Amblyopia in Astigmatic Infants and Toddlers

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ABSTRACT

Purpose. To determine whether reduced astigmatism-corrected acuity for vertical (V) and/or horizontal (H) gratings and/or meridional amblyopia (MA) are present before 3 years of age in children who have with-the-rule astigmatism.

Methods. Subjects were 448 children, 6 months through 2 years of age with no known ocular abnormalities other than with-the-rule astigmatism, who were recruited through Women, Infants and Children clinics on the Tohono O’odham reservation. Children were classified as non-astigmats (≤2.00 diopters) or astigmats (>2.00 diopters) based on right eye non-cycloplegic autorefraction measurements (Welch Allyn SureSight). Right eye astigmatism-corrected grating acuity for V and H stimuli was measured using the Teller Acuity Card procedure while children wore cross-cylinder lenses to correct their astigmatism or plano lenses if they had no astigmatism.

Results. Astigmatism-corrected acuity for both V and H gratings was significantly poorer in the astigmats than in the non-astigmats, and the reduction in acuity for astigmats was present for children in all three age groups examined (6 months to <1 year, 1 to <2 years, and 2 to <3 years). There was no significant difference in V-H grating acuity (no evidence of MA) for the astigmatic group as a whole, or when data were analyzed for each age group.

Conclusions. Even in the youngest age group, astigmats tested with astigmatism correction showed reduced acuity for both V and H gratings, which suggests that astigmatism is having a negative influence on visual development. We found no evidence of orientation-related differences in astigmatism-corrected grating acuity, indicating either that MA does not develop before 3 years of age, or that most of the astigmatic children had a type of astigmatism, i.e., hyperopic, that has proven to be less likely than myopic or mixed astigmatism to result in MA.

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in vernier acuity in 5- to 11-year-old children who had astigmatism early in life that had emmetropized by the time the children were tested. The meridional differences in acuity were most highly correlated with the amount of astigmatism present late in the first year up to 2 years of age, and the meridional differences in acuity were least correlated with the amount of astigmatism early in the first year of life and between 2 years of age and their age at the time of test (5 to 11 years). These results suggest that uncorrected astigmatism begins to influence visual development by 2 years of age. Consistent with these data are the results of several studies that have documented MA in 3- to 5-year-old astigmatic children. The purpose of this study was to compare astigmatism-corrected acuity for V and H grating stimuli in WTR astigmatic vs. low/non-astigmatic infants and toddlers. The goal was to determine if reduced acuity for V and/or H gratings and/or MA is present before 3 years of age and to determine if astigmats younger than 3 years of age show evidence of MA.

METHODS

Subjects

Subjects were children, 6 months through 2 years of age, who were recruited through Women, Infants and Children clinics on the Tohono O’odham reservation from September 2005 through December 2008. Children in the study are followed longitudinally; however, this report is based only on data from each child’s first test session. This research followed the tenets of the Declaration of Helsinki and was approved by the Tohono O’odham Nation and by the Institutional Review Board of the University of Arizona. Parents provided written informed consent before testing.

Apparatus

The apparatus for grating acuity testing consisted of a Teller acuity card (TAC) stage (Vistech Consultants, Inc., Dayton, OH) and two sets of 15 acuity cards, one set containing 14 V grating stimuli and a blank card, and one set containing 14 H grating stimuli and a blank card. V and H stimulus orientations were selected because nearly all astigmatic Tohono O’odham preschool children have WTR astigmatism (plus cylinder axis near 90°), and, therefore V and H gratings would be parallel to the astigmatic axes in nearly all astigmatic children in this population. Each card was constructed of gray cardboard, 25.5 × 56 cm in size and contained two 12-cm diameter circular openings located with the innermost edge 9 cm to the left and right, respectively, of a central 4-mm peephole. Behind one opening was a black-and-white square-wave grating and behind the other opening was a luminance-matched gray field, both of which were cut from a single unmounted TAC (Stereo Optical, Co., Inc, Chicago, IL). Grating spatial frequencies in each card set ranged from 0.32 to 38 cycles/cm in approximately half-octave steps. Ambient illumination was supplemented with a clip lamp attached to the stage to provide uniform illumination of the stage and cards that was >10 cd/m² in luminance.

