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Surveillance of severe chemical corneal injuries in the UK

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1 **Title: Surveillance of Severe Chemical Corneal Injuries in the UK**

2
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21
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23 Injuries, Limbal Stem Cell Deficiency, Surveillance

24
25 **Word Count of Paper 2486**

27 **ABSTRACT**

28

29 **Aim**

30 To estimate the incidence of severe chemical corneal injuries in the United
31 Kingdom and describe presenting clinical features and initial management.

32

33 **Methods**

34 All patients with severe chemical corneal injury in the UK from December
35 2005 to November 2006 inclusive were prospectively identified using the
36 British Ophthalmological Surveillance Unit. Reporting ophthalmologists
37 provided information regarding presentation and follow up.

38

39 **Results**

40 Twelve cases were identified, giving a minimum estimated incidence in the UK
41 of severe chemical corneal injury of 0.02 per 100 000. 66.7% of injuries were
42 in males of working age, 50% occurred at work, alkali was causative in 66.7%.
43 Only one patient was wearing eye protection at the time of injury, 75%
44 received immediate irrigation. Six patients required 1 or more surgical
45 procedures; most commonly amniotic membrane graft. At 6 months follow-up
46 best corrected visual acuity was 6/12 or better in five patients, and worse than
47 6/60 in two.

48

49 **Conclusion**

50 The incidence of severe chemical corneal injury in the UK is low. The cases
51 that occur can require extended hospital treatment, with substantial ocular

52 morbidity and visual sequelae. Current enforcement of eye protection in the
53 workplace in the UK has probabaly contributed to a reduced incidence of
54 severe ocular burns.

55

56 **Abstract Word Count 200**

57

58 **INTRODUCTION**

59

60 Chemical injuries to the eye represent an ophthalmic emergency that can
61 result in extensive damage with significant ocular morbidity.¹ Most are mild
62 and accidental and do not result in lasting ocular morbidity. Severe chemical
63 eye injuries are rare, but can cause significant visual impairment.¹

64

65 The extent of ocular injury depends on many factors, including the strength of
66 the chemical agent, duration of exposure, concentration, volume and
67 penetration of the solution.^{2,3} Severe chemical injury to the cornea is usually
68 secondary to strongly acidic and alkaline substances.⁴ Alkali injuries more
69 commonly cause significant damage.⁴ Epidemiological data shows severe
70 chemical eye injuries are more common in males, particularly those aged
71 between 16 and 45 yrs.^{1,5,6,7} Most happen accidentally at work or at home, or
72 deliberately from assault.⁴

73

74 Various classification scales exist to grade the severity of ocular chemical
75 injuries. The Hughes classification, modified by Ballen and Roper-Hall,
76 recognised the relationship between initial appearance and prognosis.¹
77 Subsequent classification schemes have been developed, but the Hughes-
78 Roper-Hall classification remains simple and popular.^{8,9} In the acute stage it
79 describes the clinical signs and severity in 4 grades. Grade 1 injuries show
80 corneal epithelial damage, with no limbal ischaemia and a clear cornea.
81 Grade 2 injuries show less than a third of limbal ischaemia and a hazy cornea
82 through which iris details can be seen. Grade 3 involves total loss of corneal

83 epithelium, stromal haze obscuring iris details, and ischaemia between one
84 third and one half of the limbus. Grade 4 injuries represent an opaque cornea
85 and ischaemia of more than half of the limbus. Grades 3 and 4 are recognised
86 as severe.¹

87

88 Acute chemical eye injury treated immediately with expedient irrigation and
89 removal of trapped debris is associated with a significantly better visual
90 outcome.⁴ Early management endeavors to preserve the globe integrity by
91 healing of the ocular surface. Treatment is aimed at promoting ocular surface
92 epithelial recovery, augmenting corneal repair, minimising ulceration and
93 controlling the inflammatory response.¹ Surgery may be necessary in the
94 acute setting if healing of the ocular surface is inadequate. In the chronic
95 stages, features of limbal stem cell deficiency can manifest.¹⁰ Long-term
96 management aims to restore the visual function by preserving tear production,
97 managing limbal stem cell deficiency and addressing associated
98 complications such as lid malposition, cataract and glaucoma.^{10,11,12}

99

100 The UK national incidence of severe chemical corneal injuries is unclear. Two
101 previous studies in UK centers have reported eight severe chemical corneal
102 injuries presenting to a large teaching hospital over a four year period and
103 none to a district general hospital over a year.^{5,7} Based on this limited
104 evidence we anticipated a maximum of 200-300 cases annually in the UK.
105 This study aimed to estimate the true incidence of severe chemical corneal
106 injuries with a UK wide prospective survey. In addition it aimed to outline the
107 presentation and management of these cases.

