Positive significant correlation between lens power and lens thickness ($P < 0.001$).

Figure 7. Lower mean lens power in hyperopes as compared to emmetropes (analysis of variances $P = 0.132$) (95% confidence interval shown by error bars).
In adult life, when corneal power and AL have stabilized, there may be a continuous slow reduction in lens power, which in turn is responsible for well-documented hyperopic shifts found with ageing. These appear to be larger in hyperopic and emmetropic eyes than in myopic eyes, reducing the positive correlation between lens power and refractive status, while maintaining the correlation between higher lens power and shorter ALs driven by this high correlation in emmetropic eyes.

It is noteworthy that population-based studies in older adults, including the Reykjavik Eye Study, the Central India Eye and Medical Study, and the Latino Eye Study and the present one show that hyperopic eyes have lower lens power. In a population-based study of cycloplegic refractive error in adults in Tehran, the prevalence of cycloplegic hyperopia >+1 diopters increased from 10.5% to 53.3% between ages 20 and 70. This 40% increase in cycloplegic hyperopic refractive error prevalence, at an age where corneal power and AL seem to be stable, can only be attributed to a decrease in lens power.

The mechanisms involved in decreasing lens power with age are under study. It is widely believed that as new fibers grow in the surface of the lens and older ones become compacted in the deeper layers, they gain refractive index, and thus a gradient of refractive index is established from the surface to the center of the lens. This gradient gives the lens an internal power greater than that due to its surface curvature alone. This gradient index structure becomes less effective with ageing, as has been recently shown using magnetic resonance imaging, reducing the lens effective refractive index by a steepening of the gradient. In theory, a lens that loses effective refractive index could lose power, so emmetropic eyes could become hyperopic.

The positive correlation we observed between lens thickness and lens power is interesting as it is known from geometrical optics that if curvatures remain unchanged, a thinner homogeneous lens would have more power than a thicker lens, setting an inverse correlation between lens thickness and lens power in a homogeneous lens. The positive correlation between lens power and lens thickness described in this study has also been reported in children, and may be explained by lens growth and compaction. A thicker lens is undoubtedly a lens that has had a higher rate of growth than a thinner one. If compaction of lens fibers is a self-regulated process due to aging, this increased rate of growth could make the gradient index shallower, thereby increasing lens power. It is difficult to understand how lens power and AL reciprocally adapt to each other showing a high negative correlation. It is possible that changes in lens power are broad developmental phenomena, rather than finely-tuned emmetropization mechanisms. In this sense, there was no correlation between refraction and lens power in this sample. Nevertheless, differences in lens power and lens thickness appear to play a role in final determination of hyperopic spherical equivalent refraction. Myopic eyes in this study had thinner lenses than other refractive groups but did not show lower lens power.

Table 4. Comparison of ocular components for three refractive groups based on AL/CR ratio

<table>
<thead>
<tr>
<th>Refractive group</th>
<th>n</th>
<th>AL/CR</th>
<th>Mean SE</th>
<th>Keratometry</th>
<th>AL</th>
<th>Lens power</th>
</tr>
</thead>
<tbody>
<tr>
<td>Hyperopes with low AL/CR</td>
<td>75</td>
<td>2.86*</td>
<td>2.89*</td>
<td>42.48*</td>
<td>22.32*</td>
<td>23.13*</td>
</tr>
<tr>
<td>Hyperopes with normal AL/CR</td>
<td>193</td>
<td>2.99*</td>
<td>1.54*</td>
<td>43.70*</td>
<td>22.72*</td>
<td>21.96*</td>
</tr>
<tr>
<td>Emmetropes</td>
<td>528</td>
<td>3.03</td>
<td>0.22</td>
<td>43.47</td>
<td>23.11</td>
<td>22.54</td>
</tr>
</tbody>
</table>

*Significant differences when compared to emmetropes by ANOVA, P<0.05. SE, standard error; AL, axial length; CR, corneal radius
biometry, refraction and lens radii measurements.\[11,13,22]\n
Those studies have shown that there are inter- and intra-individual variations in calculated lens power and effective refractive index of the lens. Without lens radii measurements, the present study has not been able to estimate refractive index of the lens, and hence, for lens power formula, Gulstrand’s schematic eye values were used.

In summary, comparing studies in children with our findings in older adults, the general impression is that AL represents the principal variable determining refractive error, since correction of hyperopic errors in infants is largely achieved through axial elongation, and myopic refractive errors are produced by axial elongation. Corneal power stabilizes quite early in development, but major changes in lens power during childhood and during the course of adult life. In childhood, axial elongation is accompanied by reductions in lens power, which can partially compensate for the tendency toward myopic shifts, while in adult life, reductions in lens power may continue, resulting in hyperopic shifts. These reductions in lens power obviously have a major impact on final refractive status, but there is not strong correlation between lens power and refraction (when cataract subjects are excluded) and the distribution of lens power is normal rather than kurtotic, all of which suggest that these mechanisms do not refine refractions with the precision given by the regulation of AL growth.

REFERENCES


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