LETTERS TO THE EDITOR

Laceration of the eye with a fishing hook

Editor,—Perforation of the eye is a challenging emergency in ophthalmology and requires immediate treatment. Visual outcome after penetrating injuries with and without intraocular foreign bodies depends on the visual acuity after injury, age of patient, and the severity of the ocular trauma.1 7 We report a case of globe laceration following an accident with a fishing hook which was successfully treated without surgery.

CASE REPORT
A 12 year old boy presented with visual disturbance and a small lid wound on the left eye at our emergency department. He had been playing with a fishing rod while sitting on a tree and the hook jerked itself out of the upper lid of the left eye. He jumped from the tree and the hook jerked itself out of the boy’s eye. A small wound of the upper lid was found. The visual acuity was 20/20 in the right eye and 20/25 in the left eye. The anterior segment was normal. The fundus of the right eye was normal. The fundus of the left eye showed a mild localised vitreous haemorrhage superotemporally near the temporal vessels. From the scleral rupture went straight up to the equator. Partly orbital fat could be seen through the wound gap (Fig 1A). The ultrasound disclosed a partly detached vitreous with adherence at the site of penetration (Fig 1B). No foreign body was detected by the orbital computed tomograph scan. The patient was admitted to hospital for 8 days and treated with intravenous antibiotics during 1 week followed by an oral antibiotic in the second week. He also received cortisone systemically for 10 days. Three weeks later the visual acuity was 20/20. On funduscopy a sclerochoroidal scar was disclosed. The edges of the retinal tear were attached and vitreous haemorrhage was completely dissolved (Fig 2). After 1 year of follow up the situation remained unchanged.

COMMENT
Although penetrating injuries of the posterior segment often require surgical treatment (that is, pars plana vitrectomy), in this case antibiotic therapy was used in order to prevent an endophthalmitis1 in combination with orally administered cortisone to reduce the inflammatory reaction. Because of an excellent fundus view, postponing surgery seems to be more appropriate, since surgery implies additional risks (for example, cataract formation, retinal detachment, etc) for the eye.7 Significant predictors for a final visual acuities of 20/50 or better are a visual acuity of 20/800 or better and youth (< 18 years).1 In a marmalian study it was disclosed that simple penetration of the equator with vitreous loss does not lead to retinal detachment.7 An intact tamponading vitreous at the time of injury seems to prevent fibrous ingrowth due to anti-proliferative effects of the hyalocytes. The findings of this case suggest that surgery is not the first treatment strategy for similar penetrations of the posterior segment.

RALF KROTT
K ULRICH BARTZ-SCHMIDT
KAUS HEIMANN
Department of Vitreoretinal Surgery, University of Cologne, Germany

Correspondence to: Ralf Krott, MD, Department of Vitreoretinal Surgery, University of Cologne, Joseph-Stelzmann-Strasse 9, D-50924 Cologne, Germany.

Accepted for publication 11 June 1999


7 Hsu HAT, Ryan SJ. Natural history of penetrating ocular injury with retinal laceration in the monkey. Graded Arch OucEyes 1986;224:1–6.


Intraocular infestation with the worm, Thelazia callipaeda

Editor,—Ocular infections with helminthic parasites are well described. The commonest organisms are filarial worms that reside in subcutaneous tissue, and are found as skin infestations or masses in the lids. Some are known to live freely in the conjunctival sac. Worms that are visible to the naked eye are often referred to as “eyeworms”, and are in the larval or adult stage of their life cycle. Thelazia callipaeda, or the oriental eye worm, is a spiruroid nematode which is the causative organism in thelaziasis, a well described condition affecting the external eye.1 It is primarily a parasite of the conjunctiva in dogs, and is also found in rabbits and humans.2 Its presence in the conjunctival sac causes lacrimation and irritation, and its frequent excursions across the cornea may cause marked discomfort and, eventually, corneal scarring. The worm also causes paralytic ectropion through its presence in the lower fornix. At least 40 cases of infection in humans have been described from China, Japan, India, Russia, Thailand, and Korea. All of these report extraocular infection only. We report the first case of Thelazia callipaeda to cause intraocular infection.

CASE REPORT
A 21 year old Chinese woman presented to the ophthalmology department at Shantou Central Hospital, Guangdong, China, complaining of decreased vision in her right eye and a floater in the right visual field. She had no photopsia or field loss, and no pain, redness, or discharge. She had no past ocular or medical history, and was otherwise well.

On examination she had a visual acuity of 6/60 in the right eye, and 6/6 in the left eye. There was no external evidence of trauma; the lids were healthy, the conjunctiva white, the cornea clear, and the anterior chamber quiet. A mild vitreous haze obscured the foveal reflex, and clearly visible within the vitreous cavity was a live, mobile, white worm. There was no retinal abnormality.

One month later she underwent a three port pars plana vitrectomy, and the worm was coaxed into a flute needle and removed intact. The patient made an uncomplicated recovery from surgery. At 6 weeks postoperatively the eye was quiet and she had a visual acuity of 6/24. On detailed examination, the worm was identified as an adult female specimen of Thelazia callipaeda. Treatment with ivermectin is recommended for cases of thelaziasis, but in the absence of preoperative and postoperative
Thelazia callipaeda, a member of the phylum Nematoda, order Spirurida, suborder Spiruridea. Adult worms look like creamy white threads; males are 4.5–13 mm long and 0.25–0.75 mm in diameter; females are larger, ranging from 6.2 to 17 mm in length and 0.3–0.85 mm in diameter (Fig 1). It is known to infect the lacrimal sac, from where it passes into the conjunctival sac. It is probably deposited in the larval form by flies settling on the eyes and this accounts for extraocular infection. The adult worm in our patient’s eye was fully developed and healthy. It had caused little inflammatory response, and appeared to thrive in the conditions provided by the vitreous cavity, suggesting the possibility of immune privilege. We cannot explain how this worm entered the eye. We put forward the suggestion that humans may be the definitive host, with dogs and rabbits being the reservoir. For intraocular colonisation, the infective stage is probably the filariform larva, the portal of entry being the skin. An alternative mode of infection may be as the larva or embryonated egg ingested with raw drinking water.

**COMMENT**

Malattia leventinese, the maculopathy is characterised by a radial pattern of innumerable small elongated basal laminar drusen. This maculopathy has been described in a 15 year old patient. The visual acuity of patients suffering from malattia leventinese, however, remains good for quite a long time. Thus, most patients are asymptomatic until the fourth or fifth decade of life, at which point they have a variety of symptoms, including decreased visual acuity, para-central scotomas, photophobia, and metamorphopsia. The main complication is macular subretinal neovascularisation, reported in some patients. In our present case, such macular subretinal neovascularisation caused a severe decrease in visual acuity down to 20/1000 in 1981. The contralateral eye was unaffected, fortunately conserving an acuity of 20/30.

