Original Article

Risk Factors Associated with Contact Lens Related Microbial Keratitis

*Lili Asma Ismail,¹ Lekhraj Rampal,² Hejar Abdul Rahman,² Nazri Omar,³ Habshah Midi,⁴ Azrin Esmady Ariffin⁵

¹ School of Optometry, Faculty of Medicine and Health Sciences, UCSI University,
56000 Cheras, Kuala Lumpur, Malaysia.

² Department of Community Health and ³ Department of Ophthalmology,
Faculty of Medicine and Health Sciences, Universiti Putra Malaysia,
43400 UPM Serdang, Selangor, Malaysia.

⁴ Department of Mathematics, Faculty of Sciences, Universiti Putra Malaysia,
43000 UPM Serdang, Malaysia.

⁵ Faculty of Optometry and Visual Sciences, SEGi University, Kota Damansara, Selangor, Malaysia.

ABSTRACT

Introduction: Microbial keratitis is one of the most challenging complications of contact lens (CL) wear. Proper CL practice plays an important role to reduce the risk for contact lens related microbial keratitis (CLRMK). **Methods:** This multi-centre case-control study was conducted from January 2008 until June 2009 to determine the risk factors associated with CLRMK. Cases were defined as respondents who were treated for CLRMK, whilst controls were respondents who were contact lens wearers without microbial keratitis. Ninety four cases were compared to 94 controls to determine the risk factors for CLRMK. **Results:** The predictors for CLRMK were: Not washing hands with soap before handling CL (aOR 2.979, CI 1.020, 8.701 p=0.046), not performing rubbing technique whilst cleaning the CL (aOR 3.006, CI 1.198, 7.538 p=0.019) and, not cleaning the lens case with multipurpose solution daily (aOR 3.242 CI 1.463, 7.186 p=0.004). Sleeping overnight with the CL in the eye (aOR 2.864, CI 0.978, 8.386 p=0.049) and overall non-compliance with lens care procedures (aOR 2.590, CI 1.003, 6.689 p=0.049) contributed significantly to CLRMK. **Conclusion:** Health education and promotion in contact lens care are important and should be conducted by eye care practitioners to reduce the occurrence of CLRMK.

Key words: Contact lenses, Microbial Keratitis, Risk factors

INTRODUCTION

Contact lenses (CL) are widely used for refractive visual correction and are worn by approximately 140 million people around the globe (1). Microbial keratitis is an infection of the cornea due to bacteria, fungi or parasites (2). It may be associated with ulceration of the corneal epithelium with inflammation of the underlying stroma (3). It can also be related to ocular trauma, ocular surgery, ocular surface disease, bullous keratopathy, corneal anaesthesia, corneal exposure cases and dry eyes (4). Although microbial keratitis is rare³, it can cause significant ocular morbidity and thus requires prompt diagnosis and treatment to prevent devastating outcomes (5).

Before the popular usage of contact lens for the correction of ametropia, microbial keratitis was rare in normal eyes because of the eye's natural resistance to infection, and occur only in association with therapeutic and aphakic contact lens wearers (6,7). Non-compliance to lens care procedure and unhygienic contact lenses and their storage cases have been associated with microbial keratitis (8). It has been widely reported that CLRMK is often caused by *Pseudomonas aeruginosa* which can lead to significant morbidity (9). Contact lens related microbial keratitis is an important public health issue because of the large number of wearers who are at risk for the disease with the majority of them who are in their prime working years (2).

Overnight use of contact lenses, poor storage case hygiene, smoking, internet purchase of contact lenses, usage of less than 6 months and higher socio-economic status are some of the risk factors for CLRMK (10). Compliance to the prescribed regime is indeed important, and practitioners should ensure users wear their lenses according to wear schedule, are properly instructed in lens handling, lens case hygiene and replacement. They should be made aware of

*Corresponding Author: Dr. Lili Asma Ismail liliasma10@gmail.com 2

the importance of good hand hygiene and increased risk of CLRMK associated with overnight use of contact lenses (11). Studies done in Malaysia found that the factors associated to CLRMK were mainly behavioral and were related to lens care procedures and practices (12). Cases-control studies on CLRMK has not been reported in Malaysia and this has motivated us to embark on the present study. The objectives of this study are to determine the sociodemographic characteristics of patients, the proportion of CLRMK by the types of contact lens and the risk factors associated with CLRMK in Klang Valley, Malaysia.

