PERSPECTIVE

Disease and risks associated with contact lenses

Contact lens (CL) wear, as an alternative to spectacles for the treatment of low refractive errors, is widespread in developed countries. It results in a number of inevitable ocular changes which, with the exception of endothelial polymegethism, are short term and recover after lens wear has ceased; these effects are seldom of functional significance.

The risks of lens wear result from both these inevitable ocular responses and the additional potential that exists for developing a CL associated disease. These risks must be set against the optical and cosmetic advantages and convenience resulting from lens use. This risk:benefit ratio should be considered by all contact lens users and practitioners before embarking on lens wear.

Contact lens associated disease, as opposed to the inevitable consequences of wear, develops in only some contact lens users. It includes minor ocular adverse reactions such as corneal neovascularisation, representing a suboptimal ocular response to the stress of lens wear, as well as severe complications such as microbial keratitis.

This review summarises the effect of these on the community, the features distinguishing CL related disorders from other external eye diseases, their classification, and epidemiology.

The consequences of lens related disease for primary eye care

Estimates of the prevalence of lens wear are based on market research and not widely available or precise but they indicate that there are approximately 1.65 million CL users in the adult population of the United Kingdom.¹ In pooled United States Food and Drug Administration premarket approval studies the incidence of severe, sight threatening, adverse reactions has been from 1:244 per year (n=3907) in gas permeable hard CL use to 1:70 per year (n=1276) in extended wear soft CL use.² Minor complications have been reported in 20% (n=100) of daily soft contact lens users over a 3 year follow up period,³ and individual minor complications, such as red eye reactions, in 27.5% (n=400) of patients using extended wear soft lenses from 0–4 years.⁴ This represents a large potential primary eye care problem for the population.

Although most problems are treated by private contact lens practitioners some are complex or serious and others difficult to diagnose and manage. This results in substantial and increasing demands on resources in both the accident and emergency departments and eye clinics of our public hospitals; studies at the same accident and emergency department in London have shown that lens related problems accounted for 2.6% of all new patients in 1978⁵ rising to 3.8% (1104/29 242) in 1988.⁶ This proportion will vary depending on the penetrance of CL use in the catchment population of the hospital and has been as high as 10% in a study in another part of London.⁷ Similar figures have been reported from the United States.⁸

These services are often ill equipped to respond to the problems of contact lens related disease. These include both a substantial group of newly described disorders and different manifestations of established external eye diseases. This has not yet been recognised in either general textbooks of

ophthalmology or textbooks on corneal and external diseases with many disorders described only in the CL literature. In addition, further conditions are continually being described as the ocular tissues respond to challenge by new lens materials, lens care regimes, and wearing patterns. As a result the effective management of CL related disease requires specialist knowledge within the corneal and external disease services provided by ophthalmic units. The diagnosis and management of sight threatening CL related disease, however, is within the province of all ophthalmologists providing general services.

The ocular response to contact lens wear

Contact lenses have a wide range of predictable effects on the eye including the tear turnover,⁹ oxygen availability, corneal metabolism, epithelial and endothelial morphology, and sensation.¹¹ The conjunctiva may also be affected by hypoaesthesia,¹² depletion of oxygen,¹³ and altered epithelial morphology.¹⁴ With the exception of corneal endothelial polymegethism all these changes are reversible soon after discontinuing lens wear. Polymegethism is probably related to chronic corneal stromal acidosis resulting from lens wear and occurs with all current types of lenses with the possible exception of silicone rubber.¹¹ It is probably not fully reversible and in some cases may be associated with endothelial cell loss.¹⁵ Functional consequences can be measured¹⁶ and significant disease may rarely result.¹⁷

Contact lens related diseases

These anticipated short and long term effects of the CL on the eye result from the effect of the lens as a biomaterial. In common with other biomaterials, such as intraocular lenses and hip prostheses, the three principal effects on tissues in contact with the biomaterial are physiological, anatomical, and the consequences of introducing a biologically active surface. Contact lens related disease can be understood in these terms and some of the better characterised CL related diseases illustrate this concept.

