

Percutaneous transsplenic embolization of esophageal and gastro-fundal varices in 18 patients

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Abstract

AIM: Clinical application and potential complication of percutaneous transsplenic varices embolization (PTSVE) of esophageal or gastro-fundal varices in patients with hepatocellular carcinoma (HCC) complicated with portal vein cancerous thrombosis (PVCT).

METHODS: 18 patients with HCC complicated with PVCT and esophageal or gastro-fundal varices who underwent PTSVE were collected. The rate of success, complication, mortality of the procedure and postoperative complication were recorded and analyzed.

RESULTS: PTSVE were successfully performed in 16 of 18 cases, and the rate of success was 89%. After therapy erythrocyte counts decreased in all of the patients. 5 of patients needed blood transfusion, 2 patients required surgical intervention because of and 11 patients with ascites were alleviated by diuresis. Among these 18 patients, the procedure-related mortality was 11% (2/18), one died of acute hepatic failure on the fourth day after procedure, another died of acute renal failure on the fifth day. The patients were follow up for 1-12 months except one. 13 of them died of their tumors but none of them experienced variceal bleeding.

CONCLUSION: PTSVE is a relatively safe and effective method to treat esophageal or gastro-fundal varices in HCC patients with PVCT when percutaneous transhepatic varices embolization (PTHVE) of varices is impossible.

Subject headings Esophageal or gastric varices/ complications; Gastrointestinal hemorrhage/ etiology; Gastrointestinal hemorrhage/therapy; Embolization, therapeutic; Radiology, interventional/methods

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INTRODUCTION

Hepatocellular carcinoma (HCC) is common in China^[1-8] and many complications occur^[9-13] including ascites, portal hypertension and splenomegaly. Reviewed literatures, we found there are many articles about treatment of HCC^[2,4,11,14-32] or varices^[33-38], but few relating the treatment of HCC complicated varices^[39-41]. The percutaneous transhepatic variceal embolization (PT HVE) of gastroesophageal varices is a highly promising procedure for controlling acute variceal hemorrhage and decreasing the mortality in patients with portal hypertension. However there are limitations of PTHVE application, because the ligation of the main trunk of portal vein is required, which was limited its use. Thus, it is very important to find another route to catheterize into portal vein in patients whose portal vein were invaded by HCC. In order to control variceal hemorrhage in these patients it is very important to find another approach for catheterizing into their portal vein. From 1999-08 to 2001-01, 18 HCC patients with portal vein cancerous thrombosis (PVCT) underwent percutaneous transsplenic embolization (PTSVE) of esophageal or gastro-fundal varices in our institution as reported here.

MATERIALS AND METHODS

Patients

18 patients (17 male, 1 female, mean age 48 yr, aged from 29-72 yr) of HCC with PVCT enrolled in to this study. The diagnosis of HCC with esophageal or gastro-fundal varices was made by clinical history, alpha-fetoprotein (AFP), ultrasound, CT or MRI. 4 of the patients experienced variceal hemorrhage in different degree and 1 patient had seven episodes of serious variceal bleeding before therapy.

Varices classification

According to Burchard *et al*, varices were classified into 4 grades: 0 means no varices were visible; 1 grade: only tiny varices were visible; 2 grade: varices were distinctly visible, and 3 grade: severe varices were seen.

Clinical data

According to the Child-Pugh scores, 10 patients were Child A and 8 patients were Child B. Patients' average platelet counts were 83.2×10^9 , ranged from $(11 \text{ to } 195) \times 10^9 \cdot L^{-1}$, and 3 patients had accepted 4 units blood transfusion due to decreased erythrocyte before therapy. The slight or moderate ascites were observed in 10 patients.

Pre-operative preparation

The patients with ascites were given albumin and diuretics. The patients have not underwent PTSVE until the ascites were alleviated. CT angiography (CTA) or MR angiography (MRA) of portal venous system was necessary to show patients' splenic vein and its tributary^[42], main trunk of portal vein and esophageal or gastro-fundal varices in order to obtain the information of the site, direction, and depth of puncture before therapy (Figure 1).

