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**Modified endoscopic ultrasound-guided selective N-butyl-2-cyanoacrylate injections for gastric variceal hemorrhage in left-sided portal hypertension: A case report**

Yang J *et al.* EUS-guided selective injections for variceal hemorrhage

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**Abstract**

**BACKGROUND**

<sup>1</sup> Left-sided portal hypertension (LSPH), also known as sinistral portal hypertension or regional portal hypertension, refers to extrahepatic portal hypertension caused by splenic vein obstruction or stenosis. N-butyl-2-cyanoacrylate (NBC) has been widely used in the endoscopic hemostasis of portal hypertension, but adverse events, including renal or pulmonary thromboembolism, mucosal necrosis, and gastrointestinal (GI) bleeding, may occur after treatment. Herein, we report successfully managing gastric variceal (GV) hemorrhage secondary to LSPH using modified endoscopic ultrasound (EUS)-guided selective NBC injections.

**CASE SUMMARY**

<sup>7</sup> A 35-year-old man was referred to our hospital due to an upper GI hemorrhage. Gastroscopy revealed GV hemorrhage, and computed tomography venography (CTV) confirmed LSPH. The patient requested endoscopic procedures and rejected surgical therapies, including splenectomy. EUS-guided selective NBC injections were performed, and confluences of gastric varices were selected as the injection sites to reduce the injection dose. The “sandwich” method using undiluted NBC and

hypertonic glucose was applied. No complications occurred. The patient was followed up regularly after discharge. Three months later, the follow-up gastroscopy revealed firm gastric submucosa with no sign of NBC expulsion, and follow-up CTV showed improvements in LSPH. No recurrent GI hemorrhage was reported during this follow-up period.

## CONCLUSION

EUS-guided selective NBC injection may represent an effective and economical treatment for GV hemorrhage in patients with LSPH.

**Key Words:** Left-sided portal hypertension; Endoscopic ultrasound; Selective; N-butyl-2-cyanoacrylate; Gastric varices; Case report

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**Core Tip:** Gastric variceal (GV) hemorrhage caused by left-sided portal hypertension (LSPH) is a severe complication. Endoscopic ultrasound (EUS)-guided procedures for GV hemorrhage demonstrated beneficial results in reducing complication risks. Herein, we report the successful management of GV hemorrhage secondary to LSPH using EUS-guided selective N-butyl-2-cyanoacrylate injection, which proved the effectiveness and safety of this method. This case is the first report choosing confluences of gastric varices as injection sites to reduce the injection dose and postoperative complications in patients with LSPH.

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## **INTRODUCTION**

Left-sided portal hypertension (LSPH), also known as *sinistral* portal hypertension or regional portal hypertension, refers to extrahepatic portal hypertension caused by splenic vein obstruction or stenosis<sup>[1-3]</sup>. LSPH accounts for about 5% of extrahepatic portal hypertension and is characterized by isolated gastric varices (GVs) and normal liver functions<sup>[3]</sup>. Pancreatic diseases are the major causes of LSPH. Most patients with LSPH have no obvious clinical symptoms, and they are often diagnosed during the endoscopic examination after gastrointestinal (GI) bleeding. Therefore, LSPH should be considered in patients with pancreatic diseases who develop unexplained GI hemorrhage<sup>[1,4]</sup>.

Gastric variceal (GV) hemorrhage leads to significant mortality in patients with portal hypertension. Although N-butyl-2-cyanoacrylate (NBC) has been widely used in the endoscopic hemostasis of portal hypertension, the early expulsion of NBC and the resultant hemorrhage is not uncommon<sup>[5]</sup>. Compared with conventional endoscopic injection, endoscopic ultrasound (EUS)-guided procedures in patients with GV bleeding demonstrated better diagnostic capability and clinical efficacy<sup>[6,7]</sup>.

Herein, we report the successful management of GV hemorrhage secondary to LSPH using modified EUS-guided selective NBC injection.

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## **CASE PRESENTATION**

### ***Chief complaints***

A 35-year-old man was referred to our hospital due to an upper GI hemorrhage.

### ***History of present illness***

A few hours before admission, the patient had no apparent reason for sudden vomiting of blood mixed with stomach contents once, and the amount was about 50-100 mL. He denied melena and syncope.

### ***History of past illness***

Nine months previously, this patient was admitted to our hospital due to persistent upper abdominal pain. He was diagnosed with severe acute pancreatitis (SAP) and underwent EUS-guided drainage of pancreatic walled-off necrosis. He also had a six-year history of hypertension and took enalapril regularly.

#### *Personal and family history*

This patient had a 10-year smoking history (a pack per day) and has not quit smoking. He denied alcoholism and taking nonsteroidal anti-inflammatory drugs.

#### *Physical examination*

After admission, physical examination revealed no abnormality except for 130/91 mmHg blood pressure.

#### *Laboratory examinations*

No apparent abnormalities were found in the emergency blood analysis.

#### *Imaging examinations*

After admission, gastroscopy confirmed GV hemorrhage (IGV1 by Sarin classification), and no esophageal varices or portal hypertensive gastropathy was found (Figure 1). Abdominal computed tomography venography (CTV) revealed stenosis of the proximal superior mesenteric vein, invisible proximal splenic vein, and increased collateral circulations (Figure 2). Neither the portal vein thrombosis nor the splenorenal shunt was detected.

### **FINAL DIAGNOSIS**

LSPH and GV hemorrhage.

### **TREATMENT**

The patient requested endoscopic procedures and rejected surgical therapies, including splenectomy. EUS-guided selective NBC injections were performed for the treatment and prophylaxis of GV hemorrhage.