MATERIALS AND METHODS

This case-control study was carried out in the eye clinics of seven hospitals throughout the Klang Valley in Malaysia. Approval was obtained from the Ethics Committee of the Medical Faculty, University Putra Malaysia and Ethics Committee from the National Medical Research Registrar, Ministry of Health Malaysia. Informed consent was obtained from the respondents before participation in the study. All respondents with CLRMK were included as cases and controls were contact lens users without CLRMK who attended the eye clinics for reasons other than anterior eye pathology. Patients with keratoconus, bandage lenses and aphakia were excluded from this study.

Methods

A self-administered questionnaire in *Bahasa Malaysia* was used. The questionnaire was aimed to determine the following risk factors for CLRMK: Hand washing with soap while handling the contact lenses, rubbing of the lenses when cleaning them, re-disinfecting the contact lenses before resuming wear, daily cleaning of lenses with multipurpose solution, sleeping overnight with lenses in the eye, and overall non-compliance with lens care procedures defined as the inability to fully adhere to the recommended lens care procedures.

Sample size determination

The sample size was calculated using a formula that is derived from Schlesselman (1982) specifically for case control study. Equal number of cases and controls was selected for sample size in each group (n per group) as above. With an estimated proportion rate among the controls (p_0) being 71% and OR for soft disposable lens being 3.51 (Radford,1998a), the sample size was 94 per group for cases and controls. The symbol p_0 is the Estimated exposure rate (proportion exposed) in controls and R is the Relative risk corresponding to the smallest decrease or increase in risk of interest.

Data collection

Data collected included demographic data, contact lens characteristics and personal contact lens handling and care. Microbial investigations were done using culture and sensitivity technique by the hospital laboratory service.

Data Analysis

Crude ratios were calculated for each of the aforementioned factors using logistic regression. Adjusted odds ratios were done for adjustment of confounders using multiple logistic regression test. All data were analysed using SPSS 16 which included chi-square test and multiple logistic regression.

RESULTS

Out of the 100 incident cases, a total of 94 cases agreed to participate giving a response rate of 94%. Out of a total of 110 controls, a total of 94 controls agreed to participate giving a response rate of 85%. A total of 188 respondents which comprised of 94 cases and 94 controls were recruited.

Sociodemographic features

Average age of the subjects was 27 ± 7 years (n = 188). Majority of cases and controls were in the younger age group (18-29 years) being 65% and 82% respectively. Female was predominant in this study with 78% females in cases and 68% in controls. Malays made the majority for cases (75%) and controls (43%). The demographic data of the respondents is shown in Table 1.

Contact lens characteristics

Of all the cases, majority (n=84, 89.36%) were using monthly disposable lenses. Others used yearly conventional lenses (n=6, 6.38%), bi-weekly disposable lenses (n=2, 2.13%) and daily disposable lenses (n=2, 2.13%). Among

the controls, 48% were conventional RGP wearers. Contact lens characteristics, wearing mode and lens replacement schedule are shown in Table 2.

Lens Care Practices

The risk factors in relation to lens care procedures were the following: Not washing hands with soap (OR 6.415, CI 2.66, 15.47, p=0.001), not performing rubbing technique to clean lenses (OR 5.693, CI 2.829, 11.454 p=0.001), not redisinfecting lenses after lenses were left longer than indicated (OR 4.085, CI 2.078, 8.030 p=0.001) and, not using multipurpose solution containing enzymatic cleaning agent weekly or not carrying out enzymatic cleaning at least once a week (OR 2.718, CI 1.353, 5.461 p=0.005). Sleeping overnight with lens in the eye (OR 4.427, CI 1.965, 9.976) and overall non-compliance with lens care procedure (OR 3.238, CI 1.592, 6.565) increased the risk for CLRMK (Table 3). Multiple logistic regression analysis was performed to obtain the adjusted odds ratio (aOR). The significant results were found for Not washing hands with soap before handling lenses (aOR 2.979, CI 1.020, 8.701 p=0.046), not performing rubbing technique (aOR 3.006, CI 1.198, 7.538 p=0.019) and, not cleaning lens case with multipurpose daily (aOR 3.242 CI 1.463, 7.186 p=0.004). Sleeping overnight with lenses in the eye contributed to an increased risk of almost three times (aOR 2.864, CI 0.978, 8.386 p=0.049) and non-compliance with lens care procedures had an increment in the risk of up to 2.6 times (aOR 2.590, CI 1.003, 6.689 p=0.049) (Table 4).