**CASE REPORT**

In 1973, ophthalmological examination led to a diagnosis of bilateral hereditary macular dystrophy (malattia leventinese) in a woman born in 1943. Her visual acuity was then 20/20 for both eyes without any correction (Fig 1A). In 1981, subfoveal neovascularisation led to an irreversible decrease in visual acuity in her right eye, down to 20/1000 (Fig 1B). In 1996, a dense right vitreous haemorrhage led to a further decrease in acuity. After resorption, fundus examination disclosed an advanced stage of the macular disease with irregular subretinal metabolism, hyperplasia of the retinal pigment epithelium and discrete radial basal laminar drusen (Fig 2A). The fundus also showed a wedge-shaped superotemporal area with intraretinal haemorrhages, hard exudates, and sheathed vessels. The fluorescein angiogram showed telangiectatic vessels, shunt vessels, and microaneurysms, in addition to neovascularisation (Fig 2B). After laser photocoagulation, no further intra- or extraocular haemorrhage episode occurred. The left eye had a visual acuity of 20/30 and fundus examination revealed a macula identical to that of the right eye, without complication.


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**Figures**

1. Light micrograph of Thelazia callipaeda (magnification ×100). (A) Electron micrograph of tail end of Thelazia callipaeda (magnification ×1000). (B) Electron micrograph of head end of Thelazia callipaeda (magnification ×1000).

2. (A) Right eye. Fundus photograph in 1973 shows a radial pattern of innumerable small elongated basal laminar drusen. (B) Right eye. Fluorescein angiogram in 1981 shows submacular neovascularisation with fluorescein leakage.
Acute central retinal vein occlusion successfully treated with intravenous thrombolysis

EDITOR,—Central retinal vein occlusion (CRVO) is a condition which often has profound effects on vision. At present there is little to offer patients in the form of treatment to preserve vision. The visual outcome is largely determined by the severity and duration of the vein occlusion. Management is currently aimed at preventing the complications secondary to retinal ischaemia. The incidence of fellow eye involvement with CRVO is believed to be in the order of 1%. We present a patient with “second eye” CRVO, who presented with acute reduction in vision and who responded dramatically to intravenous streptokinase.

CASE REPORT

A 75 year old white man noticed a sudden reduction in vision in his right eye while walking up a hill. He already had poor vision in his left eye from a CRVO 6 years earlier. He went immediately to eye casualty and was seen within 2 hours from the onset of symptoms. His only risk factor for vein occlusion was ocular hypertension treated with timoptol 0.25% twice daily to both eyes.

On examination, visual acuities were 6/36 right eye and counting fingers left eye, there was no relative afferent pupillary defect and intraocular pressures were 21 mm Hg in both eyes. Examination of the right fundus revealed scattered tiny blot haemorrhages and tortuous veins. A fundus fluorescein angiogram showed pulsatile arterial filling with venous filling delayed until 34.6 seconds (Fig 1). A diagnosis of acute central retinal vein occlusion was made. Because of the previous left CRVO and the short history of symptoms in his right eye, the option of thrombolysis was carefully discussed with the patient, including the small risk of cerebral haemorrhage. Streptokinase, 1.5 ×10^6 units, was infused 7 hours after the onset of symptoms. Within 30 minutes, his visual acuity had improved to 6/9. The patient was empirically anticoagulated with heparin and warfarin, keeping the international normalised ratio between 2 and 3. Repeat fluorescein angiography showed marked improvement in venous filling and loss of pulsatile arterial filling. In addition, several post thrombolysis haemorrhages were evident (Fig 2). The patient took warfarin for 9 months in total and now remains on aspirin alone. His vision in the right eye remains at 6/9.

COMMENT

Central retinal vein occlusion can profoundly affect vision and lead to neovascular complications. Current therapeutic options are limited mainly to the prevention or treatment of secondary complications.

Evidence for thrombus formation in CRVO has been reported by Green et al1 in a prospective histopathological study of patients with CRVO. They demonstrated recanalised thrombus in 89.7% of eyes and fresh thrombus formation in the remaining 10.3%.2

Animal studies have shown encouraging responses to thrombolysis in experimental cases of acute central retinal vein occlusion.3

Figure 1 Fundus fluorescein angiogram of the right eye demonstrating early venous filling 34.6 seconds after injection.

Figure 2 (A) Right eye. Photograph in 1996. Advanced stage of the macular disease with irregular subretinal fibrous metaplasia, hyperplasia of the retinal pigment epithelium, and discrete radial basal lamellar drusen. Note also a wedge-shaped supertemporal area with intraretinal haemorrhages, hard exudates, and sheathed vessels. (B) Right eye. Fluorescein angiogram in 1996. Note the temporal retinal ischaemia with telangiectatic vessels, shunt vessels, and microaneurysms, in addition to the neovascularisation.
CRVO. Fibrinolysis given within 2 days of experimental vein occlusion in rabbits was shown to result in resolution of retinal haemorrhages and return of normal retinal circulation within 48 hours. Similar animal studies using recombinant tissue plasminogen activator shortly after experimentally induced vein occlusion resulted in significant retinal vein patency in treated eyes. In one randomised controlled clinical trial, patients with CRVO who received streptokinase followed by full anticoagulation within 7 days of onset of visual symptoms, showed a statistically significant improvement in visual acuity. Late presentation was identified as a possible cause of limited improvement in some cases. Reports of vitreous haemorrhage occurring during treatment (3/20 cases) together with the reported incidence of cerebral haemorrhage with streptokinase (57/13607 cases) account for the absence of a defined role of this drug in the treatment of vein occlusions.

Selective cannulation of a branch retinal vein and infusion of tissue plasminogen activator in a patient with a non-acute CRVO in the second eye has been described. Several other treatment modalities had already been tried unsuccessfully. While avoiding systemic complications, the treatment failed to show any objective improvement in visual acuity and the patient subsequently went on to develop angle neovascularisation.

Clearly many vein occlusions present late, where irreversible retinal damage has occurred. We propose that only in those circumstances where the presentation is acute should the use of intravenous thrombolysis be considered.

M T J COSTEN
W B M DONALDSON
J A OLSON
Department of Ophthalmology, University of Aberdeen, Foresterhill, Aberdeen AB25 2ZB

Correspondence to: Dr J Olson.

Accepted for publication 26 April 1999


Retinitis pigmentosa with visual fluctuations and arrestin gene mutation

Editor,—We report a case of retinitis pigmentosa with day to day visual fluctuations and a mutation in the arrestin gene.

CASE REPORT
A 45 year old Japanese man first noticed night blindness at junior high school age, followed by a slowly progressive loss of visual acuities and fields. At presentation, the best visual acuity was 10/200 in each eye. Goldmann perimetrer’s revealed generalised narrowing of the peripheral field and marked loss of central visual sensitivities. Bright flash electroretinogram (ERG) in a fully dark adapted state was not recordable in either eye. Ophthalmoscopics revealed advanced stage of retinitis pigmentosa (Fig 1). There was no Mizuo’s phenomenon in the fundus. This patient reported that he had felt relatively better vision (“good day”) on every other day that alternated with worse vision (“bad day”).

Furthermore, he claimed that bad day was repeated after physical stress or alcohol drinking, followed by a good day to return to the ordinary cycle. In our 10 years’ observation, the profile of day to day variation was reproducible. To quantitatively assess the characteristic visual symptoms, perimetric tests were performed in both the centre and periphery using an automated perimeter on eight different days in a period of 2 months, half on a good day and the other half on a bad day; ERG examinations were also performed. Figures 2 and 3 illustrate the results, demonstrating that the visual sensitivity in the peripheral field varied in a manner consistent with the patient’s reports, although not obvious in the central field. In contrast with perimetrer’s, ERG responses remained undetectable irrespective of visual fluctuations. His healthy parents were first cousins, and his elder brother had night blindness due to classic retinitis pigmentosa. Because of the mild mental retardation, the day to day visual fluctuations or other symptomatic variations could not be confirmed in his brother.

