Serpiginous choroidopathy presenting as choroidal neovascularisation

Serpiginous choroidopathy is an insidious, relentlessly progressive, idiopathic inflammatory disease affecting the retinal pigment epithelium and inner choroid. Choroidal neovascularisation (CNV) is a well recognised late complication of serpiginous choroidopathy in 10–25% of affected patients. In all previously reported cases CNV was recognised at the time of or after the diagnosis of serpiginous choroidopathy was established. We report a patient presenting with CNV who subsequently developed clinical findings characteristic of serpiginous choroidopathy.

Case report
A 31 year old man presented with decreased vision in his right eye in July 1997. Examination revealed acuities of 20/40 right eye and 20/20 left eye with normal anterior segments. The right fundus showed subretinal fluid and haemorrhage adjacent to the disc (Fig 1A). The left eye showed an irregularity superior to the optic disc (Fig 1B). Examination at the National Eye Institute in April 2000 revealed acuities of 20/40 right eye and 20/16 left eye with normal anterior segments. The vitreous contained trace cells without haze bilaterally. The right fundus showed a large peripapillary chorioretinal scar. The left fundus revealed a chorioretinal scar superior to the disc and two yellow, irregularly circumscribed, deep macular lesions (Fig 3A, B). The retinal vessels and discs were normal and no subretinal fluid, haemorrhage, or macular oedema was noted in either eye.

Fluorescein angiography revealed early hypofluorescence and late hyperfluorescence corresponding to the macular lesions in the left eye (Fig 3C, D) with no evidence of CNV in either eye. Laboratory studies were non-diagnostic. A diagnosis of serpiginous choroidopathy was made based on the clinical and fluorescein characteristics of the macular lesions in the left eye.

Comment
CNV in serpiginous choroidopathy is associated with a poor visual prognosis. In a small study CNV was reported to develop within 16 months of the serpiginous diagnosis. In a larger retrospective study of 53 serpiginous patients active CNV was found in three patients at the time of initial diagnosis and in three others within 2–17 months. Our patient differs from those previously reported in that he was diagnosed and treated for idiopathic CNV before the recognition of clinical findings.

References
1. In all previously reported cases CNV was recognised at the time of or after the diagnosis of serpiginous choroidopathy was established.
2. Laboratory studies were non-diagnostic.
3. A diagnosis of serpiginous choroidopathy was made based on the clinical and fluorescein characteristics of the macular lesions in the left eye.

Figure 1 July 1997. (A) The right eye shows subretinal fluid adjacent to the disc surrounded by subretinal haemorrhage extending to the fovea. (B) An undefined irregularity is noted superior to the optic disc of the left eye.

Figure 2 (A) Angiography of the right eye reveals a wedge shaped peripapillary CNV membrane. (B) A smaller CNV membrane is present angiographically superior to the left optic disc (late phase).

Figure 3 April 2000. (A) The right eye shows a large peripapillary chorioretinal scar resulting from previous laser photocoagulation of the initial CNV lesion. (B) A similar photocoagulation scar is present in the left eye which also shows two non-contiguous active serpiginous lesions. The fluorescein angiogram of the left eye shows early hypofluorescence (C, arrows) corresponding to the locations of the active serpiginous lesions that later become hyperfluorescent (D).
diagnostic of serpiginous choroidopathy. Other causes of posterior uveitis associated with CNV and choroidal lesions similar to those seen in our patient include acute posterior multifocal placoid papillopathy (APMPP), presumed ocular histoplasmosis (POHS), sarcoidosis, multifocal choroiditis, birdshot chorori- opathy, and toxoplasmosis. As with most cases of serpiginous choroidopathy, the CNV in these entities typically occurs late in the disease course.

The exact pathogenesis of idiopathic CNV is unknown. CNV in eyes with uveitis, however, is believed to develop in direct response to the intraretinal inflammation which may alter the balance between vascular growth factors, such as vascular endothelial growth factor (VEGF), and inhibitors. In the early stages of development active serpiginous lesions and CNV may appear as poorly defined subretinal lesions difficult to differentiate by ophthalmoscopy. Typically with fluorescein angiography classic CNV and serpiginous lesions are readily distinguished as the former shows early hyperfluorescence while the latter characteristically shows early blockage. Occult CNV, which may show subtle or less pronounced early hyperfluorescence with late leakage, however, may be more difficult to distinguish from an early serpiginous lesion.

This case illustrates that serpiginous choroidopathy may present with CNV. In contrast to idiopathic CNV, optimal treatment of CNV in patients with uveitis may require immunosuppressive treatment that addresses the underlying uveal inflammation with or without adjunctive laser therapy. Further investigation is needed to better define the role of emerging therapies for CNV such as photodynamic therapy which may offer promise for the treatment of CNV in uveitis patients.

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References

Optic neuritis in anti-GQ1b positive recurrent Miller Fisher syndrome

Only five cases of optic nerve involvement in Miller Fisher syndrome (MFS) have been documented in the literature. This report further confirms that optic neuritis may be seen in anti-GQ1b positive MFS.

Case report
This 23 year old woman presented with acute blurry vision, diplopia, and pain with eye movement. Her visual acuity was 20/20 right eye and 20/200 left eye with left relative afferent pupillary defect (RAPD). She had left sixth nerve palsies, nystagmus in all gazes, and left optic disc oedema. After 1 week her visual acuity improved to 20/20 in both eyes, but her left disc remained oedematous. She then developed right lower ataxia leading to such a degree that she was unable to walk. Dysmetria and dysdiadochokinesia were more marked in her left upper extremity. She had visual acuity and extremity weakness, absent lower extremity deep tendon reflexes, and bilateral Babinski’s. She also had tingling in her hands and feet and decreased lower extremity vibratory sensation. Her mental status was normal throughout her illness. She was not taking any drugs. A magnetic resonance image (MRI) of the brain and entire spine and MR venogram were all normal. Her cerebrospinal fluid (CSF) opening pressure was 190 mm of H2O. Her CSF protein was elevated at 70 mg/dl, but CSF glucose and cell count were normal. CSF VDRL, Gram stain, routine bacterial, viral, and fungal cultures were all negative, and the CSF oligoclonal bands were seen on CSF electrophoresis. Her visual evoked potential (VEP) revealed a delayed left P100 latency at 131 ms and her brainstem auditory evoked potential (BAEP) was normal. Electromyogram/nerve conduction study (EMG/NCV) study revealed mildly decreased visual acuity, pain with eye movement, dyschromatopsia, and optic disc oedema that resulted in good visual recovery are all indicative of the diagnosis of optic neuritis. Since high titer anti-GQ1b antibodies can cross the blood-brain barrier, the optic disc oedema in this patient could represent an early anti-GQ1b IgG deposition in the CNS. The absence of serological evidence linking this antibody to MFS, the ipsilateral delayed P100 latency is consistent with a pre-chiasmal demyelinating optic neuropathy.

Infection or her optic neuritis, this patient concomitantly demonstrated the classic features of MFS which are the acute onset of external ophthalmoplegia, ataxia of the cerebellar type, and the loss of corneal and optic nerves. MFS is considered a variant of Guillain–Barré syndrome (GBS) because some patients who present with MFS progress to GBS. High titres of anti-GQ1b IgG antibodies are present in 80% to 100% of patients with MFS. MFS may be immunologically differentiated from GBS by the presence of anti-GQ1b and anti-GM1 antibodies. Although both anti-GD1a IgG and anti-GM1 IgG are associated with GBS, anti-GM1 IgG is present in patients with typical MFS who have limb weakness, as in this patient. As further evidence linking this antibody to MFS, the decrease in anti-GQ1b antibody levels after plasmapheresis correlated with the clinical recovery in this patient. Therefore, the elevated titres of anti-GQ1b and anti-GM1 antibodies, along with the clinical trial of ophthalmoplegia, arreflexia, and ataxia in this patient all support the diagnosis of MFS, and not GBS. In rare cases, MFS has been known to recur. This patient presented with a relapse of similar clinical features 10 years from her initial episode. In the study done by Chida et al., patients with recurrent MFS appeared to have similar HLA typing characteristics as the non-recuring ones. Both types of HLA-DR2 and CW3 alleles, but the frequency of HLA-DR2 was slightly higher in the patients with recurrent MFS. Therefore, this patient’s HLA-DR2-positive status may have been a risk factor for her recurrence of MFS. This case report emphasizes that optic neuritis may be a central nervous system feature that should be recognised as part of the Miller Fisher syndrome. The presence of both anti-GQ1b IgG and anti-GM1 IgG in this patient provides immunological evidence supportive
of typical MFS. The delayed P100 latency in her VEP also provides electrophysiological evidence that the optic nerve is affected in anti-GQ1b antibody positive MFS. Furthermore, this is the first documented case known to the author of optic neuritis in the recurrent subtype of MFS which is associated with a higher frequency of the HLA-DR2 allele.