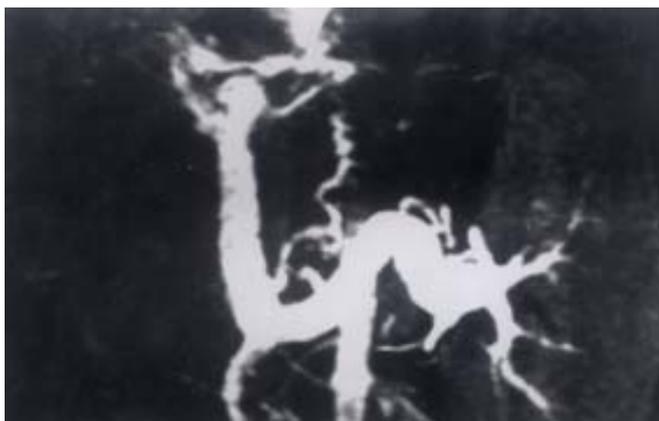


Figure 1 CTA of portal vein that show tributary of splenic vein clearly.

Procedure

For clearly visualization of portal venous system and varices, indirect or direct portography was performed on patient them.

Indirect portography

The indirect portography was performed after 5F RH catheter was catheterized into patient's splenic artery via left or right femoral artery approach. 40 mL contrast medium (Ultravist 300, Schering AG or Omnipaque 300, Nyegarrd) were injected into the splenic artery at a speed of $7 \text{ mL}\cdot\text{s}^{-1}$. The film were exposed 1 picture $\cdot\text{s}^{-1}$ for 10 s until the main stem of patients' splenic vein and portal vein were clearly displayed. The tip of catheter was to be kept in the site for localization of the puncture.

Direct portography

The puncture site was determined on the basis of CTA or MRA images of portal vein s. Usually the seventh to ninth intercostal space on the left midaxillary line was selected as the puncture site. Local anaesthesia was followed by percutaneous transsplenic puncture using a 21 G Chiba needle. The core of needle was removed and the needle was pulled back slowly with intermittent aspiration. When blood can be freely aspirated, 5 ml diluted contrast medium (1:3) was injected as test. If patient's splenic vein was displayed clearly, a 0.018" (0.457 mm) guidewire was introduced and manipulated into the main stem of portal vein through the needle. Then the needle was withdrawn and replaced by a COPE puncture cannula system. A 5F catheter's sheath was pushed into patient's splenic vein through the exchanged 0.038" (0.968 mm) guidewire. After a 5F pigtail catheter was put in patient's superior mesenteric vein (SMV), the direct portography was performed with 40 ml contrast medium injected into patient's SMV at a ratio of $7 \text{ mL}\cdot\text{s}^{-1}$.

Variceal embolization

A 5F Cobra or Simmons I catheter was catheterized into patient's varices. Absolute alcohol, gelfoam and steel coils were used as embolization material. When the variceal vessels were occluded or the blood flow in varices was very slow, the varices were considered completely embolized. Then, the catheter was removed and the sheath was pulled back to the edge of the parenchyma of spleen. The duct of puncture was embolized by gelfoam and steel coil till the main stem of splenic vein was not displayed when the contrast medium was injected.

Post-operative management

Routine post-operative management included hemostasis, hepatic

function protection and anti-infection. Hepatic function and hematocyte were examined d 2 and d 5 after procedure. Abdominal symptom and body temperature were recorded too. B ultrasound, CT or MRI was followed in 9 patients 1 wk after operation. It was considered the proof of infection that high body temperature ($> 38.5^\circ\text{C}$) and high leucocytic counts ($> 10 \times 10^9 \cdot \text{L}^{-1}$), persisted longer than 7 d.

Follow-up

Survival time and post-operative bleeding rate were recorded in 15 patients in whom the follow up was done.

RESULTS

Rate of success

The branches of splenic vein have been successfully punctured in 16 of 18 patients and esophageal or gastro-fundal varices were embolized in the patients except 1 one patient with slight varices. The success rate of PTSVE was 89% (16/18). The other 2 patients whose PTSVE was unsuccessful didn't undertake the CTA or MRA examination before treatment.

Portography

Hepatofugal blood flow in portal vein was observed in 16 patients. The main stem of portal vein obstructed and collateral circulation occurred in 14 patients. Grade 1 varices in 1 patient, Grade 2 in 4 patients, and grade 3 in 11 patients were observed (Figure 2-4). Among 3 patients whose varices fallen in grade 2, 2 patients had spontaneous splenorenal shunt and another one had spontaneous portacaval shunt.