DNA was extracted from the peripheral blood after obtaining informed consent. Exon 11 of the arrestin gene was amplified using polymerase chain reaction. The amplified product was separated on polyacrylamide gel and revealed abnormally rapidly migrated signal suggesting a small deletion. Direct sequencing using an automated nucleotide sequencer (ALF Express, Pharmacia) disclosed a homozygous deletion of adenine at nucleotide 1147 (1147delA). This mutation

Figure 2 Visual field sensitivity, determined by an automated perimeter (Interzeag Octopus 101, low vision program) on different days. Representative visual field charts (right eye) obtained on a “good day” (left chart) and a “bad day” (right chart).
Figure 4 Direct sequencing results of exon 11 of the arrestin gene in the reported case, demonstrating a homozygous deletion of adenine at nucleotide 1147 (arrowhead).

Figure 3 Mean sensitivity (dB) at 76 locations; the value in normal individuals of the same age as the case is around 30 dB. Open symbols (circles, right eye; triangles, left eye) illustrate measurements on four good days and closed symbols on four bad days. Visual sensitivity on good days is significantly better than that on bad days (paired t test, p < 0.002).

COMMENT

Some patients with advanced retinitis pigmentosa report that their vision is "crystal clear" in the morning but soon becomes fuzzy, and some others announce on a particular visit that their visual field testing either went well or poorly depending on whether they are having a good or bad day.¹ The patient described here had advanced retinitis pigmentosa and reported that he had experienced over the years alternating days of good and bad vision and that the day to day visual fluctuations had been modified by physical stress or alcohol drinking. Perimetric evaluations of this case demonstrated daily variability in visual sensitivities which corresponded to the subjective complaints. This form of visual fluctuation appears unusual, and it is remarkable that this patient had such a homoyzgous mutation in the arrestin gene that gives rise to a premature truncation of translation. The human arrestin, also known as S-antigen, has an inhibitory role in the activated phototransduction cascade. It should be mentioned that the arrestin gene is one of those expressed in the pineal gland that is considered to play a major role in the circadian rhythm.¹ Thus, it is tempting to speculate that the mutation in the arrestin gene could have modified rhythmic activities to induce daily alternating visual fluctuations in the present retinitis pigmentosa patient. The full coding sequences of the rhodopsin gene were normal in the relevant patient (data not shown). However, we could not deny the association of different polymorphisms in related gene products, or different environmental influences on the daily alternating visual fluctuations.

Recent molecular assessments revealed families with diffuse retinitis pigmentosa with or without features of Oguchi’s disease harbouirng 1147 delA in the arrestin gene. Thus the patient described here provides additional evidence that the same arrestin gene mutation is causally related not only to Oguchi’s disease but also to autosomal recessive retinitis pigmentosa. It is emphasised that our patient had classic features of autosomal recessive retinitis pigmentosa with poor visual prognosis and shown unusual visual fluctuations. In view of a rare mutation of arrestin gene, the relevant retinitis pigmentosa might be extremely rare. In a large series of white patients with aetiology of undefined retinitis pigmentosa arrestin gene mutation was not detected.¹

YASUSHI ISASHIKI
Center for Chronic Viral Diseases, Kagoshima University Faculty of Medicine, Kagoshima-shi, Japan

NORIO OHBA
KATSUAKI KIMURA
SHIGZO SONODA
TOMOKO KAKIUCHI
Department of Ophthalmology, Kagoshima University Faculty of Medicine, Kagoshima-shi, Japan

TETSUMA OZAWA
Yokohama Teishin Hospital, Yokohama, Japan

Correspondence to: Norio Ohba, MD, Department of Ophthalmology, Kagoshima University Faculty of Medicine, Kagoshima-shi 890-8520, Japan.

Accepted for publication 18 May 1999

Orbital Kimura’s disease in a white child

EDITOR.—Kimura’s disease is an uncommon, chronic inflammatory disorder of unknown aetiology which occurs predominantly in orientals and presents with tumour-like swellings mainly in the head and neck region. The condition primarily involves the subcutaneous tissues, parotid glands, and/or lymph nodes. Orbital cases are infrequent and most reported cases are in adults, with only one case in a child of Afro-Caribbean descent.¹ Our case report presents orbital Kimura’s disease in an 8 year old white child.

CASE REPORT

An 8 year old white girl presented to us with a 4 week history of a painless swelling of her right upper lid. She initially had an upper respiratory tract infection lasting a month which was treated by her general practitioner with Augmentin. She subsequently developed what appeared to be a right sided pustis. Of note, she had a strong history of allergic eye disease with intermittent conjunctivitis, and also suffers from asthma for which she takes Pulmicort and Bricanyl inhaler regularly. Examination revealed a right partial ptosis, mild right proptosis, and a palpable, painless mass in the superior aspect of her right orbit. There were no pupillary abnormalities, her extraocular movements were full, and funduscopic examination was unremarkable. There was no associated regional lymphadenopathy and the rest of the ocular and general physical examination showed no other abnormalities.

Investigations done included a full blood count, which was within normal limits. WBC 8.57 × 10^3/l with a differential of neutrophils 62%, lymphocytes 26%, monocytes 6%, eosinophils 5%, and basophils 0%. A magnetic resonance imaging (MRI) scan with...
painful and fibrosis. Haematoxylin and eosin stain. Original magnification ×50.

At presentation, the patient had marked swelling and proptosis. Computed tomography of the orbits and enhanced magnetic resonance imaging were performed which revealed an extensive, enhancing, soft tissue mass arising from the eyelid (see Fig 1). There was no evidence of extension into the orbit or brain. Orbital biopsy was performed which revealed fibrosis with a marked vascular proliferation and a prominent perivascular inflammation consisting of eosinophils and lymphocytes (see Fig 2). There were also scattered lymphocytes and eosinophils throughout the fibrotic tissue with reactive lymphoid follicles. There was no evidence of necrosis, granulomas, or malignant cells. Immunohistochemical studies revealed no cells of myogenic origin present. Studies for desmin, myoglobin, and vimentin were negative. All of the cells considered lymphoid on the haematoxylin and eosin stain also stained for the leucocyte marker common leucocyte antigen (CLA). A diagnosis of Kimura’s disease (angiolympoid hyperplasia with eosinophilia) was made.

At biopsy, the tumour was debulked and the patient was thereafter placed on a short 6 week course of oral steroids. The ptosis resolved and at 1 year follow up, there has been no clinical evidence of recurrence.

COMMENT
Kimura’s disease (or angiolympoid hyperplasia with eosinophilia) is a chronic inflammatory disorder of unknown aetiology which presents with tumour-like swellings mainly in the head and neck region. Most cases have been described in China and Japan, with relatively fewer cases reported in non-orientals.2 The disease typically affects males in the 20–40 year age group, and presents as single or multiple smooth swellings in the subcutaneous tissues, major salivary gland and/or lymph nodes in the head and neck area.

