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Risk of *Klebsiella pneumoniae* Endogenous Endophthalmitis during Bacteremia: Implications for Screening

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Dear Editor,

Endogenous endophthalmitis is caused by bacteremic spread of pathogens to the posterior segment of the eye. *Klebsiella pneumoniae* is the most frequent bacterial cause of endogenous endophthalmitis [1]. In keeping with animal models in which rapid destruction of retinal photoreceptors occurs as early as 24 - 48 hours after inoculation, *K. pneumoniae* endogenous endophthalmitis (KLEE) ocular outcomes are poor, particularly if diagnosis and treatment are delayed [2, 3]. In this regard, the role of ocular screening of asymptomatic patients with *K. pneumoniae* bacteremia is debated [4].

After obtaining the relevant ethics committee approvals (identifier numbers 14299/7985), we retrospectively identified state-wide public hospital KLEE cases via discharge summary International Classification of Diseases codes. KLEE was defined as culture of *K. pneumoniae* from an intraocular specimen, or intraocular inflammation in conjunction with *K. pneumoniae* bacteremia. By cross referencing cases against a state-wide public pathology database, we calculated the risk of KLEE during *K. pneumoniae* bacteremia, with or without concurrent liver abscess. *K. pneumoniae* bacteremia and liver abscess were defined as culture of *K. pneumoniae* from a blood or hepatic specimen respectively.

Between 2006 and 2020, we identified 11 cases of KLEE amongst 2,689 episodes of *K. pneumoniae* bacteremia (**Table 1**), giving a risk of 0.4%, which increased to 9.0% (4/44) if the patient had a concurrent liver abscess. Four patients (36%) did not report ocular symptoms at the time of presentation and for these patients (including two obtunded patients), recognition of endophthalmitis was delayed by at least 48 hours. For patients whose presentation did include ocular symptoms, intravitreal antimicrobials were administered within 24 hours.

Published rates of KLEE amongst patients with *Klebsiella* bacteremia range from 1.0% in New Caledonian [5] to 3.8% in Singapore [2]. Whilst KLEE is an emerging syndrome within Australia [3], the low frequency of hypervirulent clones (K1/K2 subtypes) amongst Australian



Author contributions

Conceptualization: RB, PG, FC. Data curation: RB, PG, PI. Formal analysis: RB, PI. Investigation: RB, BH. Methodology: RB, WC, FC. Software: RB, PI. Validation: RB, PI. Writing - original draft: RB, PI, FC. Writing - review & editing: WC, PG, BH. **Table 1.** Patient, treatment and clinical outcome data for patients with *Klebsiella pneumoniae* endogenous endophthalmitis (n = 11)

Age (years) (median, IQR)	61 (37 - 82)
Male gender (n, %)	8 (73%)
South East Asian ethnicity (n, %)	6 (55%)
Diabetic (n, %)	5 (46%)
Examination findings at presentation	
Visual acuity: hand motion or worse (n, %)	10 (91%)
Conjunctival infection (n, %)	11 (100%)
Hypopyon (n, %)	7 (64%)
Proptosis (n, %)	5 (46%)
Elevated IOP (n, %)	1 (9%)
Vitritis (n, %)	11 (100%)
Concurrent organ systems involved	
Hepatic (abscess) (n, %)	9 (82%)
Respiratory (pneumonia) (n, %)	6 (55%)
Genitourinary (abscess) (n, %)	3 (27%)
Central nervous system (meningitis/encephalitis) (n, %)	2 (18%)
Freatment	
Systemic antimicrobials (n, %)	11 (100%)
Intravitreal antimicrobials (n, %)	10 (91%)
Vitrectomy	
Primary (n, %)	1 (9%)
Secondary (n, %)	1 (9%)
Outcomes	
Evisceration or Enucleation (n, %)	6 (55%)
Visual acuity: hand movements or worse (n, %) ^a	4 (36%)
Inpatient mortality (n, %)	0 (0%)

^aAt most recent outpatient follow up assessment.

IQR, interquartile range; IOP, intra ocular pressure.

K. pneumoniae bacteremia isolates may explain the lower risk of KLEE amongst our cohort [6]. In keeping with other studies, most of our patients had epidemiological links to Asia, were diabetic, and visual acuity outcomes were poor [7].

More than one third of our patients did not report ocular symptoms at the time of presentation. As early diagnosis and therapy improves visual outcomes [7], ophthalmologic screening of bacteremic patients without (or with unrecognised) ocular symptoms may off set delays in recognition of KLEE. However universal ophthalmologic review during *Klebsiella* bacteremia may detect asymptomatic retinal lesions of uncertain significance in 43% of patients [8]. Thus, contrary to published recommendations in favour of universal ocular screening for patients with *Klebsiella* sepsis [9] (akin to standard practice for candidemia in which the risk of endophthalmitis is 10 - 37% [10]), we conclude that routine ophthalmologic review for the purposes of early detection of endophthalmitis be limited to bacteremic patients with concurrent liver abscesses or other predisposing factors such as infective endocarditis [4]. Finally, as severity of illness is a risk factor for ocular spread [9], patients obtunded due to overwhelming *Klebsiella* sepsis should be considered for routine ophthalmological evaluation.

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