Complication and mortality

All patients' erythrocyte counts were decreased after the procedure and 5 of them needed blood transfusion. 2 patients experienced hemoperitoneum and 1 patient had mild reactive pleural effusion. All of these complications were alleviated by conservative treatment. 11 patients had mild or moderate ascites because of severe hypoproteinemia which was controlled by supplying albumin and diuresis (Table 1). Hepatic functions didn't deteriorate in the patient except one who had thrombosis in the main stem of his portal vein and died on the fourth day after therapy. Another patient died of acute renal failure on the fifth day after therapy. The procedure-related mortality was 11% (2/18). No severe infection was observed. 9 patients undertook B ultrasound, CT or MRI 1 wk after procedure and no hematoma in spleen surrounding area was displayed.



Figure 2 indirect portography via transsplenic artery show splenic vein and its tributary.



Figure 3 direct portography show main portal vein obstruction and the varices.

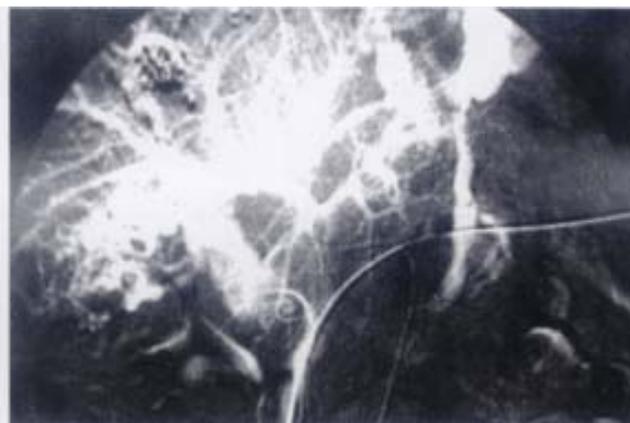


Figure 4 the same patient as figure 3, the variceal blood flow was turned slowly after embolization.

Table 1 Postoperative data of 18 HCC patients received PTSVE

Code/Survival	Reconstitu-tion of the portal vein	Operative Success	Blood infusion (Unit)	Ascites	Post-operative bleeding	Other complications
1/lost to follow up	no	yes	none	negative	unknown	none
2/dead	yes	yes	none	negative	no	pleural effusion
3/dead	yes	yes	yes/4	moderate	no	none
4/ suicide	yes	yes	none	negative	no	none
5/dead	yes	yes	none	slight	no	none
6/dead	yes	yes	none	moderate	occurred	hemoperitoneum
7/dead	no	no	none	moderate	no	none
8/alive	yes	yes	none	moderate	no	none
9/lost to follow up	no	no	none	negative	unknown	none
10/dead	no	yes	yes/2	moderate	no	none
11/alive	yes	yes	none	negative	no	none
12/dead	no	yes	yes/4	slight	no	none
13/dead	no	yes	yes/2	slight	occurred	none
14/alive	no	yes	none	negative	no	hemoperitoneum
15/dead	yes	yes	yes/2	moderate	no	Acute hepatic failure
16/dead	yes	yes	none	slight	no	none
17/dead	yes	yes	no	slight	no	none
18/dead	no	yes	no	negative	no	acute renal failure

*Patient with slight varices which needn't to be embolized.

Follow up

15 of 16 patients whose varices were successfully embolized had been followed up 1 to 12 mo. During the follow-up 13 of these 15 patients died. The cause of death was hepatic failure in 7 patients (1 in 4 d after procedure, 1 in 1 mo, 2 in 3 mo, 2 in 6 months, 1 in 8 mo). One died of acute renal failure in the fifth day after therapy, 2 died of metastasis, 2 died of ulcer bleeding which were conformed by gastroscopy, 1 committed suicide. In the 2 failed cases, one was dead, the other was beyond follow up.

7 of the Nations was died of hepatis failure which was the main ocuse of the do nth the causes of the ather 6 Natient were different. They night died of acute read failuisinth fifth day after the rany (Icase), of metastasis (2 case), of wher bleeding (2 case) and committed suicide (1 case).