Histopathologically, the features described in previous cases1 corresponded with ours; there are eosinophilic infiltrates (mainly in a perivascular pattern), vascular proliferation, fibrosis, and formation of lymphoid follicles. It is still uncertain as to whether it represents a benign lymphoid neoplastic process, a variant of haemangioma or an allergic reaction. There have been no reported cases of malignant change or fatalities. There is usually an insidious onset with a long benign course, but recurrences are common after surgical excision. Systemic associations include asthma and nephrotic syndrome.3 Regional lymphadenopathy ranges from 50–75% of cases and there is a high occurrence of peripheral blood eosinophilia.

Cases of Kimura’s disease have been reported in the orbits;4 however, there was only one previously reported orbital case in a child (of Afro-Caribbean descent).5 To our knowledge, this is the first orbital case reported in a white child. Non-orbital cases of Kimura’s disease have been treated with surgical excision, irradiation and steroid therapy. After active therapy, they tend to recur. Cases have been conservatively managed for up to 12 years successfully although the cosmetic deformity can be quite a handicap. Periorbital cases have so far been treated successfully with surgical excision and biopsy with debulking.6 Oral steroids or intrallesional injection with steroids have not been as successful. There was only one previously reported case of severe haemorrhage on surgical resection.1 Complete surgical excision appears to be the best treatment.

Amikacin retinal toxicity

EDITOR,—It is well established that early intravitreal antibiotics are the treatment of choice for bacterial endophthalmitis1 but the choice of antibiotics remains controversial.2 Most centres rely on combination therapy and aminoglycosides are often used to treat Gram negative organisms. Aminoglycosides have a narrow therapeutic index and therefore a relatively small increase in concentration can cause significant toxicity. Following concerns about the risks of retinal toxicity caused by intravitreal gentamicin many ophthalmologists now use 0.4 mg intravitreal amikacin even though its use has also been associated with retinal toxicity.3 We report a case of retinal toxicity in a patient treated with intravitreal amikacin for postoperative bacterial endophthalmitis. This case serves to remind ophthalmologists of the risk of amikacin therapy and presents an argument for an alternative strategy in the treatment of bacterial endophthalmitis.

CASE REPORT
A 60 year old highly myopic woman presented with an acute history of increasing ocular pain and redness 5 days after routine phacoemulsification cataract extraction and intraocular lens implant. She had no other ocular or systemic history.

Initial examination revealed a visual acuity of 6/18, significant anterior chamber inflammation with a fibrin deposit over the pupil. A diagnosis of bacterial endophthalmitis was made and the patient was commenced on intravenous ciprofloxacin and hourly gentamicin, vancomycin, and dexamethasone eye drops. A vitreous biopsy was performed and 2 mg of intravitreal vancomycin (0.1 ml of 20 mg/ml) and 0.4 mg amikacin (0.1 ml of 4 mg/ml) were administered. The intravitreal antibiotics were prepared by a senior pharmacist using a typewritten protocol. The following day vision had deteriorated to no perception of light and the patient had developed a
relative afferent pupillary defect. Microscopy and culture failed to demonstrate any causative organism. The patient was started on 40 mg oral prednisolone but failed to manifest any improvement in acuity. Subsequently she was transferred to a specialist vitreoretinal unit where a diagnosis of retinal amikacin toxicity was made. Fluorescein angiogram taken 16 days after initial presentation confirmed the presence of retinal haemorrhage and a discrete area of capillary non-perfusion with significant late fluorescein leakage (Figs 1 and 2).

COMMENT

Intravitreal aminoglycosides are still recommended for use in the treatment of bacterial endophthalmitis despite the risk of retinal toxicity from intravitreal gentamicin and reports that amikacin in doses of 0.2–0.4 mg can also cause toxicity. The low incidence of bacterial endophthalmitis means that physicians and pharmacists do not routinely prepare intravitreal antibiotics. Because of the low therapeutic index of aminoglycosides even relatively small dilution errors can cause retinal toxicity. For these reasons it is our practice to use a third generation cephalosporin (0.1 ml of ceftazidime 20 mg/ml) instead of an aminoglycoside. Primate studies have shown that intravitreal ceftazidime is non-toxic in doses of 2.25 mg and ceftazidime has also been found to be effective against the Gram negative organisms responsible for postoperative endophthalmitis. Over the past 18 months we have used intravitreal ceftazidime (2 mg) and vancomycin (2 mg) in 21 patients with good therapeutic effect and with no adverse reactions. Owing to the risk of severe retinal toxicity with intra- vitreal aminoglycosides we recommend the use of ceftazidime instead of amikacin or gentamicin.

TIMOTHY L JACKSON

TOM H WILLIAMSON

St Thomas’s Hospital, London

Correspondence to: Tim Jackson, Vitreoretinal Unit, Department of Ophthalmology, St Thomas’s Hospita, Lambeth Palace Road, London SE1 7EH.

Accepted for publication 18 May 1999


Simultaneous administration of hepatitis B and polio vaccines associated with bilateral optic neuritis

EDITOR,—Immunisation against hepatitis B is recommended when there is an increased risk of contracting the virus because of lifestyle, occupation, or factors such as close contact with a case. Immunisation against poliomyelitis is routine for infants in the UK with reinforcement during childhood and then again in the teenage years. For those individuals at continued risk of infection, further reinforcement may be required. Both are commonly used vaccines and serious adverse reactions are extremely rare. We describe a case of severe bilateral, progressive optic neuritis occurring 1 week after vaccination against hepatitis B and poliomyelitis.

CASE REPORT

A 44 year old female health worker presented with gradual reduction of vision in both eyes associated with retrobulbar discomfort exacerbated by ocular movement, 7 days after vaccination against hepatitis B and poliomyelitis. Ophthalmological findings revealed visual acuities of 6/18 right eye, 6/12 left eye associated with bilateral optic nerve swelling. Within the next 48 hours the visual acuities dropped to perception of light in both eyes with absent direct and indirect pupillary light responses. This was despite commencement of therapy with intravenous methylprednisolone. Systemic examination revealed no other abnormalities. All haematological and biochemical investigations were normal and no infective cause was isolated. Computed tomograph imaging was normal and there was no evidence of demyelination on magnetic resonance imaging. Cerebrospinal fluid (CSF) examination revealed no abnormality other than biochemical, after culture, or on electroencephalograms. In addition, CSF pressure was within normal limits. Visually evoked potentials revealed absent responses. Despite 5 days of intravenous methylprednisolone (1 g per day) followed by a slow tapering of oral prednisolone (1 mg/kg/day) her vision remained poor (counting fingers at 1 metre in both eyes) after 3 months.

COMMENT

The recombinant hepatitis B vaccine has been associated with a diverse range of isolated adverse reactions but ocular complications are exceedingly rare. Granell et al attributed four cases of central retinal vein occlusion in patients under 50 years of age to the vaccine, and associations with multiple evanescent white dot syndrome (MEWDS) and acute posterior multifocal placoid pigment epitheliopathy (APMPPE) have been described. Stimulating vaccine derived antigen and normal or altered host tissue proteins, immediate hypersensitivity reactions, and stimulation of a pathologic allergic response. Ocular complications following vaccination are uncommon but various hypotheses exist including immune complex mediated demyelination or neurotoxicity, antigen mimicry between the stimulating vaccine derived antigen and neuronal or altered host tissue proteins, immediate hypersensitivity reactions, and stimulation of a pathologic allergic response.

