Liver Abscess Metastatic Syndrome Caused by Hypermucoviscous *Klebsiella* Pneumoniae in a Canadian Patient of Vietnamese Origin

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Abstract

*Klebsiella pneumoniae* liver abscess syndrome (KLAS) is an emerging infection caused by hypermucoviscous strains (K1, rmpA, mgA) with a particular virulence at risk of metastatic dissemination. We describe a case of metastatic KLAS in a Canadian immunocompetent patient of Vietnamese origin who presented with fever and abnormal liver function tests. Imaging studies revealed unique liver and pulmonary abscesses. Blood and liver abscess cultures showed colonies of *K. pneumoniae* with hypermucoviscous phenotype, a K1 serotype and the presence of a rmpA gene confirming biomolecular features of the invasive syndrome. Mostly reported in patients of Asian origin, KLAS has been reported in Canada since 2007. Prompt identification and treatment prevents severe complications such as endophthalmitis, meningitis, lung abscess and spondylodiscitis.

Résumé


**Keyword:** liver abscess metastatic syndrome, hypermucoviscous *klebsiella pneumoniae*
In March 2015, a 34-year-old patient presented at the emergency department of CHU de Québec (CHUL) and was admitted to the internal medicine clinical teaching unit. He reported one week of fever and chills, profuse sweating, temporal headache, loss of appetite, and myalgia. The patient emigrated from Vietnam to Canada 11 years earlier and had not returned there since then. His only recent trip had been to Boston in 2011. He completed a treatment for latent tuberculosis in 2005. He worked as an informatician, had no allergies, and did not take medication. The patient reported no infectious contact, illicit drug abuse, or other at-risk exposures for malaria, viral hepatitis, Q fever, leptospirosis, or brucellosis. The patient did not complain of rash, weight loss, or any abdominal, pulmonary, urinary, or neurological symptoms.

The patient was diaphoretic without rash or jaundice. Physical findings included a blood pressure of 90/58 mmHg, a temperature of 38.9°C, a heart rate of 100 beats per minute, non-painful cervical lymphadenopathy (less than 2 cm), and hepatomegaly (2 cm below costal margin). The remainder of the physical examination was normal without meningeal signs, abdominal mass, tenderness, or ascites. Vital signs stabilized rapidly with treatment and intravenous fluids.

Laboratory studies showed hyponatremia at 131 mmol/L (135–145 mmol/L), white blood cell count at 9.9 × 10⁹, thrombocytopenia at 50 × 10⁹ g/L (150–400 × 10⁹ g/L), abnormal liver and pancreas functions tests: aspartate aminotransferase 0.68 µkat/L (<0.67 µkat/L), gamma-glutamyl transferase 3.57 µkat/L (<0.83 µkat/L), total bilirubin 31 µmol/L (0–21 µmol/L), direct bilirubin 16 µmol/L (<4 µmol/L) amylase 2.99 µkat/L (<1.67 µkat/L), lipase 4.21 µkat/L (< 0.92 µkat/L). Tests for amebiasis, echinococcosis, human immunodeficiency virus, and hepatitis B–C were negative and tests for Epstein-Barr virus and cytomegalovirus showed an ancient exposition.

Abdominal ultrasound, contrast-enhanced computerized tomography (CT) scan, and magnetic resonance imaging (MRI) showed a unique hepatic abscess with a diameter of 3.3 cm in the eighth segment of the liver (Figure 1). A chest CT revealed an asymptomatic pulmonary abscess in the right lower lobe (Figure 2).

Blood cultures and cultures from percutaneous drainage of the hepatic abscess grew Klebsiella pneumoniae with hypermucoviscous phenotype, defined by a positive ‘string test’ (viscous string > 5 mm when bacterial colonies on an agar plate are stretched by an inoculation loop) as shown in Figure 3. The Canadian Science Centre for Human and Animal Health (Winnipeg, Canada) confirmed a K1 serotype and the presence of a rmpA gene confirming biomolecular features of the invasive syndrome.