Table 1. Demographic characteristics data and the association between contact lens related microbial keratitis

	Case n=94	Control n=94	X ²	df	p
Total					
Age (years)					
18-29	61 (64.9)	77 (81.9)	8.69	3	0.034*
30-39	24 (25.5)	13 (13.8)			
40-49	6 (6.4)	1 (1.1)			
50-59	3 (3.2)	3 (3.2)			
Gender					
Male	21 (22.34)	30 (32)	2.18	1	0.140
Female	73 (77.66)	64 (68)			
Ethnicity					
Malay	70 (74.47)	40 (42.55)	23.7	3	0.001*
Chinese	13(13.83)	30 (31.91)		-	
Indian	6 (6.38)	21 (22.34)			
Others	5 (5.32)	3 (3.20)			
Education level					
Primary	4 (4.26)	2 (2.13)	9.4	4	0.049*
Secondary	33 (35.11)	51 (54.26)			
Diploma/Cert	26 (27.63)	15 (15.96)			
Degree	30 (31.94)	23 (24.46)			
Master/PhD	1 (1.06)	3 (3.19)			
Income level					
< RM1000	19 (22.21)	10 (10.64)	17.02	5	0.004*
RM1000-1999	21 (22.34)	16 (17.02)	- / · · · · =	-	0.00.
RM2000-2999	28(29.79)	18 (19.15)			
RM3000-3999	11	21 (22.34)			
RM4000-4999	(11.70)	8 (8.51)			
>RM5000	7(7.45) 8 (8.51)	21 (22.34)			

^{*} Significant at p < 0.05

[†] Chi square bivariate analysis

Table 2. The association between contact lens related microbial keratitis (CLRMK) and contact lens characteristics

	Case n (%)	Control n (%)	X ² †	p
Lens type				
Disposable Soft	88(93.62)	46(48.94)	59.16	0.001*
Conventional Soft	6(6.38)	3(3.20)		
Conventional RGP	0(0)	45(47.86)		
Wearing mode				
Daily wear	65(69.15)	76 (80.85)	3.46	0.181
(< 12 hours) Daily wear	27(28.72)	17 (18.09)		
(13-24 hours)	27(20.72)	17 (10.07)		
Extended wear	2 (2.13)	1 (1.06)		
(≥ 24 hours)	, ,	,		
Replacement				
schedule				
Daily	2 (2.13)	7 (7.45)	53.12	0.001*
Two weeks	2 (2.13)	1 (1.06)		
Monthly	84 (89.36)	38 (40.43)		
Yearly	6 (6.38)	48 (51.06)		

Table 3. The risk factors for CLRMK related to lens care

Risk factor	Crude Odds ratio -	95% confidence interval, CI		р
		lower	upper †	_
Not washing hands with soap	6.415	2.660	15.470	0.001*
No rubbing technique	5.693	2.829	11.454	0.001*
No redisinfecting lenses	4.085	2.078	8.030	0.001*
Not cleaning case daily with MPS	4.086	2.191	7.617	0.001*
Sleeping overnight with lens in the eye	4.427	1.965	9.976	0.001*
Overall non- compliance to lens care procedure	3.238	1.592	6.565	0.001*

^{*} Significant at p < 0.05

^{*} Significant at p < 0.05 † Chi square bivariate analysis

^{†95%} Confidence intervals lower and upper value

Table 4. The predictors for contact lens related microbial keratitis

Predictor	Adjusted odds ratio	95% confidence interval, CI		р
		lower	upper †	
Ethnicity				
Chinese	0.126	0.045	0.355	0.001*
Indian	0.295	0.088	0.989	0.048*
Other	0.414	0.066	2.611	0.348
Not washing hands	2.979	1.020	8.701	0.046*
Not performing rubbing	3.006	1.198	7.538	0.019*
Not cleaning lens case daily with MPS Sleeping	3.242	1.463	7.186	0.004*
overnight with lens Overall non	2.864	0.978	8.386	0.049*
compliance to lens care rocedures	2.590	1.003	6.689	0.049*
Constant	0.001			0.001