A standard set of astigmatism-correcting Jackson Cross Cylinder spectacles (spherical equivalent = 0), with right eye (RE) cylinder power between 0 and 8.25 diopters (D) in 0.75 D steps, and with axis at 30°, 60°, 75°, 90°, 105°, 120°, 150°, and 180°, was available at each test session. An example would be spectacles that are −1.50 + 3.00 × 90, which correct 3.00 D of astigmatism but have spherical equivalent of 0. The frames (Solo Bambini, Burlingame, CA) are specially designed for infants and young children, are made of soft plastic, and have straight temples connected by an elastic strap that can be adjusted to fit the child. We chose to measure grating acuity with eyeglasses that correct cylinder only, because previous studies of infants indicate that most infants and toddlers are hyperopes and can accommodate to focus at near and because at the test distance of 50 cm, only myopia >2.00 D, which is rare in this age group, would affect acuity. Data from subjects for whom SureSight measurements indicated myopia >2.00 D in the most positive/least negative RE sphere measurement were excluded from analyses. The most positive/least negative measurement was used to estimate myopia because, without cycloplegia, measurements are likely to over-estimate myopia due to accommodation.

Procedure

Before acuity testing, each child had refractive error assessed with the Welch Allyn SureSight autorefractor (software versions 2.16 and 2.20; Welch Allyn Medical Products, Skaneateles Falls, NY). The purpose of SureSight testing was to measure the magnitude and axis of any astigmatism the child had so as to determine the appropriate spectacles to use to correct the astigmatism during assessment of grating acuity. SureSight testing was conducted in a dimly-lit room, away from direct sunlight. Subjects sat on the parent’s lap facing the tester, who attempted to take three measurements of the child’s RE and three measurements of the child’s left eye.

Following SureSight measurements, a pair of astigmatism-correcting spectacles was selected that had a cylinder power and axis closest to the median cylinder power and axis measured by the SureSight for the RE. If no astigmatism was present, spectacles containing a plano lens were selected. If the SureSight displayed a value of 9.99 (indicating that the cylinder measurement was beyond the instrument’s measurement range of 3.00 D), spectacles were selected that corrected 3.75 D of astigmatism, with axis determined by the SureSight reading. If the SureSight would not take a reading in a cooperative child (which has been shown to indicate a cylinder value likely to be >3.00 D), and therefore no estimate of axis was available, we excluded these children from further testing. For all other children, an over-refraction through the spectacles was then performed to verify the accuracy of the correction. For the over-refraction, the left lens in the spectacles was occluded, the spectacles were placed on the child, and the tester attempted to make three more SureSight measurements of the RE through the spectacles. If the median of the measurements of cylinder through the spectacles was <1.00 D, the spectacles were judged to be appropriate and the child wore these spectacles during assessment of grating acuity. If the median of the measurements showed ≥1.00 D of cylinder, the median value was used to select a pair of spectacles that left the child with <1.00 D of residual astigmatism, and these spectacles were worn during acuity testing. If the child would not cooperate for over-refraction, acuity testing...
Parents of children who met any of the following criteria were offered the opportunity to have the child examined, with cycloplegic refraction, by a pediatric ophthalmologist.

<table>
<thead>
<tr>
<th>Reason for referral for examination</th>
<th>Age</th>
<th>Criterion (in one or both eyes)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Astigmatism</td>
<td>6 to &lt;12 months</td>
<td>≥3.00 D</td>
</tr>
<tr>
<td></td>
<td>12 to &lt;24 months</td>
<td>≥2.50 D</td>
</tr>
<tr>
<td></td>
<td>24 to &lt;36 months</td>
<td>≥2.00 D</td>
</tr>
<tr>
<td>Anisometropia</td>
<td>6 to &lt;36 months</td>
<td>&gt;1.50 D SE</td>
</tr>
<tr>
<td>Hyperopia</td>
<td>6 to &lt;36 months</td>
<td>≥4.00 D in the most hyperopic meridian</td>
</tr>
<tr>
<td>Myopia</td>
<td>6 to &lt;36 months</td>
<td>≥5.00 D in the most myopic meridian</td>
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<tr>
<td>High refractive error</td>
<td>6 to &lt;36 months</td>
<td>&quot;Out of range&quot; results on all SS measurements</td>
</tr>
<tr>
<td>Strabismus or cataract</td>
<td>6 to &lt;36 months</td>
<td>Cover test, red reflex</td>
</tr>
</tbody>
</table>

aBased on median SS cylinder measurement for each eye.
bBased on SE for each eye calculated using most hyperopic sphere measurement across all SS measures and median cylinder measurement across all SS measurements for each eye.
cBased on most hyperopic sphere measurement and median cylinder measurement across all SS measures for each eye.
dBased on most myopic sphere measurement for each eye.

SE, spherical equivalent; SS, SureSight.

was attempted using the spectacles chosen based on the initial SureSight measurements.