The pulmonary metastatic lesion was the hallmark for the diagnostic of Klebsiella pneumoniae liver abscess metastatic syndrome (KLAS). Based on in vitro susceptibilities, piperacillin-tazobactam, empirically given, was replaced by ceftriaxone for six weeks of IV ambulatory treatment. The liver abscess, of less than 5 cm, was entirely drained with a single percutaneous needle aspiration, without surgery. Drainage was not indicated for the uncomplicated pulmonary abscess. A cerebral MRI did not show endophthalmitis, abscess or meningeal anomalies.
Lumbar puncture was not initially performed because of the thrombocytopenia.

Followed until July 2015 by CT scans, hepatic and pulmonary lesions disappeared, and liver tests normalized after antibiotic treatment. The patient has not relapsed.

Discussion

*Klebsiella pneumoniae* is a well-known gram-negative bacillus with a thick polysaccharide. Standard clinical diseases are related to *K. pneumoniae* and *K. oxytoca* such as pneumonia, urinary tract infections, abdominal cavity surgery-related infections and several nosocomial syndromes often associated with a history of alcohol abuse or diabetes mellitus.

KLAS is a distinctive infection related to specific subcapsular serotypes and a marked virulence that produces liver abscesses and several distant metastatic infections associated with a history of alcohol abuse or diabetes mellitus.

KLAS is a distinctive infection related to specific subcapsular serotypes and a marked virulence that produces liver abscesses and several distant metastatic infections. Described mostly in the Asian population since the 1980s, cases have been diagnosed in North and South America, Europe, Canada, and Australia. A liver abscess caused by this particular microbiological strain with extra-hepatic infections should be recognized as KLAS. The clinical signs of KLAS are fever, chills, headache, and abdominal pain. Nausea and vomiting occur in about 25% of patients. Half of the patients have jaundice and hepatomegaly. Blood test findings are leucocytosis, increased C-reactive protein, abnormal liver function tests, and thrombocytopenia. Metastatic spread, unusual for most enteric gram-negative bacilli, is a hallmark of the hypermucoviscous Klebsiella pneumonia strain infecting immunocompetent hosts. From 8 to 30% of metastatic disease in KLAS, a case series of 23 patients with metastatic lesions reported mostly endophthalmitis, uveitis, pulmonary abscess, and purulent meningitis.

Hypervirulent *Klebsiella pneumoniae* (hvKP) is directly linked to the hypermucoviscous phenotype, related to the k1, k2 serotypes and the regulator of mucoid phenotype A (rmpA) gene. This feature gives this type of pathogen the ability to produce lethal extra liver infections in non-immunocompromized patients. Numerous virulence factors have been elucidated and implicate the presence of mucoviscosity genes like magA, rmpA, aerobactin but the exact mechanism by which this spreading take place is unknown. Shon et al. observed that hvKP was resistant to complement- and neutrophil-mediated bactericidal activity in a rat subcutaneous abscess model and produces more biofilm than others strains. This capacity increases intestinal colonization. In this case, this patient developed KLAS even if he had been colonized in an endemic area several years ago. Colonization usually happens when there is a disruption of the natural barriers (ascension into the bladder, aspiration into the respiratory tract, gastrointestinal colonizers) but KLAS has been reported in people, like this patient, without evidence of altered mucosal barriers.

Even though the majority of patients with hvKP infection are healthy, there is a significant mortality rate between 3 and 42% in part following necrotizing fasciitis and severe community-acquired pneumonia. Moreover, survivors can suffer appalling consequences such as blindness and neurologic sequelae. Even if most hvKP infections are treatable with common antibiotics, long-term follow-up is necessary because of the high risk of relapsing. Several cases of resistant strains with extended spectrum B-lactamases and carbapenemases have already been reported. All these characteristics could make hvKP one of the next “superbugs.”

Conclusion

To our knowledge, this is the first report of a complete metastatic syndrome by HvKp in Canada, following the first published case of pyogenic liver abscess caused by hvKP in Manitoba in 2007. Compelling questions about hvKP remain unanswered. Its natural course is barely understood. Diagnostic features such as a positive ‘string test’ and capsular serotypes like K1 or K2 are not always distinctive of hvKP. Its capacity for metastatic spread, an unusual trait for an enteric gram-negative, is a worrying characteristic of this strain of Klebsiella that is no longer confined to Asia.

References

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