108 **METHODS**

109

110 This was a prospective population-based study performed in association with
111 the British Ophthalmological Surveillance Unit (BOSU) monthly reporting
112 system as previously described.^{13,14} All ophthalmologists with clinical
113 autonomy in the UK receive a monthly reporting card with definitions of the
114 conditions currently under surveillance. They indicate how many new cases of
115 each disorder they have seen or confirm they have seen no new cases. For
116 the 12-month study period December 2005 to November 2006 inclusive,
117 ophthalmologists were asked to report any case of severe chemical corneal
118 injuries presenting to them. This was defined as any person sustaining a injury
119 with all of the following:

- 120 1. Total loss of the corneal epithelium, and
- 121 2. Corneal haziness obscuring iris detail or worse, and
- 122 3. Over 120° of limbal ischaemia

123 This equated to Grade 3 or 4 on the Hughes-Roper-Hall classification scale.

124

125 Every ophthalmologist who notified a patient to the BOSU was sent an initial
126 questionnaire requesting information regarding demographics, aetiology,
127 protective eyewear worn, presenting features and initial management. A 6
128 month follow-up questionnaire was also sent to ascertain subsequent
129 management and outcome. Ophthalmologists who did not return a
130 questionnaire were sent a reminder letter.

131

132 Multi-Centre Research Ethics approval was obtained (REC reference number
133 05/SO801/111).

134

135 **RESULTS**

136

137 **Reports to BOSU leading to estimates of the UK incidence of severe** 138 **chemical corneal injury.**

139 During the study period the BOSU used a reporting base of 1100
140 ophthalmologists. Each month a mean of 78% of these ophthalmologists
141 provided surveillance information (range 75% - 81%). From December 2005
142 to November 2006 inclusive BOSU received 37 reports of patients with severe
143 chemical corneal injury from 26 ophthalmology consultants. Initial
144 questionnaires were received from 31/ 37 reports (83.8% response rate). Of
145 these 19 were excluded (9 did not meet the inclusion criteria, 5 occurred
146 outwith the study period, 2 were reporting errors and 3 were duplicate cases).
147 Initial questionnaire data was therefore received in 12 true reported cases
148 within the study period. Follow up questionnaires were returned in 10/ 12
149 (83.3%) cases.

150

151 The current UK population is estimated to be approximately 60 million¹⁵. The
152 reported annual incidence of severe chemical corneal injury is therefore
153 estimated by this study at 12 per 60 million or 0.02 per 100 000. Eight of the
154 cases occurred in males of working age (16-65yrs), for whom the current UK
155 population estimate is 20 million¹⁵. From this study the estimated incidence of
156 severe chemical corneal injury in this group is therefore 8 per 20 million or
157 0.04 per 100 000.

158 **Patient characteristics and presenting details**

159 Initial details received on the 12 patients showed nine were male (75%). The
160 mean age at time of injury was 33.8 yrs (median 38.5yrs, range 10-59 yrs).
161 Eight were males of working age, 66.7%. The injury occurred during paid
162 employment in six cases (50%), following assault in 4 (33.3%) and during
163 another activity in two. One patient had a bilateral injury. The causative
164 chemical was reported as alkaline in 8 (66.7%), acidic in 2 (16.7%), other in
165 one and unknown in one. A liquid splash was the mechanism of injury in eight
166 (66.7%), debris was causative in three (25%), there was one reported injury
167 from a gas substance. One patient was wearing goggles at the time of injury,
168 11 (91.7%) were wearing no eye protection (including prescription
169 spectacles).

