Published online 2017 October 28.

Case Report

Fasciola Hepatica in Explanted Cirrhotic Liver of a Patient with Chronic Hepatitis B Virus Infection: An Accidental Finding

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Received 2016 September 25; Revised 2017 September 02; Accepted 2017 October 08.

Abstract

Introduction: Fasciola hepatica infection is frequently asymptomatic and may be neglected. However, severe conditions may be symptomatic due to stimulatory effects of the liver fluke in the liver.

Case Presentation: The present report describes a rare case of fascioliasis accidentally discovered in the explanted liver of a patient with cirrhosis due to chronic hepatitis B virus (HBV) infection.

Conclusions: Liver fluke infection may occur in cirrhotic livers. Thus, minute evaluation of liver explants for possible parasitic infestations including *F. hepatica* is suggested, particularly in endemic areas.

Keywords: Liver, Fibrosis, Transplantation, Hepatitis B, Fasciola Hepatica

1. Introduction

Fascioliasis caused by the liver fluke Fasciola hepatica is an important primary veterinary disorder as well as a human secondary disease (1). This flat-worm trematode enters the human body through digestive tract and lives in the common and hepatic bile ducts (2). The public health importance of human fascioliasis is now increasing due to its global spread as well as appearance of its considerable complications. Early infection with F. hepatica is frequently asymptomatic and may be neglected; however, in more severe conditions, stimulatory effects of the liver fluke can lead to proliferation and inflammation of biliary tracts and subsequently to hepatic fibrosis, biliary tract obstruction, cholangiocarcinoma, and even hepatocellular carcinoma (3). Furthermore, hepatic tissue damage as well as cirrhosis may appear as a result of hepatic cells injury (4). The present report describes a rare case of fascioliasis accidentally discovered in the explanted liver of a patient with cirrhosis due to chronic hepatitis B virus (HBV) infection.

2. Case Presentation

A 58-year-old female with diagnosis of cirrhosis in the background of chronic HBV infection was referred to Imam Khomeini Hospital Complex for orthotopic liver transplantation. Initial laboratory assessment showed a marked increase in alkaline phosphatase (ALP) as well as increased serum total and direct bilirubin level along with slightly elevated alanine aminotransferase (ALT) (aspartate aminotransferase (AST): 33 IU/mL, ALT: 84 IU/mL, ALP: 565 IU/mL, total bilirubin: 3.7 mg/dL, direct bilirubin: 2 mg/dL) and MELD (the model of end-stage liver disease) score of 17. Pre-transplantation evaluations showed a mass (suspected of hemangioma) in segment 4b and a fibrocalcified focus in segment 4a without other remarkable findings (Figure 1). Therefore, the patient underwent orthotopic liver transplantation. The histopathological assessment of explanted liver indicated active cirrhosis with large cell changes in some nodules. Further assessment revealed a necrotic hyalinized nodule harboring non-viable parasite in the peripheral regions of segment 4a (Figure 2). Also, small bile duct hamartoma was identified in a microscopic focus in segment 4a. A hemangioma measuring 1 cm was also found in segment 4b. The extrahepatic biliary tracts were unremarkable. No evidence of malignant change was detected. A demised degenerated worm that was morphologically consistent with F. hepatica was identified in the necrotic hyalinized nodule (Figure 2A and B). Eggs of this parasite were also found in the surrounding tissue (Figure 3). Follow-up of the patient showed good recovery after treatment with triclabendazole. The patient is alive and healthy without any major problems 2 years after trans-

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plantation.



Figure 1. CT Scan of Liver Reveals (A) a Small Linear Subcapsular Hypodense Lesion (Arrowhead) (B) Associated with Adjacent Linear Calcification (Arrow)



Figure 2. A, Demised fluke is evident in the necrotic background, separated by dense fibrotic tissue from liver parenchyma. H&E, X100. B, Schematic drawing of the worm.

Figure 3. Degenerated Egg of the Fluke in a Necrotic Background



H and E stain, X400.

3. Discussion

Recent studies showed an elevation in the incidence of liver fibrosis, cholangitis, cholecystitis, cholelithiasis, cholangiocarcinoma, and even hepatocellular carcinoma following liver fluke infection (5, 6). A systematic review showed an eightfold risk of cholangitis, fifteen-fold risk of cholecystitis, threefold risk of cholelithiasis, and fivefold risk of hepatocellular carcinoma following fascioliasis. However, the current study could not indicate a powerful relationship between this infection and cirrhosis. In fact, discovering liver fluke infection in patients with cirrhosis is a condition rarely reported (7).

