

Hepatic Resection for Hepatocellular Carcinoma in Patients With Portal Hypertension

A Long-Term Benefit Compared With Transarterial Chemoembolization and Thermal Ablation

Hua Xiao, MD, Binhao Zhang, MD, Bin Mei, MD, Chaohui Zuo, MD, Gang Wei, MD, Rui Wang, MD, Bixiang Zhang, MD, and Xiaoping Chen, MD

Abstract: The optimal treatment for hepatocellular carcinoma (HCC) in cirrhotic patients with portal hypertension (PHT) is still controversial. The objective of this study is to compare HCC patients with PHT treated with hepatic resection to those treated with transarterial chemoembolization (TACE) or thermal ablation.

A series of 167 cirrhotic patients with HCC undergoing hepatic resection or TACE/ablation from 2001 to 2008 were retrospectively analyzed. Cirrhotic patients with HCC were divided into 3 groups: hepatic resection in HCC patients with PHT (PHT-R group, $n = 58$), without PHT (NPHT-R group, $n = 67$), and TACE or thermal ablation in HCC patients with PHT (PHT-O group, $n = 42$). The short-term and long-term outcomes of liver function, operative mortality and morbidity, and survival rate were compared.

Baseline characteristics were similar among the 3 groups, except for patients in the PHT-R group had larger spleen (16.0 vs 11.4 cm, $P = 0.001$) and smaller tumor size (4.8 vs 7.1 cm, $P = 0.001$) in comparison with those in the NPHT-R group. The PHT-R group had better liver function compared with those in the PHT-O group (patients had Child–Turcotte–Pugh class B liver function: 5.2% vs 31%, $P = 0.001$). There was no significant difference of operative mortality and morbidity in all groups. The 1-, 3-, 5-year survival rates were 80.4%, 55.6%, and 28.1% in the PHT-R group; 79.1%, 64.2%, and 39.8% in the NPHT-R group (vs PHT-R, $P = 0.313$); and 60.7%, 24.4%, and 7.3% in the PHT-O group (vs PHT-R, $P < 0.001$).

Hepatic resection shows better long-term results for cirrhotic HCC patients with PHT than TACE and thermal ablation.

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Abbreviations: AASLD = American Association for the Study of Liver Diseases, AFP = α -fetoprotein, BCLC = Barcelona Clinic Liver Cancer, CTP = Child–Turcotte–Pugh, EASL = European Association for the Study of the Liver, HBsAg = hepatitis B surface antigen, HCC = hepatocellular carcinoma, HCVAb = hepatitis C virus antibody, INR = international normalized ratio, MELD = Model for End-Stage Liver Disease, PHT = portal hypertension, TACE = transarterial chemoembolization.

INTRODUCTION

Hepatocellular carcinoma (HCC) is the sixth most common malignancy and the third leading cause of cancer-related deaths in the world, and usually arises in the setting of cirrhosis.¹ Esophageal varices is considered a sign of portal hypertension (PHT) and observed in up to 67% of cirrhotic patients.² Although hepatic resection for HCC has become a safe procedure due to improved surgical techniques and pre- and post-surgery care, PHT is still considered a contraindication for liver resection according to the EASL (European Association for Study of Liver)³ and AASLD (American Associations for Study of Liver Diseases)⁴ published guidelines for HCC management in 2001 and 2011, respectively. Other treatments such as liver transplantation, thermal ablation, and transarterial chemoembolization (TACE) are recommended in these patients.

However, several studies demonstrated that if patients had a similar preoperative liver function, assessed by Child–Turcotte–Pugh (CTP) score or Model for End-Stage Liver Disease (MELD) score,^{5–8} the short- and long-term outcomes were similar in patients with or without PHT. And to the best of our knowledge, whether TACE or thermal ablations for HCC with PHT are superior to hepatic resection is still controversial. The aim of this study is to compare the therapeutic value of hepatic resection with TACE or thermal ablation for HCC in cirrhotic patients with PHT and to reevaluate the current guidelines for HCC management.

METHODS

From May 2001 to December 2008, 167 cirrhotic patients with or without PHT were retrospectively analyzed at the Hepatic Surgery Center, Tongji Hospital, Huazhong University of Science and Technology. This study was approved by the Ethics Committee for Clinical Pharmacology in Tongji Medical College, and all the information of patients were kept private. Patients were divided into 3 groups: presence (PHT-R group, $n = 58$) or absence (NPHT-R group, $n = 67$) of PHT at the time

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From the Hepatic Surgery Center (HX, Binhao Z, BM, GW, RW, Bixiang Z, XC), Tongji Hospital, Tongji Medical College, Huazhong University of Science and Technology, Wuhan; and Department of Gastrointestinal and Pancreatic Surgery (HX, CZ), the Affiliated Tumor Hospital of Xiangya Medical School of Central South University, Changsha, China.

