from presentation was considered a good result, and longer healing times a poor result. Logistic regression was used to predict a good/poor result, the primary outcome, using log2-transformed MIC as a covariate. The regression model was analysed using Pearson’s goodness of fit. A Fisher’s exact test was used to correlate genus (Aspergillus vs Fusarium) to good/poor result. All analyses were performed using STATA 9.2.

**RESULTS**

The baseline characteristics and MICs of 54 patients with fungal corneal ulcers are described in table 1.

A lower MIC was significantly associated with a good outcome, as was *Fusarium* species (as opposed to *Aspergillus* species) (table 2). When restricted to a subgroup of organisms, for example *Fusarium* cases, the relationship between MIC and clinical outcome was similar in magnitude, but no longer statistically significant (OR=0.51, 95% CI 0.12 to 2.05, p=0.54). In addition, when restricted to only *Aspergillus* cases, the relationship was neither significant nor similar in magnitude (OR=1.37, 95% CI 0.39 to 4.88, p=0.63).

**COMMENT**

Many factors contribute to the success or failure of fungal keratitis management, including ulcer size, ulcer location, organism, penetration of antimicrobial agent and susceptibility of the organism to treatment. In bacterial keratitis, studies suggest that susceptibility of the organism to the agent in vitro correlates with outcome. It remains unclear whether susceptibility correlates with outcome in fungal keratitis. In systemic fungal disease, researchers suggest that the role of susceptibility testing may be similar to that of bacterial susceptibility testing, where approximately 90% of susceptible cases and 60% of resistant cases respond to therapy. Antifungal susceptibility testing is associated with outcome in mucosal candidiasis and candidaemia, and antifungal susceptibilities influence treatment recommendations.

In fungal keratitis, in vitro susceptibility did correlate with outcome. A twofold increase in MIC was associated with a 47% reduction in the odds of healing. In addition, the organism is associated with outcome. Since only 54 of 90 cases with completed susceptibility testing had clinical data available, the study had limited generalisability. Further prospective studies would be necessary to assess whether MIC provides information useful to the clinician once the organism species has been identified, as well as the effect of other covariates such as toxicity, prior medications, age and sex.

**Table 1** Fungal organisms isolated from retrospective review of corneal ulcers from March—June 2004 (n=54)*

<table>
<thead>
<tr>
<th>Organisms</th>
<th>n (%)</th>
<th>MIC&lt;sub&gt;C50&lt;/sub&gt; (µg/ml)</th>
<th>MIC&lt;sub&gt;C90&lt;/sub&gt; (µg/ml)</th>
<th>MIC range (µg/ml)</th>
</tr>
</thead>
<tbody>
<tr>
<td><em>Aspergillus</em> species</td>
<td>24 (44)</td>
<td>32</td>
<td>64</td>
<td>8–64</td>
</tr>
<tr>
<td><em>Aspergillus</em> flavus</td>
<td>18 (33)</td>
<td>32</td>
<td>64</td>
<td>16–64</td>
</tr>
<tr>
<td><em>Aspergillus</em> niger</td>
<td>2 (4)</td>
<td>—</td>
<td>—</td>
<td>—</td>
</tr>
<tr>
<td><em>Aspergillus</em> terreus</td>
<td>3 (6)</td>
<td>—</td>
<td>—</td>
<td>—</td>
</tr>
<tr>
<td><em>Aspergillus</em> fumigatus</td>
<td>1 (2)</td>
<td>—</td>
<td>—</td>
<td>—</td>
</tr>
<tr>
<td><em>Fusarium</em> species</td>
<td>23 (43)</td>
<td>8</td>
<td>16</td>
<td>4–16</td>
</tr>
<tr>
<td>Unidentified hyaline species</td>
<td>3 (6)</td>
<td>—</td>
<td>—</td>
<td>8–64</td>
</tr>
<tr>
<td><em>Acremonium</em> species</td>
<td>1 (2)</td>
<td>—</td>
<td>—</td>
<td>4</td>
</tr>
<tr>
<td><em>Bipolaris</em> species</td>
<td>1 (2)</td>
<td>—</td>
<td>—</td>
<td>32</td>
</tr>
<tr>
<td><em>Curvularia</em> species</td>
<td>2 (4)</td>
<td>—</td>
<td>—</td>
<td>4</td>
</tr>
</tbody>
</table>

*For the complete baseline characteristics and MICs for all 98 isolates, see previously published data. MIC, minimum inhibitory concentration.

<table>
<thead>
<tr>
<th>Covariate</th>
<th>OR (95% CI)</th>
<th>p Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>MIC (µg/ml) vs natamycin</td>
<td>0.53 (0.32 to 0.86)*</td>
<td>0.01</td>
</tr>
<tr>
<td><em>Fusarium</em> vs <em>Aspergillus</em> species</td>
<td>4.94 (1.17 to 22.3)</td>
<td>0.01</td>
</tr>
</tbody>
</table>

*OR per twofold dilution in MIC.
†Pearson’s goodness of fit: p=0.33.
MIC, minimum inhibitory concentration.