Immediately after the spectacles were selected and placed on the child’s face, RE grating acuity for V and H gratings was measured with the TAC procedure21,22 at a distance of 50 cm. The grating orientation (V or H) tested first was determined according to a counterbalanced order. The acuity cards in each orientation set were presented in order from lower (coarser) to higher (finer) spatial frequencies, beginning with the card containing a 1.3 cy/cm (1.2 cy/deg) grating. The tester, who was masked to the location of the grating on each card, used the child’s eye and head movements to repeated presentations of each card to decide whether the child could discriminate the location of the grating on the card.21,22 On the basis of the child’s responses to repeated presentations of each card, the tester determined the highest spatial frequency (finest grating) that the child could resolve for each grating orientation. Threshold grating acuity scores (cy/deg) were transformed to log values for data analyses.

Parents of the subset of children whose results met any of the criteria summarized in Table 1 were offered a follow-up eye examination, with cycloplegic refraction, conducted by a pediatric ophthalmologist (JMM) who was masked to the results of the SureSight screening. The purpose of the follow-up examination was to determine whether any ocular abnormality or significant refractive error requiring further follow-up or spectacle correction was present. In a given area of the reservation, examinations were offered approximately six times per year, and parents were notified whenever an exam date was scheduled in their area.

Data Analysis

Data from children with known ocular abnormalities or developmental disabilities and from children who did not provide both SureSight measurements and grating acuity data for the RE were excluded. In addition, data from children who completed grating acuity testing but in whom the most positive/least negative SureSight RE sphere measurement indicated ≥2.00 D of myopia, or who had RE astigmatism (>0.00 D) that was not WTR (plus cylinder axis 90° ± 15°, on the basis of SureSight median axis) were excluded.

For each eye, the estimate of cylinder magnitude and axis was obtained by converting measurements to vector notation,23 calculating the median J0 and J45, and then converting the medians back into clinical notation. In cases where the SureSight did not provide cylinder magnitude (due to out of range cylinder measurement indicated by a value of 9.99) but did provide axis measurements, axis was used as the simple median of all measurements of axis obtained. Data were excluded in cases where the SureSight did not provide axis measurement (instrument turned off without making a measurement in a cooperative child, indicating a high likelihood of astigmatism >3.00 D).20)

A previous comparison of SureSight cylinder values with cylinder values obtained with the Retinomax K-Plus autorefractor indicated that SureSight cylinder values ≤3.00 D correspond to true cylinder values of ≤2.00 D in 97% of cases, and SureSight cylinder values of 9.99 or failure to provide a cylinder correspond to true cylinder values >2.00 D in 88% of cases.20 Therefore, in this study, we used the median SureSight cylinder value for the RE to classify children in this study as low/non-astigmatic (≤2.00 D of astigmatism) or highly astigmatic (>2.00 D of astigmatism). Children with SureSight values ≤3.00 D were classified as having low/no astigmatism and those with a SureSight value of 9.99 were classified as high astigmats. Those in whom the SureSight turned off without providing a reading were excluded from the data analysis, because axis of astigmatism was not known.

RESULTS

Subjects

The final sample included 448 children: 238 at 6 months to <1 year, 121 at 1 to <2 years, and 89 at 2 to <3 years of age. Although 786 children were initially recruited, data from 338 were excluded for the reasons summarized in Table 2. This included eight children in whom the initial SureSight readings indicated out-of-range
astigmatism by turning itself off after measurement in a cooperative child.

On the basis of the median SureSight reading for the RE (calculated using vector methods), 112 children (25.0%) had >2.00 D of astigmatism; this included 29.8% (238/786) of the 6 months to <1 year age group, 14.9% (18/121) of the 1 to <2 year age group, and 25.8% (23/89) of the 2 to <3 year age group.

In 417 of the 448 children (93.1%) (93.3% 222/238 of the 6 months to <1 year age group, 91.7% 111/121 of the 1 to <2 year age group, and 94.4% 84/89 of the 2 to <3 year age group), the spectacles worn during acuity testing corrected astigmatism to within 1.00 D, based on the SureSight over-refraction. The remaining 31 children did not cooperate for over-refraction with the SureSight, and a χ² test indicated that the percentage of uncooperative children did not differ across the three age groups.

There were 192 children (24.4%) who were excluded solely for inability to cooperate for grating acuity testing. A χ² test showed no difference in the prevalence of high astigmatism in this group of children [39/192 (20.3%)] vs. the prevalence of high astigmatism in the group of children who completed acuity testing and were included in the final analysis [112/448 (25.0%)].

