Does transient increase in axial length during accommodation attenuate with age?

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Background: The aim was to profile transient accommodative axial length changes from early adulthood to advanced presbyopia and to determine whether any differences exist between the responses of myopic and emmetropic individuals.

Methods: Ocular biometry was measured by the LenStar biometer (Haag-Streit, Switzerland) in response to zero, 3.00 and 4.50 D accommodative stimuli in 35 emmetropes and 37 myopes, aged 18 to 60 years. All results were corrected to reduce errors arising from the increase in crystalline lens thickness with accommodation. Accommodative responses were measured sequentially by the WAM 5500 Auto Ref/Keratometer (Grand Seiko, Hiroshima, Japan).

Results: Axial length increased significantly with accommodation (p < 0.001), with a mean corrected increase in axial length of 2 ± 18 μm and 8 ± 16 μm observed at 3.00 and 4.50 D, respectively. The magnitude of accommodative change in axial length was not dependent on refractive error classification (p = 0.959); however, a significant reduction in the magnitude and variance of axial length change was evident after 43 to 44 years of age (p < 0.002).

Conclusion: The negative association between transient increase in axial length and age, in combination with reduced variance of data after age 43 to 44 years, is consistent with a significant increase in posterior ocular rigidity, which may be influential in the development of presbyopia.

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The amplitude of accommodation decreases with age, from approximately 15 D in early infancy to 1.00 D at 60 years of age,1,2 when presbyopia is well established. Significant age-related structural changes in the accommodative apparatus (for example, increased lenticular stiffness) are thought to be responsible for the attenuation of the amplitude of accommodation and therefore, the development of presbyopia.3–7 While age-related accommodative changes within the anterior segment have been investigated (for example, ciliary muscle morphology and mobility),8,9 accommodative changes within the posterior segment with respect to age have yet to be reported.

Indeed, a significant transient elongation of axial length during short periods of accommodation has been observed in young adults.10–14 It has been hypothesised that this axial length elongation during accommodation may be produced by the force applied to the equatorial choroid by ciliary muscle contraction, thus necessitating posterior pole elongation to maintain a constant ocular volume.10,11 In support of this theory, Croft and colleagues15 observed the equatorial choroid and retina moved centripetally during Edinger-Westphal stimulated accommodation in rhesus monkeys. Due to the persistence of ciliary muscle contractility into advanced presbyopia,16 it is feasible that transient increase in axial length may also be present in presbyopic individuals; however, an in vivo anterior scleral study17 and in vitro posterior choroidal18,19 and scleral20 studies have reported ocular stiffness increases with age. Van Alphen21 suggested the ciliary muscle and choroid behave as a single elasto-muscular sheet around the eye, moulding the sclera to accommodate ciliary muscle tonus and therefore, it is conceivable that age-related increases in ocular stiffness may mute transient axial length elongation during accommodation. Nevertheless, the relocation of the anterior uveal tract after cataract surgery to a more youthful position in humans suggests the choroid may retain its elasticity with age,22 despite the sclera and choroid becoming thicker23 and thinner24 with age, respectively. Furthermore, evidence reported by Hollins (cited by Atchison25) of transient axial length elongation in one advanced presbyopic patient during near vision also suggests that accommodation-induced transient axial length elongation may persist into advanced presbyopia.

Additionally, it has been hypothesised that high levels of transient axial length
Axial elongation during accommodation may precipitate a myopic shift in refraction in susceptible eyes. Therefore transient axial length elongation during accommodation is one potential factor responsible for the connection between high levels of near work and the onset and progression of myopia in both children and adults. Mallen, Kashyap and Hampson hypothesised that myopic eyes may be more amenable to transient axial length elongation during accommodation due to reduced ocular rigidity (present due to the stretched, elongated eye characteristic of myopia). Yet, despite reduced posterior choroidal and scleral thickness in myopic eyes compared to emmetropic eyes, it remains unclear whether a correlation exists between in vivo anterior ocular rigidity and myopia. Moreover, previous research in young adults has produced contradictory results; Drejser and colleagues reported the magnitude of axial length elongation was larger in emmetropic eyes than myopic eyes, whereas Mallen, Kashyap and Hampson reported greater elongation in myopic eyes and Read and colleagues found no difference in ocular axial elongation between myopes and emmetropes during accommodation.

The aim of this study was to profile accommodative changes in axial length from early adulthood to advanced presbyopia and to determine whether any differences exist between the responses of myopic and emmetropic individuals.