Correspondence: Xiaoping Chen, Hepatic Surgery Center, Tongji Hospital, Tongji Medical College, Huazhong University of Science and Technology, 1095 Jie Fang Avenue, Wuhan, China 430030 (e-mail: chenxp@medmail.com.cn).

Hua Xiao and Binhao Zhang are co-first authors.

These authors contributed equally to this work.

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of hepatic resection, and those with PHT who were performed other treatments including TACE and thermal ablation (PHT-O group, $n=42$). Data were recruited consecutively to address potential sources of bias, and were analyzed retrospectively. The presence of preoperative PHT was defined as esophageal varices detectable at endoscopy or splenomegaly with a platelet count $<100 \times 10^3/\text{mm}^3$ according to the Barcelona Clinic Liver Cancer (BCLC) group criteria.⁹ The characteristics evaluated to define PHT are described in Table 1. The diagnosis of HCC was confirmed by histological examination or at least 2 radiologic imaging showing characteristic features of HCC, or persistent raised α -fetoprotein (AFP) $>400 \text{ ng/mL}$ together with 1 radiologic imaging showing characteristic features of HCC.³

Except for endoscopy, platelet count, and spleen major diameter, the following clinical and biochemical features were collected for each patient: age, gender, level of hepatitis B surface antigen (HBsAg), hepatitis C virus antibody (HCV-Ab), AFP (ng/mL), total bilirubin ($\mu\text{mol/L}$), albumin (g/L), prothrombin time, and international normalized ratio (INR). The CTP score was obtained according to described classification proposed by Pugh et al¹⁰ and the MELD score was calculated by the formula described by Freeman et al.¹¹ Patients with tumors involving the main portal vein or inferior vena, liver function classified as CTP class C, extrahepatic metastasis, or other major systemic disease were excluded from this study.

Most procedures were performed anatomically and all obtained a tumor-free margin.¹² Major hepatic resection was defined as removing more than 2 segments⁸; the extent of the hepatectomy was based on the International Hepato-Pancreato-Biliary Association classification.¹³ Pringle maneuver was commonly used for bleeding control during the resection. In the PHT-R group, for patients with a platelet count of $<50 \times 10^3/\text{mm}^3$, we additionally performed concomitant splenectomy with liver resection to reduce the risk of hemorrhagic complications.¹⁴ For patients who had history of variceal bleeding, pericardial devascularization surgery first described by Qiu¹⁵ was concomitantly performed. For patients who refused surgery or in whom complete removal of the tumors was considered impossible, if patients with single nodule $<5 \text{ cm}$ or 2 to 3 nodules each $<3 \text{ cm}$, radiofrequency or microwave ablation was performed, and TACE was proposed for patients with single nodule $>5 \text{ cm}$ or more than 3 nodules.

Perioperative mortality was defined as death in the hospital during the initial admission for surgery or death within 30 days after hepatic resection. Postoperative liver dysfunction was assessed by using the "50–50 criteria"¹⁶: serum bilirubin exceeding $50 \mu\text{mol/L}$ (3 mg/dL) and prothrombin time $<50\%$ on

postoperative day 5. After surgery, the majority of patients received adjuvant regional chemotherapy via the portal vein or hepatic artery through a subcutaneously implanted port at intervals of about 4 weeks by using 5-fluorouracil, mitomycin C, or epirubicin in the first 6 months following surgery. All patients were followed every 3 to 6 months, consisting of serum AFP level, biochemical liver function, and US or CT. The mean follow-up time after surgery was 35.8 months (range 1–120 months). When recurrence occurred beyond 1 year after resection, repeated liver resection was performed provided that the liver function permitted. For recurrence occurred within 1 year after resection or inappropriate for reoperation, radiofrequency or microwave ablation, ethanol injection, or TACE was performed.

STATISTICAL ANALYSIS

Continuous variables were expressed as mean and standard deviation and differences between groups were compared by the independent-samples *t* test or Mann–Whitney U-test, as appropriate; categorical variables were reported as number of cases and prevalence, and differences between groups were compared by the chi-square test or Fisher exact test, as appropriate. The patients' overall survival was calculated by using the Kaplan–Meier method, and their comparison was performed using the log-rank test. The multivariate analysis was performed using the Cox regression model. *P* value <0.05 was considered significant in all analysis. Statistical analysis was carried by SPSS 17.0 software.

