Imaging of Biliary Infections

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Abstract

Biliary tract infections cover a wide spectrum of etiologies and clinical presentations. Imaging plays an important role in understanding the etiology and as well as the extent of disease. Imaging also plays a vital role in assessing treatment response once a diagnosis is established. This article will review the imaging findings of commonly encountered biliary tract infectious diseases.

Keywords

- biliary infections
- cholangitides
- parasites
- immunocompromised
- echinococcal

Infections of the biliary tree can have a myriad of clinical and imaging manifestations depending on the infectious etiology, underlying immune status of the patient and extent of involvement.1,2 Bacterial infections account for the vast majority of infectious cholangitides though parasitic, fungal, and viral infections can also affect the biliary system. Clinically, the range of clinical manifestations of bacterial infections extends from an acute medical emergency with clinical symptoms of right upper quadrant (RUQ) pain, fever, and jaundice to indolent and chronic, low-level infections.

Bacterial Cholangitides

Bacterial cholangitis is a clinical syndrome that arises when infected bile enters the circulatory system.3–5 Cholangitis typically manifests as fever, jaundice, and RUQ pain (Charcot’s triad) and may or may not result in biliary sepsis. Reynolds pentad describes the combination of fever, jaundice, and RUQ pain with hypotension and confusion/lethargy. Commonly encountered microorganisms include Escherichia coli, Enterococcus, Klebsiella, and Pseudomonas, as well as anaerobes.5,6 The combination of microbiological contamination of bile in the setting of increased biliary pressure can lead to the clinical manifestation of acute cholangitis.5,7 Elevated biliary pressures directly impact the severity and mortality of acute cholangitis.9 Interruption of normal flow of bile into the duodenum can lead to a cascade of changes to the host immune defense mechanisms of chemotaxis and phagocytosis.7 The resultant lack of bile and secretory immunoglobulin A from the gastrointestinal tract lead to changes of the normal bacterial flora, disrupted mucosal integrity, and diminished inactivation of endotoxins resulting in sepsis.

Imaging Features of Acute Cholangitis

Diagnostic cross-sectional imaging of the liver and biliary tree is necessary for evaluation of the patient with clinical features of cholangitis.9–11 Ultrasound (US) is a valuable diagnostic tool for the evaluation of the liver and bile ducts and is associated with high sensitivities and specificities for evaluation of biliary obstruction.10 Although it is widely available, noninvasive, and cost-effective, it has limited evaluation of the distal common bile duct, a common site for stones. Computed tomography (CT) and magnetic resonance imaging (MRI) offer improved anatomic resolution and assessment of extrahepatic structures than US and therefore are more frequently utilized to assess biliary obstruction and its potential causes (►Fig. 1).12–16 Biliary obstruction can be diffuse or only involve segmental or centrally located ducts and is frequently associated with enhancement of the bile duct wall.9 CT offers the ability to assess the extent of obstruction and parenchymal changes that may be associated with cholangitis.17 Arai et al

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compared hepatic parenchymal patterns on dynamic CT of patients with cholangitis to those without and found that inhomogeneous enhancement was significantly higher in patients with cholangitis and that this finding resolved following treatment.

Parasitic Infections

Parasitic infections have a spectrum of clinical manifestations depending on their underlying etiology. When combined with bacterial infections, there is much overlap with bacterial cholangitis. Eosinophilia is a common clinical manifestation of parasitic infections and can be helpful in the diagnostic evaluation of suspected biliary inflammation.

Fascioliasis

_Fasciola hepatica_ is the causative parasite responsible for fascioliasis. _Fasciola hepatica_ is a liver fluke that is commonly encountered in underdeveloped countries. The pathway to human infection is somewhat circuitous that results from accidental ingestion of vegetables or water sources that are infected with metacercariae. Once ingested, the metacercariae penetrate the upper gastrointestinal tract and eventually migrate into the peritoneal cavity. From the peritoneal cavity, they penetrate the liver capsule and infect the liver where they can remain for extended periods of time. CT imaging during the hepatic phase shows irregular areas of hypoattenuation, often in the periphery of the liver (Fig. 2). Eventually, the flukes enter the biliary tree where they cause inflammation, ductal thickening, and localized obstruction.

Echinococcal Infections

Echinococcosis results from _Echinococcus multilocularis_ and _Echinococcus granulosus_ infestation. Both parasites are cestodes in which foxes and dogs act as definitive hosts. Human infection arises from ingestion of vegetables contaminated with parasitic eggs. After ingestion, embryos release eggs into the intestine, which then penetrate the bowel wall and are delivered to the liver via the portal venous system where they encyst within the hepatic parenchyma. Imaging findings of _E. granulosus_ hepatic infections depend on the different growth stages of the hepatic cyst. Early infections are characterized by a simple cyst on CT or MRI. The development of daughter cysts within the primary cyst gives a characteristic spoke wheel pattern (Fig. 3). Without therapy, the echinococcal cysts can enlarge and extend into the biliary tree leading to filling defects and obstruction.