A linear Pentax echoendoscope (Hoya co., Tokyo, Japan) and the color Doppler flow imaging were employed to determine the puncture site. EUS revealed an enlarged portal vein without cavernous transformation (Figure 3A). The confluences of GVs were selected as the injection sites to reduce the injection dose. A 22-gauge needle (Boston Scientific co., Natick, American) was used to perform the puncture into the selected GVs (Figure 3B). The “sandwich” method using undiluted NBC (0.5 mL/ampoule; Beijing Compont Medical Devices Co., Beijing, China) and hypertonic glucose was applied (Figure 3C). A total of 2 mL of NBC was injected into three different confluences of GVs. Hyperecho fillings and decreased blood flow signals were observed after injections (Figure 3D).

### **OUTCOME AND FOLLOW-UP**

The patient fasted for one day after the procedure. No complications, including ectopic embolism, fever, and post-injection GI bleeding, occurred. The patient was followed up regularly after discharge. Three months later, the follow-up gastroscopy revealed no sign of NBC expulsion (Figure 4), and follow-up CTV showed improvements in LSPH (Figure 5). No recurrent GI hemorrhage and other complications were reported during his three-month follow-up.

### **DISCUSSION**

Pancreatic diseases such as pancreatitis and pancreatic tumors are the most common etiology of LSPH<sup>[3,8]</sup>. The anatomical proximity between the splenic vein and the pancreas makes the splenic vein more susceptible to pancreatic diseases. When pancreatic disease obstructs the splenic vein flow, the pressure of the left portal vein system increases, and blood flows retrogradely through the short and posterior gastric veins and the gastroepiploic veins, which would lead to GVs. In patients with acute

pancreatitis, infected walled-off necrosis was one of the risk factors for LSPH, and early anticoagulation could not wholly prevent its occurrence<sup>[8]</sup>. In this case, the patient had a history of SAP and infected pancreatic necrosis, which may be responsible for his LSPH. About 20% of patients with portal hypertension may develop GV<sup>s</sup><sup>[9]</sup>, and although LSPH is a rare cause of upper GI hemorrhage, GV hemorrhage in patients with LSPH secondary to pancreatic disease is not uncommon. Liu *et al*<sup>[10]</sup> reported that about 15.3% of LSPH patients complicated bleeding GV<sup>s</sup>, and the death risk is relatively higher when recurrent GV hemorrhage occurs, so it is worthy of attention.

It is well known that splenectomy is the most effective treatment for LSPH. However, transjugular intrahepatic portosystemic shunt, balloon retrograde transvenous obliteration, endoscopic injection sclerotherapy (EIS), and endoscopic NBC injection were reported effective for patients who are not suitable or unwilling to choose surgery<sup>[11]</sup>. Although endoscopic NBC injection therapy has been proven minimal invasive and effective<sup>[12]</sup>, conventional endoscopic NBC injections may also cause severe complications, including renal or pulmonary thromboembolism, fever, severe pain caused by intraperitoneal injection, mucosal necrosis at the injection site, and GI bleeding<sup>[13]</sup>. As reported in recent years, EUS-guided hemostasis treatment, including injection of NBC or combination with coils, injection of thrombin or absorbable gelatin sponge, and clip-assisted endoscopic NBC injection, demonstrated promising results in reducing complication risks<sup>[14,15]</sup>.

Modified EUS-guided selective NBC injection was applied for three advantages in this present case. First, a reduced NBC dose may result in a lower occurrence of post-operational GI bleeding and ectopic embolism. EUS can provide detections of submucosal GV<sup>s</sup>, their confluences, and real-time effectiveness evaluation of GV obliteration<sup>[16]</sup>. These advantages make it possible to identify and select confluences of gastric varices, which were in the source direction of bleeding gastric vessels, as injection sites to reduce the injection dose. Although EUS-guided coil injection is reported superior to conventional NBC injection in terms of rebleeding after treatment<sup>[17]</sup>, it was believed that a reduced dose of NBC would be injected into GV<sup>s</sup> in



modified EUS-guided selective NBC injection, which would lead to lower chances of post-injection ulcer and GI hemorrhage. Besides, reduced NBC dose may result in a similar lower occurrence of ectopic embolism in selective NBC injection as in the coils-combined injection method and clip-assisted injection method. Second, there would be no additional risk of radioactive exposure; coils and metal clips were not used in this modified injection procedure, which decreased the cost of endoscopic procedures. Third, selective NBC injection demonstrated a faster and firmer obliteration effect in GV hemorrhage than thrombin and absorbable gelatin sponge injections, making NBC injection more suitable than other procedures for acute GV bleeding. NBC rarely causes vascular necrosis and was reported superior to EIS in the hemostasis rate for GV bleeding<sup>[18]</sup>. Thus, EUS-guided selective NBC injection was performed for this patient based on the above factors, and the result was adequate. Despite all these advantages, the operation time of EUS-guided selective NBC injection seemed a little longer than that of conventional endoscopic NBC injection due to time consumption to confirm confluences of GVs during the EUS procedure. Additional cases are needed to verify our findings and compare the efficacies and complications of different embolization methods guided by EUS. Currently, this described technique is recommended to be used only in hemodynamically stable patients. <sup>2</sup> To the best of our knowledge, this case is the first report choosing confluences of gastric varices as injection sites to reduce the injection dose and postoperative complications in patients with LSPH.

## **CONCLUSION**

Modified EUS-guided selective NBC injection may represent an effective and economical treatment for GV hemorrhage in patients with LSPH.



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