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References

Ocular myasthenia gravis and inflammatory bowel disease: a case report and literature review
Myasthenia gravis has been reported to be associated with both ulcerative colitis (UC) and Crohn’s disease (CD). The link between inflammatory bowel disease (IBD) and myasthenia gravis (MG) is thought to be related to the production of autoantibodies. Myasthenia gravis is also associated with other autoimmune diseases including alopecia, lichen planus, vitiligo, and systemic lupus erythematosus. Similarly, IBD frequently presents with other autoimmune disorders. One study demonstrated a 9.4% prevalence of autoimmune disorders in patients with UC including sclerosing cholangitis, thyroid disorders, vitiligo, insulin dependent diabetes mellitus, thyroid disease, pernicious anaemia, scoliosis, and spondyloarthropathic arthritis. Despite the association between MG and other autoimmune disorders, there are relatively few reports of ocular findings as the presenting sign of MG in patients with IBD.

Case report
A 21 year old African-American male, with a medical history of biopsy proved ulcerative colitis diagnosed in 1995, focal segmental glomerular sclerosis determined by renal biopsy in 1995, and primary sclerosing cholangitis determined by liver biopsy in 2000 presented to the neuro-ophthalmology service with complaints of binocular diplopia and ptosis of the left upper eyelid. Both the diplopia and the ptosis were better in the morning and worsened during the course of the day. His ulcerative colitis had been in remission for the past 3 years without medication.

Corrected visual acuity was 20/25 in each eye. The external examination revealed ptosis of the left upper eyelid that worsened in sustained upgaze. He had limited extraocular motility in all fields of gaze (Fig 1). The remainder of the neuro-ophthalmic examination was normal and he had no difficulty with speech or swallowing.

Laboratory evaluation revealed a positive acetylcholine receptor antibody and normal thyroid function studies. There was no evidence of a thymic mass on magnetic resonance imaging of the chest.

The patient returned to the emergency room 1 week later with difficulty swallowing and shortness of breath. He was hospitalised for plasma exchange and upon discharge treated with imuran, prednisone, and mexitone. One month later his ptosis resolved and his extraocular motility was normal.

Comment
Autoimmune disorders, including MG, occur more frequently in UC than in CD. It is not clear how many other cases of IBD manifested with ocular presentations as the initial finding of MG as in our case report. Our literature review revealed only one other purely ocular presentation of myasthenia associated with ulcerative colitis; however, details of the ocular examination were not included. Another report, of a 21 year old woman with a 3 year history of Crohn’s disease, documented diplopia and unilateral ptosis as the initial findings of MG. She was found to have acetylcholine receptor antibodies and her ocular findings improved with pyridostigmine.

Because of the relatively few reports of ocular myasthenia in patients with IBD we reviewed the English literature and found four additional reports of MG in patients with IBD. Based on these four reports and the three (including the present report) with ocular MG in patients with IBD (Table 1), the mean duration of IBD before the diagnosis of MG was 10 years.

Autoimmune dysregulation is the central defect in both MG and IBD. Both IBD and MG may be associated with an elevated carcinoembryonic antigen (CEA) and decreased peripheral lymphocyte counts that subsequently normalise following thymectomy. Some studies have shown abnormal thymic involution and the presence of an abnormal ratio of T suppressor to T helper cells in both MG and UC, while others have noted a decline in suppressor T cells and an increase in

Table 1 Previous reports of myasthenia gravis occurring in patients with inflammatory bowel disease

<table>
<thead>
<tr>
<th>Reference</th>
<th>Age (years)</th>
<th>Sex</th>
<th>IBD</th>
<th>Duration of IBD before diagnosis of MG (years)</th>
<th>AchR antibody reactivity</th>
</tr>
</thead>
<tbody>
<tr>
<td>Miller 1971</td>
<td>35</td>
<td>Male</td>
<td>UC</td>
<td>13</td>
<td>Unknown</td>
</tr>
<tr>
<td>Tan 1972</td>
<td>38</td>
<td>Male</td>
<td>UC</td>
<td>12</td>
<td>Unknown</td>
</tr>
<tr>
<td>Martin et al, 1991</td>
<td>63</td>
<td>Male</td>
<td>CD</td>
<td>15</td>
<td>Positive</td>
</tr>
<tr>
<td>Gower-Rousseau et al, 1993</td>
<td>27</td>
<td>Female</td>
<td>UC</td>
<td>10</td>
<td>Positive</td>
</tr>
<tr>
<td>Finnie et al, 1994</td>
<td>21</td>
<td>Female</td>
<td>CD</td>
<td>3</td>
<td>Positive</td>
</tr>
<tr>
<td>Lossos et al, 1995</td>
<td>11</td>
<td>Male</td>
<td>CD</td>
<td>9</td>
<td>Unknown</td>
</tr>
<tr>
<td>Present report</td>
<td>21</td>
<td>Male</td>
<td>UC</td>
<td>7</td>
<td>Positive</td>
</tr>
</tbody>
</table>

IBD = inflammatory bowel disease, MG = myasthenia gravis, AchR = acetylcholine receptor, UC = ulcerative colitis, CD = Crohn’s disease.
immature helper T cells suggesting migration without normal maturation.

The immunological link between MG and IBD is highlighted by two reports of patients undergoing surgical treatment. One report of a patient with both MG and CD documented improvement in perineal and perianal disease following thymectomy for severe uncontrolled MG. Another patient with both MG and UC demonstrated regression of the myasthenia following proctectomy.

Although the simultaneous occurrence of these two autoimmune disorders is uncommon, it is important to understand that ocular findings may be the initial manifestation of MG in patients with IBD.

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References


Magnetic resonance imaging findings in malignant melanoma of the lacrimal sac

A case of primary malignant melanoma of the lacrimal sac is presented. This is the first report of the preoperative magnetic resonance imaging (MRI) findings of malignant melanoma of the lacrimal sac.

Case report

A 54 year old Chinese woman was referred to an ophthalmologist complaining of a 6 month history of left sided bloody tears and epistaxis. She had a firm, non-tender left medial canthal swelling, and syringing revealed left nasolacrimal duct (NLD) obstruction. Ocular and periorbital examination was otherwise normal. A dacryocystogram (DCG) demonstrated a filling defect in the lacrimal sac with NLD obstruction.

An ENT opinion was sought, and nasal examination revealed left sided septal deviation, with no obvious cause for the epistaxis. Computed tomography (CT) of the head and orbits demonstrated a left lacrimal sac lesion extending into the NLD with proximal dilation of the duct and no apparent bone erosion (Fig 1A). MRI confirmed the presence of a lacrimal sac lesion with intermediate signal intensity on T1 and T2 weighted images (Fig 2A, B). The lesion enhanced with intravenous gadolinium.

An incisional biopsy of the lacrimal sac (Fig 1B) under frozen section control, and paraffin sections, confirmed malignant melanoma.

A full medical review, including MRI of the chest and abdomen, and liver function tests, excluded tumour elsewhere. However, abdominal MRI and ultrasound revealed a co-incident polycystic liver.