170
171 The time from injury to first irrigation was less than 1hr in 9 (75%) cases. One
172 case was irrigated at 1-3hrs and one at greater than 12 hrs after the injury (the
173 time to irrigation remained unknown in one patient). Review by an
174 ophthalmologist occurred in two (16.7%) patients within 1 hour of the injury, in
175 five (41.7%) cases between 1-3 hrs and in two eyes at 3-12hrs. Initial review
176 by an ophthalmologist was greater than 12 hrs after the injury in two cases
177 and unknown in one patient. Nine (75%) had received immediate irrigation at
178 the time of injury (including 5 of the 6 cases occurring in the workplace), seven
179 were irrigated by the primary care department. In one case the initial irrigation
180 was delayed until review by an ophthalmologist, unfortunately one of the
181 cases taking greater than 12 hrs to present. Ten eyes were re-irrigated by
182 ophthalmology. Six underwent eyelid eversion with removal of debris as

183 appropriate. Table 1 shows best corrected visual acuity (BCVA) in the
184 affected eye at initial presentation. In a quarter this was worse than 6/60,
185 although one third retained a BCVA of 6/12 or better.

186
187

188 Table 1. The Best Corrected Visual Acuity (BCVA) for Reported Patients with
189 a Severe Chemical Corneal Injury at Presentation Compared to 6 months
190 Following Injury

191

Patient	BCVA at Presentation	BCVA 6 mths Following Injury
1	6/9	6/12
2	CF	Not available
3	6/9	6/9
4	HM	Not available
5	6/18	6/18
6	6/60	6/60
7	6/18	6/5
8	6/12	6/6
9	6/9	6/36
10	POL	CF
11	6/18	6/12
12	6/24	POL

192

193

194 Details of the presenting clinical features in the 12 eyes is illustrated in Figure
195 1. For inclusion corneal epithelial loss was total in all patients, 58.3% had 240-
196 360° of limbal ischaemia and two-thirds showed 50-100% of conjunctival
197 epithelial loss. Four were referred from the presenting unit for a specialist
198 corneal and external eye disease opinion.

199

200 **Management by ophthalmology within the first 10 days**

201 Table 2 summarises the non-surgical initial management of these cases.

202

203 Table 2. The Initial Non-Surgical Management Reported in Patients with a
204 Severe Chemical Corneal Injury

Non- Surgical Treatment		Number of patients (n=12)
Topical	Steroid preserved	6
	Steroid unpreserved	6
	Antibiotic preserved	6
	Antibiotic unpreserved	6
	Citrate	3
	Ascorbate	6
	Anti- glaucoma	2
	Dilatation	8
Systemic	Vitamin C	9
	Tetracycline derivatives	4

205 All eyes received topical steroid and antibiotic either preserved or
206 unpreserved. Topical ascorbate and/or systemic vitamin C was used in
207 83.3%, five received both. Five eyes received surgical treatment in the first 10
208 days. Two underwent division of symblepharon with a glass rod, two had an
209 amniotic membrane graft (AMG). One eye was managed with debridement of
210 the conjunctival tissue encroaching on the cornea.

211

212 **Follow Up and Management by ophthalmology 10 days- 6mths.**

213 Follow up data from the 6 mth questionnaire was received in 10 patients. The
214 BCVA at final follow up is shown in Table 1. Half of patients had a BCVA of
215 6/12 or better 6 mths post injury. In 20% BCVA was worse than 6/60. Figure 2
216 shows the recorded complications during the six months follow injury. Central
217 corneal pannus or scar was the most frequent problem in 70%. In two cases 6
218 mth BCVA was 6/5. In the remainder the reason for the reduction in vision
219 was reported as being secondary to the associated corneal pathology. Table 3
220 summarises the non-surgical and surgical management modalities used
221 between 10 days and final follow up. AMG was used in 50% of eyes between
222 10 days and 6 months. Limbal stem cell graft was performed in 20%. In total
223 three (30%) eyes had required two surgical procedures at final follow up, three
224 had required one.

225

226 Table 3. The Non-Surgical and Surgical Management Reported from 10 days
227 Following Injury to Final Follow-Up at 6 months in Patients with a Severe
228 Chemical Corneal Injury.

