Sebaceous carcinoma of the eyelid: a clinicopathological study

Marlis Zürcher, Christoph R Hintschich, Alec Garner, Catie Bunce, J Richard O Collin

Abstract

Background—Sebaceous carcinoma of the eyelid is rare. The diagnosis might be difficult because of its ability to masquerade as other periocular lesions. Prognosis is still regarded as being poor compared with most other malignant eyelid tumours with a mortality second only to malignant melanoma. The present study retrospectively analyses clinical and histopathological findings and outcome in a series of patients with sebaceous carcinoma of the eyelid in Britain.

Methods—43 patients with histologically confirmed sebaceous carcinoma treated at Moorfields Eye Hospital between 1976 and 1992 were subjected to retrospective analysis. Clinical data of all patients were reviewed from the charts; all surviving patients except four cases lost for follow up were re-examined. Histological specimens were reviewed in 41 cases.

Results—23 females and 20 males, mean age 63 years (range 37–79), were treated. Primary therapy was surgery in 37 and radiotherapy in six cases. After a median follow up of 40 months (range 1–148) 30 patients were alive without recurrences, four patients had died from the tumour, and one was alive with local recurrence and distant metastases. Four patients had died of non-tumour related causes. Histologically, unfavourable outcome was correlated with poor tumour differentiation and extensive invasion.

Conclusion—Early diagnosis and consequent surgical therapy of sebaceous carcinoma of the eyelid leads to a better outcome and higher survival rates than generally assumed. Even local recurrences can be treated successfully. However, sebaceous carcinoma remains a threatening disease, which leads to death in 9% and to mutilating exenteration in 23% of our patients.

Sebaceous carcinoma of the eyelid may arise from the diverse sebaceous glands of the ocular adnexa.1 Owing to its rarity and its ability to masquerade as other periocular lesions, diagnosis of the disease might be difficult.5–7 Prognosis is still regarded as being poor compared with most other malignant eyelid tumours with a mortality second only to malignant melanoma.1–5,9–11 Comprehensive studies, however, are few and there is a substantial difference between surveys from America, where the tumour is rare with an incidence ranging from 0.2% to 4.7% of malignant epithelial eyelid tumours,1–5,9–11 and China, where the tumour is reported to occur in 28% of the lid cancers.12

The purpose of our study was to analyse the natural history of the tumour and its response to treatment in a series of patients in Britain.

Patients and methods

From 1976 to 1992, 43 patients with histologically confirmed sebaceous carcinoma were treated at Moorfields Eye Hospital in London. Data on patients’ history, clinical features, therapy, and outcome were collected retrospectively. In addition, all surviving patients except four, who were lost to follow up, were re-examined clinically. In 41 cases the histological specimens were reviewed and assessed. Histological evaluation was based on the study of conventional paraffin embedded sections supplemented in most cases by oil red O stained frozen sections.

The binomial method was used to assess evidence that sebaceous carcinoma is more likely to occur in left versus right eye or upper versus lower lid. We did not conduct any multivariate analyses when studying survival time and time to recurrence, bearing in mind that the power of such tests relates to the number of events not to the number of subjects.11

Results

Characteristics of our study population are presented in Table 1; patient details including sex, race, age at diagnosis, tumour position and size, time between first presentation and treatment, modalities of therapy, complications, and outcome are shown in Table 2.

Patients’ history

Twenty three women and 20 men were treated. Patients’ age at time of correct diagnosis ranged from 38 to 79 years (mean age 62.6 years). Forty two patients were white with two Indian, one Arab, one Maltese, and one Spanish patient, and one patient was Asian from China. The Asian, Maltese, and Spanish patients had lived in their countries of birth for at least 10 years.

Details of the medical history were obtained in 33 patients. One patient had received radiotherapy for facial acne (dose unknown), one had a history of multiple basal cell carcinomas on his back, and one had been treated surgically for an osteosarcoma of the femur 25 years earlier. No patient had been exposed to chemotherapy for any reason before manifestation of the sebaceous carcinoma.

Thirteen of 29 patients interviewed gave a positive family history of cancer with four of them having more than one affected relative.
The tumours included four cases of gastric carcinoma, two of breast carcinoma, and one each of cancer of the skin, brain, lung, prostate, throat, and liver.

