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**Collagenous nodule mixed simple cyst and hemangioma coexistence in the liver**

Zheng ZJ *et al.*Atypical masses in the liver

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**Abstract**

A 20-year-old female patient presented with two masses located in the left liver. In this patient, a computed tomography (CT) scan revealed a hypodense mass and a second well-defined mass with a calcified nodule in the left hepatic lobe. No enhancements were apparent in or around the masses. A laparotomy was performed due to the patient’s symptoms, namely the atypical CT findings and a risk of rupture of the subcapsular lesion. The operation revealed two masses in the left hepatic lobe and a left liver resection was subsequently performed. One of the masses involved segment III and the other mass was located in segment IV. The histopathologic findings supported a diagnosis of collagenous nodule mixed simple cyst and hemangioma. A diagnosis of collagenous nodule mixed simple hepatic cyst is extremely rare and radiologically mimics a teratoma, hepatolithiasis, parasitic cyst, or hemangioma. Although hepatic hemangiomas are the most common benign tumors found in the liver, the present case showed atypical radiographic features.

**Key words:** Collagenous nodule; Hepatic hemangioma; Liver; Mass; Simple cyst

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**Core tip:** This case represents a rare presentation of a patient with multiple liver masses. A definitive diagnosis depends on the histopathologic findings. Collagenous nodules are a rare clinicopathologic finding in the liver. This paper discusses the [pathogenesis](javascript:void(0);) of a collagenous nodule found in the liver. Although hepatic hemangiomas are the most common benign tumor in the liver, this case showed atypical radiographic features.

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**INTRODUCTION**

Collagenous nodule mixed simple hepatic cyst is an extremely rare mass that has not been reported previously in the publically available literature. Hyaline degeneration of the connective tissue is the predominant pathologic feature of a collagenous nodule. This form of degenerative alteration has been found in atrophic breast tissue, scar tissue, atheromatous plaques, and organized tissue. In contrast, hepatic hemangiomas are the most common benign tumor in the liver[1]. Diagnosis of a hepatic hemangioma is generally straightforward and accurately diagnosed by the various imaging modalities. The case described here, however, presented with atypical radiographic features. The diagnosis of hepatic hemangioma was considered due to the pathologic findings. Here, we present a case of hepatic hemangioma in conjunction with a collagenous nodule mixed simple hepatic cyst in a 20-year-old female patient.

**CASE REPORT**

A 20-year-old female patient presented with a two-month history of recurrent right epigastric pain. She had no history of exposure to parasites. Serum alpha-fetoprotein and eosinophil levels were normal. Serological testing for hydatid antibody was negative. A computed tomography (CT) scan revealed a hypodense mass and a second well-defined mass with a calcified nodule in the left hepatic lobe. Enhancements were not apparent in or around the masses (Figure 1). The chief radiographic differential diagnosis included teratoma, hepatolithiasis, parasitic cyst, or hemangioma. Both the patient and her relatives consented to the laparotomy. The operation revealed two masses in the left hepatic lobe and a left liver resection was performed. One of the masses involved segment III and another was localized to segment IV. The segment III mass was a tender, rufous and subcapsular mass (diameter: 9 cm) while the mass in segment IV was described as a cystic mass (diameter: 5 cm). A hard, gray-white, translucent, and inelastic nodule (diameter: 2 cm) was observed in the inner wall of the cystic mass (Figure 2, blue arrow). Histologically, the segment III mass had dilated vascular channels, fibrous stroma, and dark venous blood. The segment IV mass revealed thickened, banded and sheet-like collagenous fibers (Figure 3). No parasites were observed in the specimen obtained from the cystic mass. After a histopathologic analysis, the diagnosis of collagenous nodule mixed simple cyst and hemangioma was confirmed. The patient made an uneventful recovery.

**DISCUSSION**

Prior to surgery, high-density nodules were classified as hepatic calcifications. Bezerra *et al*[2] retrospectively analyzed 1362 consecutive CT scans of the abdomen and identified intrahepatic calcifications in 3.6% (49/1362) of the patients. Calcifications that develop within a hepatic mass are either produced by the tumor itself or, more commonly, represent dystrophic calcification secondary to necrosis and/or hemorrhage within the mass[3]. Calcifications may be found in inflammatory hepatic lesions and in benign and malignant liver masses[4]. The clinical appearance of a cystic mass with a high-density nodule is nonspecific and can mimic a teratoma, hepatolithiasis, parasitic cyst, or hemangioma. Although supporting evidence was lacking, a diagnosis of hepatic hydatidosis was primarily considered, as it is endemic in the Sichuan Province. The patient presented with no personal history of potential exposure to parasites, a normal level of eosinophils, and a negative hydatid antibody test. Thus, the patient underwent a left liver resection to avoid a misdiagnosis that may have potential deleterious consequences. A pathologic examination subsequently confirmed that the mass was a collagenous nodule mixed simple hepatic cyst, which has not been reported previously.

The predominant pathologic feature of a collagenous nodule is hyaline degeneration of connective tissue. This kind of degenerative alteration can be found in atrophic breast tissue, scar tissue, atheromatous plaques, and organized tissue. The cause of a collagenous nodule in a liver simple cyst is unclear. It is possible that the cyst was infected or bleeding, after which the pus or blood clot was replaced by granulation tissue, which is known as organization. This granulation tissue may then have been replaced by fibrous connective tissue. The process is analogous to wound healing and scar tissue formation. In this case, the collagenous nodule appeared as a high-density shadow on a CT scan. There are no specific radiologic features that differentiate a collagenous nodule from calcification. A definitive diagnosis is dependent on histopathologic findings. Had the diagnosis of a collagenous nodule mixed simple hepatic cyst been considered preoperatively, conservative observation may have been an option.