Management	Number of patients (n=12)
Artificial tears	6
Puncal occlusion	1
Therapeutic contact lens	5
Botulinum toxin ptosis	2
Tarsorrhaphy	0
AMG	5
Limbal Stem Cell Graft	2
Oral Mucosal Graft	1
Fornix Reconstruction	1

229

230

231 **Clinical Details of the Patients Excluded who Met the Inclusion Criteria**
 232 **but Occurred Outwith the Study Period.**

233 Five patients were excluded from analysis as the reported injury had occurred
 234 out with the study period. Their injury had occurred 1 mth- 2yrs prior to the
 235 study period. All were males of working age, 4 injuries occurred during paid
 236 employment, 1 following assault. None were wearing eye protection at the
 237 time of injury, 4 received immediate irrigation. The BCVA at presentation
 238 ranged from 6/12 to 6/60, 1 had 240° –360° limbal ischaemia, 2 had 50-100%
 239 conjunctival loss. All received initial treatment with topical steroid or antibiotic,
 240 3 received topical ascorbate and/or systemic vitamin C. An AMG was used to
 241 treat 1 patient, 2 received a limbal stem cell graft. Six month BCVA was 6/6 or

242 6/9 in all patients except for one with a BCVA of hand-movements. Corneal
243 pathology was the reported cause of reduced vision.

244

245

246 **DISCUSSION**

247

248 This study estimates the UK annual incidence rate of severe chemical corneal
249 injury is 0.02 per 100 000. This incidence is low. A degree of under-
250 ascertainment is a feature of studies of this nature, however, active
251 surveillance as practiced by the BOSU has been shown to be more effective
252 than other methods.^{16,17} Similar to previous surveillance studies the mean
253 card return rate was high (78%) and questionnaire response rates were above
254 80%. Furthermore, the BOSU is generally well supported by reporting
255 ophthalmologists, with previous estimated ascertainment rates of 75-95%.¹³
256 Based upon response rates it is safe to assume that ascertainment in this study
257 is likely to be similar to previous surveillance studies using the BOSU
258 reporting system. The incidence rate reported is a minimum rate and it is
259 possible that the true incidence may be up to 25% greater than this. Under-
260 reporting is usually attributed to random error (eg forgetting to report a case,
261 misunderstanding the case definition), reluctance to participate, or
262 management of cases by ophthalmologists who do not receive BOSU
263 reporting cards^{13,14} rather than systematic error which would bias the
264 representative nature of this population based co-hort. Even accounting for
265 any potential under-ascertainment the reported incidence is lower than might

266 be anticipated from the limited previous data^{5,7} and probably reflects a true
267 reduction in the UK incidence of severe chemical corneal injury

268

269 Due to the small number of reported cases in this study it is difficult to come to
270 firm conclusions regarding the current presentation and management of
271 severe chemical corneal injury in the UK. In accordance with epidemiological
272 data from other studies, this survey confirmed that severe chemical corneal
273 injury is seen more commonly in men (75% of cases) mainly of working age
274 (66.7%), with the injury most often occurring at work (50%) or following
275 assault (33.3%) and an alkaline substance most commonly implicated.
276 Previous studies have show severe chemical eye injuries are more common
277 in males between 16 and 45 yrs, occurring accidentally at work or at home, or
278 as the result of deliberate assault.^{1,4,5,6} A retrospective study of 221 chemical
279 injuries by Morgan et al ⁵ in the UK showed alkali injuries were twice as
280 common as acid injuries and men were affected 75.6% of the time. In his
281 study, 63% of injuries occurred in the workplace and 33% at home, 10.6%
282 were secondary to assault. None of their reported cases were classed as
283 severe. Kucklehorn published a subsequent series of 236 eyes from Germany
284 in which 70% of chemical injuries occurred in males between aged 16-45yrs.⁶
285 Industrial accidents accounted for 61% of injuries and household accidents
286 37%. A second series was limited to severe chemical injuries and showed
287 these most often resulted from industrial accidents in males, aged 20-40
288 years.⁶