Three weeks after biopsy, a wide local excision including the medial upper and lower eyelids, dacryocystectomy and medial maxillectomy was performed. A tumour, confined to the sac, and invasion through the medial wall of the upper NLD, into the lateral wall of the nose, and apposing nasal septal mucosa, was seen peroperatively and confirmed histologically.

She underwent postoperative adjuvant radiotherapy (55 rads) and to date, 4 months later, remains well.

Comment

Malignant melanoma of the lacrimal sac is rare accounting for 3% of lacrimal sac tumours. It has an unfavourable prognosis compared with other causes of lacrimal sac tumour, and is considered more aggressive than cutaneous malignant melanoma. Response to treatment is generally poor, with up to 80% of cases recurring within 2 years.

Radiological features of lacrimal sac tumours include filling defects on DCG and mass lesions on CT. However, to the authors’ knowledge, this is the first report of the MRI findings of malignant melanoma of the lacrimal sac.

Owing to the paramagnetic properties of melanin, malignant melanoma appears hyperintense on T1 weighted imaging, and hypointense on T2 weighted imaging. A study of six mucosal melanomas of the head and neck found that on T1, five lesions were hyperintense and one was isointense. On T2, five were of mixed intensity and one was iso-intense. They concluded that hyperintensity on T1 of mucosal melanomas was characteristic but not universal.

The majority of malignant lacrimal sac tumours are epithelial in origin. Imaging features suggesting malignancy include invasion of bone, rapid growth, and irregular margins with skin fixation. On MRI, the majority of epithelial tumours have intermediate signal intensity on T1 and high T2 signal intensity. High tumour cellularity is associated with intermediate to low T2 signal intensity.

High signal intensity on T1 is not specific for malignant melanoma. Subacute haemorrhage caused by the presence of methaemoglobin is more likely and although melanoma may undergo intratumoral haemorrhage, other tumours with a tendency to bleed include small cell lung carcinoma, choriocarcinoma, and renal cell carcinoma metastases. Less likely causes include fat containing tumours (lipoma, dermoid, and teratoma).

Figure 1 (A) Coronal CT scan demonstrating a solid mass of the left lacrimal sac with proximal dilation of the nasolacrimal duct (arrow). (B) Incisional biopsy with lacrimal sac opened and melanoma visible.

Figure 2 (A) T1 weighted sagittal MRI demonstrating intermediate signal intensity mass lesion of the lacrimal sac and proximal nasolacrimal duct (arrow). (B) T2 weighted axial MRI demonstrating intermediate signal intensity mass lesion of the left lacrimal sac (arrow).
requiring MRI fat suppression methods, paramagnetic material (manganese, iron, and copper), and very high (non-paramagnetic) intratumoral protein concentration.

MRI has been reported as a useful investigative tool in the assessment of lacrimal disease owing to its ability to delineate soft tissues. Intravenous and intracanalicular gadolinium adds useful information on lesion enhancement and lacrimal apparatus structure and function. The predictive value of MRI for lacrimal sac melanoma, however, appears to be variable. Hyperintensity on T1 relies on the paramagnetic properties of melanin, the presence of which is variable in amelanotic melanoma. This is supported by our case, where only moderate T1 hyperintensity with contrast enhancement was demonstrated.

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References

Photodynamic therapy for recurrent myopic choroidal neovascularisation after limited macular translocation surgery

Limited macular translocation (LMT) is one of the treatment options for subfoveal choroidal neovascularisation (CNV) resulting from pathological myopia. The fundamental surgical principle involves the transposition of the foveal neurosensory retina to a new site with more healthy underlying retinal pigment epithelium. Direct laser photocoagulation is usually employed as an adjunct measure in eradicating the original CNV after the surgery. It has been observed that geometrically sizeable translocation is a prerequisite for a long term surgical success. The degree of translocation is, however, not often predictable and any ineffective displacement may render the subsequent laser photocoagulation extremely difficult or even impossible to perform. As a result, the recurrent or persistent CNV intruding the newly relocated fovea may jeopardise the final visual outcomes. Photodynamic therapy (PDT) may be considered a viable adjunct treatment option in such circumstance.

Case report
A 41 year old woman with pathological myopia of −11.0 dioptres in both eyes presented with a subfoveal CNV and subretinal haemorrhage in her right eye in July 2000. The best corrected visual acuity (BCVA) was 20/200 in the right eye and 20/30 in her left eye. LMT with superotemporal 6 mm scleral iridectomy was performed in July 2000. The operation was uneventful and an inferior displacement of the fovea by 600 µm was achieved. The CNV, however, was still located in the vicinity of the juxtafoveal area and therefore laser photocoagulation, bearing the potential risk of late creeping scar, was not suggested. At the 4 months postoperative visit, her left BCVA was 20/200 and the original CNV became more fibrotic with minimal leakage upon fluorescein angiogram. Nevertheless, she came back at 5 months with a return of metamorphopsia and a drop in her right vision from 20/200 to 10/200. Dilated fundus examination showed a tiny patch of submacular haemorrhage in direct continuity with the old fibrotic scar (Fig 1A). Fluorescein angiogram of the early phase demonstrated a fresh recurrent CNV budding out from the original fibrotic CNV and extending to the centre of the foveal avascular zone (Fig 1B). Moderate fluorescein leakage could be seen in the late phase (Fig 1C). Treatment comprising revision macular translocation surgery, submacular surgery, photodynamic therapy, and observation had been thoroughly discussed with the patient. In view of minimal invasiveness and comparatively better preservation of surrounding neurosensory retinal tissue, PDT was adopted in treating the CNV recurrence. PDT with verteporfin infusion and laser delivery was performed in accordance with the standard protocol. After the treatment, the blood clot in the fovea was gradually resorbed and the vision improved to 20/200 at 3 months of follow up. Complete regression of the recurrent CNV at the fovea without angiographic leakage was documented over the follow up angiogram at 3 months and subsequently (Fig 1D). The vision remained stable at 20/200 in the latest visit at 24 months after the PDT.

Comment
It has been shown that significant visual improvement may be achieved by LMT for the treatment of subfoveal CNV associated with age related macular degeneration (AMD) or pathological myopia. However, the surgical techniques are demanding and the potential complications are not unusual. One of the late postoperative visually important complications is recurrence of the CNV and this is partially caused by an ineffective translocation of the fovea or a large lesion size of CNV. The incidence of persistent or recurrent CNV after limited LMT has been reported to be 40% and 35% respectively in age related macular translocation and being 21% and 14% respectively in pathological myopia. Not many treatment options are available once the fovea is involved. Viable surgical options including repeated LMT, full 360 degree retinotomy MT, or submacular surgery may be considered but the surgical risk may be inadvertently higher in the redetachment of the neurosensory retina. PDT induces a selective thrombosis of the abnormal CNV and has been proved to be an effective treatment in preventing a significant loss of vision in patients with CNV secondary to AMD or pathological myopia.
Its clinical indications and applications are expanding. Its minimal invasiveness and clinical efficacy make it a safer and visually desirable supplementary treatment in recurrent CNV after LMT. In our patient, the complete closure of CNV was achieved with concomitant vision improvement after a single session of PDT without evidence of recurrence at 24 months.

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References

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References

Acquired Glanzmann’s thrombasthenia causing prolonged bleeding following phacoemulsification

Phacoemulsification under topical anaesthesia using clear corneal incision is not a challenging procedure for the haemostatic system. In patients with known bleeding diathesis, this may be the procedure of choice to remove cataract. We report a patient who bled continuously for 36 hours following phacoemulsification under topical anaesthesia through a clear corneal incision. This was managed by using a topical haemostatic agent that has not been used in ophthalmic surgery before. Extensive haematological evaluation revealed the underlying cause to be an acquired form of Glanzmann’s thrombasthenia, a very rare condition.1

Case report
A 79 year old woman underwent left phacoemulsification with intraocular lens implantation through a clear corneal temporal incision. The procedure was uneventful but she was seen to bleed from the operated eye in the recovery room. The eye was patched but the bleeding continued soaking the pads. When re-examined 2 hours later, as there was continuous bleeding, the eye was patched with gentle pressure. Examination the next day showed that the bleeding was persistent. Pressure bandage was reapplied. Examination in the operating theatre confirmed the conjunctival origin of the bleeding from the site where the left handed surgeon held the conjunctiva during surgery. Cauterization and an attempt to suture the conjunctiva were unsuccessful. It was decided that the safest option was to use a small piece of oxidised regenerated cellulose (Surgicel, Ethicon) on the bleeding site and patch the eye.

