Multi-Organ Failure Secondary to a *Clostridium Perfringens* Gaseous Liver Abscess following a Self-Limited Episode of Acute Gastroenteritis

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**Patient:** Female, 81

**Final Diagnosis:** Liver abscess

**Symptoms:** Diarrhea • jaundice • vomiting • weakness

**Medication:** —

**Clinical Procedure:** CT scan guided drainage

**Specialty:** Gastroenterology and Hepatology

**Objective:** Rare disease

**Background:** *Clostridium perfringens* is an unusual pathogen responsible for the development of a gas-forming pyogenic liver abscess. Progression to septicemia with this infection has amplified case fatality rates.

**Case Report:** We report a case of an 81-year-old lady with pyogenic liver abscess with gas formation that was preceded by an acute gastroenteritis. The most common precipitating factors are invasive procedures and immunosuppression. *Clostridium perfringens* was unexpectedly isolated in the drained abscess, as well as blood. It is a normal inhabitant of the human bowel and a common cause of food poisoning, notoriously leading to tissue necrosis and gas gangrene.

**Conclusions:** We report a case of gas-forming pyogenic liver abscess and bacteremia progressing to fatal septic shock, caused by an uncommon *Clostridium perfringens* isolate.

**MeSH Keywords:** *Clostridium perfringens* • Gas Gangrene • Liver Abscess, Pyogenic
Background

Pyogenic liver abscess is a critical disease with high morbidity and mortality rates. *Clostridium perfringens* is an unusual pathogen responsible for the development of gas-forming pyogenic liver abscesses. Progression to septicemia with this infection has amplified case fatality rates.

Case Report

We report a case of an 81-year-old Greek lady who was brought to the emergency department with complaints of vomiting and diarrhea starting a few days before. The patient had previous cerebrovascular stroke, for which she was being treated with aspirin and clopidogrel. She was prescribed anti-depressant medications: Paroxetine, Clonazepam, and Olanzapine. She used fesoterodine for overactive bladder. Other medications included atorvastatin, levetiracetam, levothyroxine, and lisinopril and laxatives as needed. She has relevant surgical history of cholecystectomy performed 10 years ago and a remote history of thyroidectomy. On physical exam, she was febrile and icteric. Rest of the exam was unremarkable. Laboratory work revealed leukocytosis and transaminitis. Viral hepatitis panel was negative. Unfortunately, stool culture was not obtained as diarrhea had already resolved promptly at the beginning of hospitalization. The patient had an esophago-gastro-duodenoscopy for evaluation of dysphagia and positive occult blood in stool, which was unrevealing except for antral hernia. Colonoscopy was deferred for outpatient follow-up. Ultrasound abdomen and CT abdomen and pelvis with contrast showed a 2-mm punctate focus of calcification was found within the right hepatic lobe, which may be a small calcified granuloma. A 5-mm low-attenuation lesion was found within the right hepatic lobe, too small to further characterize (Figure 1A–1C). The patient improved and diagnosis of an acute self-limited gastroenteritis was hypothesized without any need for further treatment with antibiotics.

Five days later, her condition worsened and she was brought to the hospital in a state of lethargy, fever, and deep jaundice. She was hypotensive and tachycardic with a BP of 87/29 mm hg. Rapid workup demonstrated leukocytosis with a WBC of 22×10^9/L. In addition, serum creatinine was 1.89 μmol/L and BUN was 17.5 mmol/L. Liver function progressively deteriorated from her recent hospitalization. AST was 318 U/L, ALT 231 U/L, ALP 494 U/L, and bilirubin total 7.2 μmol/L/direct 5.6 μmol/L. Diagnosis of septic shock was made.

Intravenous fluids were initiated, in addition to empirical antibiotics (cefepime, metronidazole, and vancomycin). The patient was transferred to the ICU, and was intubated thereafter for respiratory distress, then was started on mechanical ventilation.

![Figure 1. CT scan of the abdomen and pelvis with intravenous contrast (A) Liver: The liver is mildly enlarged measuring up to 18.1 cm along the mid-clavicular plane. A 2 mm punctate focus of calcification is present within the right hepatic lobe, which may be a small calcified granuloma. A 5 mm low-attenuation lesion was found within the right hepatic lobe, too small to further characterize (Figure 1A–1C). The patient improved and diagnosis of an acute self-limited gastroenteritis was hypothesized without any need for further treatment with antibiotics.](image-url)
Liver abscesses are the most common type of visceral abscess; pyogenic liver abscesses account for 48% of visceral abscesses and 13% of intra-abdominal abscesses [1]. The annual incidence of liver abscess has been estimated at 2.3 cases per 100,000 population and is higher among men than women, indicating it is still an uncommon disease [2]; higher rates have been reported in Taiwan [3].

Bacterial pathogens causing liver abscess are usually mixed and depend on the precipitating cause. *Staphylococcus aureus, Streptococcus pyogenes*, and other Gram-positive cocci are seen after invasive liver procedures like trans-arterial embolization for hepatocellular carcinoma [4]. Hepato-splenic candidiasis can occur in patients after chemotherapy [2]. Diabetes or impaired glucose tolerance are the most common risk factors for Klebsiella primary liver abscess (KLA) [5–8]. Hyperglycemia is an important factor for GFPLA (gas-forming pyogenic liver abscess). Poor control of DM leads to neutrophil dysfunction and chemotaxis failure. Local tissue damage is caused by gas-forming bacteria and compounded by the diabetic micro-angiopathy [9]. Our reported patient had uncontrolled hyperglycemia that could be from stressful the disease process. Uncommon organisms could cause liver abscess, like tuberculous [8] and amebic liver [11] in endemic areas.