### METHODS

The study was approved by the Aston University Research Ethics Committee and was conducted in accordance with the tenets of the Declaration of Helsinki. Written informed consent was obtained from all the participants after an explanation of the nature and possible consequences of the study. Participants were included in the study, if they were aged between 18 and 70 years; could achieve 0.0 logMAR visual acuity, were classified as emmetropic (spherical equivalent between −0.75 and +0.50 D) or myopic (spherical equivalent more myopic than −0.75 D) and had 0.75 DC or less of uncorrected astigmatism. All participants were screened to exclude those with a positive history of ocular or systemic disease or prescribed medications known to affect the accommodative response.

### Sample size calculation

The calculated total sample size for repeated measures analysis of variance (ANOVA) testing (within and between interaction), including an effect size (f) of 0.25, an error probability (α) of 0.05 and required power (1 − β) of 0.80 for three measurements between the two groups, was 28 participants.

### Amplitude of accommodation

The right eye of all the myopic participants was fitted with a soft daily disposable spherical contact lens (Focus Dailies, nelficon A, 69 per cent water content; Ciba Vision, Duluth, Georgia, USA). The power of the contact lens was chosen following a subjective refraction. A minimum settling period of 10 minutes was permitted prior to over-refraction to ensure the contact lens prescription fully corrected the ametropia. The subjective amplitude of accommodation of the right eye was measured using a Royal Air Force Rule (Richmond Products, New Mexico, USA) via the push-up/pull-down method. The left eye of each participant was occluded. Using their right eye, participants were asked to attempt maintaining clear focus of the high-contrast N5 print, as it was slowly pushed toward them until the point where they first reported sustained blur (push-up amplitude). Subsequently, the first point at which the N5 print became clear when moved away from the eyes (pull-down amplitude) was measured. If a patient was unable to achieve the minimum amplitude of accommodation measurable (2.00 D), a +3.00 D full-aperture trial lens was introduced in front of the right eye. The power of the lens was subsequently subtracted from the push-up/pull-down amplitude value obtained. Push-up/pull-down amplitude was measured three times and averaged to calculate the mean amplitude of accommodation for the right eye.

### Accommodative stimulus response

Change in objective spherical equivalent refractive error during accommodation was measured by the binocular open-field WAM-5500 autorefractor (Grand Seiko Co. Ltd., Hiroshima, Japan), with an external fixation target (a Maltese cross) viewed through a +5.00 D Badal lens attached to the instrument. The fixation target was placed 20 cm, 8.0 cm and 2.0 cm away from the Badal lens to stimulate zero, 3.00 and 4.50 D of accommodation, respectively. Each participant was asked to focus on the centre of the Maltese cross as accurately as possible and was exposed to the stimulus for 20 seconds prior to the acquisition of data. A one minute distance viewing break was permitted between accommodative levels. The presentation order of each accommodative level was randomised between participants. Three consecutive measurements of refraction were acquired at each accommodative level.

### LenStar biometry

LenStar biometry was measured while stimulating accommodation with an external target (viewed through a pelvic beam splitter and a +10.00 D Badal lens). To provide zero, 3.00 and 4.50 D accommodative stimuli, the Badal lens system letter target was placed 10 cm, 7 cm and 5.5 cm away from the +10.00 Badal lens, respectively. The average of three repeat measures of anterior chamber depth (ACD), crystalline lens thickness (LT) and axial length (AXL) were recorded at the three accommodative levels (zero, 3.00 and 4.50 D). Corneal thickness (CT) was measured at zero dioptres only because it has been well-established the human corneal thickness does not change during accommodation. Anterior segment length (ASL = CT + ACD + LT) and vitreous chamber depth (VCD = AXL − ASL) were calculated from these values. If the LenStar was unable to acquire lens thickness measurements for a participant, these data were excluded from the analysis.

### Biometric error calculations

The LenStar uses an average ocular refractive index to convert an optical path length (OPL) to a geometric axial length. Therefore, to correct for the potential over-estimation of axial length due to an increase in lens thickness with accommodation and age, the individual error (E) was calculated for each participant using Equations 1 to 6 as proposed by Atchison and Smith. In line with previous publications, the relative proportions of the crystalline lens taken up by the anterior cortex, nucleus and posterior cortex during accommodation were kept constant because, despite the well-established thickening of the nucleus during accommodation, the exact nature of the change in refractive index during accommodation is not fully understood. To compensate for age-related changes in anterior cortex thickness (ACT), nucleus...
thickness (NT) and posterior cortex thickness (PCT), Equations 1, 2 and 3 were used to modify the segmentation of the crystalline lens according to age (in years), derived by Dubbelman and colleagues.\(^4\) Optical path length and the average refractive index of the eye (\(n_{\text{av}}\)) were calculated using the refractive indices specified by Gullstrand’s No. 1 (exact) eye with shell lens (Equations 4 and 5).\(^4\) Equation 6 was used to calculate the error (E), which is subtracted from the geometric axial length reported by the LenStar to provide corrected axial length values.

