Diagnosis and Therapy in Ophthalmology

Hand-held optical coherence tomography imaging in children with anterior segment dysgenesis

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In this study, we investigated the potential of hand-held optical coherence tomography (HH-OCT; Fig. 1) to improve diagnosis in anterior segments (AS) by visualizing the anterior and posterior eye structures without general anaesthetic (GA) or sedation.

Patients were prospectively recruited from paediatric clinics at the Leicester Royal Infirmary (Table 1).

Case 1: A 5-year-old patient was seen because of conjunctivitis. An irregular circular white structure located underneath of the endothelium anteriorly to the limbus was found incidentally (Fig. 2). Diagnosis of Axenfeld’s anomaly was made. Optical coherence tomography (OCT) angle scans demonstrated a prominent anteriorly displaced Schwalbe line. The patient had several small (less than 1 mm) zones of the iridocorneal adhesions originating from the angular part of the iris and fixed behind the displaced Schwalbe line (Fig. 2E, right) in both eyes.

Case 2: A 5-day-old child was referred because of suspected left cataract. Anterior segments (AS) examination showed bilateral paracentral corneal opacities (Fig. 2D). A tentative diagnosis of Peter’s anomaly was made. At the last assessment, biomicroscopy showed structural changes of iris and a round white structure located underneath the endothelium anteriorly to the limbus bilaterally. No cataract was seen. At this point, diagnosis was revised to Axenfeld’s anomaly. Optical coherence tomography (OCT) showed a heterogeneous cornea (Fig. 2F) bilaterally with a smooth epithelial contour and uneven endothelium corresponding to the severe corneal opacities on slit-lamp examination (Fig. 2D). The patient had a ‘band’ around the entire iris bilaterally forming a ‘ring’ located 1–2 mm from the trabecular meshwork.

The AS OCT findings allowed diagnosis of iridocorneal endothelial syndrome (ICE) because of endothelium irregularity and the absence of iridocorneal adhesions and embryotoxon.

Case 3: A 2-month-old baby was referred due to a left corneal opacity. Investigation under GA showed iridocorneal and corneolenticular adhesions associated with corneal/lens opacity.

Optical coherence tomography (OCT) at the age of 6 years showed an irregular corneal structure with pericentral corneal thickening on the affected side (Fig. 3). An iridocorneal adhesion began at the thickest part of corneal endothelium and entered into the pupillary part of the iris. Typical
Table 1. Age, gender, ethnicity and clinical data of the patients with congenital anterior segment abnormalities on the date of optical coherence tomography examination.

| Patient’s number | Pathology                              | Age (years) | Gender | Ethnicity | LogMar visual acuity | Refraction (spherical equivalent) | Intraocular pressure | Associated ophthalmic pathology | Associated general pathology | Clinical findings                                                                 | OCT findings                                                                 |
|------------------|----------------------------------------|-------------|--------|-----------|----------------------|-----------------------------------|----------------------|--------------------------------|---------------------------------|---------------------------------------------------------------------------------|--------------------------------------------------------------------------------|
| 1                | Axenfeld’s anomaly                      | 5           | F      | C         | 0.1                  | +1.0                              | 13                   | 14                             | –                               | –                                                                               | Irregular circular white structure underneath of the endothelium                | Prominent anteriorly displaced Schwalbe line                                   |
| 2                | ICE syndrome                            | 5           | F      | C         | 0.3                  | 0.9                               | 14                   | 14                             | Left esotropia, MLN              | Arthrogryposis multiplex Congenital Hypothyroidism                              | Bilateral paracentral corneal opacities; round white structure underneath the endothelium | Smooth epithelial contour and uneven corneal endothelium; a ‘ring’ located 1–2 mm from the trabecular meshwork in BE |
| 3                | Peter’s anomaly                         | 6           | M      | C         | 0.4 POL              | +1.25                             | 13                   | 12                             | Left esotropia, MLN              | –                                                                               | Iridoconal and corneolenticular adhesions associated with corneal/lens opacity | Irregular corneal structure with pericentral corneal thickening on the affected side with iridocorneal adhesion; no iris folds on the LE; RE cup diameter/depth smaller than in control |
| 4                | PAX6 mutation with aniridia             | 14          | M      | C         | 0.5                  | 0.4                              | 15                   | 15                             | Infantile nystagmus              | –                                                                               | Aniridia and small central lens opacities bilaterally                          | Missing central part of the iris with hypoplastic stump 360°; foveal hypoplasia with continuation of RNFL, ganglion cell, inner nuclear and inner plexiform layers |
| 5                | Iris, optic nerve coloboma              | 12          | M      | C         | 0.85                 | 0.1                              | 14                   | 14                             | –                               | –                                                                               | LE iris and choroidal coloboma                                                 | Hypoplastic irides with a remaining stump; large ON with RNFL thinning; large LE foveal pit |
| 6                | Primary childhood glaucoma              | 4           | M      | C         | 0.9                  | 0.1                              | 9                    | 12                             | –                               | –                                                                               | Large RE corneal diameter with haze and Haab’s stria; abnormal iris insertion on gonio; cup/disc ratio of 0.6 | Additional tissue on the endothelium RE; abnormal iris insertion; increased cup/disc ratio with RNFL thinning |

BE = both eyes, F = female, ICE = iridocorneal endothelial syndrome, LE = left eye, M = male, MLN = manifest latent nystagmus, OCT = optical coherence tomography, ON = optic nerve, POL = perception of light, RE = right eye, RNFL = retinal nerve fibre layer.