Corneal neovascularisation is principally a *physiological* effect resulting from reduced corneal oxygen availability, due to the relatively low oxygen transmission of some contact lenses. This is most often seen at the upper limbus, in soft contact lens wearers, where the cornea is already exposed to reduced oxygen tension levels, because of upper lid cover. These levels are further reduced by the presence of a lens.¹¹

A type of corneal dellen, 3 and 9 o'clock keratopathy (peripheral stain),¹⁸ is an *anatomical* effect resulting from interference by a hard CL with tear resurfacing by the lid. This is probably due to a combination both of physical separation of the lid from the corneal limbus by the lens edge, and from the inhibitory effect of the lens on the blink reflex.^{18 19}

Contact lens associated papillary conjunctivitis (giant papillary conjunctivitis) is the result of the CL developing a *biologically active surface* in which there is an allergic response to ocular deposits that have accumulated on the lens surface. In common with other biomaterials the lens surface rapidly

becomes coated with materials derived from its environment including bacteria, cell debris, mucus, and proteins.²⁰

Many lens related disorders may be the result of *complex interactions* in which two or more of these effects are combined in the pathogenesis. An example is bacterial keratitis. Important factors are the increased susceptibility of the CL wearing eye to bacterial infection resulting both from epithelial trauma and/or physiological stress, particularly in extended wear,^{21 22 23} the exposure of the eye to pathogenic bacteria contaminating the lens case^{24 25} and adhering to the lens,^{26 27 28} or directly colonising the lens surface^{29 30 31 32} and reduced clearance of organisms from the ocular surface due to interference with tear flow and tear resurfacing by the lids.³³

Classification of lens related disease

Our current understanding of the pathogenesis of CL related disease is adequate for a clinical classification based on the adverse ocular effects of CL wear. This approach to the classification of CL related disorders assists in understanding both their development and management unlike the commonly used anatomical classifications. The clinical classification of disease described here was developed for clinical research projects^{6 7} and is based on both the concept of the CL as a biomaterial and the probable pathogenesis. Twenty of the more common CL related diseases, their clinical characteristics, and probable pathogenesis are summarised in Table 1.

The risks of lens related disease

There has been some recent interest in the use of formal epidemiological studies to provide more information about some aspects of CL related disease. These studies have aimed both to assess the effect of some contact lens related diseases on the population as a whole, to establish if the incidence and/or risk of some complications was greater for some types of contact lens and patterns of wear than others, and to assess potential causes with multivariable analysis of associated factors.

MICROBIAL KERATITIS

The impetus for these studies has been principally related to widespread concern about CL wear as a predisposing factor for microbial keratitis, the major sight threatening complication of CL use, and in which bacteria, particularly *Pseudomonas*, have been the major cause.^{31 50 51} Attention was drawn to this problem by the publication of simple descriptive studies and keratitis case series. In these the number of CL users suffering from keratitis were seen to be overrepresented, as were extended wear soft contact lenses in comparison to other lens types.^{50 52} The studies carried out for licensing in the United States before the introduction of new lenses there, as well as post marketing studies of CL users, were uncontrolled, often small, and carried out on carefully selected users.² As a result these did not predict the problems that have become apparent now that these lenses have increased in popularity and are being used by millions of young adults. The principal findings of the recent case control^{53 54} and incidence studies⁵⁵ can be reasonably extrapolated to all countries where contact lenses have a major share of the optical appliance market. The case control study of microbial keratitis in the United Kingdom has shown that CLs are now the major associated cause of microbial keratitis in London with a risk that is significantly higher than that for corneal trauma.⁵⁴ The relative risk (RR) with 95% confidence limits (given in brackets) of keratitis associated with these causes, compared with cases without an identifiable predisposing

factor (the referent with a baseline risk of 1.0) was 80 times (38–166) higher for CL wear and 14 times (6–32) for trauma. Contact lens wear, principally of soft CLs, was also shown to be responsible for 65% of all new microbial keratitis cases at this centre where no serious cases attributable to this cause had been reported a decade earlier. This study also showed that, compared to hard CLs, the risk for extended wear soft CLs was 21 times (7–60) and daily wear soft CLs 3.6 times (1–14). Continuous periods of extended wear of more than 6 days were associated with a further increase in the risk of keratitis. This study confirmed the results both of a previous pilot study carried out in London⁷ and of an independent case control study of ulcerative keratitis, carried out in multiple centres in the United States, which showed that the risk of extended wear soft CL use was 9–15 times higher than that for daily wear soft CLs and that the risk was incrementally related to the period of extended wear; the RRs for hard CLs could not be assessed.⁵³ An incidence study in New England, carried out at the same time as this case control study estimated an annual incidence for ulcerative keratitis in the United States of 20.9:10 000 (15.1–26.7), for extended wear soft CL use compared to 4.1:10 000 (2.9–5.2), for daily wear soft CLs.⁵⁵ Applying these incidence figures to estimates of the size of the CL using population in the United States gave an estimated incidence of 12 000 new cases per year in the soft CL using population. Extrapolation of these incidence data to the relative risks shown in the United Kingdom study for gas permeable hard CLs gives an estimate of the annual incidence of 1:10 000 for this lens type. Similar application of these incidence estimates to the data in the London study using the figure of 1.65 million for the estimated population of CL users in the United Kingdom gives a rough annual estimate of 840 new cases of CL related microbial keratitis per year. Although the incidence of soft CL wear associated keratitis defined in the New England study is the most precise currently available one large retrospective survey of contact lens practitioners, including 196 000 daily wear soft lens users and 72 100 extended wear lens users, in the United States produced estimates as high as 350:10 000 for extended wear soft CLs and 50:10 000 for daily wear soft CLs; at the upper level of the confidence limits in the New England study.⁵⁶