DISCUSSION

Interventional treatments include two different way, one is shunt (for example: transjugular intra-he patic portosystemic shunt, TIPSS), and the other is varices embolization^[33-36]. There are some defects of TIPSS: proximal shunt can decrease hepatic blood flow in portal vein, which makes hepatic function deteriorated and restenosis rate high. Traditional embolization route includes transhepatic and transumbilical vein and celiotomy incision to puncture branches of SMV. The latter

two have been abandoned because of difficult operation and traumatic problem.

Percutaneous transhepatic variceal embolization (PTHVE) may be useful to control acute variceal bleeding and decreasing mortality. Since PTHVE is less traumatic and it s success rate is high, some authors apply PTHVE as a prophylactic management for the patients wit h severe varices in whom the rate of bleeding is high. However, the premise to perform PTHVE is that main portal vein must be patent and there is no tumor on the route of puncture. HCC has a high in cidence in our country. Advanced HCC usually complicated with portal vein cancerous thrombosis (PVCT) which deteriorated the previous portal hypertension. In such patient there was a high incidence of esophageal or gastro-fundal variceal bleeding. Since main portal vein is invaded and obstructed by HCC and the tumor always impedes the puncture route, the utility of PTHVE is limited. Basing on PTHVE and percutaneous transsplenic portography^[43], we designed PTVSE to avoid puncturing obstructed main stem of portal vein and to treat embolus in portal vein in addition.

In 16 of 18 patients, the splenic vein was punctured, and superselective catheterization into varices was done successfully. In order to determine the puncture site, most of the patients undertook pre-operative CTA or MRA of portal vein and indirect portography via splenic artery. The rate of success of PTSVE in this group was 89%

(16/18), similar to PTHVE and Liang's report in which 16 of 17 transsplenic portography were done successfully under B ultrasound guidance^[43]. We find that only Rasinska has reported one case treated by PTVSE in 1987 through our review.

It is suggested that transsplenic puncture might lead to hemorrhage because of decrease of erythrocyte counts after procedure in all patients. But in most of cases, it didn't need blood transfusion except 5 patients who accepted 2 to 4 units of blood transfusion. Hemoperitoneum occurred in two cases and slight reactive pleural effusion occurred in one. Both of the complications could be alleviated by conservative treatment. Portal vein thrombosis developed in 1 case within 4 d and led to acute hepatic failure. 1 case experienced acute renal failure within 5 d. Both of them died. Procedure-related mortality was 11% (2/18). Examination in 9 patients by imaging modalities 1 wk after procedure, no hematoma was found around spleen. In addition, 11 patients developed slight or moderate ascites after procedures that were alleviated by heteropathy. No severe infection occurred in all 18 patients. It is suggested that PTSVE be a safe technique relatively^[43]. However PTSVE is not the first choice of treatment when PTHVE can be performed because its mortality was higher and may lead to hemorrhage compared with.

No esophageal or gastro-fundal variceal bleeding occurred in the 15 patients during follow-up. Only two cases had bleeding due to gastric ulcer verified by gastroscop. It is also suggested that PTSVE be relatively effective in short term and similar to PTHVE. Since the life expectancy of patients of HCC complicated with PVCT is short, the long-term follow up seems to be unnecessary.