\[
ACT = LT \times \frac{(0.51 + 0.012 \times \text{age})}{(0.51 + 0.012 \times \text{age}) + (2.11 + 0.003 \times \text{age}) + (0.33 + 0.0082 \times \text{age})}
\]

\[
NT = LT \times \frac{(2.11 + 0.003 \times \text{age})}{(0.51 + 0.012 \times \text{age}) + (2.11 + 0.003 \times \text{age}) + (0.33 + 0.0082 \times \text{age})}
\]

\[
PCT = LT \times \frac{(0.33 + 0.0082 \times \text{age})}{(0.51 + 0.012 \times \text{age}) + (2.11 + 0.003 \times \text{age}) + (0.33 + 0.0082 \times \text{age})}
\]

\[
\text{OPL} = (CT \times 1.376) + (ACD \times 1.336) + (ACT \times 1.386) + (NT \times 1.406) + (PCT \times 1.386) (\text{VCD} \times 1.336)
\]

\[
n_{\text{av}} = \frac{\left(\frac{CT}{\text{AXL}} \times 1.376\right) + \left(\frac{ACD}{\text{AXL}} \times 1.336\right) + \left(\frac{ACT}{\text{AXL}} \times 1.386\right) + \left(\frac{NT}{\text{AXL}} \times 1.406\right)}{\left(\frac{PCT}{\text{AXL}} \times 1.386\right) + \left(\frac{VCD}{\text{AXL}} \times 1.336\right)}
\]

\[
E = \frac{\text{OPL}}{n_{\text{av}}} - \text{AXL}_{\text{unaccommodated}}
\]

### Statistical analysis

To assess whether biometry changed significantly with accommodation (within-subject factor) and to determine its dependency on refractive error (between-subject factor), a repeated measures ANOVA was performed (SPSS, SPSS Inc, Chicago, Illinois, USA). The relationship between biometry and age was determined by linear regression analysis.

To compare the variance of the data between the younger and older participants, a two-tailed variance test was used, which accounts for differences in group sizes. The ANOVA residual mean square indices produced by individual linear regression analysis of the younger and older groups are divided to produce an F ratio. The significance of the difference in variance is determined by a two-tailed F look-up table, according to the degrees of freedom in each group.

### RESULTS

#### Participants

Seventy-two participants aged 18 to 60 years (median age 33.60 years) were recruited from an Aston University staff and student volunteer call. Participants were classified as emmetropic (spherical equivalent from -0.50 to +0.50 D) or myopic (spherical equivalent -0.75 D or greater) and all subjects had less than 0.75 DC astigmatism. Of the total cohort, 37 were myopic emmetropes 7.26 ± 2.83 D). As expected, right eye amplitude of accommodation decreased significantly with age (r = 0.918, p < 0.001; Figure 2). Baseline uncorrected spherical equivalent was not significantly correlated with age in either group (myopes r = 0.063, p = 0.701; emmetropes r = 0.084, p = 0.621).

#### Ocular biometry

The disaccommodated vitreous chamber depth was not significantly correlated with age in either group (myopes r = 0.093, p = 0.584, emmetropes r = 0.277, p = 0.108); however, the disaccommodated anterior chamber depth was significantly smaller with increasing age (myopes r = 0.316, p = 0.001, emmetropes r = 0.347, p = 0.041) and lens thickness (myopes r = 0.707, p < 0.001, emmetropes r = 0.602, p < 0.001) and anterior segment length (myopes r = 0.417, p = 0.010, emmetropes r = 0.375, p = 0.027) were significantly larger with increasing age. Older emmetropic individuals had significantly longer axial lengths than younger emmetropic individuals (myopes r = 0.045, p = 0.790, emmetropes r = 0.437, p = 0.009).