ACANTHAMOEBA KERATITIS

Acanthamoeba keratitis is a rare, painful, and disabling cause of keratitis⁵⁷ which may also result in scleritis⁵⁸ and chorioretinitis.⁵⁹ Contact lens wear is associated in 85% of cases but no differences in risk have yet been identified for the different types of lens,⁶⁰ although a recent case series has disposable soft lens wear as a risk factor in the United Kingdom.⁶¹ The use of home made saline solutions has been identified as a risk factor in the United States.⁶² The prognosis is good following early diagnosis and the use of effective medical therapy.^{63 64} However treatment is often delayed because of misdiagnosis, usually as herpes simplex virus or fungal keratitis.⁶⁵ Currently this disease is apparently increasing in frequency in the United Kingdom⁶¹ and familiarity with the early signs of disease, punctate keratopathy, pseudodendrites, epithelial infiltrates, diffuse or focal subepithelial infiltrates,⁶⁶ and radial keratoneuritis⁶⁷ is important to reduce the morbidity. The ring infiltrate and corneal ulceration are usually late signs of disease.⁶⁸ *Acanthamoeba* must be considered in the differential diagnosis of contact lens users with an atypical keratitis in whom the diagnosis of herpes keratitis should be treated with scepticism.

OTHER COMPLICATIONS

The risks of other adverse responses of CL wear have been

Table 1 Classification of lens related disorders

Classification	Disease	Probable aetiology	Symptoms	Corneal signs	Conjunctival signs	Associated lens
Metabolic (hypoxia, hypercapnia and related effects): Epithelial	Acute epithelial necrosis ^{4a} (overwear syndrome)	Epithelial cell necrosis separation of cells due to hypoxia ¹¹	Often blurred vision before onset due to corneal oedema Delayed pain and epiphora from necrosis. Resolves in hours (days if severe) Starting during or after overnight wear. Vision usually affected	Central punctate epithelial erosions may coalesce into an ulcer. Involved area is larger in SCL users. Stromal oedema in severe cases	Ciliary injection	PMMA-HCL and SCL ^b
	Tight lens syndrome ^{3a}	Lens tightening precipitated by hypoxia and reduced pH ¹⁰ also other factors ¹¹		As above but stromal oedema and an epithelial defect common	Ciliary injection and limbal indentation from a tight lens	EW SCL ^c
	Microcystic epitheliopathy ^{11, 21}	Impaired epithelial metabolic activity	Asymptomatic or minor discomfort	Mini erosions during symptomatic episodes. Clear or opaque epithelial cysts and punctate keratitis	None	All lens types
Stromal	Epithelial oedema ^{3a} (Sartler's veil or central corneal clouding)	Hypoxia ¹⁰ and tear hypotonicity ¹¹	Blurred vision after some hours of wear. May recover on lens removal/progress to acute epithelial necrosis	Dull corneal reflex from central epithelial oedema	None	All lens types
	Stromal oedema ¹⁰ (striae keratopathy)	Stromal lactate accumulation, tear hypotonicity causing corneal swelling ¹¹	Blurring of vision in some cases only	Deep stromal folds from corneal oedema occurring in severe acute epithelial necrosis	None except when associated with acute epithelial necrosis	Usually EW SCL
	Neovascularisation: superficial and deep ¹¹ Deep stromal opacity ^{4b}	Hypoxia causes stromal oedema and release of vasogenic mediators Probably due to prolonged hypoxia and hypercapnia	None unless lipid keratopathy results from deep vessels, causing loss of vision Asymptomatic or reduced acuity	Superficial/deep stromal vessels. Lipid keratopathy associated with deep vessels Pre-Descemet's opacity in central cornea	None	Rare with HCL ^d Common with SCL Rare with HCL
