**Leukocoria Caused by Intraocular Heterotopic Brain Tissue**

Leukocoria caused by diverse ocular conditions is distinguishable by history, characteristic clinical findings, and ancillary imaging. Atypical cases pose a diagnostic dilemma necessitating enucleation if retinoblastoma or malignancy cannot be excluded. We report a newborn with leukocoria in a microphthalmic eye containing an uncalcified subretinal mass. Teratoid medulloepithelioma was suspected. Instead, benign heterotopic brain tissue was found, which to our knowledge has only been reported once before, due to aberrant differentiation of the neuroectoderm into cerebral gray matter instead of the retina.

**Report of a Case.** A 5-day-old, healthy baby girl with leukocoria of the microphthalmic right eye was seen in the Retinoblastoma Program at the Hospital for Sick Children, Toronto, Ontario. Perinatal and family histories were unremarkable. The left eye was normal, but the right eye appeared blind. Under general anesthesia, intraocular pressure and ocular movements were normal. The 8-mm-diameter right cornea was clear and the anterior chamber, deep. The iris was hypoplastic with persistent tunica vasculosa lentis bridging the pupil. The irregularly shaped lens was small but clear. Zonules were incomplete (Figure 1). A large, nasal, creamy-white, elevated subretinal mass, obscuring the retina and optic nerve, extended into the clear vitreous until just behind the lens.

B-scan ultrasonography results revealed a homogeneous, uncalcified tumor filling two thirds of the eye (Figure 2). Sclera underlying the tumor bulged like a staphyloma. The medial rectus muscle moved freely. Computed tomography results showed an abnormal-shaped, small eye with ectatic or deficient sclera underlying the uncalcified tumor and medial rectus (Figure 3). Coronal T2-weighted magnetic resonance imaging results showed no melanin or blood in the tumor, which was isodense with cerebral cortex (Figure 4). The tumor border was smooth and concave with a vitreous band, possibly persistent hyaloid, extending across the vitreous/subretinal space to the sclera. There was no extraocular extension. The optic nerves and brain appeared normal.

Malignant teratoid medulloepithelioma with persistent hyperplastic primary vitreous was suspected. Serial B-scans during 2 weeks suggested growth and change in vitreous density. Eye Cancer Network and International Tumor Board (New York, NY) consultations all recommended enucleation of this blind eye with possible malignancy. The 16 × 17 × 16-mm eye showed no scleral defects. On opening the eye through an inferior, horizontal pupil-optic nerve incision, a solid, uncalcified, homogeneously grayish-white tumor unlike retinoblastoma was found arising in a staphylomatous choroidal coloboma, filling much of the vitreous and abutting the lens anteriorly (Figure 5). Where imaging suggested deficient sclera was a choroid coloboma with scleral ectasia.

The tumor, eosinophilic on hematoxylin-eosin stain, resting di-
directly on ectatic sclera in the anterior retina/ciliary region, extended backward across the choroid to the optic nerve (Figure 6A). Areas of dysplastic retina were thrown into focal pseudorosettes in their periphery (Figure 6A). The tumor was composed of brain tissue strongly positive for synaptophysin (Figure 6B). Large, mature, crystal violet–positive neurons (Figure 7A, B, and D); glial, fibrillary, acidic, protein–positive, plump oligodendrocytes; and spindle-shaped astrocytes (Figure 7A and C) were admixed with neuropil containing rare focal calcification. The cornea and anterior chamber were unremarkable. Lens epithelial and bladder cells had migrated into the posterior subcapsular area. The ciliary body was attenuated. The tumor appeared to arise from the retina in some areas (Figure 8A) and the retinal pigment epithelium in others (Figure 8B). The 6-mm, tumor-free optic nerve was atrophic, drawn into the globe under traction (Figure 8B).

Comment. Normal brain tissue not communicating with the central nervous system (brain heterotopia) has been reported in the head and neck, rarely in the orbit, at the limbus and in 1 case in the anterior segment of an infant’s eye. Intraocular heterotopic brain tissue arising from the retina or anterior retinal pigment epithelium was reported recently. To our knowledge, this is the second report.