Repeated measures ANOVA testing showed the decrease in anterior chamber depth (\(F = 95.994, p < 0.001\)) and the increase in lens thickness (\(F = 142.410, p < 0.001\)), anterior segment length (\(F = 8.299, p = 0.001\)), vitreous chamber depth (\(F = 5.955, p = 0.006\)) and axial length (\(F = 11.104, p < 0.001\)) was statistically significant during accommodation. No differences in the accommodative response emerged based on the grouping of ametropia (anterior chamber depth \(F = 0.507, p = 0.562\); lens thickness \(F = 3.482, p = 0.053\); anterior segment length \(F = 1.342, p = 0.263\); vitreous chamber depth \(F = 1.330, p = 0.266\); axial length \(F = 0.277, p = 0.730\)). As expected, anterior chamber depth (\(F = 7.653, p = 0.007\)), vitreous chamber depth (\(F = 68.321, p < 0.001\)) and axial length (\(F = 65.741, p < 0.001\)) values were consistently longer in myopic eyes when compared to emmetropic eyes.\(^4\)\(^5\) No significant differences emerged between myopic and emmetropic lens thickness (\(F = 0.885, p = 0.395\)) and anterior segment length (\(F = 2.340, p = 0.131\)). Considering the whole cohort, a non-statistically significant, possible negative association emerged between the change in anterior chamber depth
Axial elongation during accommodation

Laughton, Sheppard, Mallen, Read and Davies

Figure 1. Schematic diagram of the LenStar accommodative stimulus setup. The retro-illuminated target (T) was placed at the zero, 3.00 and 4.50 D stimulus levels via the +10.0 D Badal lens (B), accordingly. T was viewed via a 92 per cent transmission/eight per cent reflection pellicle beam-splitter (BS₁).

Figure 2. Right eye objective accommodative response to the 3.00 D target (triangles; $y = -0.0026x^2 + 0.1441x + 0.0691$, $R^2 = 0.539$), 4.50 D target (circles; $y = -0.003x^2 + 0.134x + 1.8137$, $R^2 = 0.679$) and right eye subjective amplitude of accommodation (squares; $y = -0.257x + 15.99$, $R^2 = 0.829$) according to age. The dashed horizontal line represents 4.50 D of accommodation. The solid vertical line represents the intercept of the subjective accommodation regression line and the dashed 4.50 D of accommodation line ($y = 44.9$ years). The objective accommodative response reaches zero at 56.00 years of age.

(r = 0.128, p = 0.285), lens thickness (r = 0.067, p = 0.577) and anterior segment length (r = 0.020, p = 0.093) with the change in axial length observed at 4.50 D.

The relationship between the accommodation-induced axial length changes and age are illustrated in Figure 3. After visual inspection of the graphical data in Figure 3, the cohort was split into under 43 years (59 participants) and 43 years and older (13 participants), owing to the dramatic reduction in data spread after age 43 years. The difference in variance of the corrected axial length data between the groups was significant at the 3.00 D ($F = 2.983$, $p < 0.020$) and 4.50 D ($F = 8.852$, $p < 0.002$) accommodative levels. To validate the choice of 43 years as the fulcrum point, the spread of data was analysed in one-year increments either side of age 43 years at 3.00 D (41 years $F = 1.848$, $p > 0.100$; 42 years $F = 2.204$, $p < 0.100$; 44 years $F = 2.983$, $p < 0.020$; 45 years $F = 2.454$, $p < 0.050$) and 4.50 D (41 years $F = 2.736$, $p < 0.020$; 42 years $F = 5.055$, $p < 0.002$; 44 years $F = 8.852$, $p < 0.002$; 45 years $F = 8.312$, $p < 0.002$). Between the ages of 43 to 44 years was therefore selected as Gibson⁷⁰ reported that the posterior movement of the crystalline lens is negligible until the accommodative demand reaches approximately 2.00 D (eliciting an accommodative response of approximately 1.50 D). Therefore, only participants who demonstrated an accommodative response greater than 1.50 D to the 4.50 D accommodative target were included in calculations to determine the changes in ocular biometry per dioptre of accommodation exerted. The top age limit included in the biometric response per dioptre of accommodation measurements was consequently readjusted to 44 years; however, younger participants who failed to accommodate more than 1.50 D (Figure 2) were excluded, resulting in an amended cohort size of 56 participants. Despite a significant attenuation of the accommodative change in anterior chamber depth, lens thickness and axial length with age according to the magnitude of the accommodative stimulus, the changes per dioptre of accommodation exerted were not dependent on age (Table 1). The decrease in anterior chamber depth per dioptre of accommodation exerted was 0.081 ± 0.053 mm/D, lens thickness increased by 0.109 ± 0.052 mm/D, anterior segment length increased by...
Axial elongation during accommodation Laughton, Shephard, Mallen, Read and Davies

0.028 ± 0.057 mm/D, vitreous chamber depth decreased by 0.025 ± 0.059 mm/D and axial length increased by 0.003 ± 0.006 mm/D.