The piece of Surgicel with clotted blood that was lying loose on the conjunctiva was removed at review 24 hours later. The conjunctival site had stopped bleeding with evidence of altered inflammation though a clear corneal temporal incision. The piece of Surgicel had been applied (Fig 1A). At her last review 8 weeks later, she was found to have a corrected visual acuity of 6/18 due to pre-existing macular changes secondary to retinal detachment that was reattached in 1976. The conjunctiva had healed well (Fig 1B). The patient had previously undergone an uneventful phacoemulsification and intraocular lens implantation in her right eye under sub-Tenon’s anaesthesia.

The patient’s recent medical history was significant for recurrent admissions elsewhere for investigation of severe anaemia following gastrointestinal bleeding. Platelet count and clotting screen had been normal. Angiodysplasia of stomach and duodenum were treated with laser and angiodysplasia of colon was treated by hemicolecotomy. Three episodes of epistaxis and an episode of vaginal bleeding were managed conservatively. She had received 60 units of blood transfusion over a period of 1 year. Interestingly, she had appendicectomy and multiple dental extractions elsewhere many years previously without any significant bleeding. She has not been on any antiplatelet agents or anticoagulants. There was no family history of bleeding disorders.

A defect in the platelet function was suspected, as her coagulation screening including the platelet count was normal. Platelet aggregation tests showed no aggregation against any agonists other than ristocetin, which is dependent on platelet glycoprotein Ib. The platelet showed normal surface expression of glycoprotein antigens Ib/IIa and Ib. The patient’s serum showed presence of inhibitory antibody against glycoprotein Ib/IIa. This led to a diagnosis of acquired Glanzmann’s syndrome, an extremely rare condition of autoimmune thrombasthenia. No underlying malignant, autoimmune, or lymphoproliferative disorder had been identified as a cause for this patient’s acquired Glanzmann’s thrombasthenia.

Comment
The patient described had uncontrollable bleeding for 36 hours following a procedure, which is generally considered safe in the absence of a bleeding disorder. She developed bleeding from the conjunctival site where the surgeon grasped the conjunctiva during certain stages of the procedure. One would usually not expect any significant bleeding from this site; however, in a patient with compromised haemostasis the bleeding may be prolonged. Although the bleeding was no more than a gentle ooze at any point in time it was persistent enough for 36 hours before the topical haemostatic material Surgicel had been put to use. The consequences of an intraocular bleed may have seriously threatened her sight.1

We are not aware of any reports of the use of Surgicel in ophthalmic surgery. All reports of its use are in other fields of surgery.2 This material is supposed to swell up with blood and form a gelatinous mass that aids in the formation of clot. It acts as a haemostatic adjunct. The exact mode of its action in this patient with antiplatelet antibodies is unclear. Our experience shows that oxidised regenerated cellulose (Surgicel) may have a role in ophthalmic surgery especially in lacermal and orbital surgery, when faced with bleeding that is difficult to stop. Various cautionary tales associated with use of Surgicel have been reported.3

Our report suggests that in the presence of a severe bleeding disorder, clear corneal phacoemulsification under topical anaesthesia may not be totally safe. When performing such a procedure in a patient with known bleeding disorder it may be safe to take all the necessary precautions in consultation with a haematologist to avoid a serious bleed that may be sight and life threatening. There may be a role for haemostatic agents like Surgicel.

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Figure 1  [A] Conjunctival site immediately after removal of Surgicel. [B] Healed conjunctival bleeding site.
Institute for further evaluation. Vitreal analy-
sis revealed acuities of 20/25 in the right eye. A portion of the vitreous specimen was cultured for fungi, aerobic and anaerobic micro-organisms were negative.

The right eye was unresponsive to corticosteroid treatment. A 78 year old man with a history of vitreous aspirate vitreous cells and a choroidal naevus. A left eye was normal with the exception of flare, dilated iris vessels, an IOL without significant for no anterior chamber cells or intraocular pressures. The right eye was suspected of delayed postoperative endophthalmitis diagnosed by histopathological technique of microdissection and PCR.

The diagnosis of delayed postoperative endophthalmitis is suspected. In this case, although the chronic inflammation and intracytoplasmic Gram positive bacilli in a few macrophages suggested an infectious process, the negative cultures precluded the diagnosis of an infectious endophthalmitis. To further investigate the possibility of a bacterial infection nested PCR was performed on the micro-dissected bacilli.

Molecular analysis verified the presence of P acnes and a diagnosis of delayed postoperative endophthalmitis was confirmed.

Propionibacterium acnes endophthalmitis diagnosed by microdissection and PCR

Although Propionibacterium acnes, a Gram positive anaerobic bacillus, is the most commonly identified cause of delayed onset postoperative endophthalmitis, routine vitreous cultures are frequently inadequate for its diagnosis. This case describes the utility of the histopathological technique of microdissection and polymerase chain reaction (PCR) for the diagnosis of delayed postoperative endophthalmitis.

Case report

A 78 year old man with a history of vitreous floaters, a coronary bypass, and aortic valve replacement underwent an uncomplicated cataract extraction with intraocular lens (IOL) implantation in the right eye. Three months later, he developed increasing floaters in the right eye and was diagnosed with vitritis unresponsive to corticosteroid treatment. Examination revealed acuities of 20/25 in the right eye and 20/20 in the left with normal intraocular pressures. The right eye was significant for no anterior chamber cells or flare, dilated iris vessels, an IOL without deposits, 3+ vitreous cells with trace haze, and peripheral pigmentary degeneration. The left eye was normal with the exception of trace vitreous cells and a choroidal naevus. A diagnostic vitrectomy was performed in the right eye. A portion of the vitreous specimen was cultured for fungi, aerobic and anaerobic bacteria, and the remainder was processed for cytopathological examination. All cultures for micro-organisms were negative.

The vitreous supernatant and unstained cytology slides were sent to the National Eye Institute for further evaluation. Vittreal analysis for interleukin 2 (IL-2), IL-4, IL-6, IL-10, IFN-γ, and TNF-α using ELISA (Endogen, Woburn, MA, USA), revealed undetectable cytokine levels. The vitreous slides were stained with Giemsa, Gram, and immunohistochemical stains for T cells, B cells, and macrophages. Cytopathological examination showed clusters of macrophages admixed with CD4+ and CD8+ T cells and B cells (Fig 1A). Gram positive bacilli were seen in the cytoplasm of a few macrophages (Fig 1B). The engulfed bacilli were microdissected under a microscope with a 30 gauge needle and submitted for PCR. Nested PCR with P acnes specific oligodeoxynucleotide primers complementary to regions of 16S rDNA was used. The primers were Pa1, AAG GCC CTG CTT TTG TGG; Pa2, TCC ATC CGC AAC GGC CGA A; and rPa3, ACT CAC GCT TGG TCA CAG. Nested-PCR analysis revealed P acnes (Fig 2). A diagnosis of delayed postoperative endophthalmitis was made.

The most common causes of vitritis in elderly patients are acquired or postoperative infections, sarcoidosis, and intraocular malignancies masquerading as uveitis. An early diagnostic procedure is indicated if postoperative endophthalmitis is suspected. In this case, although the chronic inflammation and intracytoplasmic Gram positive bacilli in a few macrophages suggested an infectious process, the negative cultures precluded the diagnosis of an infectious endophthalmitis. To further investigate the possibility of a bacterial infection nested PCR was performed on the micro-dissected bacilli.