**CLINICAL FEATURES**

Twenty-seven of the carcinomas were found on the left, 16 on the right side (p=0.12; binominal test). The tumour was localised in the upper eyelid in 26 patients, and less frequently in the lower eyelid in 14 patients (p=0.08). Both lids were involved in three cases. The conjunctiva was affected in 14 patients, three of them additionally had corneal epithelial changes. At the time of diagnosis two patients showed intraorbital tumour, two had local lymph node metastases, and one had a distant metastasis.

Tumours were graded as small (<10 mm) in 11 cases, of medium size (10–20 mm) in 14 cases, and large (>20 mm) in six cases. In 10 patients, the tumour was diffuse or multifocal, imitating a blepharoconjunctivitis, and could not be measured accurately. In two cases no measurements were available.

The diagnoses were made on histological examinations based on specimens obtained by incisional biopsy in 14 cases (one combined with conjunctival biopsy), full thickness eyelid biopsies in 23 cases (mainly excisional biopsies), and three excisional biopsies which were not further specified. A conjunctival biopsy only was performed in two eyes. In one patient the diagnosis was made from the exenteration specimen.

At the time of referral a correct initial diagnosis had been established in only eight of 43 patients (18.6%). The most frequent clinical misdiagnosis was chalazion (19 patients) (Fig 1), followed by chronic blepharoconjunctivitis (eight patients) (Fig 2), basal cell carcinoma (four patients) (Fig 3), epithelial carcinoma in situ (four patients), and squamous cell carcinoma (three patients). Other isolated misdiagnoses at presentation included leucoplakia, sweat gland carcinoma, and papilloma. More than one misdiagnosis during the course of the disease until the definitive histological diagnosis was made was found in five patients.

The time interval between first documented symptoms and diagnosis ranged from 1 month to 15 years (median 12 months). Only 13 patients (30%) were diagnosed within 6 months of the first symptoms. After the definitive diagnosis had been established, therapy was instituted at a median time of 1 month.

The follow up beginning after the first treatment at Moorfield’s ranged from 1 month to 12.3 years (median 3.3 years). Four patients were lost to follow up after 1, 2, 60, and 87 months, respectively.

**THERAPY**

The primary treatment at Moorfields Eye Hospital was mainly a surgical approach. Thirty-seven patients were operated on, and only six were treated by radiotherapy. Surgery consisted of extensive excision with a macroscopically tumour free margin of at least 4 mm. The surgical margin was evaluated histologically by frozen section in 10 patients and by Mohs’ technique in one patient. Primary orbital exenteration had to be performed in eight patients.

**OUTCOME**

At the end of the survey 30 patients were alive without detectable tumour recurrences. One patient was alive but had a local recurrence and distant metastases. Eight patients had died during the period of follow up. In four there was a tumour related cause. One patient died from metastatic disease of her eyelid tumour (no 29), and three from causes related to their lid carcinoma, but which were not proved by necropsy (patient nos 19, 31, and 41). In the other four cases death was due to broncho-genic carcinoma in two and to unknown causes in the other two. All four patients who died from sebaceous carcinoma had received palliative radiotherapy as primary treatment. The two patients dying from unknown causes had undergone orbital exenteration. None of the patients treated surgically died from tumour related causes.

Four patients were lost to follow up—two immediately after the treatment, one after 5 years, and another after 7 years of recurrence-free follow up.

Twelve patients (28%) suffered from either local recurrences and/or distant metastases. Six (14%) of them developed only a local tumour recurrence after 2 to 48 months (median 9 months). Preauricular lymph node metastases were seen in six patients 6–24 months (median 14 months) after initial treatment. Three patients showed both local recurrence and preauricular lymph node involvement. One of these patients (no 26) developed histologically confirmed distant subcutaneous metastases as well.