Endothelial Trauma:	Endothelial polymegethism ¹¹	Prolonged hypoxia and hypercapnia	None	Endothelial polymegethism	None	All
	Corneal abrasion ^{3a}	Trauma during lens handling or trapped foreign bodies behind lens, deposits on lens or poor CL fitting	Sudden onset of pain and epiphora. Resolves in hours	Linear or sharply circumscribed epithelial defect	Hyperaemia	Commoner with HCL
Toxic and allergic disorders:	Toxic keratopathy ⁴¹	Exposure to compounds adsorbed onto or absorbed by lens	Pain arising after inserting a lens soaked in proteolytic enzyme/chemically preserved soaking solution	Widespread punctate stain	Ciliary injection	SCL
	Thiomersal keratopathy ⁴²	Preservatives act as haptens causing a delayed hypersensitivity response	Irritation and redness soon after inserting lenses. Vision affected in severe cases	Superior limbal injection, neovascularisation, keratopathy affecting superior quadrant in severe cases None	Intense hyperaemia with lens. Little except follicular changes when lens out	SCL Rare with HCL
	Contact lens related papillary conjunctivitis ⁴³ (giant papillary conjunctivitis)	Multifactorial aetiology: immune response to antigenic proteins on lenses, mechanical effect of lens edge	Increased discharge and greasing of lenses. Itching on lens removal, in early stages, later severe irritation		Upper tarsal hyperaemia, mucus and fine papillary response. 'Giant' (compound) papillae in advanced disease	Common in SCL use
Sterile keratitis:	Contact lens intolerance ⁴⁴	Due to lens spoilation	Chronic redness, discomfort and loss in tolerance. Vision may be blurred	Punctate stain common	Hyperaemia. Papillae and follicles common	All lens types
	Sterile corneal infiltrates ^{44, 45}	Inflammatory response in absence of infecting organism. Hypersensitivity to disinfectants, lens fit, hypersensitivity to bacterial products implicated	Discomfort, redness and discharge	Appearances similar to marginal keratitis with or without ulceration	Hyperaemia	Commoner with SCL
Microbial keratitis:	Microbial keratitis ⁴⁶	Multiple factors including increased ocular susceptibility, and exposure, to pathogens	Rapid onset and progression of pain, hyperaemia and discharge	Epithelial ulcer with underlying stromal infiltrate. <i>Pseudomonas</i> , <i>Staphylococcus</i> and <i>Acanthamoeba</i> , associated	Ciliary injection	Most common with EW SCL Rare in HCL
	Tear surfacing disorders (Dellen): 3 and 9 o'clock stain ^{48, 49}	Drying of corneal surface adjacent to lens edge and abnormal blink	Interpalpebral redness. Rarely discomfort	Punctate keratopathy at 3 and 9 o'clock, rarely vascularised opacities	Interpalpebral hyperaemia	HCL
Corneal distortion:	Inferior corneal stain ⁴⁷ Dimple veil ⁴⁸	Incomplete blinking Static air bubbles under lens	Inferior limbal redness and discomfort Asymptomatic or blurred vision	Inferior/interpalpebral punctate stain Fluorescein pooling in epithelial depressions	Inferior limbal hyperaemia None	SCL HCL
	Corneal warpage ⁴⁹	Mechanical and metabolic factors	Irregular astigmatism, vision good in contact lenses and poor in spectacles	Irregular keratometry and topography	None	PMMA-HCL

^a Polymethylmethacrylate hard contact lenses.
^b Soft contact lenses.
^c Extended wear soft contact lenses.
^d Hard and gas permeable hard contact lenses.