Previous authors proposed that when embryonal neurectodermal
stem cells segregated outside the central nervous system during development, teratomatous proliferation or true astrocytomas or benign brain heterotopia could result. In our patient, the tumor appeared to arise from both the retina and retinal pigment epithelium within a coloboma, suggesting abnormal neurectoderm dif-

Figure 7. A, High-power photomicrograph of the tumor demonstrating mature neurons (arrow), oligodendrocytes (diamond-headed arrow), and astrocytes (open arrow) interspersed in neuropil, typical of cerebral gray matter (hematoxylin-eosin). B, Crystal violet stain of mature neurons within the tumor (arrow). The stain bound very strongly to RNA in the rough endoplasmic reticulum of the neuron. Note the prominent nucleolus. C, Glial fibrillary acidic protein in astrocytes (arrows), the most common glia in the tumor. D, A single neuron within the tumor immunolabeled with neurofilament antibody. Note the dendritic processes (D) and large central nucleus (N) typical of most neurons (all images original magnification ×400).

Figure 8. A, Relationship between the tumor (T), choroid (C), and normal retina (R) with the tumor appearing contiguous with the outer retina (arrow) (hematoxylin-eosin, original magnification ×40). B, The tumor (T) appeared to have drawn the optic nerve (ON) into the eye. Note the displacement of the retina (R) and apparent continuity of the tumor with the retinal pigment epithelium (RPE) (arrows) (hematoxylin-eosin, original magnification ×100).
fication within the optic cup in the embryonic fissure, causing subse-
quently microphthalmia and anterior segment maldevelopment. The path-
genesis is uncertain; failure of a clone of pluripotent stem cells in the em-
bryonic fissure to initiate normal reti-

nal development may have oc-
curred. This may be due to a somatic mutation in a gene involved in the retinal signaling pathway, such as CHX-10, PAX6, PAX2, Rb; or sonic hedgehog, SHH, which are re-
quired for vertebrate retinal develop-
ment. These mutations potentially could leave cortical development as a default pathway for retinal differ-
entiation. Thus, a local cerebrocorti-
cal cell mass arising in the anterior retina/ciliary region may be ana-
alogous to experimental eye formation in ectopic locations when the Dros-
ophila gene, eyeless (ey), homolo-
gous to mammalian PAX6, drives eye development wherever ectopic ex-
pression occurs. A germline dele-
tion in SHH has been associated with iris and uveoretinal colobomas. We hypothesize in the present case that somatic mutation in such a gene may be causative.

The finding of leukocoria in a child requires thorough evaluation to exclude retinoblastoma or other intraocular malignancies. This neo-
natal development of an uncalcifi-
fied, homogeneous intraocular mass in a microphthalmic eye was more consistent with a developmental anomaly than retinoblastoma. With no possible useful vision but poten-
tial malignancy, including atypical teratoid medulloepithelioma or retinoblastoma, enucleation of the eye was indicated. This child contin-
tues to thrive without needing further medical tests or interven-
tions.

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Orbital Myositis in Churg-Strauss Syndrome

A case of orbital myositis in a patient with Churg-Strauss syndrome (CSS) is reported herein. To our knowledge, this association has been reported only once previously. We also describe the previously re-
ported ophthalmic manifestations of CSS.

Report of a Case. A 55-year-old woman was referred to our unit with a 1-week history of redness and pain behind the right eye, which wors-
ened on eye movement.

Her medical history included adult-onset, steroid-dependent asthma, diagnosed at 30 years of age and currently stabilized with in-
haled corticosteroid use. She also had, from allergic rhinitis, nasal pol-
yps and severe sinus disease requiring repeated surgery. She had been noted to have peripheral blood eos-
inophilia on several occasions for the last 7 years and had a recurrent rash for the last 4 years involving her trunk and back, responsive to corti-
costeroid treatment and found to be interstitial granuloma annulare on tissue biopsy results obtained 4 years earlier. A repeated biopsy 1 year later revealed an intense urticarial inflam-
matory reaction with perivascular eosinophilic and lymphocytic inflam-
mmatory infiltrates. Peripheral blood eosinophil levels, 4 and 3 years previously, were raised at 2300/µL and 2060/µL, respectively. A bone marrow biopsy obtained 3 years earlier revealed a marrow eosinophilia and an elevation of eosinophil pre-
cursor populations (eosinophil and eosinophil precursors, 10%) with normal erythropoiesis, myelopo-
iesis, marrow cells, and marrow archite-
chture. Findings from bone mar-
row lymphocyte surface marker analysis and gene rearrangement studies were normal. Allergies in-
cluded sulfur, aspirin, and certain foods, for which she had under-
gone desensitization therapy.