Molecular analysis verified the presence of P acnes and a diagnosis of delayed postoperative endophthalmitis was confirmed.

Vitreous cultures are positive in less than 50% of postoperative endophthalmitis cases. In a study of 25 patients with delayed onset endophthalmitis aqueous culture and microscopy were diagnostic in 0% of cases, vitreous culture was positive in 24% and PCR from the aqueous and vitreous yielded a positive diagnosis in 84% and 92%, respectively.

Treatment of P acnes endophthalmitis includes intravitreal vancomycin plus consideration of pars plana vitrectomy with or without capsulotomy with or without IOL removal. Although aggressive surgical intervention eradicates the infection similar visual outcomes are reported with more limited surgical treatment.

In our case the intracytoplasmic bacteria in the macrophages were the only evidence of a bacterial infection. To detect the presence of P acnes we referenced the PCR method described by Hykin that used 150 µl of the vitreous culture and 100 µl for PCR. Using the technique of microdissection and PCR with a similar volume of vitreous we additionally performed cytology and cytokine analysis which are helpful in the diagnosis of other causes of vitritis.

This case further illustrates the benefits of molecular analysis for the diagnosis of culture
negative delayed onset endophthalmitis. It also describes for the first time microdissec-
tion and PCR for the evaluation of endoph-
thallitis. Advantages of this technique are
that it allows for a more comprehensive
pathological examination on a limited speci-
men and provides the option of having the
molecular studies being performed elsewhere.

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References
dNA analysis of specific cell populations in
lesions <1 mm in size. Am J Pathol 1995; 146:955–9

2 Hykin PG, Tobal K, McIntyre G, et al. The
diagnosis of delayed post-operative
endophthalmitis by polymerase chain reaction
of bacterial DNA in vitreous samples. J Med

3 Berrocal AM, Davis JL. Uveitis following
intraocular surgery. Ophthalmol Clin North
Am 2002; 15:357–64

4 Lohmann CP, Linde HJ, Reischl U. Improved
detection of microorganisms by polymerase
chain reaction in delayed endophthalmitis
after cataract surgery. Ophthalmolgy
2000; 107:1047–51; discussion 1051–2

5 Aldave AJ, Stein JD, Deramo VA, et al. Treatments strategies for postoperative
Propionibacterium acne endophthalmitis.
Ophthalmol 1999; 106:2395–401

6 Buggage RR, Whitcup SM, Nussenblatt RB, et al. A rising interleukin 10 to interleukin 6 ratio
to distinguish primary intraocular lymphoma and uveitis. Invest Ophthalmol Vis Sci
1999; 40:2462–3

Interferon treatment of
childhood conjunctival
lymphoma

Mucosa associated lymphoid tissue (MALT) lymphoma is the most common ocular adni-
canal or lacrimal gland neoplasm. These neoplastic lesions have a more indolent course than
non-MALT lymphomas, are usually found in the older age groups (50–70 years), are
usually limited to localised (stage I) disease at presentation, and radiotherapy and che-
motherapy have been the mainstay of treatment.1

Case report
A 15 year old male was referred by an ophthalmologist after an 8 month history of
unusual painless follicles at both nasal for-
nices (Fig 1A). There were no visual symp-
toms and, based on a working diagnosis of an
atypical conjunctival lesion, the patient was referred to the Department of Ophthalmology,
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References
1 Blasi MA, Gherlinzoni F, Calvisi G, et al. Local chemotherapy with interferon-α
for conjunctival mucosa-associated lymphoid tissue lymphoma. Ophthalmology
2001;108:559–62

aspects of 117 cases and relationship to systemic lymphoma. Ophthalmology
2001; 108:979–84

3 Akpek EK, Polcharoen W, Ferry JA, et al. Conjunctival lymphoma masquerading as
conjunctival cytomegaly. Ophthalmology
1999; 106:757–60

4 Hearing AH, Franke FE, Hütz WW. Conjunctival CD5+ MALT lymphoma. Br J
Ophthalmol 2001; 85:498–9

5 Cahill MT, Moirarity PA, Kennedy SM. Conjunctival “MALToma” with systemic recurrent.


7 Bessel EM, Henk JM, Whitecock JF, et al. Ocular morbidity after radiotherapy of orbital
and conjunctival lymphoma. Eye 1987;1:90–6

lymphoid tissue lymphoma with intralymphatic injection of interferon alfa-2b.
Arch Ophthalmol 2000; 118:284–9

in an HIV+ patient. Eur J

www.bjophthalmol.com

Unilateral corneal anaesthesia and ulceration following squint surgery in a child with Pendred syndrome and bilateral sixth nerve palsy

We present a 4 year old child with Pendred syndrome and bilateral sixth nerve palsy. To our knowledge this association has not been previously reported. In addition, this patient developed unilateral corneal ulceration with associated corneal anaesthesia following squint surgery. We will discuss the pathophysiology of this unusual complication following squint surgery.

Case report

This patient presented when he was 6 months old with right congenital divergent squint. He was diagnosed with Pendred syndrome (sensorineural hearing loss and thyroid dysfunction) by the paediatricians and the otolaryngologists following abnormal thyroid function tests and a computed tomograph (CT) scan of the temporal bones showing Mondini malformations of both cochleae. At presentation his visual acuities were 6/60 right eye and 6/36 left eye using the Cardiff acuity cards. He had bilateral alternating esotropia with an inability to abduct either eye. There was no globe retraction or abnormal lid movements and a magnetic resonance imaging (MRI) scan had shown congenital absence of the auditory nerves but no other abnormality. A diagnosis of bilateral sixth nerve palsy was made. The squint was cosmetically poor and measured at 45 prism dioptries in the distance and near. He had low hypermetropia with no significant anisometropia. Funduscopic was normal. He was reviewed regularly in the paediatric eye clinic over the next 3 years during which time his visual acuities were within normal limits, the best recorded acuity being 6/9 right eye and 6/9 left eye using singles.

When he was 4 years old, he underwent bilateral superior rectus and inferior rectus lateral transpositions under general anaesthesia, which was uneventful with no immediate postoperative complications, and a cosmetic result of stable alternating convergent squint of 15 prism dioptries for distance and near.

Two months later he developed a left inferior corneal ulcer (Fig 1) with surrounding punctate epitheliopathy which surprisingly did not seem to cause him as much distress as expected. The left corneal sensation was definitely reduced compared to the right which appeared normal. Sensation was assessed clinically (an anaesthesiometer was not available), and was consistently reproducible by different ophthalmologists. There was no exophthalmos or any other sign of thyroid orbitopathy. The right eye remained asymptomatic. Empirical therapy with topical olofoxacin and lubricants was unhelpful. He proceeded to have glue tarsorrhaphy which transiently aided the healing of the corneal ulcer. However, the ulcer quickly recurred when the tarsorrhaphy reversed. He subsequently had left inferior lid shortening with a canthal sling to elevate the lower lid to protect the corneal epithelium. The ulcer resolved leaving an area of corneal scarring. He is being reviewed regularly in the eye clinic.

Comment

Pendred syndrome is an autosomal recessive disorder characterised by congenital deafness and thyroid goitre with a high incidence of deafness is usually severe and is present at birth, and the goitre generally appears at puberty or later but may be present in early childhood with an associated euthyroid or hypothyroid state.1 11-12 Affected individuals are reported to be otherwise normal.

The pathophysiology of the corneal anaesthesia and ulceration in this patient is unclear. There are several reasons for the corneal anaesthesia. They include herps simplex keratitis, postoperative anterior segment ischaemia, surgical trauma to the long posterior ciliary nerves or ciliary ganglion, congenital absence of sensation, and surgery reducing Bell's phenomenon.