Local tumour recurrences were managed by repeated local excision in six and orbital exenteration in two patients. Preauricular lymph node metastases were treated by radical neck dissection, usually combined with partial...
Table 2  Clinical features of all 43 patients

<table>
<thead>
<tr>
<th>Patient No</th>
<th>Sex</th>
<th>Race</th>
<th>Age at presentation (months)</th>
<th>Tumour size (mm)</th>
<th>Tumour position</th>
<th>Time: first symptom to diagnosis (months)</th>
<th>Time: diagnosis to therapy (months)</th>
<th>First therapy</th>
<th>Complications</th>
<th>Time: therapy to recurrence (months)</th>
<th>Therapy of recurrence</th>
<th>Follow-up (months)</th>
<th>Death</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>F</td>
<td>W</td>
<td>64</td>
<td>5-10 (small)</td>
<td>L, upper lid</td>
<td>30</td>
<td>1</td>
<td>Excision (wide)</td>
<td>?</td>
<td>18</td>
<td>Excision (local), neck dissection</td>
<td>2</td>
<td></td>
</tr>
<tr>
<td>2</td>
<td>F</td>
<td>W</td>
<td>55</td>
<td>5-10 (small)</td>
<td>L, lower lid, conj/cornea</td>
<td>21</td>
<td>1</td>
<td>Excision (wide)</td>
<td>?</td>
<td>2</td>
<td></td>
<td>82</td>
<td></td>
</tr>
<tr>
<td>3</td>
<td>M</td>
<td>W</td>
<td>63</td>
<td>5-10 (small)</td>
<td>L, lower lid</td>
<td>9</td>
<td>2</td>
<td>Excision (wide)</td>
<td>?</td>
<td>87</td>
<td></td>
<td>87</td>
<td></td>
</tr>
<tr>
<td>4†</td>
<td>F</td>
<td>W</td>
<td>37</td>
<td>5-10 (small)</td>
<td>L, upper lid</td>
<td>12</td>
<td>1</td>
<td>Excision (wide)</td>
<td>?</td>
<td>2</td>
<td></td>
<td>74</td>
<td></td>
</tr>
<tr>
<td>5</td>
<td>M</td>
<td>W</td>
<td>79</td>
<td>5-10 (small)</td>
<td>R, upper lid</td>
<td>29</td>
<td>2</td>
<td>Excision with Mohs' technique</td>
<td>?</td>
<td>2</td>
<td></td>
<td>2</td>
<td></td>
</tr>
<tr>
<td>6</td>
<td>F</td>
<td>W</td>
<td>62</td>
<td>5-10 (small)</td>
<td>R, upper lid</td>
<td>50</td>
<td>0</td>
<td>Excision (wide)</td>
<td>?</td>
<td>113</td>
<td></td>
<td>1</td>
<td></td>
</tr>
<tr>
<td>7</td>
<td>M</td>
<td>W</td>
<td>62</td>
<td>5-10 (small)</td>
<td>L, lower lid</td>
<td>1</td>
<td>1</td>
<td>Excision with frozen section</td>
<td>?</td>
<td>1</td>
<td></td>
<td>17</td>
<td></td>
</tr>
<tr>
<td>8†</td>
<td>F</td>
<td>W</td>
<td>44</td>
<td>5-10 (small)</td>
<td>L, lower lid</td>
<td>18</td>
<td>0</td>
<td>Excision (wide)</td>
<td>Tumour recurrence local, metastases</td>
<td>18</td>
<td>Excision (local), neck dissection</td>
<td>87</td>
<td></td>
</tr>
<tr>
<td>9†</td>
<td>M</td>
<td>W</td>
<td>77</td>
<td>5-10 (small)</td>
<td>L, upper lid</td>
<td>6</td>
<td>2</td>
<td>Excision (wide)</td>
<td>Tumour recurrence local</td>
<td>2</td>
<td>Excision (local)</td>
<td>135</td>
<td></td>
</tr>
<tr>
<td>10†</td>
<td>M</td>
<td>W</td>
<td>73</td>
<td>5-10 (small)</td>
<td>L, lower lid</td>
<td>60</td>
<td>9</td>
<td>Radiotherapy</td>
<td>?</td>
<td>1</td>
<td></td>
<td>1</td>
<td></td>
</tr>
<tr>
<td>11</td>
<td>M</td>
<td>Ch</td>
<td>50</td>
<td>5-10 (small)</td>
<td>L, upper lid</td>
<td>2</td>
<td>1</td>
<td>Excision with frozen section</td>
<td>?</td>
<td>2</td>
<td></td>
<td>27</td>
<td></td>
</tr>
<tr>
<td>12†</td>
<td>F</td>
<td>W</td>
<td>68</td>
<td>10-20 (intermediate)</td>
<td>L, upper lid</td>
<td>12</td>
<td>1</td>
<td>Radiotherapy</td>
<td>Tumour recurrence local</td>
<td>48</td>
<td>Excision (local)</td>
<td>52</td>
<td></td>
</tr>
<tr>
<td>13</td>
<td>M</td>
<td>W</td>
<td>63</td>
<td>10-20 (intermediate)</td>
<td>L, lower lid, conj/cornea</td>