Mixed infections may be found in 14–55% of cases of routine pyogenic liver abscesses, but KLA cases are almost uniformly mono-bacterial [14,17]. Prior to the era of rapid patient assessment and expeditious surgery, appendiceal pathology was the most common source of liver abscesses [12,13]. In the modern era, biliary disease is the most common etiology [12,15]. Other potential sources include penetrating trauma, distant sources (i.e., outside the abdomen), and contiguous spread from lung, kidney, colon, or stomach. Still, many are deemed cryptogenic (40–99%) [12,14].

GFPLA (gas-forming pyogenic liver abscess) accounts for 7–32% of PLA (pyogenic liver abscess) cases. It was reported in one of the studies that of the 69 patients included who had PLA, 22 had GFPLA, and 21 had DM. *Klebsiella* primary liver abscess (KLA) [5–8]. Hyperglycemia is paired glucose tolerance are the most common risk factors for hepatocellular carcinoma [4]. Hepato-splenic candidiasis can occur in patients after chemotherapy [2]. Diabetes or impaired glucose tolerance are the most common risk factors for KLA cases [5–8]. Prior to the era of rapid patient assessment and expeditious surgery, appendiceal pathology was the most common source of liver abscesses [12,13]. In the modern era, biliary disease is the most common etiology [12,15]. Other potential sources include penetrating trauma, distant sources (i.e., outside the abdomen), and contiguous spread from lung, kidney, colon, or stomach. Still, many are deemed cryptogenic (40–99%) [12,14].

Most common pathways through which bacteria can reach the liver and start forming the abscess is through the portal venous system, hepatic artery, biliary tract, and direct spread. Among these, the biliary tract is the most frequent source, occurring in 60% of PLA cases [14].

A report of 52 cases treated at a single institution showed that the pathogenesis of liver abscesses in one group of patients was mainly malignant biliary obstruction and spontaneous or iatrogenic necrosis of primary hepatic neoplasms with superimposed bacterial infection. In contrast, the pathogenesis of the abscesses in the second group of patients included portal venous suppuration secondary to colorectal cancer,
post-gastrectomy for gastric cancer, and hepatic artery seeding with uncertain infectious nidi following systemic anticancer treatment [16]. Pyogenic liver abscess can be a presentation of underlying hepato-pancreato-biliary malignant disease at a pre-terminal stage and carries a grave prognosis. In contrast, patients who have pyogenic liver abscess and non-malignant disease have a favorable outcome [16].

As regards our case, no malignancy was identified from history or on work-up done, including the CT abdomen. Unfortunately, colonoscopy was not done on either admission to exclude malignancy.

The major presenting symptoms were fever, chills, and abdominal pain; however, a few patients presented with only altered mental status, dizziness, or general malaise [17].

Recently, the introduction and refinement of percutaneous drainage techniques have dramatically improved the treatment success rate [18]. However, these refined techniques seemed not to influence the outcome of critically ill patients with PLA in one study [19]. Given that most patients in that study had severe sepsis, treatment not only should have focused on a local inflammation or infection, but also should have regulated a systemic, complex immunologic reaction [19].

Among patients with PLA, the incidence of septic shock and bacteremia is higher in GFPLA patients compared to non-GFPLA patients. Septic shock is noted in 32.5% of patients with GFPLA and in 11.7% of patients with PLA patients [20]. GFPLA also has a high fatality rate, which is around 27.7% to 37.1%. GFPLA also ruptures easily because of tissue invasion and fragility of abscess wall, and further gas formation increases the internal pressure of the abscess. In previous studies, spontaneous rupture of PLA had occurred in 7.1–15.1% of the cases and 20 of 24 patients with ruptured PLA had GFPLA [17,21,22].

Conclusions

Our reported case is a presentation of gas-forming pyogenic liver abscess with mixed bacteria, but we wish to emphasize *Clostridium perfringens*, which is rare without an invasive procedure or immunosuppression. Bacterial seeding could happen from the GI tract to the liver, causing liver abscess. Our patient developed septic shock with multi-organ system failure and with no signs of disseminated intravascular coagulopathy or intra-vascular hemolysis.

Learning objective

Liver abscesses can be a serious life-threatening condition. Treatment should be very early and aggressive, which usually includes appropriate antibiotics and surgical drainage. *Clostridium perfringens* is one of the uncommon organisms involved.

Statement

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Clostridium perfringens is a Gram-positive, rod-shaped, anaerobic, spore-forming bacterium of the genus *Clostridium* [20]. It is part of the intestinal normal flora in humans and known to cause tissue necrosis and gas gangrene by producing alpha toxin [23]. It is a common cause of food-borne diseases in the USA [24].

After a literature review, only a few cases were found to have gas-forming pyogenic liver abscess secondary to *C. perfringens*. One of them occurred spontaneously in an immune-competent patient with intravascular hemolysis [25]. Another case was reported after pancreatectomy [26]. The third one was reported after laparoscopic cholecystectomy [27].
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