Imaging features of hepatic infections

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Learning objectives

The purpose to describe the different causes of hepatic infections and review their imaging findings.

Background

Liver infections include bacterial, viral, fungal, and parasitic diseases. Some are common entities, such as viral hepatitis, hepatic abscess and hydatid disease. Other diseases are more rare and difficult to diagnose, presenting a lower level of suspicion, such as tuberculosis or fascioliasis.

Background

Imaging techniques having a high accuracy, plays a crucial role in the diagnosis of hepatic infections. Even so, the imaging findings are generally not specific, with differential diagnostic difficulties with hepatic cysts and necrotic tumors, which is why the clinical context is extremely important.

The radiologist has an important role beyond diagnosis, monitoring and even treatment with percutaneous image-guided drainage.

Imaging findings and procedure details

Hepatic Abscess

Liver abscess

- Pyogenic
- Amoebic
- Fungal

Microabscess - when smaller than 2cm.

Liver abscess

Although imaging findings are generally non-specific, there are signs that suggest the presence of an abscess:

- Cluster sign;
- Wall enhancement;
- Double target sign;
- Perilesional edema;
- Gas.

Cluster sign

Grouping of small lesions with low attenuation or high T2 signal intensity, that gradually coalesce into a larger. Especially suggestive of a pyogenic abscess.



Liver abscess at contrast-enhanced CT

• Grouping of low attenuation lesions in the liver - cluster sign.



Liver abscess at contrast-enhanced CT - same patient, three weeks later. Coalescence of the small lesions into a larger one.

Wall enhancement

Identified on contrast studies, virtually diagnostic of an abscess in the appropriate clinical setting.



Contrast-enhanced CT - Liver abscess

• Low attenuation liver lesion with evident wall enhancement.

Double target sign

Zone of external low attenuation, besides the enhancement of the wall.



Contrast-enhanced CT - Amoebic abscess.

 Thick enhancing wall (black arrow) and outer hypodense rim (white arrow) -Double target sign.

Perilesional edema

Helps differentiate an abscess from a benign cystic lesion. A minority of malignant lesions may also show this abnormality.

в A C D

Liver abscess at MRI.

 Irregular lesion, hypointense at T1 (A), hyperintense at T2 (B), with defined wall that enhances with contrast (C - precontrast T1 with fat suppression, D - portal phase). Note perilesional edema (B arrow).



A classic and specific sign but only present in a minority of cases. It's more common in *Klebsiella* and anaerobic infections.



Contrast-enhanced CT. Polymicrobial abscess on the caudate lobe with presence of gas (arrow). Identified agents were Streptococcus viridans and Bacteroides (anaerobe).

Liver abscess

- Pyogenic
- Amoebic
- Fungal

The most common abscesses in developed countries. Usually caused by biliary obstruction or hematogenous spread from a intestinal infection.

<u>Causative agent</u>: *Escherichia coli*, other gram-negative bacteria, anaerobes. Half are polymicrobial.

<u>**Clinical and laboratorial findings:**</u> Pain in the right upper quadrant, fever and jaundice. May be clinically occult. Leucocytosis and abnormal liver function. Blood cultures are positive in half.

Imaging Findings:

- Cross sectional imaging techniques detect more than 90% of pyogenic abscesses. Multiple in 50% of cases, with a coalescent pattern with gram negative bacterial infection.
- Appearance varies with pathologic stage. In the more acute stage they have an indistinct appearance, progressing to a more welldefined, cystic appearance, as necrosis and liquefaction ensues.

Imaging Findings:

- US: most often a complex cystic lesion, but may also appear solid.
- CT: most commonly a thick-walled lesion with low attenuation.
- MRI: variable signal intensities on T1 and T2, depending on protein content.

Accompanying abnormalities: hepatomegaly, elevation of the right hemidiaphragm, pleural effusion and basal lung atelectasis or infiltrates.



Liver abscess. Complex cystic appearance at US, with thickened wall.



Two cases of liver abscesses with hypoechoic solid appearance on US.

- Special cases:
 - Staphylococcal infection: Multiple abscess with a milliary pattern of distribution, similar to fungal abscesses.

Klebsiella pneumoniae abscess:

- Gas is more common.
- Septal breakage sign arborizing patterns of septa in the abscess cavity.
- Turquoise sign numerous septal breakages.
- Hair-ball sign tangled pattern of hairline content in the abscess fluid.



Contrast-enhanced CT. Klebsiella abscess in the right liver lobe. Discontinuous septa (arrow) - septal breakage sign.

Treatment:

- Medical therapy alone may be successful in lesions with less than 3-5cm of diameter.
- Image-guided drainage combined with antibiotics is the treatment of choice, even if multiple or plurilocular lesions, with success rates higher than 90%. Drainage is considered more effective than aspiration, although it has some disadvantages, like catheter blockage or displacement.
- Some patients may need surgical drainage.



Percutaneous CT-guided drainage of the abscess

Amoebic abscess

Causative agent: *Entamoeba histolytica*. It is the most frequent cause of liver abscess worldwide. The most common extra-intestinal complication of amebiasis.