<td>9</td>
<td>0</td>
<td>Excision (wide)</td>
<td>Tumour recurrence local</td>
<td>48</td>
<td>Excision (local)</td>
<td>148</td>
<td></td>
</tr>
<tr>
<td>14</td>
<td>F</td>
<td>W</td>
<td>66</td>
<td>10-20 (intermediate)</td>
<td>L, upper lid, conj/cornea</td>
<td>48</td>
<td>0</td>
<td>Excision with frozen section</td>
<td>?</td>
<td>1</td>
<td></td>
<td>29</td>
<td></td>
</tr>
<tr>
<td>15</td>
<td>M</td>
<td>W</td>
<td>67</td>
<td>10-20 (intermediate)</td>
<td>L, upper lid</td>
<td>1</td>
<td>0</td>
<td>Excision (wide)</td>
<td>Tumour recurrence local</td>
<td>6</td>
<td>Excision (local)</td>
<td>45</td>
<td></td>
</tr>
<tr>
<td>16</td>
<td>F</td>
<td>W</td>
<td>64</td>
<td>10-20 (intermediate)</td>
<td>L, lower lid</td>
<td>6</td>
<td>0</td>
<td>Excision (wide)</td>
<td>Tumour recurrence local</td>
<td>6</td>
<td></td>
<td>18</td>
<td></td>
</tr>
<tr>
<td>17</td>
<td>M</td>
<td>W</td>
<td>47</td>
<td>10-20 (intermediate)</td>
<td>L, lower lid</td>
<td>6</td>
<td>0</td>
<td>Excision with frozen section</td>
<td>?</td>
<td>50</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>18</td>
<td>M</td>
<td>W</td>
<td>71</td>
<td>10-20 (intermediate)</td>
<td>L, upper lid</td>
<td>48</td>
<td>0</td>
<td>Excision with frozen section</td>
<td>?</td>
<td>50</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>19</td>
<td>F</td>
<td>W</td>
<td>63</td>
<td>10-20 (intermediate)</td>
<td>L, lower lid, conj/cornea</td>
<td>14</td>
<td>1</td>
<td>Radiotherapy</td>
<td>Tumour recurrence local</td>
<td>6</td>
<td></td>
<td>103</td>
<td></td>
</tr>
<tr>
<td>20</td>
<td>M</td>
<td>W</td>
<td>65</td>
<td>10-20 (intermediate)</td>
<td>L, upper lid</td>
<td>1</td>
<td>1</td>
<td>Excision with frozen section</td>
<td>?</td>
<td>38</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>21</td>
<td>M</td>
<td>W</td>
<td>61</td>
<td>10-20 (intermediate)</td>
<td>L, upper lid, conj/cornea</td>
<td>10</td>
<td>0</td>
<td>Excision (wide)</td>
<td>Tumour recurrence local</td>
<td>6</td>
<td></td>
<td>134</td>
<td></td>
</tr>
<tr>
<td>22</td>
<td>M</td>
<td>I</td>
<td>66</td>
<td>10-20 (intermediate)</td>
<td>L, upper lid</td>
<td>12</td>
<td>1</td>
<td>Excision (wide)</td>
<td>?</td>
<td>62</td>
<td></td>
<td>62</td>
<td></td>
</tr>
<tr>
<td>23</td>
<td>M</td>
<td>I</td>
<td>78</td>
<td>10-20 (intermediate)</td>
<td>R, lower lid</td>
<td>180</td>
<td>0</td>
<td>Excision (wide)</td>
<td>?</td>
<td>52</td>
<td></td>
<td>52</td>
<td></td>
</tr>
<tr>
<td>24</td>
<td>F</td>
<td>W</td>
<td>38</td>
<td>10-20 (intermediate)</td>
<td>L, upper lid</td>
<td>12</td>
<td>2</td>
<td>Excision (wide)</td>
<td>?</td>
<td>91</td>
<td></td>
<td>91</td>
<td></td>
</tr>
<tr>
<td>25</td>
<td>M</td>
<td>W</td>
<td>67</td>
<td>20-30 (large)</td>
<td>L, upper lid</td>
<td>1</td>
<td>3</td>
<td>Excision with frozen section</td>
<td>?</td>
<td>32</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>26</td>
<td>M</td>
<td>W</td>
<td>75</td>
<td>20-30 (large)</td>
<td>L, upper lid, conj/cornea</td>
<td>6</td>
<td>1</td>
<td>Excision (wide)</td>
<td>Tumour recurrence local, metastases</td>
<td>6</td>
<td>Exenteration, radiotherapy</td>
<td>17</td>
<td></td>
</tr>
<tr>
<td>27</td>
<td>F</td>
<td>W</td>
<td>71</td>
<td>20-30 (large)</td>
<td>L, upper lid</td>
<td>17</td>
<td>3</td>
<td>Excision with frozen section</td>
<td>Metastases</td>
<td>12</td>
<td>Neck dissection</td>
<td>15</td>
<td></td>
</tr>
<tr>
<td>28</td>
<td>M</td>
<td>W</td>
<td>62</td>
<td>20-30 (large)</td>