On examination, she had swell-
ing and tenderness of the right up-
per lid and bony tenderness over the right cheek. Her right conjunctiva was inflamed and edematous with a
initially referred for monocular hyphema.

**Report of a Case.** A 25-year-old white man was initially seen with a 1-week history of floaters and decreased vision in his left eye. The patient had been diagnosed with Tourette syndrome at age 7 years, obsessive-compulsive disorder at age 11 years, and depression at age 24 years. His motor tics involved excessive blinking, blepharospasm, clapping, jabbing his fingers into his eyes, and punching himself in the periocular area. The patient was taking buspirone hydrochloride (10 mg twice a day) and clomipramine hydrochloride (25 mg twice a day). On examination, the patient was alert and oriented, and he had no evidence of cognitive impairment. Visual acuity was 20/200 OD and hand motion OS. There was no afferent pupillary defect. Intraocular pressures were 18 OD and 16 OS. Slit-lamp examination findings of the anterior chamber, as well as a dense posterior subcapsular cataract, appeared as follows: Sarit Patel, MD, Madison, Wis; Joanne Dondey, MD, Helen Eliott, MD, Detroit, Mich

The left eye had a less than 1-mm hyphema and many (4+) circulating red blood cells in the anterior chamber, as well as a dense posterior subcapsular cataract. The left eye had a less than 1-mm hyphema and many (4+) circulating red blood cells in the anterior chamber, as well as a dense posterior subcapsular cataract. The left eye had a less than 1-mm hyphema and many (4+) circulating red blood cells in the anterior chamber, as well as a dense posterior subcapsular cataract. The left eye had a less than 1-mm hyphema and many (4+) circulating red blood cells in the anterior chamber, as well as a dense posterior subcapsular cataract.

Funduscopescopy revealed a retinal dialysis from the 1:30 to the 4:30 clock position with a macula-on-retinal detachment in the right eye. Vitreous hemorrhage was present centrally in the left eye, and there were nasal and temporal giant retinal tears. The right eye was re- paired with a scleral buckling procedure. The left eye underwent anterior segment washout, pars plana lensectomy, pars plana vitrectomy, endolaser, and silicone oil injection. Intraoperatively, the giant retinal tears were found to extend from the 12:30 to the 4:30 clock position with 4 long radial extensions to the temporal macula and from the 6-o'clock to the 11-o'clock position with 1 long radial extension to the optic disc. There was an additional radially oriented posterior retinal break. Postoperatively, the retinas were attached in both eyes. One month later, the left eye developed proliferative vitreoretinopathy with retinal detachment and underwent reoperation. At 6 months, the retinas remained attached and the visual acuity was 20/100 OU.

**Comment.** Ophthalmic manifestations of Tourette syndrome include frequent blinking and blepharospasm, gaze deviations and abnormal saccades, and accidental and self-inflicted ocular injuries.3-5 The retinal detachments in our patient were most likely the result of repeated, self-induced finger jabbing to the eyes since the patient had no other risk factors for retinal detachment. In patients with retinal detachment, factors suggesting a traumatic etiology typically include unilateral vitreoretinal findings, retinal dialysis or giant retinal tear, and age younger than 40 years.6,7 However, in patients with self-induced or repeated trauma, the vitreoretinal pathologic features may be bilateral, as demonstrated by our patient. To prevent further self-injury, patients should wear protective polycarbonate goggles, and they should be monitored closely in conjunction with the psychiatry service. Treatment of the underlying disorder with behavior modification and pharmacotherapy is essential, and pharmacological agents that antagonize dopamine are most effective in reducing the severity of motor and vocal tics.

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