Counselling for people with sight loss in the UK: the need for provision and the need for evidence

For adults of any age the diagnosis of a visual impairment can be traumatic, and timely referral to informal peer support and/or
professional counselling may be both beneficial and appropriate. It is estimated that 45/115 (40%) of UK voluntary organisations for people with sight loss provide professional counselling (n=17) or ‘informal support’ (n=28), such as peer support groups, telephone helplines and befriending. However, what is the evidence that these services help people adjust emotionally to their acquired vision loss and the consequences that flow from this?

During the spring/summer of 2008, we conducted a follow-up to the scoping survey reported earlier to assess the evidence for effectiveness of professional counselling services for people with acquired sight loss. We contacted the 17 counselling services previously identified by Rees and further services via Vision 2020 UK, the National Association of Local Societies for Visually Impaired People, the Visual Impairment Network for Counselling and Emotional Support, and specialist ophthalmic nurses via the Royal College of Nursing. To evaluate the evidence for counselling services we requested the documentation used to support their development and copies of any evaluation reports.

We identified 28 services providing professional counselling in the UK for people with sight loss, 25 of which provided a service to a specific geographical area and three nationally by telephone. Counselling was rarely provided through the NHS or via social services (8/28), and whilst free telephone-based counselling was available across the UK, the provision of free face-to-face counselling was patchy.

Six organisations initiated their services in response to a range of policy and research reports: a policy document outlining the provision of social services for visually impaired adults (n=1) plus a low vision consensus forum document (n=1), a research report by the Royal National Institute of Blind People (RNIB) (n=2), and an evaluation report by the RNIB showing promise for a face-to-face formal counselling service pilot service (n=2). Of the 28 counselling services, three were in the process of being evaluated, 11 had not been evaluated, 11 had collected client satisfaction data that would need to be updated and enhanced with validated scales, and three provided evaluation reports. Two of the three reports were of cross-sectional evaluations using client satisfaction data, but one report found that 100% of clients on completion of face-to-face counselling had reliably and significantly improved in emotional well-being, including a 41% reduction in mild risk of suicide. This pilot could be built upon with trials using more stringent controls of confounding variables and longer-term follow-up.

Our scoping survey was limited in that some services operating in the UK may not have been captured by our recruitment strategy. It is unclear whether these services provided formal counselling as 4/28 services were not provided by qualified counsellors, 16/28 organisations could not detail the training of their counsellors, and 11/28 could not characterise the type of counselling provided (eg humanistic, psychodynamic, etc). Our findings suggest that there is inequity in the provision of free face-to-face counselling to people with vision loss in the UK and that there has been little systematic evaluation of the counselling services available. The RNIB’s pilot counselling services show promise, but their evaluations have yet to recruit control groups or assess long-term outcomes. We call upon researchers to evaluate emotional support services for people with sight loss to provide an evidence-base for their effectiveness in enhancing psychosocial well-being and to inform how these services can be improved. With this evidence voluntary organisations would receive greater recognition and funding for their emotional support services, thereby enhancing the quality of life of people with vision loss.

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REFERENCES


VSX2 in microphthalmia: a novel splice site mutation producing a severe microphthalmia phenotype

Microphthalmia shows great genetic and clinical heterogeneity, whether as part of a syndrome or an isolated ocular phenotype. Chromosomal or single-gene disorders and teratogens may all cause microphthalmia. Associated syndromic features include cardiac problems, clefting, microcephaly and learning disabilities. Microphthalmia is frequently bilateral, but commonly asymmetrical in severity.

Homozygous mutations in VSX2/CHX10 have been demonstrated in human and murine microphthalmia. VSX2 is thought to act principally as a repressor of transcription, particularly of the genes encoding cyclin-dependent kinase inhibitor (p27Kip1) and microphthalmia transcription factor (MITF). These repressive roles enable cell proliferation by preventing retinal progenitor cells from exiting the cell cycle, and by maintaining neuroretinal cell identity. Loss of these functions therefore causes failures in eye development. Other genes implicated in microphthalmia include SOX2, PAX6, sonic hedgehog (SHH), RAX, OTX2, CRYBA and FOXE3. Additional loci with no gene identified include 15q12–q15, 14q32 and Xq27–q28.

Our patient has healthy first-cousin Turkish parents with no ocular anomalies, and an unaffected brother. Her very small eyes were noted at birth, but no other congenital anomalies. Her karyotype demonstrated 46, XX. Cranial MRI (see figure 1) confirmed severe microphthalmia and small optic nerves. Growth and development, given complete absence of vision, have progressed normally to her current age of 3.5 years. Neonatally, the right vestigial eye development, given complete absence of vision, have progressed normally to her current age of 3.5 years. Neonatally, the right vestigial eye remained severely microphthalmic (axial length 12.2 mm) with no useful vision. The cornea remained clear, intraocular pressure was normal (10 mm Hg), but leucocoria suggested a retrorenal plaque. B-scan ultrasound revealed total retinal...
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