Others such as the trivalent measles, mumps, and rubella (MRR) vaccine2 and the monovalent rubella vaccine,3 have also been associated with bilateral optic neuritis. The exact mechanism behind neurological complications following vaccination are unknown but various hypotheses exist including immune complex mediated demyelination, immune complex mediated demyelination or neurotoxicity, antigen mimicry between the stimulating vaccine derived antigen and neuronal or altered host tissue proteins, immediate hypersensitivity reactions, and stimulation of a pathologic allergic response. Ocular complications generally occur 1–3 weeks following vaccination which supports the claim that vaccination was the aetiological factor in this case. To the best of our knowledge, this is the first report of this complication with either of these vaccines. The fact that there are many case reports describing a diverse variety of systemic reactions of an autoimmune nature associated with both vaccines also adds to the claim. Those cases of optic neuritis attributed to other vaccines and to post infectious optic neuritis generally did better with corticosteroid therapy than in our case but high dose corticosteroids are generally considered to be the treatment of choice in these rare but potentially devastating complications.

OWN STEWART

BERNARD CHANG

JOHN BRADBURY

Department of Ophthalmology, Bradford Royal Infirmary, Bradford, West Yorkshire

Correspondence to: Mr Owen Stewart, Department of Ophthalmology, St James’s University Hospital, Leeds, West Yorkshire, LS9 7TF.

Accepted for publication 18 May 1999

Letters

Retinopathy after long term, standard doses of hydroxychloroquine

EDITOR,—While the antimalarial drug chloroquine has frequently been reported to cause retinopathy, there have been very few documented cases occurring with hydroxychloroquine (Plaquenil, Winthrop Pharmaceuticals, New York, USA). Patients may tolerate large cumulative doses (up to 3923 g) of hydroxychloroquine without developing retinopathy. Doses of ≤400 mg/day and ≤6.5 mg/kg of body weight/day of hydroxychloroquine have been used safely and some authors have suggested that ophthalmic screening is not necessary for patients on these doses. We present a case of hydroxychloroquine toxicity that developed in a patient after long term use of 400 mg (6.3 mg/kg) of hydroxychloroquine daily.

CASE REPORT
A 61 year old white woman presented with a 1 year history of increased glare in both eyes without change in visual acuity. She had a history of rheumatoid arthritis for which she took chloroquine and had no history of hepatitis B. Her total hydroxychloroquine dose was 1460 g. The patient had never taken chloroquine and had no history of macular disease. Her weight had been stable over the 10 year period. Family history was negative for macular dystrophy or retinal degeneration. Visual acuity at presentation was 20/20 in both eyes. Colour vision testing with pseudoisochromatic plates was normal in both eyes. Fundus examination showed subtle retinal pigment epithelium (RPE) pigmentary changes in a crescent pattern around the inferior fovea of both eyes (Fig 1A). Fluorescein angiogram confirmed the fundus findings (Fig 1B) which were felt to be early bull’s eye maculopathy. No drusen or signs of macular degeneration were appreciated in either eye. A central visual field performed with Humphrey automated static threshold perimeter using a white test object revealed bilateral paracentral ring scotoma corresponding to the macular pigmentary changes (Fig 2, upper). A diagnosis of hydroxychloroquine retinopathy was established and the drug was discontinued.

Ten months later the patient’s vision remained 20/20 with normal colour vision in both eyes. Her symptom of glare in both eyes had resolved. The RPE changes in the retina were unchanged and visual field testing showed some improvement of the paracentral scotoma (Fig 2, lower).

COMMENT
Hydroxychloroquine retinopathy is a rare condition characterised by bull’s eye maculopathy or pigmentary changes in the macula, corresponding scotomas on visual field testing, and irreversible signs and symptoms once hydroxychloroquine is stopped. Cases of preretinopathy have been described in which visual field defects were elicited using a red test object but not a white test object on Humphrey perimetry. All cases of preretinopathy completely resolved after the discontinuation of hydroxychloroquine. Five cases of true hydroxychloroquine retinopathy reported in the literature demonstrated that discontinuation of hydroxychloroquine resulted in stabilisation but not resolution of the retinopathy. Unlike the other reported cases of true hydroxychloroquine retinopathy, our patient had some resolution of visual field loss.

Much controversy exists as to whether the daily or cumulative dose of hydroxychloroquine contributes the greater risk for retinopathy. While there have been reported cases of hydroxychloroquine toxicity at cumulative doses similar to our patient’s (1460 g over a 10 year period), these patients had (1) received chloroquine therapy, (2) were treated with daily doses similar to our patient’s (1460 g over a 10 year period), (3) had renal impairment, (4) were treated with lower cumulative doses of hydroxychloroquine, (5) had no history of macular disease, and (6) had a lower cumulative dose of hydroxychloroquine.

REFERENCES

JENNIFER E THORNE
Department of Ophthalmology, Scheie Eye Institute, The Hospital of the University of Pennsylvania, University of Pennsylvania School of Medicine, Philadelphia, USA
The first report of post filtration hypotony and scleral rigidity. To our knowledge this is associated with disc swelling. We feel it may be more common in myopes as they have gested that hypotony related complications surgery, reversible and profound vision loss in maculopathy are known complications of such surgery. Although ocular hypotony and bleb revision with scleral patch grafts, with drainage into "large eyes" as a young child, as shown in this photograph taken at age 2 years (top left). Reversal of transient visual obscurations which suggested critically impaired perfusion of the disc. To prevent even further vision loss, a lumbar puncture to reduce the CSF pressure was suggested, to provide short term improvement in disc perfusion, which it was hoped in the longer term would be helped by raising her intraocular pressure. She underwent further surgery to both her eyes for hypotony, later that evening, which comprised revision of both blebs with scleral patch grafts. This was preceded by a lumbar puncture under general anesthesia at which there was an opening pressure of 22.5 cm H2O. A volume of 30 ml of CSF was removed and the following day was recorded as 5.5 cm H2O with microbiological and biochemical analysis of CSF revealing no abnormality. The day after surgery, the fourth day after initial presentation, examination revealed improved visual acuities of HM in both eyes with IOPs of 19 and 23 mm Hg in the right and left eye respectively. The disc swelling was felt to be reduced, and over the next few days improved gradually. Seven days after presenta- tion, her IOPs had fallen to 0 and 8 mm Hg, right and left, respectively, and she was taken back to theatre for anterior chamber refilling with Healon-GV. Her IOPs remained at around 20 mm Hg again 10 days later, with decreasing disc swelling, and a repeat ultrasound showed reduced choroidal detachments and axial lengths of 24.8 mm in the right and 24.6 mm in the left.

She underwent one other further surgical procedure, 7 weeks after presentation, when her IOPs became elevated again at 35 and 28 mm Hg right and left respectively, to a degree where it was felt her optic discs would be compromised. Treatment consisted of needling procedures to both blebs with subconjunctival injections of 5-fluorouracil to reduce scar formation. Her IOPs responded very quickly to this final procedure and she remains on no antiglaucoma medication with normal intraocular pressures (less than 20 mm Hg) now 7 months after presentation. Both drainage blebs are Seidel negative with good evidence of aqueous drainage into the filtration blebs (Fig 1, top right). Her visual acuities and visual function have now recovered quite substantially. However she does has a relative afferent pupillary defect in the left eye, and her current refraction and visual acuities are right eye +1.0/−4.0 × 05° (RVA 20/30) and left eye −1.75/−6.5 × 180° (LVA 20/120). Her optic discs show no evidence of swelling, and she has neither choroidal detachments nor macular folds (Fig 1, bottom left and right) as confirmed on ultrasound.