^{*} Significant at p < 0.05

DISCUSSION

A compliant CL user is a person who washes his hands before handling contact lenses, uses an FDA (Food and Drug Administration) approved care system and in agreement with the published guidelines of the manufacturer and good hygiene while adhering to the recommended wear schedule (13). Non-compliance with contact lens care procedures is defined as failure to adhere to the proper technique of contact lens maintenance. There have been relatively few studies conducted on the predisposing factors leading to CLRMK in Malaysia. This is the first local multi-centre case control study on the risk factors for CLRMK. The present study showed that the predictors for CLRMK were: Not washing hands with soap before handling lenses (aOR 2.979, CI 1.020, 8.701 p=0.046), Not performing rubbing technique (aOR 3.006, CI 1.198, 7.538 p=0.019) and Not cleaning lens case with multipurpose daily (aOR 3.242 CI 1.463, 7.186 p=0.004). In addition, it was discovered that sleeping overnight with the lens in the eye (aOR 2.864, CI 0.978, 8.386 p=0.049) and overall non-compliance with lens care procedures (aOR 2.590, CI 1.003, 6.689 p=0.049), were predictors for CLRMK.

Not washing hands before handling lenses

Omitting hand washing increased risk for CLRMK by 3 folds (aOR 2.979, CI 1.020-8.701, p=0.046). Previous compliance studies have revealed that 16-50% of subjects did not pay proper attention to hand washing prior to lens insertion (13). By diligently adhering to the cleaning guidelines such as washing hands with soap, most if not all organisms will be removed from the hands and fingernails, thus preventing contamination of the lens (14). Previous studies have found patients practiced poor hand washing or incomplete rinsing of hands, did not clean lens case regularly, did not replace lens case regularly, did not observe cleanliness with bottle caps, kept contact lens solution bottle caps open and kept lens cases open, did not rinse lenses after taking them out of the cases and topping off solution in the lens cases (15).

Not performing the rubbing technique

In this study, after adjusting for confounders, not performing the rubbing technique was found to increase in the risk for CLRMK by three folds (aOR 3.006, CI 1.198, 7.538 p=0.019). Cleaning and rinsing contact lenses immediately after lens removal with the cleaner/multipurpose solution and saline is more effective than cleaning before inserting

^{† 95%} Confidence intervals lower and upper value

the lens in the eye (16). This is because all the tear deposits, contaminants and microorganism will be removed from the lens before they are placed in the lens case. Physical cleaning of lenses and lens case is helpful in reducing microorganisms before chemical disinfection (17). Complacency to lens hygiene and the emphasis of convenience which has been encouraged by the availability of no-rub solutions may have contributed to this behaviour. Performing the rubbing technique improves the effectiveness of lens cleaning and failure to perform this technique increases the risk of CLRMK (18). If the lens rubbing is not performed properly, deposits will accumulate at the midperiphery area of the lens, which will subsequently lead to an allergic response caused by the friction between the lens and the palpebral aperture when the eye blinks (19).

Not cleaning lens case daily with multipurpose solution

Cleaning the lens case daily with a multipurpose solution and disinfecting the lens case weekly are recommended hygiene practices (20). Non-compliance with care of lens case has been associated with contamination of the case and infection in contact lens wearers (21) and increase risk for microbial keratitis including Acanthamoeba (5,9,22). Contact lens users risk up to 50-fold increase in the risk of *Acanthamoeba* keratitis, largely attributable to repeated wear of lenses, irregular or lack of disinfection and use of saline and chlorine-based solutions (16).

Overnight wear of contact lens

Overnight wear of lens is known to be a major risk factor for microbial keratitis (23). In an eye with contact lenses, there are greater levels of hypoxia and hypercapnia compared to open eye. Coupled with tear stagnation beneath the lenses in a closed lid environment it can lead to epithelial compromise, oedema and superficial punctate keratitis (SPK) (24-26). In this study, overnight wear of contact lens increased the risk of CLRMK by 3 folds (aOR 2.86, CI 0.978-8.386, p=0.049). The results supported other studies which reported an increased risk of CLRMK when contact lenses were worn overnight: (OR 3.95, CI 1.02, 15.26) for daily wear soft contact lens (27).