The clinical course was not typical of herpetic simplex and there was no previous history of corneal pathology. Postoperative anterior ischaemic syndrome was unlikely as only two recti muscles were operated on and no anterior uveitis was observed. To our knowledge there are no reported cases of corneal anaesthesia after squint surgery. There was no evidence of pre-existing involvement, which one may expect with trauma to the long posterior ciliary nerves or ciliary ganglion.

Congenital absence of corneal sensation was the most likely cause, especially in view of his unusual cranial nerve anomalies, and we believe he had pre-existing corneal anaesthesia before squint surgery despite the absence of any other fifth cranial nerve signs. Following the lateral transposition of the superior rectus his Bell's phenomenon was noted to be absent thereby compromising his corneal protection. In addition, he was observed to have significant lagophthalmos while asleep. We believe that the combination of corneal anaesthesia, abolished Bell's phenomenon, and lagophthalmos compromised his corneal integrity resulting in corneal ulceration.

This case highlights the importance of determining corneal sensation before transposition surgery on the superior rectus as Bell's phenomenon may be abolished therefore compromising corneal protection. This is especially relevant in patients with unusual cranial neuropathy and lagophthalmos.

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References


Gemella haemolytica acute postoperative endophthalmitis

Endophthalmitis is perhaps the most feared complication of cataract surgery, with a reported incidence beyond 0.13%.1 2 The most common organisms reported in previous studies are Gram positive staphylococci and streptococci.1 3 We report a case of severe endophthalmitis with an unusual Gram positive organism, after uncomplicated phacoemulsification, with foldable intraocular lens implantation.

Case report

A 66 year old white man underwent routine phacoemulsification cataract extraction with posterior chamber lens implantation (Acrylic, Model Hydroview Hero, Bausch & Lomb) to the right eye in January 2002. The left eye had previously undergone similar surgery in September 2001. He was generally in good health, and on no medication. There was a past medical history of sarcoidosis treated with oral prednisolone in 1970, which has since been in remission, and an episode of staphylococcal septicemia in 1987, without sequelae.

On the first postoperative day, visual acuity measured 6/9 unaided and ocular examination was unremarkable. That same afternoon the patient developed ocular pain, initially relieved by paracetamol (acetaminophen), which however, worsened during the night with progressive deterioration of vision. He presented to the ophthalmic emergency department the following morning with the aforementioned symptoms. Visual acuity was reduced to hand movements right eye and 6/9 left eye. Slit lamp examination revealed an oedematous cornea with Descemet's folds. The anterior chamber was hazy, with 1 mm hypopyon and the intraocular pressure measured 38 mm Hg.

There was no red reflex. B-scan ultrasound examination showed extensive vitreous debris with attached retina. The left eye was pseudoephelitic with no abnormalities of note. A diagnosis of acute postoperative endophthalmitis was made. Anterior chamber and vitreous samples were obtained for aerobic and anaerobic culture/sensitivity and Gram staining. Intravitreal vancomycin 2 mg and amikacin 300 µg, each in 0.1 ml of balanced salt solution and subconjunctival cefoxime 125 mg were administered. Oral ciprofloxacin 500 mg twice daily, prednisolone 40 mg once a day, topical gentamicin hourly, ofloxacin hourly, and atropine 1% twice a day were commenced.

Preliminary Gram staining suggested a Gram positive coccus sensitive to ciprofloxacin—oral and topical antibiotics were therefore continued. Owing to difficulty in identifying the nature of the organism, the samples were sent to a regional reference laboratory, which identified G. haemolytica from both anterior chamber and vitreous aspirates. The organism was reported to be sensitive to gentamicin, ciprofloxacin, laevofloxacin, amoxicillin/clavulanate, chloramphenicol, and resistant to trimethoprim.

Endophthalmitis due to Gemella haemolytica was diagnosed. The patient had mild hypertension and the intraocular pressure remained 38 mm Hg. The visual acuity reduced to hand movements right eye and 6/9 left eye. Slight improvement of vision occurred despite a relapse of endophthalmitis. The anti-inflammatory therapy and follow-up was satisfactory and the patient was discharged.


gemella haemolytica; endophthalmitis; phacoemulsification; intraocular lens implantation
The patient continued to make steady progress; 2 months later vision had improved to 6/9 unaided. The patient at that time was troubled by floaters secondary to considerable vitreous debris. At last review in September 2002, visual acuity had further improved to 6/4 with −0.75DS ph correction.

Comment

Gemella haemolysans is an aerobic or facultative anaerobic, Gram positive coccus, a normal commensal of the oral cavity and upper respiratory tract of low virulence. It is a non-pathogenic anaerobic, Gram positive coccus, a normal commensal of the oral cavity and upper respiratory tract of low virulence. Systemic infection may lead to septic shock, meningitis, arthritis, or pneumonia, all of which are rare. Identification is difficult. Though Gram positive, the cocci are easily decolorised and hence may appear Gram variable or even negative. Identification is difficult. Though Gram positive, the cocci are easily decolorised and hence may appear Gram variable or even negative.

Gemella was included under the genus Neisseria but is now classified as a separate genus within the family Streptococcaceae. No studies on susceptibility to antiseptics have been published, though there is no reason to believe that it may be resistant to povidone-iodine preparations. The organism is susceptible in vitro to penicillin, streptomycin, vancomycin, chloramphenicol, and tetracycline. A literature search revealed only one previously reported case of infection by Gemella haemolysans, with keratitis and consecutive endophthalmitis. Interestingly this patient was reported to have active sarcoidosis on systemic steroid therapy, whereas our patient had a past history of sarcoidosis. This possible association between sarcoidosis and infection by Gemella may be purely coincidental, as no such association has been reported with systemic infection.

Gemella haemolysans is difficult to identify, because of its close resemblance to viridans streptococcus and Neisseria. As diagnostic technology improves, Gemella haemolysans endophthalmitis may be described more often in the future. This report highlights the importance of infection with rare commensal organisms in healthy, immunocompetent individuals after uneventful phacoemulsification cataract surgery.

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References


Does topical brimonidine tartrate help NAION?

There is no proved treatment for non-arteritic anterior ischaemic optic neuropathy (NAION). Topical brimonidine tartrate has been reported to have a neuroprotective benefit for retinal ganglion cells following experimental elevation of intraocular pressure and optic nerve injury in the rat, which is blocked with coadministration of the α2 antagonist, rauwolscine. Increased retinal ganglion cell survival has also been shown to occur following oral administration of brimonidine in monkeys with experimental glaucoma. These results were the basis of the recently aborted clinical trial of topical brimonidine furate for acute NAION and our retrospective study of 31 patients with NAION, who were evaluated within 3 weeks of the onset of visual loss and followed up for a minimum of 8 weeks. During 2001–2, we treated all (14) patients with brimonidine tartrate within 14 days (mean 3.3, SD 3.52) of the onset of visual loss. Five patients were treated after 1 day of symptoms. Three were taken four times a day in 11, three times a day in one, and twice a day in two patients. All (17) untreated patients were evaluated the year before and were matched to the treated group for age, sex, cardiovascular risk factors, previous aspirin use, and previous first eye NAION.

Snellen visual acuity and colour vision, using the Ishihara colour plates, were documented and expressed as a decimal equivalent (for acuity: 20/60 = 0.33 and light perception = 0.001; for colour vision: the number of correctly identified plates/the total number of plates). The visual fields (Humphrey or tangent perimetry) were analysed and defects were graded according to the following scale: 0 = normal, 1 = accurate nerve fibre bundle defects, 2 = retinal nerve fibre bundle defects (6 degrees), 3 = arcuate, and 4 = sectoral defects, 5 = altitudinal defect plus additional loss, 6 = no light perception. A third examiner, who was unaware of the dates of the visual fields and the patients’ treatment status, also evaluated all visual fields and determined, in each patient, whether the field was better or worse than or equivalent to the other field. The intraocular pressure was noted in all patients except two patients. The pressure was 25 mm Hg in one patient in the treated group and 24 mm Hg in one patient in the untreated group.