RESULTS

Preoperative Characteristics and Intraoperative Data

The clinical characteristics and intraoperative data among the 3 groups are summarized in Tables 2 and 3. In comparison with patients in NPHT-R group, those in the PHT-R group had significant larger spleen ($16.0 \text{ vs } 11.4 \text{ cm}$, $P=0.001$), smaller tumor size ($4.8 \text{ vs } 7.1 \text{ cm}$, $P=0.001$), lower rate of Pringle maneuver use ($46.6\% \text{ vs } 70.1\%$, $P=0.007$), and higher rates of concomitant splenectomy ($24.1\% \text{ vs } 0$, $P<0.001$) or pericardial devascularization surgery ($17.2\% \text{ vs } 0$, $P<0.001$). The CTP classification, MELD score, mean surgical time, blood loss, transfusion rate, and in-hospital stay were all similar between the 2 groups. In the PHT-O group, 24/42 patients underwent microwave ablation, 15/42 patients underwent TACE, and the remaining 3 patients were performed radiofrequency ablation. And patients in the PHT-O group had worse liver function (patients had CTP class B liver function: $31\% \text{ vs } 5.2\%$, $P=0.001$) compared with those in the PHT-R group.

TABLE 1. Clinical Characteristics Related to Presence of Portal Hypertension as Defined by the Barcelona Clinic Liver Cancer Group Criteria

Variables	PHT-R (n = 58)	NPHT-R (n = 67)	PHT-O (n = 42)
Esophageal varices	28 (48.3%)	0	22 (52.4%)
Platelet count $<100,000/\text{mm}^3$	49 (84.5%)	5 (7.5%)	37 (89.0%)
Platelet count ($\times 10^3/\text{mm}^3$; median; range)	79.0 (21–205)	146.8 (30–294)	66.6 (17–309)
Splenomegaly	53 (91.4%)	19 (28.4%)	38 (90.5%)
Esophageal varices without splenomegaly + platelet count $<100,000/\text{mm}^3$	9 (15.5%)	0	6 (14.3%)
Esophageal varices + splenomegaly + platelet count $<100,000/\text{mm}^3$	19 (32.8%)	0	16 (38.1%)
Splenomegaly + platelet count $<100,000/\text{mm}^3$ without esophageal varices	30 (51.7%)	0	20 (47.6%)

NPHT = ??, PHT = portal hypertension.

TABLE 2. Baseline Characteristics of the Whole Study Population

Variables	PHT-R (n = 58)	NPHT-R (n = 67)	PHT-O (n = 42)	P	
Age (y)	49.9 ± 11.8	46.9 ± 10.0	47.7 ± 9.4	0.207*	0.312 [†]
Male gender	52 (89.7%)	64 (95.5%)	38 (90.5%)	0.207	0.893
Hepatitis serology				0.618	0.658
Hepatitis B	54 (93.1%)	61 (91.0%)	37 (88.1%)		
Hepatitis C	2 (3.4%)	1 (1.5%)	3 (7.1%)		
Hepatitis B + C	0	1 (1.5%)	0		
Negative	2 (3.4%)	4 (6.0%)	2 (4.8%)		
Albumin (g/L)	38.1 ± 4.4	41.0 ± 4.0	34.9 ± 6.0	0.355	0.108
Total bilirubin (μmol/L)	18.8 ± 8.4	15.4 ± 6.4	24.9 ± 11.5	0.175	0.178
CTP classification				0.336	0.001
Class A	55 (94.8%)	66 (98.5%)	29 (69.0%)		
Class B	3 (5.2%)	1 (1.5%)	13 (31%)		
α-Fetoprotein (ng/mL)				0.471	0.425
<200	34 (58.6%)	32 (47.8%)	27 (64.3%)		
200–400	5 (8.6%)	8 (11.9%)	1 (2.4%)		
>400	19 (32.8%)	27 (40.3%)	14 (33.3%)		
MELD classification				0.815	0.649
Score <9	48 (82.8%)	58 (86.6%)	32 (76.2%)		
Score 9–10	5 (8.6%)	5 (7.5%)	6 (14.3%)		
Score >10	5 (8.6%)	4 (6.0%)	4 (9.5%)		
Tumor size (cm)				0.005	0.664
<3	9 (15.5%)	7 (10.4%)	9 (21.4%)		
3–5	23 (48.3%)	11 (16.4%)	18 (42.9%)		
>5	25 (36.2%)	49 (73.1%)	15 (35.7%)		
Number of nodules				0.756	0.063
Single	48 (82.8%)	54 (80.6%)	28 (66.7%)		
Multiple	10 (17.2%)	13 (19.4%)	14 (33.3%)		
Adjuvant regional chemotherapy	35 (60.3%)	46 (68.7%)	—	0.332	—

Continuous variables are reported in mean and standard deviation. CTP = Child–Turcotte–Pugh, MELD = model for end-stage liver disease, NPHT = ??, PHT = portal hypertension .