COMMENT

This young girl presented to us with severe ocular hypotony following bilateral primary glaucoma filtration surgery with mitomycin C. She developed vision loss, with episodes of transient obscurations attributable to hypotony and disc swelling. Her response to the initial surgery and subsequently to the corrective Visual loss after primary trabeculectomy with mitomycin C in Ehlers-Danlos syndrome

CASE REPORT

A 12 year old girl was referred to Moorfields Eye Hospital for a second opinion regarding her deteriorating vision following bilateral trabeculectomies with adjuvant mitomycin C for raised intraocular pressure.

At the age of 3 years, she was diagnosed as having megalocornea with ocular hypertension. Two years later, because of the persistent degree of IOP elevation (between 30–40 mm Hg), she was commenced on bilateral medical treatment. However, at the age of 11, she developed such severe problems with her topical treatment that surgery was considered necessary. In May 1996 right and subsequently left trabeculectomies were performed and during the procedures subconjunctival mitomycin C (0.3 mg/ml) was applied for 5 minutes. Her postoperative recovery was complicated by the development of ocular hypotony.

She was seen at Moorfields for the first time with progressive bilateral vision loss and recent onset of transient obscurations, 5 months after her filtration surgery and the onset of hypotony. Her visual acuities had fallen from right eye 20/20 (−0.5/−0.25 × 180°) to hand movements 5 months after presentation. Counting fingers (CF). Her visual fields on confrontation with a red target showed an enlarged blind spot. Both eyes had large, draining, diffuse and avascular filtration blebs with signs of hypotony as evidenced by bilateral IOPs of 0 mm Hg, superficial corneal epithelial staining, striae, and macular and choroidal folds. Both optic discs were swollen although no haemorrhages or cotton wool spots were observed. The patient was sensitive to light and good quality photographs could not be obtained. An ultrasound showed bilateral anterotemporal choroidal detachments involving the ciliary body (right larger than left) and axial lengths recorded as 23.0 mm in both eyes.

She underwent several surgical procedures. The day after she first presented, she underwent a right subconjunctival autologous blood injection to the right bleb under general anaesthesia. The following day, however, her vision was worse with visual acuities being recorded as patchy areas of perception of light on the right and hand movements temporally on the left. Her IOPs were 4 and 0 mm Hg in the right and left respectively with the former degree of disc swelling (as judged by further elevation of the disc) was noted bilaterally. It was considered by a neurologist (GTP) that the disc swelling may itself be contributing to the visual loss, particularly as the patient had transient visual obscurations which suggested critically impaired perfusion of the disc. To prevent even further vision loss, a lumbar puncture to reduce the CSF pressure was suggested, to provide short term improvement in disc perfusion, which it was hoped in the longer term would be helped by raising her intraocular pressure.

She underwent further surgery to both her eyes for hypotony, later that evening, which comprised revision of both blebs with scleral patch grafts. This was preceded by a lumbar puncture under general anaesthesia at which there was an opening pressure of 22.5 cm H2O. A volume of 30 ml of CSF was removed and the following day was recorded as 5.5 cm H2O with microbiological and biochemical analysis of CSF revealing no abnormality.

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The development of severe vision loss in our patient was associated with the occurrence of transient obstructions and hypotonous disc swelling. Resolution of the disc swelling was 4–6 weeks after correction of the hypotony, and improvement in visual acuity was noted thereafter. Its natural history—that is, the pattern of resolution and visual recovery, is compatible with disc swelling. Visual loss associated with hypotonous maculopathy, however, may never recover despite reversal of hypotony.10 Prolonged disc swelling in ocular hypotony is a recognised complication that is fortunately very rare. Experimental models have suggested that axoplasmic transport is delayed in cases of hypotony induced optic disc swelling giving rise to the same features histologically as seen with raised intracranial pressure papilloedema—for example, axonal swelling, accumulation of mitochondria, and cytoid body (cotton wool spots) formation.11 The pathogenesis of disc swelling in ocular hypotony may be related to that occurring in raised intracranial pressure if it is postulated that in both situations the CSF pressure in the subarachnoid space around the optic nerve exceeds the perfusion pressure of the optic disc. Therefore in our patient, it was considered that lowering the CSF pressure to well below normal levels (<20 mm H₂O) might provide temporary improvement in disc perfusion—hence our target for a very low closing pressure of 5.5 mm H₂O in the lumbar puncture. Two previous reports of unilateral disc swelling following trabeculectomy also suggest that optic disc oedema could result from a disturbed equilibrium between ICP and IOP—though none of the patients described in these papers had hypotony or profound vision loss.12,13

The development of profound vision loss in association with hypotony induced optic disc swelling in our patient, was as a complication of mitomycin C assisted glaucoma filtration surgery. However, the degree of hypotony was most probably related to reduced scleral rigidity. We believe that her abnormal wound healing response and defective connective tissue contributed to the severity of the hypotony that followed initial surgery with mitomycin C. We suggest that antiproliferatives be used with extreme caution in patients who you suspect may have a connective tissue disorder. A quick screening procedure might include a joint hypermobility score as shown in Table 1. Patients with Ehlers-Danlos syndrome, such as our patient, may be at high risk of post filtration surgery.

Table 1 Screen for joint hypermobility

<table>
<thead>
<tr>
<th>Action</th>
<th>Score (maximum = 9)</th>
</tr>
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<tbody>
<tr>
<td>Passive hyperflexion of 5th metacarpophalangeal joint beyond 90°</td>
<td>1 point each hand (max = 2)</td>
</tr>
<tr>
<td>Passive hyperextension of thumb to flexor aspect of wrist</td>
<td>1 point each hand (max = 2)</td>
</tr>
<tr>
<td>Hyperextension of elbow beyond 10°</td>
<td>1 point each arm (max = 2)</td>
</tr>
<tr>
<td>Hyperextension of knee beyond 10°</td>
<td>1 point each leg (max = 2)</td>
</tr>
<tr>
<td>Forward flexion of trunk, palms flat on floor in front, knees in extension</td>
<td>1 point</td>
</tr>
</tbody>
</table>

Letters Addressed to the Editor

Figure 2 Our patient demonstrates joint hypermobility, a feature of Ehlers-Danlos syndrome. She can oppose both thumbs to the flexor aspect of her wrist (top left), fold her fingers one over the other (top right), bend over to place her palms flat on the floor keeping her knees extended (bottom left), and put her ankles around her head (bottom right).

surgical procedures made us suspect that she had an inherent problem in wound healing and scleral rigidity, especially as she displayed such dramatic changes in axial lengths (1.6–1.8 mm difference in pre- and postoperative lengths). We therefore wondered if she had an underlying connective tissue disorder.

She was found to have Ehlers-Danlos syndrome type II (EDS II). Examination revealed her to have joint hypermobility (Fig 2), with a high joint hypermobility score (Table 1) of 7/9. In addition, she showed evidence of skin striae in the lumbar region, thighs and breasts with a pappery scar on her right knee, and gave a history of bruising easily with prolonged bleeding when cut. Echocardiography however was normal, with a normal palate, no spinal defects, no arachnodactyly, and a normal arm span to height ratio. However, there was a strong family history of joint hypermobility.

