Primary *Klebsiella pneumoniae* Liver Abscess with Metastatic Spread to Lung and Eye, a North-European Case Report of an Emerging Syndrome

Sara K Sobirk¹,* , Carsten Struve² and Sanda G Jacobsson³

¹Clinical Microbiology, University and Regional Laboratories Region Skåne s-221 85 Lund, Sweden
²Department of Bacteriology, Mycology and Parasitology, Statens Serum Institut, 2300 Copenhagen S, Denmark
³Department of Infectious Diseases, County Hospital of Helsingborg, s-251 87 Helsingborg, Sweden

Abstract: A syndrome of community acquired liver abscess caused by highly virulent hypermucoviscous, *rmpA* positive *Klebsiella pneumoniae* strains occurs with high incidence in Asia. We here report a case complicated by metastatic infection, to our knowledge, the first one described in Northern Europe. This and other recently reported cases indicate the emergence of this severe syndrome outside of Asia.

Keywords: *Klebsiella pneumoniae*, metastatic infection, hypermucoviscous, *rmpA*.

INTRODUCTION

In recent years a high incidence of community-acquired primary liver abscess caused by highly virulent strains of *Klebsiella pneumoniae* has been documented in Taiwan and other Asian countries. [1-4]. A syndrome, with metastatic spread to eyes, meninges, brain and other sites, has been described, and is strongly associated with *K. pneumoniae* strains of capsular serotype K1 of a characteristic hypermucoviscous phenotype [4, 5]. Two plasmid encoded virulence genes, *rmpA* (regulator of the mucoid phenotype) and the iron siderophore aerobactin, as well as two chromosomal encoded virulence genes, *kfu* encoding an iron uptake system and *allS* associated with allatonin metabolism are significantly correlated with the high virulence of the invasive strains [6, 7]. These highly virulent strains are so far rare outside of Asia, but case reports from USA, Canada, Spain and Belgium have been published lately [8-14].

We here report a Swedish patient with this emerging syndrome caused by a highly virulent *K. pneumoniae* isolate. This is, to our knowledge, the first case described in Northern Europe.

CASE REPORT

A 45-year old man of Vietnamese origin was admitted with a ten day history of fever, anorexia, fatigue and right-sided abdominal pain. He had no previous history of disease and no medication. At examination, he had signs of peritonitis, hypotension and a red and swollen left eye. He was initially treated with intravenous fluids and imipenem for suspected septicemia. Pulmonary X-ray and CT-scan revealed right-sided pulmonary infiltrates with pleural effusion and a large mass in the liver with the appearance of multiple cysts.

(Fig. 1) Examination of the eye showed marked chemosis, absence of light reflex and intravitreal hemorrhage. Empiric treatment for endophthalmitis was given intravitreously (cefazidime and vancomycin). Cultures of blood, urine, aspirates from liver and pleural effusion all grew *K. pneumoniae* sensitive to imipenem and cefazidime. Cultures that were taken from the eye, after initiation of intravenous therapy, were negative. Despite adequate treatment and continuous drainage of both liver abscess and empyema, repeated cultures from these sites showed the same mucoid strain of *K. pneumoniae* up to 23 days after initiation of intravenous antibiotics. Two repeated blood cultures taken one week after initiation of therapy were negative. Following three weeks of intravenous therapy, the patient was treated with oral ciprofloxacin for another five weeks. His recovery, although slow, corresponded well with repeated CT-scans of lungs and liver showing regress of the infection. Although the pain in his left eye decreased, his vision is permanently damaged, and he is now blind on the affected eye. The patient had immigrated to Sweden 8 years previously, passing a refugee-camp in Hong Kong, and had been back to visit Vietnam once, 10 months before start of symptoms. Initial investigations of the patient therefore included TB-analyses and a broad parasitic screen, all found to be negative.

The *K. pneumoniae* isolate that caused the infection was found to express the hypermucoviscous phenotype as shown by the formation of mucoviscous strings when a loop was passed through a colony (Fig. 2). Capsular serotyping at the International *Eschericia coli* and *Klebsiella* Reference Centre (WHO), Statens Serum Institut revealed that the isolate belongs to the K1 serotype which was confirmed by detection of the K1 serotype specific gene *magA* as previously described (Fig. 3) [15]. Furthermore the isolate was positive for all four virulence genes *rmpA*, aerobactin, *kfu* and *allS* as revealed by polymerase chain reaction using specific primers (Fig. 3) [5]. Thus the isolate exhibited similar characteristics.
as the highly virulent *K. pneumoniae* isolates that have so far mainly been observed in Asia.