REFERENCE

- 1 Wu MC. Clinical research advances in primary liver cancer. *World J Gastroenterol*, 1998;4: 471-474
- 2 Tang ZY. Advances in clinical research of hepatocellular carcinoma in China. *World J Gastroenterol*, 1998;4(Suppl 2):4-7
- 3 Liu E, Zhang QN, Li WG. Effect of various drinking water on human micronucleus frequency in high-risk population of PHC. *World J Gastroenterol*, 1998;4:183-184
- 4 Jia YC, Tian JM, Wang ZT, Chen D, Ye H, Liu Q, Yang JJ, Sun F, Lin L, Lu JP, Wang F, Cheng HY. A retrospective review on interventional treatment of 10000 cases of liver cancer. *Huaren Xiaohua Zazhi*, 1998;6:2-3
- 5 Gu GW, Zhou HG. New concept in etiology of liver cancer. *Huaren Xiaohua Zazhi*, 1998; 6:185-187
- 6 Yu SZ, Dong CH. Risk identification, assessment and control of primary hepatocellular cancer. *Huaren Xiaohua Zazhi*, 1998;6:1026-1029
- 7 Liu WW. Etiological studies of hepatocellular carcinoma. *Shijie Huaren Xiaohua Zazhi*, 1999;7:93-95
- 8 Schafer DF, Sorrell MF. Hepatocellular carcinoma. *Lancet*, 1999;353: 1253-1257
- 9 Chao TC, Jeng LB, Jan YY, Wang CS, Chen MF. Spontaneous gastroduodenal perforation in cancer patients receiving chemotherapy. *Hepatogastroenterology*, 1998;45:2157-60
- 10 Koda M, Murawaki Y, Idobe Y, Horie Y, Suou T, Kawasaki H, Ikawa S. Is choledocholithiasis a late complication of nonresectional therapies for hepatocellular carcinoma? *Hepatogastroenterology*, 1999;46:3091-3094
- 11 Yoshida H, Onda M, Tajiri T, Umehara M, Mamada Y, Matsumoto S, Yamamoto K, Kaneko M, Kumazaki T. Treatment of spontaneous ruptured hepatocellular carcinoma. *Hepatogastroenterology*, 1999;46:2451-2453
- 12 Sithinamsuwan P, Piratvisuth T, Tanomkiat W, Apakupakul N, Tongyoo S. Review of 336 patients with hepatocellular carcinoma at Songklanagarind Hospital. *World J Gastroenterol*, 2000;6:339-343
- 13 Chen CY, Chau GY, Yen SH, Hsien YH, Chao Y, Chi KH, Li CP, Chang FY, Lee SD. Life-threatening haemorrhage from a sternal metastatic hepatocellular carcinoma. *J Gastroenterol Hepatol*, 2000;15:684-687
- 14 Tang ZY. Clinical research of hepatocellular carcinoma in the 21st century. *China Natl J New Gastroenterol*, 1995;1:2-3
- 15 Wang JH, Lin G, Yan ZP, Wang XL, Cheng JM, Li MQ. Stage II surgical resection of hepatocellular carcinoma after TAE: a report of 38 cases. *World J Gastroenterol*, 1998;4:133-136
- 16 Qian SB, Chen SS. Transduction of human hepatocellular carcinoma cells with human γ -interferon gene via retroviral vector. *World J Gastroenterol*, 1998;4:210-213
- 17 Fan J, Ten GJ, He SC, Guo JH, Yang DP, Weng GY. Arterial chemoembolization for hepatocellular carcinoma. *World J Gastroenterol*, 1998;4:33-37
- 18 Li L, Wu PH, Li JQ, Zhang WZ, Lin HG, Zhang YQ. Segmental transcatheter arterial embolization for primary hepatocellular carcinoma. *World J Gastroenterol*, 1998;4:511-512
- 19 Chen ZN, Bian HJ, Jiang JL. Recent progress in anti-hepatoma monoclonal antibody and its application. *Huaren Xiaohua Zazhi*, 1998;6:461-462
- 20 Gu GW, Zhou HG. Relationship between pathology and prognosis of liver cancer. *Huaren Xiaohua Zazhi*, 1998;6:632-633
- 21 Wu MC. Progresses in surgical treatment of primary hepatocellular carcinoma. *Huaren Xiaohua Zazhi*, 1998;6:921-923
- 22 Livraghi T, Goldberg SN, Lazzaroni S, Meloni F, Solbiati L, Gazelle GS. Small hepatocellular carcinoma: treatment with radio-frequency ablation versus ethanol injection. *Radiology*, 1999;210:655-661
- 23 Iida K, Kadota J, Kawakami K, Shirai R, Abe K, Yoshinaga M, Iwashita T, Matsubara Y, Ishimatsu Y, Ohmagari K, Kohno S. Immunological function and nutritional status in patients with hepatocellular carcinoma. *Hepatogastroenterology*, 1999;46:2476-82