Non-compliance with lens care procedures

Non-compliance with lens care procedures, and contaminated lenses and lens cases were associated with an increased risk for microbial keratitis (8). A study in Hong Kong found non-compliance with contact lens care procedures to be a risk factor for CLRMK (OR 11.04, CI 1.87, 135.56, p=0.003) (28). The specific areas of non compliance that are frequently discussed are: Lack of hand hygiene, lack of case hygiene, prolonged contact lens use, inadequate use of lens care maintenance systems and infrequent follow-up visits (29). In this study, overall non-compliance was a risk factor for CLRMK and contributed to an increased risk of 2.59 (aOR 2.59, CI 1.003-6.689, p=0.049). Thus, it is important to improve patient compliance with contact lens care procedures as it has been shown in our study that non-compliance increased the risk for CLRMK. Our study also supports studies conducted elsewhere which found *Pseudomonas aeruginosa* as the most common microorganism in CLRMK in developing worlds. It was also in agreement with the findings that *Pseudomonas* was the most frequent pathogen found in corneal ulcers of soft contact lens wearers (30,31,32,33).

Limitations

Although this was the first case-control study on the risk factors for CLRMK to be conducted in Malaysia to the knowledge of the authors, there were unavoidable limitations. Among the limitations was not using matching in the design stage of the study. The reasons were due to inability to find suitable hospital matched controls in our setting. The use of multiple logistic regression in the analytical part of the study seemed more appropriate for the case control study design besides controlling for confounders. In addition, by using hospital based controls, it only provided internal validity to the study. The fact that a high proportion of hospitals based controls were RGP users may not actually reflect the real situation in the community. In view of that, the authors proposed that future case control studies be conducted using population based controls.

CONCLUSION

Health promotion and education should be conducted by eye care practitioners, doctors and optometrists for the public specifically for the contact lens users. With improved public health promotion, this will increase awareness on the risks for CLRMK and minimize the occurrence of CLRMK.

ACKNOWLEDGEMENT

The authors thank colleagues in the participating hospitals and others for their cooperation and contribution in any possible way to the successful completion of this research.

REFERENCES

- 1. Stapleton F, Keay L, Jalbert I, Cole N. The epidemiology of contact lens related infiltrates. Optom Vis Sci 2007; 84 (4): 257-722.
- 2. Dart JKG. Disease and risk associated with contact lenses. Br J Ophthalmol 1993; 77: 49-53.
- 3. Keay L, Edwards K, Stapleton F. Signs, symptoms, and comorbidities in contact lens related microbial keratitis. Optom Vis Sci 2009; 86 (7): 803-809.
- 4. Weismann BA, Mondino BJ. Microbial keratitis. In: Efron N (ed.) The cornea and its examination in contact lens practice. Oxford: Butterworth Heinemann; 2001. p 50-85.
- 5. Schein OD, Glynn RJ, Poggio EC. The relative risk of ulcerative keratitis among users of daily wear and extended wear soft contact lenses. A case control study. Microbial Keratitis Study Group. N Engl J Med 1989; 321:773-778.
- 6. Dart JKG. Predisposing factors in microbial keratitis: the significance of contact lens wear. Br J Ophthalmol 1988; 72:926-930.
- 7. Dart JKG, Stapleton F, Minassian D. Contact lenses and other risk factors in microbial keratitis. Lancet 1991; 338: 650-653.
- 8. McLaughlin-Borlace L, Stapleton F, Matheson M, Dart JKG. Bacterial biofilm on contact lenses and lens storage cases in wearers with microbial keratitis. J Appl Microbiol 1998; 84: 827-838.
- 9. Houang E, Lam D, Fan D, Seal D. Microbial keratitis in Hong Kong: relationship to climate, environment and contact-lens disinfection. Trans R Soc Trop Med Hyg 2001; 95: 361-367.
- Stapleton F, Keay L, Edwards K. The incidence of contact lens related microbial keratitis in Australia. Ophthalmol 2008; 115 (10): 1655-1662.
- 11. Dart JKG, Radford CF, Minassian D. Risk factors for microbial keratitis with contemporary contact lenses: a case control study. Ophthalmol 2008; 115: 1647-1654.
- 12. Norhalwani H. Contact lens induced microbial keratitis in Hospital Kuala Lumpur. An evaluation of its contributing factors, causative agents and final outcome. Masters of Surgery (Ophthalmology) Dissertation, University Kebangsaan Malaysia, 2007.
- 13. Donshik PC, Ehlers WH, Anderson LD, Suchecki JK. Strategies for better engage; educate & empower patient compliance and safe lens wear: Compliance: What we know, what we do not know & what we need to know. Eye Contact Lens 2007; (33) 6: 430-433.
- 14. Yung MS, Boost M, Cho P, Yap M. The effect of a compliance enhancement strategy (self review) on the level of lens care compliance and contamination of contact lenses and lens care accessories. Clin Exp Optom 2007; 90 (3): 190-202.
- 15. Levy B, Orsborn G. Clinical risks: Myths and truths Interpreting the evidence-based data about contact lens care. Clin Ref Optom 2008; 19 (6): 165-171.
- 16. Radford CF, Bacon AS, Dart JKG, Minassian DC. Risk factors for Acanthamoeba keratitis in contact lens users : A case control study. Br J Ophthalmol 1995; 310: 1567-1570.