Clinical and laboratorial findings: Patients are sicker than with pyogenic abscess. Mucous diarrhoea. Laboratory results similar to pyogenic infection. Serology positive > 90%.

Amoebic abscess

Imaging findings: Lesions are often solitary and near the liver capsule. An abscess with a thick wall and perilesional edema may suggest this diagnosis.

Combination with diaphragmatic disruption is quite specific. Other associated findings include thickening of the wall of the cecum, pleural and perihepatic fluid.

Amoebic abscess

 Treatment: Medical therapy is highly effective. Aspiration only in a minority of cases.



Contrast-enhanced CT - Amoebic abscess. Thick enhancing wall (black arrow) and outer hypodense rim (white arrow) - Double target sign.



Contrast-enhanced CT - Amoebic abscess _ Double target sign.



Causative agent: Candida albicans is the most common causative organism.

Clinical and laboratorial findings: Complication of immunosuppression and hematologic malignancies. Patients present with fever that does not respond to large spectrum antibiotics.

Fungal abscess

Imaging findings: Typically, there are multiple microabscesses in the liver and spleen. On US, the lesions may have a "wheel-within-wheel" or "bulls- eye" appearance, but uniformly hypoechoic nodules is the most common pattern. After treatment, foci of scarring and calcification may persist.

Treatment: Antifungal agents.


Contrast-enhanced CT. Multiple hepatic microabscesses in a immunosuppressed patient with leukemia.



Same patient - Liver US shows small hypoechoic nodules.

Parasitic disease

Hydatid Disease:

 Endemic in the Mediterranean and other sheep-raising countries. Acquired by ingestion of contaminated food or contact with dogs. Liver affected in 55-75% of cases.

Hydatid	cyst	is	composed	d of	three	layers:	
-			outer			pericyst	
-						ectocyst	
-	inner	endoc	cyst o	r ۽	germinal	layer.	
Maturation of a cyst is characterized by development of daughter							
cysts.							

Hydatid Disease

Causative agent: *Echinococcus granulosus*

Clinical and laboratorial findings:

- Generally asymptomatic, expansion of a cyst may cause pain and its rupture can lead to anaphylaxis.
- Eosinophilia. Serology tests have a wide range of sensitivity and specificity.

Hydatid Disease

- Imaging findings:
 - Variable
 - There are various classification systems:
 - Gharbi classification
 - OMS classification

WHO classification of cystic echinococcosis (CE) / Gharbi classification International consensus classification

CE 1 / Type 1	Unilocular simple cysts	Active
CE 2 / Type 3	Multivesicular multiseptated cyst	Active
CE 3 / Type 2	Floating membrane (water lily sign)	Transitional
CE 4 / Type 4	Heterogeneous degenerative contents	Inactive
CE 5 / Type 5	Thick calcified wall	Inactive



Hydatid cysts.

- A Type 1 hydatid cyst at US.
- B Another patient. Simple appearing hydatid cyst at contrast-enhanced CT, except for a defined wall. This cyst was however complex at US.



Liver hydatid cyst.

• A - Type 2 cyst at US, with detachment and colapse of the germinal layer - water lily sign.

MRI:

- B Pericyst is hypointense and matrix is hyperintense at T2.
- C Pericyst and matrix are hypointense at T1.
- D After contrast injection, portal phase, there is no internal enhancement.



Hyperechoic liver lesion, solid appearing. At CEUS (contrast-enhanced ultrassound), there is no internal enhancement. Case of type 4 hydatid cyst.



Massively calcified hydatid cysts. A - Type 5 lesion at US. B - CT.



Massively calcified hydatid cysts - Type 5

Hydatid Disease

US:

There is an additional sign - the snowflake sign - caused by freefloating protoescoleces or hydatid sand.

CT:

 50% have wall calcifications and 75% have detectable daughter lesions.



Contrast-enhanced CT. Cystic lesion of the liver with wall calcifications. Case of hydatid cyst.



Hydatid cyst

- A Pre-contrast CT axial plane.
- B Contrast-enhanced CT, coronal reformat. Presence of smaller hypodense lesions inside the cyst (arrow) - daughter cysts.

Hydatid Disease

MRI:

Best demonstrates the pericyst, the matrix and the daughter cysts.

- Pericyst hypointense rim both at T1 and T2.
- Matrix (hydatid sand) hypointense at T1 and hyperintense at T2.
- Daughter cysts hypointense relative to the matrix at T2.

Hydatid Disease

Treatment:

 Includes drainage with instillation of a scolicidal agent, complemented with medical therapy with albendazole.

Echinococcus Multilocularis Cyst

Endemic to central and northern Eurasia and North America, less common but more invasive that hydatid disease. Caused by direct contact with definitive hosts (foxes) or ingestion of contaminated food.

Echinococcus Multilocularis Cyst

Imaging Findings:

- Multilocular alveolar cysts scattered throughout the liver, the most frequent site of involvement.
- Hailstorm pattern (US) multiple echogenic nodules with irregular and indistinct margins.
- In latter stages, central calcifications develop on regions of necrosis.
- Hepatic hilar involvement up to 50% of cases, causing dilatation of intra-hepatic ducts and atrophy by hypoperfusion of affected liver segments.