- 24 Zhou XD. Prevention and treatment of recurrences and metastases of hepatocellular carcinoma. *Shijie Huaren Xiaohua Zazhi*, 1999;7:260-261
- 25 Meng ZH, He ZP. Current situation of gene therapy studies in inhibition of liver cancer. *Shijie Huaren Xiaohua Zazhi*, 1999;7:350-352
- 26 Wu MC, Shen F. Progress in research of liver surgery in China. *World J Gastroenterol*, 2000;6:773-776
- 27 Fan J, Wu ZQ, Tang ZY, Zhou J, Qiu SJ, Ma ZC, Zhou XD, Ye SL. Multimodality treatment in hepatocellular carcinoma patients with tumor thrombi in portal vein. *World J Gastroenterol*, 2001;7:28-32
- 28 Miyoshi S, Minami Y, Kawata S, Imai Y, Saitoh R, Noda S, Tamura S, Tarui S. Changes in hepatic functional reserve after transcatheter embolization of hepatocellular carcinoma. *J Hepatol*, 1998;6:332-336
- 29 Clip Group (Cancer of the liver Italian programme). Tamoxifen in treatment of hepatocellular carcinoma: a randomised controlled trial. *Lancet*, 1998;352:17-20
- 30 Koike Y, Shiratori Y, Sato S, Obi S, Teratani T, Imamura M, Yoshida H, Shiina S, Omata M. Des- γ -carboxy prothrombin as a useful predisposing factor for the development of portal venous invasion in patients with hepatocellular carcinoma. *Cancer*, 2001;91:561-569
- 31 Nakamoto T, Inagawa H, Takagi K, Soma GI. A new method of antitumor therapy with a high dose of TNF perfusion for unresectable liver tumors. *Anticancer Res*, 2000;20:4087-4096
- 32 Caturelli E, Siena DA, Fusilli S, Villani MR, Schiavone G, Nardella M, Balzano S, Florio F. Transcatheter arterial chemoembolization for hepatocellular carcinoma in patients with cirrhosis: evaluation of damage to nontumorous liver tissue-long-term prospective study. *Radiol*, 2000;215:123-128
- 33 Hirota S, Matsumoto S, Tomita M, Sako M, Kono M. Retrograde transvenous obliteration of gastric varices. *Radiology*, 1999;211:349-56
- 34 Matsumoto A, Hamamoto N, Nomura T, Hongou Y, Arisaka Y, Morikawa H, Hirata I, Katsu KI. Balloon-occluded retrograde transvenous obliteration of high risk gastric fundal varices. *Am J Gastroenterol*, 1999; 94:643-649
- 35 Ahari HK, Feldman L, Kaufman JA, Gianturco LE. Vascular and interventional case of the day. Peristomal varices. *AJR*, 1999;173:829, 831-832
- 36 Taniai N, Onda M, Tajiri T, Yoshida H, Mamada Y. Interventional radiology and endoscopic therapy for recurrent esophageal varices. *Hepatogastroenterology*, 2001;48:133-136
- 37 Sniderman KW. Hepatocellular carcinoma with portal vein tumor thrombus. *Radiology*, 1998;207:552-553
- 38 Imazu H, Matsui T, Noguchi R, Asada R, Miyamoto Y, Kawata M, Nakayama M, Matsuo N, Matsumura M, Fukui H. Magnetic resonance angiography for monitoring prophylactic endoscopic treatment of high risk esophageal varices. *Endoscopy*, 2000;32:766-772
- 39 Desautels SG, Slivka A, Schoen RE, Carr B, Rabinovitz M, Silverman W. Gastrointestinal bleeding in cirrhotic patients with hepatocellular carcinoma undergoing intrahepatic artery chemotherapy. *Gastrointestendosc*, 1997;46:430-434
- 40 Yamakado K, Tanaka N, Nakatsuka A, Matsumura K, Takase K, Takeda K. Clinical efficacy of portal vein stent placement in patients with hepatocellular carcinoma invading the main portal vein. *J Hepatol*, 1999;30:660-668
- 41 Letier MH, Krige JEJ, Lemmer ER, Terblanche J. Injection sclerotherapy for variceal bleeding in patients with irresectable hepatocellular carcinoma. *Hepatogastroenterology*, 2000;47:1680-1684
- 42 Zhang XL, Qiu SJ, Chang RM, Zou CJ. Animal experiments and clinical application of CT during percutaneous splenoportography. *World J Gastroenterol*, 1998;4:214-218
- 43 Liang HL, Yang CF, Pan HB, Chen CKH, Chang JM. Percutaneous transsplenic catheterization of the portal venous system. *Acta Radiol*, 1997;38:292-295