- 17. Ifejika CP, McLaughlin-Borlace L, Lucas VJ, Roberts ADG, Walker JT. Efficacy of a contact lens cleaning device and its enhancement of the performance of contact lens care products. Br J Ophthalmol 2000; 84: 539-541.
- 18. Rosenthal RA, Henry CL, Schlech BA. Contribution of regimen steps to disinfection of hydrophilic contact lens. Cont Lens Anterior Eye 2004; 27 (3): 149-156.
- 19. Terry RL, Schnider CM, Holden BA, Cornish R, Grant T, Sweeney D, La Hood D, Back A. Standards for Success of Daily and Extended Wear contact lenses. Optom Vis Sci 1993; 70 (3): 234-243.
- 20. Hughes R, Kilvington S. Comparison of hydrogen peroxide contact lens disinfection systems and solutions against Acanthamoeba polyphaga. Antimicrob Agents Chemother 2001; 45 (7): 2038-2043.
- 21. Devonshire P, Munro FA, Abernethy C, Clark BJ. Microbial contamination of contact lens cases in the west of Scotland. Br J Ophthalmol 1993; 77: 41-45.
- 22. Stapleton F, Dart JKG, Minassian D. Risks factors with contact lens related suppurative keratitis. CLAO J 1993; 19 (4): 204-210.
- 23. Schein OD, Buehler PO, Stamler JF. The impact of overnight wear on the risk of contact lens associated ulcerative keratitis. Arch Ophthalmol 1994; 112: 186-190.
- 24. Ang JH, Efron N. Corneal hypoxia and hypercapnia during contact lens wear. Optom Vis Sci 1990; 67: 512-521.
- 25. Tabbara KF, El-Sheikh HF, Aabed B. Extended wear contact lens related bacterial keratitis. Br J Ophthalmol 2000; 84 (3): 327-328.
- 26. Dart JK, Badenoch PR. Bacterial adherence to contact lenses. Eye Contact Lens 1986; 12: 220-224.
- 27. Radford CF, Minassian DC, Dart JKG. Disposable contact lens use as a risk factor for microbial keratitis. Br J Ophthalmol 1998; 82 (11): 1272-1275.
- 28. Lam DS, Houang E, Fan DS. Incidence and risk factors for microbial keratitis in Hong-Kong; Comparison with Europe and North America. Eye 2002; 16: 608-618.
- 29. Claydon BE, Efron N, Woods C. A prospective study of non-compliance in contact lens wear. Cont Lens Anterior Eye 1996; 19: 133-140.
- 30. Dang YNT, Rao A, Kastl PR, Blake RC, Schurr MJ, Blake DA. Quantifying Pseudomonas Aeruginosa adhesion to contact lens. Eye Contact Lens 2003; 29:65-69.
- 31. Fleiszig SMJ. The pathogenesis of contact lens related keratitis. Optom Vis Sci 2006; 83: 866-873.
- 32. Holden B, De La Jara PL. Contact lenses: Optimal vision-sub optimal carrier? Optom Vis Sci 2007; 84 : 365-367.
- 33. Willcox MDP. Pseudomonas Aeruginosa: Infection and Inflammation during contact lens wear: A review. Optom Vis Sci 2007; 84(4): 273-278.