Visual acuity in the affected eye is shown in bold.
contraction folds of the iris were missing (Fig. 3B). The anterior angle in the affected right eye (RE) was open and the iris root appeared thinner (Fig. 3C) than in the left eye (LE) (Fig. 3E) and healthy control (Fig. 3I). On the unaffected side, in the patient with transparent cornea, cup diameter and cup depth were smaller as compared to the control.

**Case 4:** A 6-month-old child presented due to a family history of PAX6 gene mutation. Examination showed the presence of aniridia and small central lens opacities bilaterally.
On OCT (Fig. 4), aniridia was visualized by a missing central part of the iris. A small flat hypoplastic iris stump was detected 360° around the anterior chamber. The patient had clearly defined scleral spurs (Fig. 4B, dotted arrow). Posterior examination showed severe foveal hypoplasia with continuation of retinal nerve fibre layer (RNFL), ganglion cell, inner nuclear and inner plexiform layers in the fovea bilaterally (Fig. 4E).

Case 5: A 3-month-old patient presented because of irregular pupil. Clinical examination showed left iris and choroidal coloboma. Optical coherence tomography (OCT) showed hypoplastic irides with a remaining iris stump at the area of the coloboma.

A horizontal tomogram through the centre of the coloboma (Fig. 5E) showed the optic nerve (ON) with a significantly larger distance between the edges of RPE. Optic nerve (ON) cup diameter and depth in the affected eye were larger and nasal and temporal RNFL were thinner than in the fellow/control eye. Foveal pit was also larger in the unaffected eye (Fig. 5F).

Case 6: A 4-month-old patient was referred because of a larger RE. Eye examination under GA revealed asymmetrical horizontal corneal diameters with right corneal haze and Haab’s striae. Gonioscopy showed abnormal iris insertion with iris root seen in front of the trabecula meshwork. Optic nerve (ON) assessment showed an enlarged cup in the RE with a cup/disc ratio of 0.6. Diagnosis of primary congenital glaucoma (PCG) was established. Hand-held optical coherence tomography (HH-OCT) at the age of 4 years showed the presence of an additional tissue on the endothelium in the centre of the right cornea (arrow on the Fig. 6A) that was not seen on clinical assessment. Optical coherence tomography (OCT) also confirmed abnormal iris insertion (Fig. 6B). The horizontal cup diameter and maximal cup depth were larger in the glaucoma eye. Retinal nerve fibre layer (RNFL) thickness was approximately 15 µm thinner in the eye with glaucoma.

Anterior OCT assessment enabled the pathology of the cornea to be determined in 4 of 6 patients that was not found clinically. In case 2, OCT findings dictated the necessity of long follow-up of this patient as ICE is slowly progressive condition with extension of the endothelial membrane over the angle and high risk of subsequent glaucoma (Chandran et al. 2015).

Lai et al. (2013) described good visualization of the structures of the angle with swept-source OCT and showed that it is possible to use OCT for risk assessment of adult angle-closure glaucoma (Lai et al. 2013; Radhakrishnan & Yarovoy 2014; Sharma et al. 2014). We also found the OCT to be a useful technique in the
assessment of the iridocorneal structure in case 6 with PCG, who had a large iridocorneal angle with insertion of the iris root in front of the Schlemm canal on OCT. To our knowledge, there are no data describing angle structure using OCT in patients with PCG. In PCG, patients often have some corneal opacification and Haab’s striae (Patil et al. 2015). Abnormal iris insertion with the iris root starting in front of the Schlemm canal was clearly visualized on OCT. This indicates that OCT can be helpful in selecting the best surgical techniques using goniotomy and trabeculotomy if angle structure is preserved and drainage devices in severe angle dysgenesis.

Optical coherence tomography (OCT) findings in cases 4 and 5 indicate that subclinical maldevelopment can be present in retinal structures in clinically unaffected eyes. Important clinically relevant information helping diagnosis and management can be obtained non-invasively without GA or sedation with anterior and posterior OCT to assist diagnosis and management of anterior segment dysgenesis (ASDs) especially PCG.

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Fig. 6. Spectral domain optical coherence tomography of case 6 with primary childhood glaucoma in the right eye (A–C) and the healthy control (D–F). The white arrow on figure A shows additional tissue on the endothelium in the centre of the cornea; arrow on figure B indicates abnormal iris insertion (iris root in front of the Schlemm canal) when compared to control; the horizontal tomogram of the optic nerve in the patient with glaucoma showed a larger horizontal diameter and depth of the cup as compared to the healthy control (F).