assessed in small uncontrolled studies and in some larger retrospective studies. Overall risks of lens related disease for hard contact lenses have been infrequently reported. Analysis of the pooled results of 48 consecutive premarketing studies on 22 739 CL users for the Food and Drug Administration in the United States have given annualised incidence rates for severe adverse reactions (including keratitis, uveitis, abrasion, infiltrate, oedema, vascularisation, and scarring) of 1:244 (41:10 000) in gas permeable daily CL wear, 1:189 (53:10 000) for daily wear soft CL wear, and 1:70 (143:10 000) for reusable extended wear soft CL wear.² These studies are carried out carefully but uncontrolled and on volunteer users. For these reasons they may not represent the real post-marketing situation. A large retrospective Japanese study on 66 218 CL users gave annualised incidence rates for a variety of ill defined adverse reactions as 16:1000 (160:10 000) for polymethylmethacrylate hard CLs, 6:1000 (60:10 000) for gas permeable hard CLs, and 12:1000 (120:10 000) for low water content soft CLs.⁶⁹ Other smaller prospective and retrospective uncontrolled studies have been carried out in daily wear soft CL use giving incidences of all adverse reactions of 20% (n=100 followed for 3 years an annualised incidence of 6.3% [630:10 000]).³ For extended wear soft lenses studies are available from which annualised incidence rates can be calculated. These range around 1.81% (181:10 000 [n=1099])⁷⁰ for serious complications. Other studies with useful data on the rate of CL complications do not permit the calculation of annualised incidence rates from the data presented.^{4 71 72} These data are difficult to interpret in a way which allows meaningful comparison of the risks associated with the wear of different lens types. Because of this, case control studies have been used to estimate the risks for the development of acute and subacute complications of lens wear. Compared to gas permeable hard contact lenses (the referent with a baseline risk of 1.0), extended wear soft lenses had the largest overall risk for any complication at 2.7 times (1.73–4.16) followed by daily wear soft CLs at 1.3 times (1.0–1.72). The greatest differences in risk were for metabolic disorders [2.1 times (1.28–3.4)] and sterile infiltrates [2.4 times (1.22–4.84)] in extended wear soft CL use and for toxic/allergic disorders [5.9 times (3.27–10.49)] in daily wear soft CL use. Corneal abrasion was the only complication to occur more often in gas permeable hard CL use at 2.9 times more compared to daily wear soft CL use [0.34 times (0.24–0.47)] and 4.4 times more compared to extended wear soft CL use [0.226 (0.13–0.40)].⁶ These results confirmed the findings of previous pilot studies on complications⁷ and sterile corneal infiltrates.⁴⁴

Disposable contact lenses

Disposable soft contact lenses have only become widely available in the last 4 years and experience with these is limited. Enthusiasm for this type of lens wear is based on the theoretical advantages due to the potential for elimination of problems relating to surface deposits, in particular contact lens associated papillary conjunctivitis, and solution reactions.⁷³ The potential reduction of other complications, in particular microbial keratitis, was less certain and metabolic complications would be expected to be identical.^{1 74} Several studies have reported a low incidence of adverse reactions.^{75 76 77} However these study designs are unlikely to identify less common but serious disorders, such as keratitis, for which a relatively low incidence only becomes important when there is a large population at risk. Case series and reports have shown that keratitis may occur in disposable CL wear.⁷⁴ A case control study design is ideally suited to investigating whether there are differences in risks between lens types, allowing comparison of new types of lens and lens wearing regime with those for which the level of risk is better

established.⁶ Small case control studies using this methodology suggest that risks for both sterile and microbial keratitis may be as great or greater than those for conventional soft lens wear^{78 79}; failure of compliance with recommended lens care and wear regimes may be one cause of this. It is to be hoped that disposable lenses will deliver the reduced risks of allergic and toxic adverse reactions that was expected and apparent in pilot studies. This may not be the case.⁸⁰ Although the disposable concept has potentially much to offer in terms of safety and convenience these lenses should be treated with the same caution as other types of lens for the present.

The prevention and management of contact lens related disease

That contact lenses have optical, occupational, sporting, and cosmetic advantages for millions of individuals is clear. However the individual lens user who develops a serious complication is often surprised to discover that risks are attached to their use. Also the size of this population now at risk, of even infrequent serious adverse reactions, has resulted in an increasing problem for those delivering primary eye care. This situation has arisen because of the success of contact lenses as a form of optical correction. Emphasis on their convenience and, for extended wear and disposable lenses, their carefree aspects has led to their widespread introduction and the tendency to trivialise their use.⁸¹ The clear demonstration of differences in risk for different lens types, and the increased risks associated with the use of extended wear, should be understood by all contact lens practitioners in the context both of the population of contact lens wearers and their own practice. The contact lens user must also be educated about the risks of lens wear, and the importance of compliance with the appropriate hygiene and lens wear regimes, so that an informed choice can be made; this is now a complicated area which should not be the province of the inadequately trained. A better understanding of the nature of adverse reactions is needed to provide safer contact lens wear.

When adverse reactions do occur their morbidity can be limited if the problem is rapidly identified and appropriately managed. This is particularly important for keratitis which the general ophthalmologist in the United Kingdom must be competent to deal with. The remaining adverse reactions are usually self limiting if lens wear is discontinued. Referral to a contact lens practitioner who is competent to deal with this specialised area of external and corneal disease is then required. An increased level of education, for both ophthalmologists and contact lens practitioners, is needed to deal more effectively with this problem.

JOHN K G DART

Moorfields Eye Hospital,
City Road,
London EC1V 2PD

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