**DISCUSSION**

The rising incidence of *K. pneumoniae* community acquired primary liver abscess in otherwise healthy individuals caused by highly virulent strains and often complicated by severe metastatic infections is disturbing [2, 3, 14]. Whereas this emerging infectious disease so far has been observed mainly in Asia, a rising number of cases from other geographic regions indicate that these infections may develop into a world wide problem. It is remarkable that the
majority of cases outside Asia have involved patients of Asian ethnicity. The reasons for the predominance of this syndrome in patients of Asian ethnicity are not known, but suggest that genetic susceptibility may play an important role. Alternatively, it could be speculated that there may be a food source of the virulent bacteria. Cases with no history of traveling to Asia have been reported demonstrating that highly virulent K. pneumoniae strains are present outside of Asia [10, 12, 13]. However, although our patient had only had Swedish and Vietnamese contacts, he may have been colonized with the strain in Vietnam 10 months before onset of symptoms.

Patients with a recent history of travelling in foreign countries are often thoroughly investigated for infections with virus, bacteria or parasites that are typical for the region visited. Bacterial species that are common all over the world can, with the same antibiotic susceptibility-pattern, vary geographically in regard to virulence-factors corresponding to clinical picture and outcome. A connection between the causative agent, the region visited and the clinical picture is easily overlooked if analyses of these bacterial strains are restricted to identification to species level and susceptibility-testing. With this case report, we would like to highlight an emerging syndrome caused by a bacterial species common in Europe, but of a serotype and with specific virulence genes still rare outside of Asia.

ACKNOWLEDGEMENTS

We thank Dr Johan Rydberg at the Department of Clinical Microbiology and Immunology at the University Hospital in Lund, for his support and valuable comments.

REFERENCES

[1] Wang JH, Liu YC, Lee SS, et al. Primary liver abscess due to Klebsiella pneumoniae in Taiwan. Clin Infect Dis 1998; 26: 1434-8.

[2] Tsai FC, Huang YT, Chang LY, Wang JT. Pyogenic liver abscess as endemic disease, Taiwan. Emerg Infect Dis 2008; 14: 1592-600.

[3] Chung DR, Lee SS, Lee HR, et al. Emerging invasive liver abscess caused by K1 serotype Klebsiella pneumoniae in Korea. J Infect 2007; 54: 578-83.

[4] Fang CP, Chang FY, Lee SC, et al. A global emerging disease of Klebsiella pneumoniae liver abscess: is serotype K1 an important factor for complicated endophthalmitis? Gut 2002; 50: 420-4.

[5] Fang CT, Lai SY, Yi WC, Hsieh PR, Liu KL, Chang SC. Klebsiella pneumoniae genotype K1: an emerging pathogen that causes septic ocular or central nervous system complications from pyogenic liver abscesses. Clin Infect Dis 2007; 45: 284-93.

[6] Yu VL, Hansen DS, Ko WC, et al. Virulence characteristics of Klebsiella and clinical manifestations of K. pneumoniae bloodstream infections. Emerg Infect Dis 2007; 13: 986-93.

[7] Yu WL, Ko WC, Cheng KC, Lee CC, Lai CC, Chuang YC. Comparison of prevalence of virulence factors for Klebsiella pneumoniae liver abscesses between isolates with capsular K1/K2 and non-K1/K2 serotypes. Diagn Microbiol Infect Dis 2008; 62: 1-6.

[8] Nadasy KA, Domiati-Saad R, Tribble MA. Invasive Klebsiella pneumoniae syndrome in North America. Clin Infect Dis 2007; 45: e25-e8.

[9] Fang FC, Sandler N, Libby SJ. Liver abscess caused by magA+ Klebsiella pneumoniae in North America. J Clin Microbiol 2005; 43: 991-2.

[10] Frazier BW, Hansen S, Lambert L. Invasive Infection With Hypermucoviscous Klebsiella pneumoniae: multiple cases presenting to a single emergency department in the United States. Ann Emerg Med 2009; 53: 639-42.

[11] Keynan Y, Karlofsky JA, Walus T, Rubinstein E. Pyogenic liver abscess caused by hypermucoviscous Klebsiella pneumoniae. Scand J Infect Dis 2007; 39: 828-30.

[12] Karama EM, Willermain F, Janssens X, et al. Endogenous endophthalmitis complicating Klebsiella pneumoniae liver abscess in Europe: case report. Int Ophthalmol 2008; 28: 111-3.

[13] Gomez C, Broseta A, Otero JR, Chaves F. Primary pyogenic liver abscesses caused by magA+ Klebsiella pneumoniae in Spain. Clin Microbiol News 2007; 29: 100-2.

[14] Lederman ER, Crum NF. Pyogenic liver abscess with a focus on Klebsiella pneumoniae as a primary pathogen: an emerging disease with unique clinical characteristics. Am J Gastroenterol 2005; 100: 322-31.

[15] Struve C, Bojer M, Nielsen EM, Hansen DS, Krogfelt KA. Investigation of the putative virulence gene magA in a worldwide collection of 495 Klebsiella isolates: magA is restricted to the gene cluster of Klebsiella pneumoniae capsule serotype K1. J Med Microbiol 2005; 54: 1111-3.