Imaging features of _E. multilocularis_ differ from _E. granulosus_. _Echinococcus multilocularis_ hepatic infection manifests as a multilocular cystic structure that induces intense fibrosis and that can extend out of the liver into surrounding structures, including the biliary tree and hepatic vessels.

Schistosomiasis

The trematodes _Schistosoma mansoni_ and _Schistosoma japonicum_ are venoinvasive parasites that infect humans through contact with contaminated water. The parasites are capable of cutaneous penetration after which they migrate to pulmonary parenchyma and eventually invade branches of
the portal venous system via branches of the superior and inferior mesenteric veins. Although in the portal veins, the eggs grow to sexual maturity where male and female worms mate before migrating to the bowel wall. Some eventually infect the liver via the portal vein and once deposited within the liver, the eggs activate an immunologic response that stimulates granulomatous response and eosinophilia that can lead to changes of the biliary tree that manifests as fibrosis. Posterior acoustic shadowing is not a typical sonographic feature. Although endoscopic retrograde cholangiopancreatography is the diagnostic gold standard, cross-sectional imaging with CT or MRI/magnetic resonance cholangiopancreatography (MRCP) is useful adjuncts to US. CT shows hyperattenuating linear structures within the common bile duct or gallbladder. MRCP findings include low signal intensity on T2-weighted imaging.

**Biliary Infections in the Immunocompromised**

Immunocompromised patients are at risk of hepatobiliary tract infections. The human immunodeficiency virus can involve the hepatobiliary tree, especially in individuals with severe immunosuppression (CD4 counts < 100/mm^3). A wide variety of causative organisms have been associated with acquired immune deficiency syndrome (AIDS)-related cholangiopathy, including Cryptosporidium, Cryptococcus, cytomegalovirus (CMV), Mycobacterium, and histoplasmosis. Clinically, AIDS-related cholangiopathy presents as RUQ pain, usually in the setting of serologic manifestations of liver function abnormalities. Imaging features of AIDS-related cholangiopathy resemble findings of sclerosing cholangitis. Cholangiographic abnormalities often demonstrate a beaded appearance of the bile ducts with segmental strictures. MRCP shows characteristic ductal abnormalities such as papillary stenosis, intraductal strictures, and extrahepatic long-segmental strictures or a calculous cholecystitis.

Patients who have undergone liver transplant represent another group at risk of biliary infections. Cholestatic liver disease is the most commonly encountered problem following orthotopic liver transplant (OLT) and can result from several causes. Viral infection, rejection, or ischemic injury to the bile ducts have been reported in OLT patients. Percutaneous biopsy is often necessary to elucidate the etiology of cholestasis, especially when ischemic, rejection, or drug effects are suspected. However, there can be significant overlap with obstructive and nonobstructive causes of cholestatic liver injury OLT patients such that an accurate diagnosis often requires cross-sectional imaging. MRCP and transhepatic cholangiography are more sensitive than US for the assessment of biliary stasis in OLT patients. Liver transplant recipients are also prone to opportunistic viral infections, of which CMV is most common. CMV-infected immunocompetent individuals are usually asymptomatic. In the immunosuppressed liver transplant recipient, however, CMV infection can present with
severe clinical symptoms including fever, mononucleosis, gastritis, colitis, abdominal pain, and diarrhea. Liver involvement of CMV includes hepatitis with a cholestatic presentation based on liver function abnormalities. Biopsy is often necessary to excluded tissue rejection. In addition, biologic markers of CMV pp65 or CMV DNA by polymerase chain reaction are detectable at very early stages of infection. To combat the effects of CMV infection, aggressive antiviral prophylaxis with valganciclovir or ganciclovir for up to 3 months following liver transplant is often employed. This approach, however, may result in delayed onset CMV disease once antivirals are discontinued. Another approach is to carefully monitor serologic CMV viral loads or pp65 antigenemia for signs of infection and initiate antiviral therapy. Ganciclovir is a highly effective antiviral for CMV infections and must most liver transplant recipients show good treatment response after initiation of therapy.

**Conclusion**

Infections of the biliary tree comprise a wide range of etiologies and clinical manifestations. The role of diagnostic imaging is critical to help identify its causes, to determine the extent of disease, and to assess for infection-related complications.

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