EDS is a heterogeneous group of genetically determined disorders of connective tissue affecting skin, ligaments, joints, blood vessels, and internal organs, of which at least nine subtypes exist, with the following diagnostic triad: extensible skin (extraordinary being described in fairground artistes, such as the “elastic skin man” of the 19th century), connective tissue fragility, and hypermobile joints.2 Although ocular complications in EDS are mostly associated with EDS type VI, a few ocular problems in EDS II have been documented such as prominent epicanthal folds, redundant skin on the upper eyelid, blue sclera, and the absence of peribulbar fat.

The wound healing process in EDS II is known to be abnormal in the skin because of the anomalous development of scar formation due to aberrant collagen fibrils. An abnormal healing response to the initial filtration surgery in this young girl might have been exaggerate with the use of mitomycin C. Cutaneous wounds in EDS II initially appear with gaping edges—so called “fish mouths” and heal slowly, and when scar formation eventually occurs, are characteristically shiny and stretched—papperyaceous. Our findings on initial exploration of the surgical site in the eye, in this case, revealed a friable, disintegrating scleral flap which would have permitted unguarded passage of aqueous into the subconjunctival space—hence the occurrence of hypotony when combined with a non-healing conjunctiva.

Ocular hypotony is associated with various complications3 including hypotonous maculopathy, resulting in a permanent reduction in vision and acuity. This occurrence has previously been attributed to reduced scleral rigidity, as seen in young and myopic eyes where a reduction in axial length has been documented following filtration surgery.4 Another mechanism put forward for the vision loss in these eyes, has been compression of the submacular sclera due to the resting tone of the inferior oblique muscle and decreased scleral rigidity.5

The incidence of post filtration ocular hypotony following full thickness procedures is 17–41%.6 This incidence was reduced significantly with the advent of guarded, partial thickness techniques, but the recent introduction of antiscarring agents such as mitomycin C and 5-fluorouracil7 have led to its increase once again. We believe that our patient had a complex and extreme response to filtration surgery with adjuvant mitomycin C complicated by her reduced scleral rigidity secondary to EDS II, as evidenced by her large eyes and marked changes in axial length in association with reduction in intraocular pressure.

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Supported in part by the Wellcome Trust (grant no. 048474) (to MFC), Guide Dogs for the Blind (MFC), and the Medical Research Council (to PTK).

M FRANCESCA CORDEIRO Wound Healing Research and Glaucoma Units, Institute of Ophthalmology and Moorfields Eye Hospital, London

GORDON T PLANT Moorfields Eye Hospital and National Hospital for Neurology and Neurosurgery, London

ANNE CHILD St George’s Hospital Medical School, London

BARRY JONES Moorfields Eye Hospital, London

PENG T KHAW Wound Healing Research and Glaucoma Units, Institute of Ophthalmology and Moorfields Eye Hospital, London

Figure 2 Our patient demonstrates joint hypermobility, a feature of Ehlers-Danlos syndrome. She can oppose both thumbs to the flexor aspect of her wrist (top left), fold her fingers one over the other (top right), bend over to place her palms flat on the floor keeping her knees extended (bottom left), and put her ankles around her head (bottom right).
COMMENT

The early recognition of oculomotor nerve compression by a posterior communicating artery aneurysm is essential given that the mean interval between the onset of diplopia and bleeding is 4 weeks in patients who develop a subarachnoid haemorrhage. Although the retro-orbital pain was typical, this case is unusual for two reasons. Firstly the ptosis and ophthalmoplegia were notably greater degree of compression of the third nerve in this case. Furthermore, in this case upper lid retraction and, on eye movement testing, esotropia and hypertropia were observed (Fig 1). Although modest variability of symptoms (for example, intermittent diplopia) has previously been described, we can find no reports in the literature of such marked fluctuations in ptosis due to third nerve compression. Additionally, in this case upper lid retraction and, on eye movement testing, esotropia and hypertropia—that is, excessive activity in third nerve innervated muscles, was observed.

A similar phenomenon of co-existing inappropriate neural discharge and block is seen in hemifacial spasm where compression of the seventh nerve by the vascular loop produces hemifacial spasm and mild facial weakness. Although there is some controversy as to whether it is the mechanical irritation of the facial nerve or secondary hyperexcitability of the facial nerve nucleus that is responsible for hemifacial spasm, it is now accepted by most authors that compression of the facial nerve is the causative factor. A precise intra-operative correlation between decompression of the facial nerve and electromyographic disappearance of normal facial muscle activity has been demonstrated although the facial weakness has been reported to take longer to resolve. An analogous situation of aneurysmal compression causing alternating ophthalmoplegia and third nerve dysfunction in this case reported here is suggested by the similar course of resolution to that seen following decompression of the facial nerve for hemifacial spasm—the third nerve hyperactivity resolved immediately following clipping of the aneurysm while the ophthalmoplegia improved rapidly but took several weeks to resolve fully. The degree of third nerve hyperactivity, however, much greater than the mild facial weakness seen in hemifacial spasm and may have been due to the greater degree of compression of the third nerve in this case.

This case is important for two reasons. Firstly, this is the first report of compression of the third cranial nerve producing alternating oculomotor hypo- and hyperfunction as a result of a compressive lesion. Secondly, it emphasizes the importance of excluding compression by aneurysmal aneurysm in a patient with variable signs which could be explained by a partial third nerve palsy.

MARGARET R DAYAN
Oxford Eye Hospital, Radcliffe Infirmary, Woodstock Road, Oxford OX2 6HE

Correspondence to: Miss Dayan
Accepted for publication 31 May 1999

Fluctuating oculomotor hyperfunction and hypofunction caused by aneurysmal compression of the third cranial nerve

EDITOR,—Aneurysms of the posterior communicating artery classically present with a painful progressive palsy of the third cranial nerve producing ptosis, ophthalmoplegia, and mydriasis. We present a case in which aneurysmal compression of the third nerve produced a variable ptosis and ophthalmoplegia without pupillary involvement in association with intermittent hyperfunction of the ipsilateral third nerve innervated muscles.

CASE REPORT

A 73 year old woman with a history of hypertension and ischaemic heart disease was referred to the eye department with a 3 week history of intermittent diplopia which she described as inconsistently horizontal, vertical, or oblique, and a 2 week history of right retro-orbital pain and difficulty in opening her right eye. The diplopia and ptosis were often worse in the mornings.

On examination the corrected visual acuities were right 6/6 and left 6/9. There was a very variable and apparently fatiguable right ptosis which was occasionally replaced by right upper lid retraction (Fig 1). The results of ocular motility testing were inconsistent, ranging from 4 prism dioptries of esotropia to 23 prism dioptries of exotropia in the primary position with between 3 and 7 prism dioptries of left hypertropia. There was variable limitation of elevation and depression of the right eye and occasional right upper lid retraction on downward gaze. On upgaze, the deviation reversed to become a right hypertropia due to right superior rectus overaction (Fig 1).