Some probable mechanisms are described for development of cirrhosis following liver fluke infection. A recent study suggested that *F. hepatica* infection was associated with the upregulation of mRNAs associated with hepatic fibrogenesis in vivo and in activated hepatic stellate cells (8). Furthermore, the liver fluke may secrete some metabolites that can play an immunogenic stimulatory role leading to strong humoral immune responses (9). In fact, the presence of parasite antigens in various hepatobiliary portions such as biliary epithelium and its surrounding tissue can facilitate inflammatory cell infiltration and also activate host immune responses (10).

According to the current study findings, the development of liver fluke infection is expected in patients with cirrhosis, and thus, minute evaluation of liver explants for possible parasitic infestations including *F. hepatica*, particularly in endemic areas is suggested.

Footnotes

Authors' Contribution: Elham Nazar, collection of clinical data, histopathologic evaluations, and drafting of the manuscript; Farid Azmoudeh-Ardalan, histopathologic evaluations and editing the manuscript; Iraj Mobedi, identification of the parasite and drawings of the worm; Seyed Habiballah Dashti, surgery and pre-operation practices on the patient; Faeze Salahshour, radiologist of the transplantation team and review of diagnostic images; Zahra Ahmadinejad, infectious disease consultant.

Implication: *F. hepatica* may infest cirrhotic livers; posttransplantation flare up of fascioliasis can also be problematic. Therefore, biliary system and liver parenchyma of explanted livers should be comprehensively evaluated for potential infestations in endemic areas.

References

- Keiser J, Utzinger J. Emerging foodborne trematodiasis. *Emerg In-fect Dis.* 2005;**11**(10):1507–14. doi: 10.3201/eid1110.050614. [PubMed: 16318688].
- Control of foodborne trematode infections. Report of a WHO Study Group. World Health Organ Tech Rep Ser. 1995;849:1-157. [PubMed: 7740791].

- Lim JH. Liver flukes: the malady neglected. Korean J Radiol. 2011;12(3):269–79. doi: 10.3348/kjr.2011.12.3.269. [PubMed: 21603286].
- Villegas F, Angles R, Barrientos R, Barrios G, Valero MA, Hamed K, et al. Administration of triclabendazole is safe and effective in controlling fascioliasis in an endemic community of the Bolivian Altiplano. *PLoS Negl Trop Dis.* 2012;6(8):e1720. doi: 10.1371/journal.pntd.0001720. [PubMed: 22880138].
- Marcos LA, Yi P, Machicado A, Andrade R, Samalvides F, Sanchez J, et al. Hepatic fibrosis and Fasciola hepatica infection in cattle. J Helminthol. 2007;81(4):381–6. doi: 10.1017/S0022149X07850231. [PubMed: 17958928].
- Sripa B, Brindley PJ, Mulvenna J, Laha T, Smout MJ, Mairiang E, et al. The tumorigenic liver fluke Opisthorchis viverrini-multiple pathways to cancer. *Trends Parasitol.* 2012;28(10):395–407. doi: 10.1016/j.pt.2012.07.006. [PubMed: 22947297].
- 7. Marcos LA, Bussalleu A, Terashima A, Espinoza JR. Detection of

antibodies against Fasciola hepatica in cirrhotic patients from Peru. J Helminthol. 2009;**83**(1):23–6. doi: 10.1017/S0022149X08067205. [PubMed: 18817587].

- Marcos LA, Terashima A, Yi P, Andrade R, Cubero FJ, Albanis E, et al. Mechanisms of liver fibrosis associated with experimental Fasciola hepatica infection: roles of Fas2 proteinase and hepatic stellate cell activation. J Parasitol. 2011;97(1):82-7. doi: 10.1645/GE-2420.1. [PubMed: 21348611].
- Perrigoue JG, Marshall FA, Artis D. On the hunt for helminths: innate immune cells in the recognition and response to helminth parasites. *Cell Microbiol.* 2008;10(9):1757–64. doi: 10.1111/j.1462-5822.2008.01174.x. [PubMed: 18505479].
- Xia J, Jiang SC, Peng HJ. Association between Liver Fluke Infection and Hepatobiliary Pathological Changes: A Systematic Review and Meta-Analysis. *PLoS One*. 2015;10(7):e0132673. doi: 10.1371/journal.pone.0132673. [PubMed: 26186510].