DISCUSSION

The present study is the first to document the effect of age on axial elongation during accommodation. In accordance with previous studies investigating accommodative changes in axial length in young adults,10–14 a statistically significant axial elongation with accommodation was observed in the current investigation, which was proportional to the magnitude of the accommodative stimulus. The change in ocular biometry per dioptré of accommodation exerted remained invariant between the ages of 18 to 44 years, as previously reported;3,6 however, beyond the age of 43 to 44 years, the change in axial length was negligible and the variance of data reduced significantly (Figure 3).

Due to the persistence of ciliary muscle contractility into advanced presbyopia,9,16 it is feasible that the reduced variance and magnitude of change in axial length with accommodation may signify in vivo posterior ocular rigidity increases significantly during the presbyopic years. Indeed, previous research has also relied on changes in axial length to determine changes in posterior ocular rigidity; Ebneter, Wagels and Zinkernagel51 reported ocular rigidity is higher in patients with glaucoma due to a smaller decrease in axial length following a pharmacologically induced (500 mg oral acetazolamide) fall in intraocular pressure when compared to a healthy control group. However, changes in axial length alone are not able to disentangle the relative contributions of each ocular layer to changes in rigidity and thickness, only to infer their cumulative effect. Currently, in vitro studies give the best indication of how the properties of the individual ocular layers may vary with age.

In support of the data presented here, an age-related decrease in the elasticity of choroidal and scleral sections has been found by multiple in vitro studies.19,20,52 It is feasible that a reduction in the flexibility of the posterior pole with age may contribute to the development of presbyopia; reducing the ability of the choroid and posterior zonules to restore the accommodative apparatus to an disaccommodated position.53 Studies of in vitro isolated choroidal and scleral sections do not account for the impact of vascular rigidity, perfusion pressure, intraocular pressure, extraocular muscle action, post-mortem structural changes and how these factors may vary with age.20 Indeed, the choroid24 and its blood vessels54 thin and the blood flow reduces with age,55 whereas the sclera becomes thicker with age.23 Unfortunately, it is not possible to measure posterior ocular rigidity in vivo directly and hitherto, the literature has not reached a consensus as to whether anterior eye rigidity increases with age.17,32

Furthermore, it is feasible the hypothesised increase in ocular shell rigidity with age is accompanied by a reduction in the capacity of the choroid to thin during accommodation, which has been suggested to contribute to the magnitude of axial elongation in young adults.14 The cumulative effect of these age-related changes may be responsible for the abrupt loss of axial elongation during accommodation after 43 to 44 years of age. It is possible to extract choroidal thickness measurements from further analysis of the LenStar A-scan peaks, which allows choroidal thickness to increase with age; however, these data are beyond the scope of the present study.

**Table 1.** Dependency of accommodative changes in biometry on participant age (56 participants). Vitreous chamber depth (VCD) and axial length (AXL) values have been corrected for the increase in lens thickness (LT) with age and accommodation.

| Parameter (mm) | 0.00 to 3.00 D accommodative stimulus | 0.00 to 4.50 D accommodative stimulus | Per dioptré of accommodation exerted* |
|---------------|-----------------------------------|-----------------------------------|-----------------------------------|
|               | r       | p       | r       | p       | r       | p       | r       |
| ACD           | 0.271   | 0.021   | 0.404   | <0.001  | 0.028   | 0.833   |        |
| LT            | 0.412   | <0.001  | 0.509   | <0.001  | 0.051   | 0.700   |        |
| ASL           | 0.105   | 0.378   | 0.184   | 0.121   | 0.021   | 0.875   |        |
| VCD           | 0.087   | 0.467   | 0.154   | 0.196   | 0.030   | 0.821   |        |
| AXL           | 0.129   | 0.279   | 0.264   | 0.025   | 0.105   | 0.431   |        |

ACD: anterior chamber depth, ASL: anterior segment length.

*Only results from participants able to accommodate more than 1.50 D are included.
Figure 4. Diagrammatic representation of the hypothesised shape of a disaccommodated (solid line) and accommodated (dashed line) posterior pole. The x-axis describes the distance from the fovea and the y-axis describes the distance of the posterior pole from the cornea. The blue arrows demonstrate the putative accommodation-driven inward movement at the equator and the purple arrows show the potential corresponding pattern of posterior pole elongation. In this scenario, the axial length measured at the central fovea appears to have decreased during accommodation.
Axial elongation during accommodation

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