Magnetic resonance imaging and angiography were performed showing a 6 mm diameter saccular aneurysm arising from the intracranial internal carotid artery at the level of the posterior communicating artery. Attempted embolisation of the aneurysm failed owing to a focal stenosis at the origin of the right internal carotid artery and the patient underwent uneventful clipping of the aneurysm instead. There was a rapid improvement in the ophthalmoplegias following clipping and 6 weeks postoperatively all the neurological signs had resolved.

Figure 1 (Top) Right ptosis with frontal overaction and right hypertropia. (Middle) Right ptosis has resolved. (Bottom) A few minutes later there is right upper lid retraction and right hypertropia indicating right third nerve hyperfunction.


Late onset of Leber’s hereditary optic neuropathy in HIV infection

EDITOR,—We report a case of late onset of Leber’s hereditary optic neuropathy (LHON) in a 59 year old patient with HIV infection being treated long term since 1991 with zidovudine. The onset and course of the patient’s eye disease as well as the diagnostic process were protected. Molecular genetic testing revealed the mitochondrial DNA (mtDNA) mutation in nucleotide position 11778 confirming the diagnosis of LHON. In our patient late disease expression may be interpreted as a result of the rare combination of the 11778 mutation with HIV infection and long term zidovudine treatment. This is the first report on a patient with LHON suffering from additional HIV infection.
A 59 year old white man was admitted to our outpatient department in July 1997 with acute onset of visual loss in his left eye within the previous 2 weeks. The patient had been diagnosed with HIV infection in 1991. No opportunistic infections defining AIDS had been present so far. Antiretroviral therapy consisted of zidovudine since June 1991 as well as additional indinavir and nevirapine since January 1997. Best corrected visual acuity was right eye 20/20 and left eye 6/20. Further ophthalmic examination revealed a left afferent papillary defect and an absolute central scotoma in the left eye. Indocyanine green angiography and fluorescein angiography disclosed a slight oedema of the optic disc without leakage. Visual evoked potentials (VEP) of the left eye showed reduced central amplitudes and prolonged latencies. Ophthalmic examination of the right eye at the time of presentation was normal.

High dose steroid treatment on the basis of a presumptive diagnosis of optic neuritis could not prevent further progression of vision loss to hand movements in the patient’s left eye. In October 1997 the patient noticed similar symptoms in his right eye. Visual acuity was reduced to right eye 12/20.

On specific questioning the patient reported five relapses—one of his brothers and four cousins—who had lost vision in early adulthood. Molecular genetic testing of a blood sample to investigate the presence of mitochondrial DNA (mtDNA) mutations yielded a mutation at position 11778, homoplasmic to the level of detection, confirming the diagnosis of Leber’s hereditary optic neuropathy (LHON). Therefore the patient was specifically referred for neurological evaluation and lumbar puncture the next day. However, on presentation to the paediatric neurologist, the patient noted that his diplopia had gradually resolved. Lumbar puncture was not performed because neurological examination was entirely unremarkable. Follow up oculomotor examination revealed normal ocular alignment and motility.

Acute onset comitant esotropia is most commonly benign when occurring in infancy or early childhood. Examination to rule out a parietic deviation is essential. However, acute onset of comitant esotropia has also been reported in association with serious neurological disease. It has been reported in association with hydrocephalus.1 2 These patients generally have obvious neurological problems associated with hydrocephalus, and are not a diagnostic dilemma.3 They may have a meningo(myelo)cele or encephalocele, and an A-pattern esotropia.1 3 In cases of comitant esotropia associated with shunt failure, all the esotropias resolve when the shunts are revised. These patients do not have A-pattern esotropia as presenting sign of demyelinating disease.

CASE REPORT
An 11 year old boy was referred for evaluation of diplopia. The patient stated he woke up the previous day with double vision. He denied decreased visual acuity, photophobia, pain with eye movements, headache, or nausea and vomiting. One week before the onset of esotropia, he had an upper respiratory infection with a fever that resolved without antibiotics. His mother also noted that he had been very tired over the previous week with episodes of falling asleep during the day at home and school, and one episode of dizziness. He was an otherwise healthy child on no medications and with no allergies.

On examination, visual acuity without correction was 20/20 bilaterally. Pupils were equally reactive without an afferent pupillary defect. Extraocular motility was full, and saccades were grossly normal and symmetrically. Alternate cover testing revealed a comitant 16 prism dioptre esotropia for both near and distance fixation. He was able to fuse with the appropriate prism in place. Slit lamp examination was unremarkable. Dilated retinal examination revealed pink and sharp optic nerves with no papilloedema. Macula, vessels, and retinal periphery were all within normal limits.

The patient underwent magnetic resonance imaging (MRI) the next day (Fig 1). The study revealed a hyperintense signal on T1 weighted imaging in the area of the periaqueductal grey matter in the midbrain. The lesion was consistent with a demyelinating disease including acute disseminating encephalomyelitis or multiple sclerosis. The patient was referred for neurological evaluation and lumbar puncture the next day. However, on presentation to the paediatric neurologist, the patient noted that his diplopia had gradually resolved. Lumbar puncture was not performed because neurological examination was entirely unremarkable. Follow up oculomotor examination revealed normal ocular alignment and motility.

COMMENT
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esotropias. Patients may not necessarily present with papilloedema even when hydrocephalus is present. In the present case, no hydrocephalus was noted on neuroradiographic study.

Arnold-Chiari malformation has also been reported in association with acute comitant esotropia. These cases may present with an A-pattern, co-existing nystagmus and hydrocephalus. Arnold-Chiari malformations sometimes do not manifest until late childhood or adulthood, and can be mild. Neuroradiographic study did not reveal Arnold-Chiari malformation in this patient.

Central nervous system tumours have also been reported in association with acute onset comitant esotropias. Tumours in this group include cerebellar astrocytomas, and medulloblastomas, as well as pontine gliomas. An A-pattern esotropia is unusual in these patients. In a report by Williams and Hoyt, three of their six patients had some form of nystagmus associated with the comitant esotropia. They suggested neurological evaluation in any patient with both nystagmus and acute comitant esotropia. They further suggested evaluation for any patient with acute onset comitant esotropia if ocular motor fusion can not be established with prism or surgery.

Acute comitant esotropia has been associated rarely with a hypertensive haemorrhage in the thalamus. However, these patients also present with other neurological signs leading rapidly to the diagnosis. In addition, acute comitant esotropia has been reported rarely with myasthenia gravis and seizures.

This patient had a brainstem lesion that was consistent with an acute central nervous system demyelination. Its location in the midbrain may have interfered with neurons associated with horizontal gaze centres or vergence mechanisms. This is the first reported case of acute demyelinating disease presenting with acute comitant esotropia. Acute disseminated encephalomyelitis is generally post viral and is characterised by abrupt headache fever, drowsiness, and focal neurological dysfunction. Cerebrospinal fluid analysis may exhibit pleocytosis. It may resolve completely or may result in permanent impairment and seizures.

Whether this lesion represents a localised variant of acute disseminated encephalomyelitis or multiple sclerosis is not clear. However, cases of acute onset comitant esotropia with suspicious presentations warrant neuroimaging to rule out intracranial pathology including central nervous system demyelinating lesions.

BARRY N WASSERMAN
Chester County Eye Care Associates, 606 East Marshall Street, Suite 104, West Chester, PA 19380, USA

Accepted for publication 8 June 1999