289

290 Only one patient was wearing any eye protection at the time of injury similar to
291 previous reports of chemical eye injury which show the vast majority of
292 patients are not wearing any protection at the time of injury.¹⁸ Lack of eye
293 protection is a major risk for the development of a severe chemical injury
294 should exposure occur. Legislation in the UK enforces appropriate use of eye
295 protection in the workplace,¹⁴ Formal education, reinforcement, compulsory
296 use and formal legislation have been shown to be effective in improving
297 compliance.^{19,20} In addition safer working practices, hazard warnings and
298 safety advice on chemical product packaging can help.^{19,20}

299
300 Immediate irrigation following chemical eye injury is probably the single most
301 important intervention influencing outcome more than any other therapeutic
302 approach.^{1,4,18} Three-quarter of our patients received immediate irrigation at
303 the time of injury, including all but one of those that occurred in the workplace.
304 It suggests occupational and public health measures in the UK highlighting the
305 importance of this are partially effective.

306
307 Early management aims to reduce inflammatory cell infiltration and promote
308 corneal re-epithelialisation, keratocyte proliferation and collagenase
309 production.¹ Longer term management aims to restore the visual function by
310 managing the effects of limbal stem cell deficiency, preserving tear production
311 and addressing additional complicating factors.^{11,12,21} Initial treatment involves
312 a combination of preservative-free topical antibiotic, steroids, cycloplegics and
313 pressure lowering treatment as appropriate.⁴ Topical ascorbate, citrate,
314 systemic vitamin C and tetracycline derivatives may promote collagen

315 remodeling, reduce the incidence of ulceration and prevent progression of
316 established corneal ulceration.^{22,23,24} In this survey all patients received initial
317 treatment with topical antibiotic and steroid, with preservative free varieties
318 selected in half of cases. Topical citrate and systemic tetracycline derivatives
319 were used in one quarter of the patients.

320

321 Surgical procedures reported in this survey included amniotic membrane graft
322 (AMG), limbal stem cell graft, oral mucosa graft and fornix reconstruction.
323 AMG was the most commonly reported surgical procedure. AMG is
324 recognised to promote epithelisation, prevent conjunctival adhesions and aid
325 conjunctival surface reconstruction.^{6,11,12,21} Limbal stem cell graft was used in
326 only two patients, but may have been required later as inflammation of the
327 ocular surface induced by chemical burns has a negative impact in the
328 survival of limbal stem cells.²¹

329

330 Six months following injury BCVA was impaired to below 6/12 in half of
331 patients, with two suffering very severe visual loss (worse than 6/60).
332 Reduced vision was attributed in all cases to the corneal pathology indicating
333 substantial ocular morbidity and visual sequelae may be the consequence of
334 severe chemical corneal injury.

335

336 Initial questionnaire data was received in thirty one patients in total. Of these
337 nineteen were excluded from analysis for a variety reasons. Five of these
338 were non-duplicates cases, which met the inclusion criteria, but occurred out
339 with the study period. Summary of their clinical information, supplements the

340 data from the included patients and shows they were broadly similar to those
341 that occurred in the study period.

342

343 This study suggests that severe chemical injury is rare in the UK, occurring
344 less often than previously. It seems that measures to enforce safe use of
345 chemicals at work are generally enforced and effective, but when injury does
346 occur it is evident that significant ocular morbidity and visual sequelae can
347 result.

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423 Mr McKibbin, Mr Mathew, Mr Quinlan, Dr Ramaesh, Mrs Reynolds, Mr

424 Rostron, Mr. Stannard, Ms Weber.

425

426 **DECLARATIONS**

427 This work has been presented at the RCOphth Congress 2008

428

429 Competing Interests: None declared.

430 All authors declare that the answer to the questions on your competing
431 interest form are all No and therefore have nothing to declare

432

433 This research was run through The British Ophthalmological Surveillance Unit
434 The BOSU, run by the Royal College of Ophthalmologists, aims to provide a
435 methodological framework for the systematic investigation of the incidence
436 and clinical features of rare eye conditions with significant public health or
437 scientific importance. Individual investigators submit proposals and the BOSU
438 steering committee determines which conditions will be surveyed. The
439 scheme involves all consultant or associate specialist ophthalmologists with
440 clinical autonomy in the UK. Each month they receive a report card with case
441 definitions of all conditions currently under surveillance. Respondents indicate
442 how many cases of each disorder they have seen (including if they have not
443 seen any). Individual investigators are notified of positive reports and contact
444 the reporting ophthalmologist directly, using a questionnaire to collect
445 information about the reported case. This establishes the patient's eligibility
446 for inclusion and confirms the diagnosis. By collecting unique identifiers,
447 duplicate reports of the same case can be excluded.