Contrast-enhanced CT of Echinococcus multilocularis infection. Multiple lowattenuation lesions scattered throughout the right liver lobe.

Echinococcus Multilocularis Cyst

Treatment:

 Being highly invasive, patients are usually subjected to wide resections or liver transplant.

Fascioliasis

Hepato-biliary infection, suspected in endemic regions or recent travellers to Central and South America, South-eastern Asia and the Mediterranean.

Acquired by ingestion of contaminated water or aquatic plants. The parasites exit the intestine, migrate through the peritoneal cavity and invade the liver.

Causative agent: Fasciola hepatica

Fascioliasis

Clinical and laboratorial findings

Characterized by two phases:

- Hepatic phase: 1-3 months after ingestion, parasites migrate through the liver capsule and parenchyma, toward the biliary ducts. Right upper quadrant pain, fever, urticaria, hepatomegaly, eosinophilia.
- Biliary phase: intermittent right upper quadrant pain, with or without cholangitis and cholestasis.

Serology and parasitology tests are useful for confirmation.

Fascioliasis

Imaging findings:

 Multiple small nodular and branching linear lesions, predominantly subcapsular, which are hypoechoic on US, hypodense on CT, T1 hypointense and T2 hyperintense.

Other abnormalities:

Peri-hepatic lymphadenopathies, subcapsular haemorrhage and hepatic necrosis in the hepatic phase, bile duct dilatation and wall thickening in the biliary phase. The worm itself may be identified.

Treatment:

Medical therapy.



Fascioliasis at contrast-enhanced CT

- A Coronal reformat, subcapsular nodular hypodense lesions.
- B Axial plane, linear distribution of lesions (arrow).



US of the gallbladder reveals a small tubular image - the parasite.

Schistosomiasis

- Endemic to tropical and subtropical areas.
- These parasites penetrate the skin and migrate to the mesenteric vein. Their eggs may be carried to the liver.
- Imaging findings reflect periportal fibrosis, with marked enhancement after contrast.
- *S. japonicum* may cause a turtleback appearance on CT.

Other Infections

Hepatic Tuberculosis

A frequent infection worldwide, tuberculosis affects the liver in different ways. Generalized miliary tuberculosis: most common, hepatic involvement usually not detected by imaging studies, ocasionally low attenuation foci scattered throughout the liver Localized liver tuberculosis: either a disseminated nodular form or a focal tuberculous abscess or tuberculoma. In the healing stage, calcifications are detected. Imaging studies are non-specific and almost all patients require biopsy for diagnosis.



Abdominal contrast-enhanced CT in a patient with pulmonary tuberculosis. Scattered low attenuation lesions in the liver and spleen.



Contrast-enhanced CT in a immunocompromised patient. Hypodense nodule with central necrosis (arrow) -Tuberculous abscess.



Contrast-enhanced CT. Coarse liver calcification in the right lobe in a patient with previous hepatic tuberculosis.

Viral Hepatitis

Causative agents: hepatitis viruses (usually A, B, C, D, E), herpes, adenovirus. Although a common liver infection, it has often unremarkable or non-specific imaging findings in the acute phase, including hepatomegaly, peri-portal edema and lymphadenopathies. Ocasionally starry night pattern at US. Radiology has a more important role in excluding differential diagnosis. Chronic viral hepatitis resemble early stage liver cirrhosis.

HIV infection

Predisposes to opportunistic infections like *Mycobacterium avium* complex, *Pneumocystis* and cytomegalovirus.



Contrast-enhanced CT. Multiple hepatic microabscesses in a immunosuppressed patient with HIV.



Contrast-enhanced CT. Multiple hepatic microabscesses in a immunosuppressed patient with HIV

References: Medical Imaging, Faculty of Medicine of Coimbra, University Hospital of Coimbra – Coimbra, Portugal
Cat-scratch disease

Caused by *Bartonella henselae*. Dissemination occurs in 5-10% of cases with granuloma formation in the liver.

Conclusion

The role of the radiologist is essential in cases of liver infections, from diagnosis to, monitoring and treatment.

In addition to the characteristic aspects of imaging, in cases where these are less specific, the clinical context plays a fundamental role in the diagnostic aid.

References

- Benedetti NJ, Desser TS, Jeffrey RB. Imaging of hepatic infections. Ultrasound Q. 2008;24:267-78.
- Bonder A, Afdhal N. Evaluation of liver lesions. Clin Liver Dis. 201216:271-83.
 Bricault I. Infections of the right hypochondrium. Diagn Interv Imaging. 2012;93:453-65.
- Koç Z, Ulusan S, Tokmak N. Hepatobiliary fascioliasis: imaging characteristics with a new finding. Diagn Interv Radiol. 2009;15:247-51.
- Mortelé KJ, Segatto E, Ros PR. The infected liver: radiologic-pathologic correlation. Radiographics. 2004;24937-55
- Mortelé KJ, Ros PR. Cystic focal liver lesions in the adult: differential CT and MR imaging features. Radiographics. 2001;21:895-910.