<td>L, upper lid, conj/cornea</td>
<td>1</td>
<td>1</td>
<td>Exenteration</td>
<td>?</td>
<td>16</td>
<td></td>
<td>16</td>
<td></td>
</tr>
<tr>
<td>29†</td>
<td>F</td>
<td>W</td>
<td>77</td>
<td>20-30 (large)</td>
<td>R, lower lid</td>
<td>5</td>
<td>2</td>
<td>Radiation therapy</td>
<td>?</td>
<td>16</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>30</td>
<td>M</td>
<td>W</td>
<td>57</td>
<td>20-30 (large)</td>
<td>L, upper lid</td>
<td>9</td>
<td>1</td>
<td>Excision with frozen section</td>
<td>?</td>
<td>4</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>31</td>
<td>F</td>
<td>W</td>
<td>76</td>
<td>Diffuse</td>
<td>R, upper lid, lower lid, conj/cornea, diffuse</td>
<td>19</td>
<td>2</td>
<td></td>
<td>a</td>
<td>2</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>32</td>
<td>F</td>
<td>W</td>
<td>60</td>
<td>Diffuse</td>
<td>R, upper lid, conj/cornea</td>
<td>14</td>
<td>2</td>
<td>Exenteration</td>
<td></td>
<td>12</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>33</td>
<td>F</td>
<td>W</td>
<td>53</td>
<td>Diffuse</td>
<td>L, upper lid</td>
<td>7</td>
<td>1</td>
<td>Excision (wide)</td>
<td>Metastases</td>
<td>10</td>
<td>Neck dissection, radiotherapy</td>
<td>76</td>
<td></td>
</tr>
<tr>
<td>34†</td>
<td>F</td>
<td>W</td>
<td>66</td>
<td>Diffuse</td>
<td>R, upper lid, lower lid, orbit</td>
<td>1</td>
<td>1</td>
<td>Exenteration</td>
<td></td>
<td>4</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>35</td>
<td>F</td>
<td>W</td>
<td>52</td>
<td>Diffuse</td>
<td>L, upper lid, conj/cornea</td>
<td>14</td>
<td>6</td>
<td>Exenteration</td>
<td></td>
<td>2</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>36</td>
<td>M</td>
<td>W</td>
<td>68</td>
<td>Diffuse</td>
<td>R, upper lid, conj/cornea</td>
<td>18</td>
<td>1</td>
<td>Exenteration</td>
<td></td>
<td>27</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>37</td>
<td>M</td>
<td>W</td>
<td>78</td>
<td>Diffuse</td>
<td>L, upper lid</td>
<td>30</td>
<td>1</td>
<td>Excision (wide)</td>
<td>Tumour recurrence local</td>
<td>12</td>
<td>Radiotherapy</td>
<td>24</td>
<td></td>
</tr>
<tr>
<td>38</td>
<td>F</td>
<td>W</td>
<td>47</td>
<td>Diffuse</td>
<td>L, upper lid, lower lid, conj/cornea, orbit</td>
<td>60</td>
<td>0</td>
<td>Excision (wide)</td>
<td></td>
<td>70</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>39</td>
<td>F</td>
<td>W</td>
<td>67</td>
<td>Diffuse</td>
<td>L, upper lid, conj/cornea</td>
<td>12</td>
<td>2</td>
<td>Exenteration</td>
<td>Tumour recurrence local, metastases</td>
<td>15</td>
<td>Neck dissection</td>
<td>132</td>
<td></td>
</tr>
<tr>
<td>40</td>
<td>F</td>
<td>W</td>
<td>59</td>
<td>Diffuse</td>
<td>R, upper lid, conj/cornea</td>
<td>72</td>
<td>2</td>
<td>Exenteration</td>
<td></td>
<td>12</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>41†</td>
<td>M</td>
<td>W</td>
<td>58</td>
<td>?</td>
<td>L, lower lid</td>
<td>5</td>
<td>1</td>
<td>Radiotherapy</td>
<td></td>
<td>2</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>42†</td>
<td>M</td>
<td>W</td>
<td>62</td>
<td>?</td>
<td>L, lower lid</td>
<td>7</td>
<td>2</td>
<td>Exenteration</td>
<td>Tumour recurrence local, metastases</td>
<td>?</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>43</td>
<td>F</td>
<td>W</td>
<td>53</td>
<td>?</td>
<td>L, lower lid</td>
<td>36</td>
<td>0</td>
<td>Excision (wide)</td>
<td>Tumour recurrence local, metastases</td>
<td>16</td>
<td>Excision (local), neck dissection</td>
<td>60</td>
<td></td>
</tr>
</tbody>
</table>