448

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451

452 The researchers remained independent from the funders

453

454 All authors, external and internal, had full access to all of the data in the study
455 and can take responsibility for the integrity of the data and the accuracy of the
456 data analysis.

457

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464

465 **TITLES AND LEGENDS TO FIGURES**

466

467 Table 1. The Best Corrected Visual Acuity (BCVA) for Reported Patients with
468 a Severe Chemical Corneal Injury at Presentation Compared to 6 months
469 Following Injury.

470

471 Table 2. The Initial Non-Surgical Management Reported in Patients with a
472 Severe Chemical Corneal Injury.

473

474 Table 3. The Non-Surgical and Surgical Management Reported from 10 days
475 Following Injury to Final Follow-Up at 6 months in Patients with a Severe
476 Chemical Corneal Injury.

477

478 Figure 1. The Clinical Features of Reported Patients with a Severe Chemical
479 Corneal Injury at Presentation

480

481 Figure 2. The Complications Reported During 6 months Follow-Up in Patients
482 with a Severe Chemical Corneal Injury.

Figure 1

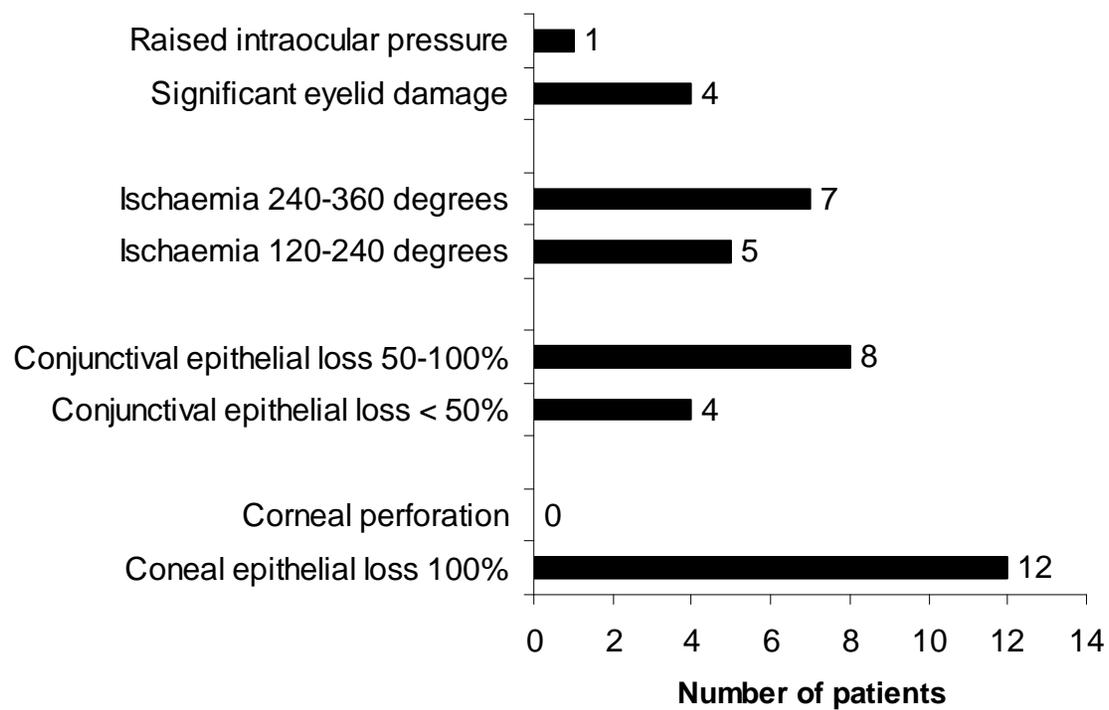


Figure 2