Cycloplegic retinoscopy data were available for 152 (33.9%) of the 448 children: 88 at 6 months to <1 year, 39 at 1 to <2 years, and 25 at 2 to <3 years. On average, the data were obtained 6.7 months (SD 6.8) after acuity testing. Among the 152 children who had a cycloplegic refraction, 28 (18.4%) had astigmatism >2.00 D in the RE based on cycloplegic retinoscopy. Analysis of agreement between SureSight classification and classification based on cycloplegic retinoscopy indicated that 58% (72/124) of children classified as low astigmas based on cycloplegic retinoscopy were also classified as low astigmas based on SureSight readings and 68% (19/28) of children classified as high astigmas based on cycloplegic retinoscopy were also classified as high astigmas based on SureSight readings. In 7 of the 28 children classified as high astigmas based on cycloplegic retinoscopy, the astigmatism was simple or compound myopic or mixed, and in the remaining 21 children, it was simple or compound hyperopic.

Grating Acuity

Fig. 1 shows mean acuity for (A) V and (B) H gratings for the astigmatic and the low/non-astigmatic groups. Analysis of variance (ANOVA) indicated significant effects of age (F(2,448) = 32.09, p < 0.001 for V stimuli; F(2,448) = 26.66, p < 0.001 for H stimuli) and astigmatism group (F(1,448) = 11.67, p < 0.002 for V stimuli; F(2,448) = 5.27, p < 0.03 for H stimuli) and no interaction between age and astigmatism group. Post hoc comparisons (with Bonferroni correction) indicated that all pairwise comparisons of age for both V and H stimuli were significant (all p values <0.004).

Fig. 2 plots the difference in acuity for V vs. H gratings, to determine whether MA is present in the astigmatic group. The values plotted are based on the absolute value of the difference between V and H acuities for each child, because in the absence of cycloplegic refraction data for sphere, it was not possible to know whether a child was a myopic, mixed, or hyperopic astigmat, and therefore to predict whether V or H gratings would be expected to yield better acuity if MA were present (i.e., in myopic/mixed, we would predict acuity for V would be better, and in hyperopic, we would predict acuity for H would be better). ANOVA indicated no significant effect of age or of astigmatism group (astigmatic vs. low/non-astigmatic), indicating no evidence of MA for the astigmatic group. Mean difference between acuity for V and H gratings was 0.26 octave (95% CI 0.20 to 0.32) for the astigmatic group and 0.21 octave (95% CI 0.18 to 0.24) for the low/non-astigmatic group.

Grating Acuity in the Subset of 152 Children Who Underwent Cycloplegic Retinoscopy

ANOVA indicated no difference in acuity for either V or H gratings across three groups of children: 124 with low/no astigmatism (≤2.00 D), 7 with myopic or mixed astigmatism >2.00 D, and 21 with hyperopic astigmatism >2.00 D. Mean acuity for V gratings was 4.36 cy/deg (SD 0.44 octave) for the low/no astigmatism group, 4.90 cy/deg (SD 0.59 octave) for the myopic/mixed...
astigmatism group, and 4.55 cy/deg (SD 0.34 octave) for the hyperopic astigmatism group. Mean acuity for H gratings was 4.45 cy/deg (SD 0.37 octave) for the low/no astigmatism group, 4.90 cy/deg (SD 0.39 octave) for the myopic/mixed astigmatism group, and 4.32 cy/deg (SD 0.35 octave) for the hyperopic astigmatism group. Fig. 3 plots the difference in acuity for V vs. H gratings for the three groups of children. An ANOVA indicated no significant difference among groups. Mean difference in acuity (V-H results) was close to 0 in all three groups.

**DISCUSSION**

The results of this study of 448 children 6 months to <3 years of age, 121 (24.3%) of whom had astigmatism >2.00 D, indicated that astigmatism-corrected acuity for both V and H gratings in children with high astigmatism was worse than that of children with astigmatism ≤2.00 D, and that the reduction in acuity was present for children in all three age groups examined (6 months to <1 year, 1 to <2 years, and 2 to <3 years). These data suggest that amblyopia is present. However, no evidence of MA was found for the high astigmatism group as a whole or when data were analyzed for each age group.

The reduction in grating acuity for both V and H gratings in astigmatism-corrected infants and toddlers in comparison with low/non-astigmatic children of the same age is a new finding, because previous studies of children in this age range have not included a non-astigmatic comparison group. However, if most of
the infants and toddlers in the astigmatic group had hyperopic astigmatism, which is suggested by cycloplegic refraction of a subset of the children, then the finding of reduced acuity for both V and H gratings is consistent with data from hyperopic astigmatic children aged 3 years and older in this population.\textsuperscript{5,8} Preliminary data from these older children suggest that, when uncorrected, young hyperopic astigmats may accommodate between the two astigmatic focal planes,\textsuperscript{24} rendering both orientations out of focus during development.

