Case report / Приказ болесника

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Not so innocent bystander – gallbladder varices without portal vein thrombosis

Не баш безазлени посматрачи – варикси жучне кесе без тромбоze портне вене

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SUMMARY
Introduction Gallbladder varices (GBV) represent a rare form of ectopic varices that usually occur in patients with portal hypertension and portal vein thrombosis.

Case outline We present a case of a 38-year-old woman with decompensated autoimmune liver cirrhosis who was referred to our institution for evaluation for liver transplantation. She was incidentally discovered to have GBV during routine B-mode abdominal ultrasonography as part of pre-transplant evaluation. GBV were confirmed by Color Doppler Sonography (CDS), and multi detector computed tomography angiography. Interestingly, portal vein was patent and without thrombus.

Conclusion Despite being asymptomatic in most cases, the presence of GBV is valuable information for a surgeon because they might be a source of potentially catastrophic bleeding, which is particularly poorly tolerated by patients with decompensated liver cirrhosis. Ultrasound has the irreplaceable role not only in discovering GBV, but in prompt diagnosis of before mentioned rare, but unpredictable and fatal complications as well.

Keywords: gallbladder varices; ectopic varices; portal hypertension

INTRODUCTION
Gallbladder varices (GBV) are rare form of ectopic varices that usually develop in patients with portal hypertension. They represent a form of porto-systemic shunting that occurs between portal vein through the cystic vein branches and the veins of anterior abdominal wall [1]. Hence, it is of no surprise that the gallbladder is directly affected by portal hypertension. Portal hypertension may lead to development of thickened gallbladder wall secondary to impaired venous drainage. GBV occur with incidence of 12–30% in patients with portal hypertension, are usually associated with portal vein thrombosis (PVT) and are characteristic feature of portal bilopathy [2, 3]. Most of the time they are asymptomatic but their spontaneous bleeding results in hemobilia, recurrent gastrointestinal bleeding (GIB) or even gallbladder perforation and hemoperitoneum [4].
We present a case of a patient with decompensated liver cirrhosis secondary to autoimmune hepatitis who was diagnosed with GBV during routine abdominal ultrasonography as a part of pre-liver transplant evaluation. The diagnosis was confirmed by Color Doppler Sonography (CDS) and abdominal MDCT.

CASE REPORT

A 38-year-old female patient was referred to the Clinic for Gastroenterology and Hepatology of the Clinical Center of Serbia for transplant evaluation due to end stage liver disease (ESLD) secondary to autoimmune hepatitis. She was diagnosed with decompensated liver cirrhosis 7 years prior to her current hospitalization and since then she has been admitted several times due to the various complications of ESLD such as recurrent ascites, jaundice, hepatic encephalopathy and recurrent GIB. During the last admission she had GIB and upper esophagogastroduodenoscopy (EGD) showed grade III varices with "red cherry spots" which were successfully treated by band ligation. Due to worsening Model of End Stage Liver Disease (MELD) score of 24 she was felt to be a transplant candidate. On admission patient was hemodynamically stable and without fever or leukocytosis. Her abdomen was distended but non-tender with palpable splenomegaly and positive fluid shift. Cardio-pulmonary exam was unremarkable and skin showed evidence of telangiectasia. Neurological exam was non focal and there was no encephalopathy. Pre transplant evaluation included routine abdominal ultrasonography that revealed an enlarged, nonhomogeneous liver with massive splenomegaly of 250 mm in craniocaudal diameter, as well as circular changes in the gallbladder wall. Doppler sonography of portal system confirmed gallbladder varices, however, without a portal vein thrombosis (Figures 1 and 2). MDCT angiography of the abdomen confirmed a thickened and delaminated wall of the gallbladder with GBV as well as dilated, but patent portal vein without thrombosis (Figures 3, 4, and 5).
DISCUSSION

Ectopic varices represent dilated splanchnic veins or dilated porto-systemic collaterals which occur along the entire gastrointestinal tract, outside the common variceal sites such as gastroesophageal varices and internal hemorrhoids [2]. GBV are form of ectopic varices seen as a complication of portal hypertension. They consist of the enlarged blood vessels in the gallbladder wall or gallbladder fossa and represent a portosystemic shunt between the cystic branches of the portal vein and the systemic veins of the anterior abdominal wall [1, 2, 3].

The incidence is similar in adult and pediatric population with portal hypertension, estimated to be up to 30% [4]. Majority of patients with GBV also have PVT, however, as our case illustrates, they might develop even in the absence of PVT. The gold standard for diagnosis is CDS, which shows the varices as venous flow in the delaminated and thickened parts of the gallbladder wall [3, 5]. If feasible, contrast-enhanced US can be a valuable further diagnostic tool, while CT scan and MRI appear to be less-sensitive compared to US.