W = white; Ch = Chinese; I = Indian; ? = unknown; a = patient died before first therapy; *lymph nodes at first presentation; †correct diagnosis at presentation; ‡death, tumour related; §death, other causes.
parotidectomy. In one case, local recurrence with lymph node metastasis was treated by excision and radiotherapy of the face and neck. A subjective assessment of the functional and cosmetic outcome was carried out in 27 patients excluding the patients subjected to orbital exenteration. The functional result was regarded as good in 19, moderate in six, and poor in two patients, whereas the cosmetic outcome was regarded as good in 23, moderate in three, and poor in one patient.

**HISTOPATHOLOGY**

A review of the histopathological specimens was possible in 41 of 43 cases. The features which were evaluated include pattern, differentiation, stromal invasion, epithelial involvement, and mitotic activity.

**Pattern**

A lobular arrangement of cells was apparent in 36 cases (Fig 4). One tumour had a predominantly papillary pattern and four were mixed. There was no example of a pure form of comedo carcinoma.

**Differentiation**

On the basis of the proportion of tumour cells which had abundant vacuolated or foamy cytoplasm and were comparable in appearance to normal sebaceous tissue, individual lesions were categorised as well differentiated in 12 cases (Fig 4), moderately differentiated in 22 and poorly differentiated in seven cases (Fig 5).

**Invasion**

Tumours which showed generally circumscribed lobules and little evidence of stromal infiltration accounted for nine cases (Fig 4). A moderate stromal invasion but with preservation of the lobular structure was observed in 21 specimens. In the remaining 11 cases a more extensive infiltration with a break up of the lobules was observed (Fig 5).

**Epithelial involvement**

Pagetoid spread within the adjacent surface epithelium was observed in 16 cases in the form of isolated or, more commonly, groups of tumour cells infiltrating and replacing the basal layers (Fig 6). In some instances, where the infiltration was extensive, this was associated with degeneration and acantholysis of the overlying epidermis and/or conjunctival epithelium. In 11 cases there was full thickness replacement of the surface epithelium by tumour tissue (Fig 7) in the absence of ulceration resembling epithelial carcinoma in situ (Bowen’s disease). In one case areas of both pagetoid and carcinoma in situ changes were visible. In the remaining 14 cases there was no apparent involvement of the surface epithelium.