Our failure to find evidence of MA in infants and toddlers is in agreement with the results of previous studies of astigmats in this age group.\textsuperscript{13–16} One explanation for the failure to find MA in infants and toddlers is that MA does not develop at these young ages. Another explanation, however, is that most astigmatic children in our sample were probably hyperopic astigmats, a condition that is not likely to result in MA in children in this population.\textsuperscript{3,8} It is also possible that, while MA is not found in children younger than 3 years, astigmatism during this age range may lead to later development of MA. This would be consistent with the finding of Gwiazda et al.,\textsuperscript{12} who tested non-astigmatic 5- to 11-year-old children whose refractive error development had been followed longitudinally and found meridional differences in vernier acuity that were correlated most highly with the amount of astigmatism present late in the first year up to 2 years of age. The meridional differences were least correlated with the amount of astigmatism early in the first year of life, and between 2 years of age and their age at the time of test (5 to 11 years).\textsuperscript{12}

The design of this study did not include cycloplegic refraction of all subjects, so we do not know what proportion of the overall sample of astigmats was hyperopic. However, a subset of the study population did receive a complete eye examination, including cycloplegic retinoscopy. This was not a random sample; instead, parents were offered an eye examination for their child on a different date if there was evidence of ocular abnormalities on cover testing or evidence of high refractive error based on SureSight measurements. Among the 152 children whose parents took them for the eye examination, 28 (18.4%) had astigmatism >2.00 D based on the cycloplegic retinoscopy, and 75.0% of these had simple or compound hyperopic astigmatism. Grating acuity results showed no evidence of MA in the group of 21 hyperopic astigmats or in the 7 myopic/mixed astigmats (Fig. 3). Because the cycloplegic retinoscopy was a clinical exam for safety purposes and referral for further care if indicated, rather than gold-standard measurement of refractive error, it is possible that there were additional children with astigmatism ≥2.00 D who were not identified and who could have provided additional information concerning grating acuity in hyperopic, myopic, and mixed astigmats. Data from a larger number of myopic and/or mixed astigmats and more precise cycloplegic measures of refractive error would help to determine whether MA does occur in children younger than 3 years. However, data from the relatively small number of hyperopic astigmats identified do support the hypothesis that the reason data in the present study show reduced acuity for both V and H gratings, without evidence for MA, is likely due to a high proportion of the astigmatic infants and toddlers being hyperopic, rather than myopic or mixed, astigmats.

One noticeable finding of the grating acuity results for both astigmatic and non-astigmatic subjects is that they are at the low end of the normal range for monocular acuity results for children of the same age.\textsuperscript{25,26} It is likely that this result is due to several sources of distraction to the child (wearing spectacles during testing, having siblings nearby during testing, and other activity going on at the Women, Infants and Children clinic). It could also reflect under-correction or over-correction of astigmatism in children who did not cooperate for over-refraction with the SureSight when the spectacles were worn. However, the percentage of subjects who failed to cooperate for the over-refraction was small, only 6.9%.

This study has both strengths and limitations. Strengths include a large number of subjects, assessment of subjects who were younger than 3 years, and the fact that all subjects, including non-astigmats, wore spectacles during testing. Limitations include the possibility that the astigmatism-correcting lenses under- or over-corrected the astigmatism in the small percentage (6.9%) of children in whom over-refraction when the spectacles were worn was not possible; the absence of cycloplegic refraction data with which to classify all children as hyperopic, myopic, or mixed astigmats; and the fact that one-fourth of the children failed to cooperate for grating acuity testing: 20% (74/361) at 6 months to <1 year, 41% (108/263) at 1 to <2 years, and 28% (46/162) at 2 to <3 years. If the relatively high proportion of children who failed to cooperate for acuity testing had included a substantial number of high astigmats, it would suggest that the acuity results obtained from the cooperative group might have underestimated the true reduction in acuity that would have been found if all high astigmats could have been tested. However, comparison of the uncooperative and the cooperative groups indicated no difference in the proportion with high astigmatism (20.3% vs. 25.0%), suggesting that the grating acuity results obtained were probably representative of results from the entire group of children.

In conclusion, the results of this study failed to find evidence of orientation-related differences in astigmatism-corrected grating acuity in astigmatic infants and toddlers younger than 3 years who were tested with spectacle correction of astigmatism. However, even in the youngest age group tested (age 6 months to <1 year), astigmats tested with spectacle correction showed reduced acuity for both V and H gratings, suggesting that astigmatism is having a negative influence on visual development.

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