Statistical analysis of the data involving comparisons of the treated and untreated groups at baseline and 8–12 weeks was performed using the two tailed t test. The Wilcoxon signed rank test was used to compare the individual visual performance from baseline to the 8–12 week examination. For visual acuity and colour vision, a positive rank indicated improvement and a negative rank indicated a worse visual outcome. For the visual field grade, the scale was reversed. Spearman correlation analysis was performed on the time to start therapy and whether worsening in any visual parameter occurred. The mean baseline acuity (0.56, SD 0.30) and visual field (1.9, SD 0.73) for the treated group was similar to the acuity (0.40, SD 0.22) and field (1.9, SD 0.73; p=0.96) for controls. The mean baseline colour vision (0.74, SD 0.44) for the treated group was higher than the colour vision (0.45, SD 0.44) for controls, but the difference was not significant (p=0.07). At the 8–12 week examination, the mean visual acuity was 0.29 (SD 0.30) for treated and 0.49 (SD 0.39; p=0.12) for untreated patients. The mean visual field grade was 2.2 (SD 0.81) for treated and 1.0 (SD 0.70; p=0.04) for untreated patients. The mean colour vision was 0.42 (SD 0.41) for treated and 0.55 (SD 0.46; p=0.43) for untreated patients.

For the masked examiner’s evaluation, the mean baseline visual field (2.0, SD 0.91) was similar to the field (1.93, SD 0.96; p=0.85) for controls. At the 8–12 week examination, the mean visual field grade was 2.15 (SD 0.99) for treated and 1.87 (SD 0.92; p=0.43) for untreated patients. This examiner further found that the outcome visual fields for the treated group were improved in two patients, worsened in six patients (50%), and unchanged in four patients. The outcome visual fields for the control group were improved in five patients, worsen in two patients (13%), and unchanged in eight patients.

The Wilcoxon signed rank analysis demonstrated that for visual acuity, two patients in the control group and 10 patients in the treated group had negative values or a worse outcome at 8–12 weeks (p=0.046). For colour vision, one patient in the control group and eight patients in the treated group had negative values or a worse outcome (p=0.013). For visual fields, one patient in the control group and four patients in the treated group had positive values or a worse outcome at 8–12 weeks (p=0.046).

The average time to start the drops was 3.5 days from the onset of visual loss in those patients who worsened. There was no correlation with a worse outcome and time to initiate therapy.

For all parameters of vision testing, there was a trend for worse visual performance at 8–12 weeks in the group treated with topical brimonidine. Although there was no significant difference for the colour vision outcome, this might reflect that the baseline colour vision value was better for the treated group. The outcome visual field grade was significantly worse in the treated group. The masked examiner’s visual field evaluations demonstrated that more treated patients worsened than in the untreated group. When the baseline and outcome of all visual parameters for each individual were compared, the treated group had a significantly worse outcome at 8–12 weeks.

Our results are not the first description of worse outcome in patients treated with α2 agonists for central nervous system ischaemic disease. Studies in animal models and clinical studies in humans suggest that certain classes of drugs, including α2 receptor agonists, may impede recovery following stroke. Clonidine administration caused recurrence of the neurological deficit in animals who had initially recovered. In a recent clinical study, the level of motor recovery of stroke patients was worse in those treated with α2 agonists than in patients not receiving these agents.

Although in experimental optic nerve injury in animal models, brimonidine appears to offer neuroprotection, our results demonstrate that brimonidine tartrate, applied topically up to four times daily, does not appear to be a beneficial treatment for acute NAION.
is possible earlier treatment might have been more effective, although patients who worsened received treatment sooner than those who did not worsen. Increased dosing frequency or using a different preparation of botulinum toxin might be more effective. Additionally, the number of subjects in the study was small and a negative trend could appear more profound.

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Comment
The pain demonstrated by these two patients is typically much worse in certain directions of gaze and particularly during prolonged gaze holding such as when reading or watching television. It had a clear precipitating event and the most remarkable feature is that it had persisted for over 2 years in each case with no significant progression or regression. No active disease process could be found to account for the continued pain. The pain is severe and responds only to high doses of analgesics, particularly NSAIDs. None of our patients felt that their pain was satisfactorily controlled by their analgesics.

We believe that there may be a process of chronic low grade inflammation affecting the extracranial muscles around the eye which is exacerbated by continued contraction and relaxation of the same muscles. Muscular spasm perhaps triggered by this inflammatory process may be the cause of the most severe pain and this could account for the exacerbations of pain in certain directions of gaze and on prolonged gaze holding activities. Ocular muscle ischaemia, perhaps caused by constricting scar tissue, remains a possibility but the onset of the pain is very fast making this less likely.

The pain relief seen in our patients may simply be the result of paralyzing an inflamed muscle but there is growing evidence for a separate antinociceptive effect of botulinum toxin.1 No direct peripheral cutaneous antinociceptive effect could be shown by Blersch et al,6 however inhibition of release of substance P has been demonstrated in vitro and it can be hypothesised that botulinum toxin treatment may reduce the local release of nociceptive neuropeptides from either cholinergic neurons or from C or A delta fibres in vivo. The mechanisms by which botulinum toxin may relieve pain, including a possible analgesic effect of botulinum toxin metabolites, are reviewed by Guyer.7

There is a growing literature on the use of botulinum for painful conditions,8 particularly those in which muscle spasm plays a part. These include writer’s cramp,9 postoperative pain in spinal cerebral palsy,10 and perhaps more surprisingly perhaps11 and painful tic convulsif.12 Many of the reported uses are single case studies and not all controlled trials have shown a positive effect of treatment.

It is not possible to rule out a powerful placebo effect in our patients but, whatever the mechanism of action, their pain was vastly improved and botulinum toxin treatment is very safe in competent hands.

In the cases described paralyzing botulinum toxin served a dual purpose in that it had the potential to improve their ocular deviation for which it is well known and it also reduced the severe ocular discomfort. Unfortunately, the resulting diplopia limited its usefulness in one case but we feel that this treatment should be considered in this unusual group of patients who present a difficult management problem even to the most experienced ophthalmologists.

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References
Intrastromal lamellar femtosecond laser keratoplasty with superficial flap

Lamellar keratoplasty has usually been performed taking a trephine to delineate the extent of the tissue to be excised, and a knife or similar instrument to remove the lamellar corneal tissue from the underlying deep corneal bed. In a similar way, the lamellar donor tissue was prepared and inserted into the recipient bed. The depth of the lamellar excision corresponded to the surgical trephine, i.e., to the superficial surface to the deep corneal stroma. Marked disadvantages of the technique have been pronounced corneal astigmatism and optical insufficiency at the interface between the lamellar graft and the recipient corneal bed caused by irregularities of both surfaces. The purpose of the present report was to describe the new femtosecond laser technology, which may enable us to perform a new type of intrastromal lamellar keratoplasty with preservation of an intact Bowman's layer and regular corneal epithelium.

Case report

Using a corneal contact lens and a femtosecond laser (20/10 Perfect Vision, Amoerfeldt 211 D-69123 Heidelberg, Germany) with a wavelength of 1060 nm, a spot size of about 10 μm, and a laser pulse duration of several hundred femtoseconds, a pre-descentemal incision running parallel to the corneal surface was created in five postmortem eyes of slaughterhouse pigs. The diameter of the deep stromal incision was 7 mm. In a second step, a circular sagittal incision was performed starting from the peripheral edge of the already existing incision in the pre-descentemal level to the superficial layer of the corneal stroma. In continuation of the latter, the corneal flap was prepared with a diameter of 7 mm, a thickness of about 100 μm, a hinge, and three positional pikes. The pikes in the flap with the corresponding markings on the back of the flap were formed to increase the rotational stability of the flap after repositioning. The height of the peaks was about 0.40 mm. After opening of the flap the intrastromal segment situated between the pre-descemetal incision and the incision in the superficial stromal level was removed and exchanged against a similar formed segment obtained from another (donor) pig eye. In all cases, the flap was repositioned.