**Mitotic activity**

Semiquantitative assessments based on the number of mitotic figures identified per high power microscopic field (×250) gave a range of 1–12 and a mean of 5.8.

**Discussion**

Although carcinoma of the meibomian glands was described more than a century ago by Fuchs, subsequent reports have been infrequent and usually limited to single cases. In 1956, Straatsma published the first extensive series, which did much to clarify the natural history and the prognosis of sebaceous carcinoma of the eyelid. Since then several other studies have appeared. Comprehensive reviews are available.
Sebaceous carcinoma is an uncommon skin tumour which accounts for less than 1% of malignancies of the skin. It occurs most frequently on the eyelids, where it comprises 4.7% of malignant epithelial tumours. The incidence on the eyelid is subject to considerable geographical variation. Among white people the tumour is rare and, according to reports from the USA, represents between 0.2% and 1.2% of all lid lesions and between 1.13% and 3.2% of all malignant lid neoplasms. In China and other Asian countries forming the western Pacific seaboard, the incidence appears to be much higher; one study from Shanghai gave an incidence of 32.7% of all eyelid malignancies. Since in our study there was only one Asian patient, who had lived for at least 10 years in the country of birth, we cannot give a statement concerning the possible influence of racial factors in the propensity to develop sebaceous carcinoma.

In the literature a higher incidence for females is suggested with a female to male ratio of 1.51 to 1.0. In our study as in some others, there was no significant sex distinction.

The median age of our patients at diagnosis was 63 years, which is similar to other studies (means between 57 and 68 years). Sebaceous carcinoma of the eyelid in younger patients is apparently a rare event and, not infrequently, appears to be associated with prior radiotherapy. Neither of the two patients in our series aged under 40 had been exposed in this way.

The tumour originates in the upper lid more often than in the lower lid. The ratio varies from 1.3 to 3.0 in the published series. In our series, the ratio of upper to lower lid was 1.8 (including three cases with involvement of both lids). These data provide some evidence (p = 0.08); predominance of the upper eyelid possibly reflects the greater number of meibomian glands in the upper eyelid.

The clinical diagnosis of sebaceous carcinoma may be difficult, partly because it is rarely encountered and partly because of its propensity to simulate other eyelid lesions. The present paper emphasises this difficulty. Chalazion, followed by blepharoconjunctivitis, was the most common misdiagnosis, with other benign and malignant neoplasms also causing significant problems. In exceptional patients the presenting sign is enlarged cervical lymph nodes, the usual primary lid lesion being inconspicuous and easily overlooked. Two of our patients presented with preauricular lymph node metastases at the time of diagnosis.

These diagnostic difficulties must be accepted and emphasise the importance of biopsies in uncertain and suspicious inflammatory
states which fail to respond to appropriate treatment. A full thickness eyelid biopsy, combined with conjunctival biopsies in certain cases, is the appropriate approach. The importance of histological examination of all suspicious chalazia must also be stressed. This is not to claim that histopathological diagnosis is always straightforward: in one of our patients, presenting as an ostensible case of blepharitis, the presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.

It might be anticipated that the sooner appropriate treatment is instituted the better the prognosis and a number of published reports bear this out, although our own data are inconclusive.

Numerous factors have been reported to influence the prognosis. Because of the small number of tumour deaths in our series one cannot really comment much on this. However, tumours in excess of 10 mm are associated with a particularly poor outcome; the patients in our current case of blepharoconjunctivitis, its presence of minor pagetoid spread in a biopsy specimen was initially overlooked contributing to a delay in diagnosis from the first symptoms of 60 months. Nevertheless, after review of the histological specimens the diagnoses on all biopsies taken at Moorfields Eye Hospital proved to be correct.
Sebaceous carcinoma of the eyelid

This paper was presented in part at the Royal College of Ophthalmologists Meeting 1994 in Guernsey, UK. This study was supported by the Verrey-Foundation, Lausanne, Switzerland.