Cavernous hemangiomas are the most common benign tumor in the liver, with a reported incidence ranging from 1% to 20%[1]. This lesion occurs more frequently in women than in men. A hepatic hemangioma originates from the proliferation of vascular endothelial cells and enlarges by ectasia rather than hyperplasia[5]. Hemangiomas are usually detected incidentally while imaging for other purposes. The classic enhanced CT scan is early peripheral enhancement of the mass, followed by centripetal fill-in of the contrast medium with persistent enhancement on delayed phase images[6]. In this case, the radiographic finding was atypical, with a lesion that showed a hypodense mass without enhancement. Slow flow in the central sinusoids, central fibrosis, central thrombosis, and hemorrhage may have contributed to the nonenhancement of the hemangioma[7]. Knowledge of the broad spectrum of atypical hepatic hemangiomas is critical and can help avoid most diagnostic errors[8]. Atypical hemangioma findings include a non-enhancing central core and septa (usually seen in large hemangiomas), calcification, fluid–fluid level, and multicystic hemangiomas[9]. Hepatic hemangiomas that are larger than 4 cm in diameter are classified as giant hemangiomas[10]. The surgical indications for liver hemangiomas are Kasabach–Merritt syndrome, symptoms of organ compression that have been well established (excluding other diseases), and traumatic or spontaneous rupture[11]. Asymptomatic or minimally symptomatic patients can be safely observed[12]. Previous studies have demonstrated that nonoperative management of a giant liver hemangioma is safe in most patients, indicating that the size of the lesion is not a criterion for surgery[13].

In conclusion, this case represents a rare presentation of a patient with multiple liver masses. Surgical resection was performed due to the symptoms, uncertain diagnosis as a result of atypical CT findings, and a risk of rupture of the subcapsular lesion. Liver resection is a safe procedure. In our case, the preoperative diagnosis remained uncertain, and the surgical resection was justified. The management of liver masses should be based on a balance between the risks that might be prevented by surgical resection and the estimated risks of surgery.

**COMMENTS**

***Case characteristics***

A 20-year-old female patient presented with a two-month history of recurrent right epigastric pain.

***Clinical diagnosis***

Tenderness in the upper abdomen.

***Differential diagnosis***

Teratoma, hepatolithiasis, parasitic cyst, or hemangioma.

***Laboratory diagnosis***

Serum alpha-fetoprotein and eosinophil levels were normal and serologic testing for hydatid antibody was negative.

***Imaging diagnosis***

A computed tomography scan revealed a hypodense mass and a second well-defined mass with a calcified nodule in the left hepatic lobe, without enhancements in or around either mass.

***Pathological diagnosis***

Histological examination revealed a collagenous nodule mixed simple cyst and a hepatic hemangioma.

***Treatment***

A left liver resection was performed.

***Related reports***

A collagenous nodule is a rare clinicopathologic characteristic of the liver. This paper discusses the pathogenesis of a collagenous nodule occurring in the liver. Although a hepatic hemangioma is the most common benign tumor of the liver, our case presented with atypical radiographic features.

***Term explanation***

The primary pathologic feature of a collagenous nodule is hyaline degeneration of connective tissue. This form of degenerative alteration can be found in atrophic breast tissue, scar tissue, atheromatous plaques, and organized tissue. The cause of a collagenous nodule in a liver simple cyst is unclear. It is possible that the cyst was infected or bleeding after which the pus or blood clot was replaced by granulation tissue, which is known as organization. This granulation tissue may then be replaced by fibrous connective tissue.

***Experiences and lessons***

The diagnosis and treatment of atypical masses in the liver should depend on a histologic examination during surgery.

***Peer-review***

The article can be published due to the rarity of the 3 diseases at the same time. The surgical procedure has been performed well due to the difficulties of the diagnosis.

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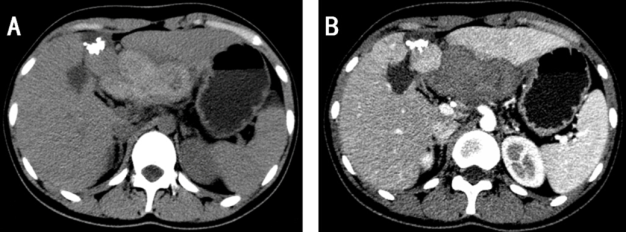
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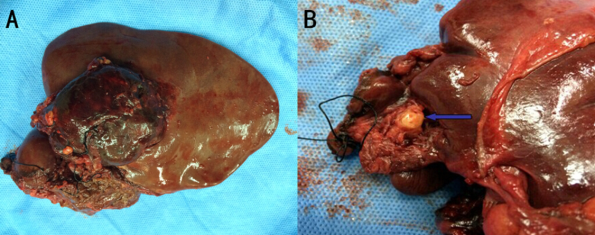
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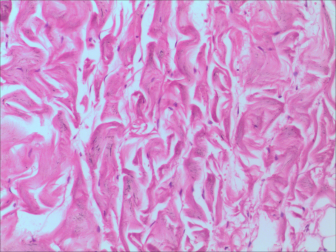
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**Figure 1 Radiographic findings**. A: A non-enhanced computed tomography (CT) scan revealed two masses in the liver, one of which contained a high-density nodule; B: A contrast-enhanced CT scan obtained during the hepatic arterial phase showed no enhancement in or around the masses.



**Figure 2 Intraoperative findings.** A: A solid mass localized in segment III; B: A cystic mass with a hard nodule (blue arrow) localized in segment IV.



**Figure 3 Histologic findings.** A histologic section from the nodule revealed thickened, banded and sheet-like collagenous fibrous tissue (HE stain, × 200).