For all eyes included in the study, the intrastromal corneal button and the superficial flap with the three positional pikes could be prepared without major difficulties. The corneal buttons could easily be repositioned into their original beds as well as into the recipient beds of other eyes in which the recipients were created with the same diameter as the donor button. The time taken for preparation of the intrastromal corneal button and the corneal flap, and for the exchange of the corneal buttons was less than 10 minutes in all cases.

Comment

Femtosecond laser technology allows a new type of intrastromal lamellar keratoplasty with removal of a mid-stromal segment and preservation of an intact Bowman's membrane. Considering the decreased amount of allogenic corneal tissue transplanted, and regarding the preservation of the original corneal surface, lamellar intrastromal femtosecond laser keratoplasty may be associated with a smaller rate of immunological graft reaction and with a lower postoperative corneal astigmatism in some eyes. Future clinical studies may show whether positional edges in the superficial flap increase its postoperative rotational stability.

Proprietary interest: none

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References


Demographic study of paediatric allergic conjunctivitis within a multiethnic patient population

From October 1999, all patients referred to the paediatric ophthalmology service in Bradford have been added to a computerised database. This is the only paediatric ophthalmology service within the city of Bradford and receives all GP referrals of this type. Patients with a clinical diagnosis of chronic allergic conjunctivitis were identified from October 1999 to July 2001. Comparisons were made if the patient had characteristic symptoms and signs based on criteria set out by Buckley in 1998. This was done to ensure accurate and consistent diagnosis of chronic allergic conjunctivitis. The patients were divided into groups by ethnic origin: white, Asian, and mixed disease. There were 24 Asians and 11 white children. For Asian patients, the mean age was 9.58 (SD 0.82) years. For the white patients, the mean age was 7.82 (SD 3.19) years. Follow up ranged from 3–14 months, mean 6 months.

The prevalence of allergic conjunctivitis in Asians was 59 per 100 000 (24 in 40 524) and in white children, 12 per 100 000 (11 in 93 998); a relative prevalence of 5 to 1 (χ2 test, p <0.001).

There was a predominance of males in both ethnic groups, 2:1 in Asians and 1:2:1 in white children. This difference in sex was not significant between both groups (Fisher's test, p =0.71).

The overall age distribution for all males was 5 to 14 years and for all females was 10 to 16 years. For Asians, the mean age for males was 9.18 years and for females was 10.57 years. For white children, the mean age was 7.00 years and for females was 9.25 years.

Comment

Various studies have reported allergic eye disease to be more common among Asian and black patients. This may be due to genetic and environmental factors.

We found allergic eye disease to be more common in Asians than white children. It is possible that ocular allergy is multifactorial but perhaps with a greater genetic predisposition in certain ethnic communities. We could not comment on the prevalence of chronic allergic conjunctivitis in the community because of referral bias since we only see patients referred by GPs. The extent to which milder cases are treated in the community is not known but we feel that the more severe cases are the ones referred to our department.

Our findings highlight that allergic eye disease appears to be more common and complicated in Asian patients in the Bradford population. This potential risk of sight threatening disease means that they are more likely to require topical steroid treatment. This has led us to recognise that appropriately aggressive treatment is essential in these patients.

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References


Rise in organ transplant numbers

According to UK Transplant, the UK has seen the highest number of organ transplants in six years.

Second Sight

Second Sight, a UK based charity whose aims are to eliminate the backlog of cataract blindness in India, has established strong links between Indian and British ophthalmologists, is regularly sending volunteer surgeons to India. Details can be found at the charity’s website (www.secondsight.org.uk) or by contacting Dr Lucy Mathen (lucymathen@yahoo.com).

Elimination of avoidable blindness

The 56th World Health Assembly (WHA) considered the report on the elimination of avoidable blindness (doc A56/26) and urged Member States to: (1) Commit themselves to supporting the Global Initiative for the Elimination of Avoidable Blindness by setting up a national Vision 2020 plan by 2005; (2) Establish a national coordinating committee for Vision 2020, or a national blindness prevention committee to help implement the plan; (3) Implement the plan by 2007; (4) Include effective monitoring and evaluation of the plan with the aim of showing a reduction in the magnitude of avoidable blindness by 2010; (5) To support the mobilisation of resources for eliminating avoidable blindness. The WHA also urged the Director-General to maintain and strengthen WHO’s collaboration with Member States and the partners of the Global Initiative for the Elimination of Avoidable Blindness as well as aid in the coordination and support of national capability.

MSc course in Community Eye Health

The International Centre for Eye Health is offering a full time MSc course in Community Eye Health from 29 September 2003 to 19 September 2004. The course is not clinical and is specifically for eye health professionals wanting to work in the field of community eye health. The course is designed in keeping with the aims, priorities, and strategies of Vision 2020—the Right to Sight. The course costs £3939 for home students and £14 110 for overseas students. Further information: The Registry, 50 Bedford Square, London WC1B 3DP, UK (tel: +44 (0)20 7927 2239; fax: +44 (0)20 7323 0638; email: Adrienne.Burrough@lshtm.ac.uk; website: www.lshtm.ac.uk).

Ophthalmic Anesthesia Society (OAS)—17th Scientific Meeting

The 17th Scientific Meeting of the Ophthalmic Anesthesia Society (OAS) will be held 3–5 October 2003 at the Westin Michigan Avenue Chicago, Chicago, USA. Programme co-chairs: Marc Allen Feldman MD MHS and Steven T Charles MD. The CME joint sponsor is the Cleveland Clinic Foundation; CME hours are pending. Fees for OAS members are $300; non-members $475; students $50. Further details: OAS, 793-A Foothill Blvd, PMB 1319, San Luis Obispo, CA 93405 USA; tel: +1 805 354 0300; fax: +1 805 354 9300; email: info@eyeanaesthesia.org; website: www.eyeanaesthesia.org.

Glaucma Society 24th Annual Meeting and Dinner

The Glaucma Society 24th Annual Meeting and Dinner will take place on 20 November 2003, from 8.30 am at the College of Physicians, London, UK. Further details: Ms Janet Flowers (email: glauсос@ukiere freeserve.co.uk).

Detachment Course with international faculty on: Retinal and Vitreous Surgery with Case Presentations preceding the Annual Meeting of Iranian Society of Ophthalmology

The detachment course with international faculty on: Retinal and Vitreous Surgery with Case Presentations preceding Annual Meeting of Iranian Society of Ophthalmology will be held...
on 29–30 November 2003 and 1–4 December 2003 respectively, at the Razi Conference Center, Hemmat Hyw, Tehran, Iran. Further details: Scientific programme: Prof Ingrid Kreissig, University of Tuebingen, Schleichstr. 12, Breuningerbau, 72076 Tuebingen, Germany (tel: +49 7071 295209; email: ingrid.kreissig@med.uni-tuebingen.de). Local organisation: Dr Arman Masheyekhi, Dr Siamak Moradian, Dept of Ophthalmology, Labanfinejad Medical Center, Pasdaran Ave, Boosten 9, Tehran, 16666, Iran (fax: +98 21 254 9039; email: labbafi@hotmail.com).

5th International Symposium on Ocular Pharmacology and Therapeutics (ISOPT)
The 5th International Symposium on Ocular Pharmacology and Therapeutics (ISOPT) will take place 11–14 March 2004, in Monte Carlo, Monaco. Please visit our website for details of the scientific programme, registration, and accommodation. To receive a copy of the Call for Abstracts and registration brochure please submit your full mailing details to http://www.kenes.com/isopt/interest.htm.


XVth Meeting of the International Neuro-Ophthalmology Society
The XVth Meeting of the International Neuro-Ophthalmology Society will take place 18–22 July 2004, in Geneva, Switzerland.
Further details: Prof. A Safran, University Hospital Geneva, c/o SYMPOG SA, Geneva (fax: +4122 839 8484; email: info@symopg.ch; website: www.symopg.ch).