It is important to consider other etiologies that might mimic GBV and present similarly. These etiologies are more common than GBV and include: acute or chronic cholecystitis, gallbladder cancer and porcelain gallbladder to name a few. The absence of mineralization and the presence of vascular enhancement rules out porcelain gallbladder, while the absence of pericholecystic fluid and inflammation make cholecystitis unlikely. Gallbladder cancer can present radiologically in similar fashion, but one would expect to see some degree of local soft tissue invasion or presence of metastatic lesions, which were absent in our case. In spite of their ability to affect the contractility of the gallbladder they are not associated with higher risk for development of cholelithiasis [6]. When present, GBV may cause hemobilia, intraabdominal hemorrhage or rupture of the gallbladder as illustrated in several case reports [7]. US has the irreplaceable role not only in discovering GBV, but in prompt diagnosis of
before mentioned rare, but unpredictable and fatal complications as well. [8]. Despite being rare, GBV are potential cause of detrimental gastrointestinal hemorrhage. The bleeding from GBV is serious because as population with portal hypertension and decompensated cirrhosis tends to be sick and poorly tolerates hemodynamic protuberances. Our case illustrates a rare entity which should be considered in any patient with planned abdominal surgery, particularly those with portal hypertension who have increased incidence of GBV. While they usually develop concomitantly with PVT, GBV might be isolated and occur in the absence of PVT, as we have shown in this report.

Considering high availability and low-cost of CDS, which is considered a gold standard for GBV diagnosis, there is no reason for careful evaluation of gallbladder not to be done in every patient with portal hypertension. If GBV are discovered, surgical team should be informed as it is pertinent information in planning and executing abdominal surgeries. By increasing awareness of this rare portosystemic shunt, we can prevent or decrease the incidence of massive bleeding from GBV, which in turn will decrease perioperative mortality [7–11].

**Ethical standards:** All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional and/or national research committee and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards. Written consent to publish all shown material was obtained from the patient.

**Conflict of interest:** None declared
REFERENCES

1. Way LW, Pellegrini CA, editors. Surgery of the gallbladder and bile ducts. Philadelphia: Saunders; 1987
2. West MS, Garra BS, Horii SC, Hayes WS, Cooper C, Silverman PM, et al. Gallbladder varices: imaging findings in patients with portal hypertension. Radiology. 1991 Apr;179(1):179–82. DOI.org/10.1148/radiology.179.1.2006274
3. Saad WE, Lippert A, Saad NE, Caldwell S. Ectopic Varices: Anatomical Classification, Hemodynamic Classification, and Hemodynamic-Based Management. Techniques in Vascular and Interventional Radiology. 2013 Jun 1;16(2):108–25. PMID: 23830673 DOI: 10.1053/j.tvir.2013.02.004
4. Kessler A, Graif M, Konikoff F, Mercer D, Oren R, Carmiel M, et al. Vascular and biliary abnormalities mimicking cholangiocarcinoma in patients with cavernous transformation of the portal vein: role of color Doppler sonography. J Ultrasound Med. 2007 Aug;26(8):1089–95. PMID:17646372 DOI: 10.7863/jum.2007.26.8.1089
5. Shah VH, Kamath PS. Chapter 92: Portal hypertension and variceal bleeding. In: Feldman M, Friedman LS, Brandt LJ (eds). Sleisenger and Fordtran's Gastrointestinal and Liver Disease, 10th edn. Saunders: Philadelphia, PA, 2016, pp 1524–52.
6. Venerito M, Mönkemüller K, Malfertheiner P, Rickes S. Hepatobiliary and pancreatic: gallbladder varices. J Gastroenterol Hepatol. 2006 Aug;21(8):1348. PMID: 16872323 DOI: 10.1111/j.1440-1746.206.04603.x
7. Temel JS, Greer JA, El-Jawahri A, et al. Effects of early integrated palliative care in patients with lung and GI cancer: a randomized clinical trial. J ClinOncol 2017; 35:834–41. PMID: 28029308 DOI: 10.1200/JCO.2016.70.5046
8. Pravisani R, Bugiantella W, Lorenzin D, Bresadola V, Leo CA. Fatal hemoperitoneum due to bleeding from gallbladder varices in an end-stage cirrhotic patient: A case report and review of the literature. Ann ItalChir. 2016;87(ePUB). PMID: 27238768
9. Gnerre J, Sun Y, Jedynak A, Gilet A. Case Report: Gallbladder Varices in a Patient with Portal Vein Thrombosis Secondary to Hepatocellular Carcinoma. J Radiol Case Rep. 2016 May 31;10(5):22–8. PMID: 27761177 DOI: 10.3941/jrcr.v10i5.2416
10. Garcia-Tsao G, Abraldes JG, Berzigotti A, Bosch J. Portal hypertensive bleeding in cirrhosis: Risk stratification, diagnosis, and management: 2016 practice guidance by the American Association for the study of liver diseases. Hepatology. 2017;65(1):310–35. PMID: 27786365 DOI: 10.1002/hep.28906
11. Clayton S, DeClue C, Lewis T, Rodriquez A, Kolhorts K, Syed R, Kumar A, Davis C, Brady P. Radiologic versus endoscopic placement of gastrostomy tube: comparison of indications and outcomes at a tertiary referral center. South Med J 2019;112:39–44. PMID: 30608631 DOI:10.14423/SMJ.00000000000000916
Figure 1. Gallbladder varices on B mode ultrasound
Figure 2. Gallbladder varices on B mode ultrasound
Figure 3. Delaminated gallbladder wall on abdominal MDCT
Figure 4. Dilated portal vein branch on abdominal MDCT
Figure 5. Dilated portal vein branch on abdominal MDCT