*This value indicates the comparison results between the patients with (PHT-R group) or without (NPHT-R group) PHT at the time of hepatic resection.

[†]This value denotes the comparison results between the patients with PHT who underwent hepatic resection (PHT-R group) or other treatments (PHT-O group).

Short-Term Results

In the PHT-R group, 1 patient underwent reoperation due to intra-abdominal bleeding 18 hours after the initial surgery. Although bleeding was successfully controlled, the patient died

of liver failure on the ninth day after surgery (Table 4). There was no operative death in the other 2 groups. Postoperative complications rate was 32.8% in the PHT-R group, which was similar compared with those in the NPHT-R (34.3%) and PHT-

TABLE 3. Intraoperative Data of Hepatectomy for Hepatocellular Carcinoma in Cirrhotic Patients With (PHT-R Group) or Without (NPHT-R Group) Portal Hypertension (n = 125)

Variables	PHT-R (n = 58)	NPHT-R (n = 67)	P
Mean surgical time (min)	209.0 ± 58.5	186.8 ± 49.3	0.197
Mean blood loss (mL)	529.3 ± 406.6	600.4 ± 547.3	0.181
RBC transfusion (mL)	19 (32.8%)	12 (17.9%)	0.055
Anatomic resection	51 (87.9%)	57 (85.1%)	0.642
Extent of hepatectomy			0.176
Wedge and segmentectomy	44(75.9%)	41(61.2%)	
Bisegmentectomy	12(20.7%)	20(29.9%)	
Major hepatectomy	2(3.4%)	6(9.0%)	
Pringle maneuver	27 (46.6%)	47 (70.1%)	0.007
Concomitant splenectomy	14 (24.1%)	0	<0.001
Concomitant pericardial devascularization	10 (17.2%)	0	<0.001

NPHT = ??, PHT = indicates portal hypertension, RBC = red blood cell.

TABLE 4. Deaths and Complications After Hepatic Resection or Other Treatments (PHT-O Group) for Hepatocellular Carcinoma in Cirrhotic Patients With (PHT-R Group) or Without (NPHT-R Group) Portal Hypertension

Variables	PHT-R (n = 58)	NPHT-R (n = 67)	PHT-O (n = 42)
Deaths	1 (1.7%)	0	0
Complications	19 (32.8%)	23 (34.3%)*	11 (26.2%) [†]
Liver dysfunction	9 (15.5%)	6 (9.0%) [‡]	2 (4.8%) [§]
Ascites	0	1 (1.5%)	0
Bile leakage	0	1 (1.5%)	0
Intra-abdominal abscess	4 (6.9%)	2 (3.0%)	2 (4.8%)
Atelectasis	0	1 (1.5%)	0
Pleural effusion	9 (15.5%)	12 (17.9%)	3 (7.1%)
Intra-abdominal hemorrhage	3 (5.2%)	1 (1.5%)	0
Wound infection	1 (1.7%)	1 (1.5%)	0
Pneumonia	0	0	1 (2.4%)
Wound dehiscence	1 (1.7%)	1 (1.5%)	0
Gastrointestinal bleeding	1 (1.7%)	1 (1.5%)	0
Pneumothorax	0	2 (3.0%)	0
Hepatic encephalopathy	0	0	1 (2.4%)
Postembolization syndrome	0	0	2 (4.8%)
Hospital stay (d)	20.7 ± 11.8	19.1 ± 8.5	16.3 ± 10.3

NPHT = ??, PHT = indicates portal hypertension.

* Compared with PHT-R group, *P* = 0.853.

[†] Compared with PHT-R group, *P* = 0.478.

[‡] Compared with PHT-R group, *P* = 0.260.

[§] Compared with PHT-R group, *P* = 0.114.

O (26.2%) groups (*P* = 0.853 and 0.478, respectively). According to the Clavien–Dindo classification of surgical complication,¹⁷ the incidence of grade IIIa or more severe morbidities was also not significantly different between the PHT-R group (17.2%) and the NPHT-R group (19.4%, *P* = 0.756).

Long-Term Outcomes

Patients with preoperative PHT who were performed hepatic surgery had similar 1-, 3-, and 5-year survival rates compared with those without PHT: 80.4% and 79.1%, 55.6% and 64.2%, and 28.1% and 39.8%, respectively (*P* = 0.313), which were significantly better than those treated with TACE or thermal ablation (60.7%, 24.4%, and 7.3%, respectively, *P* < 0.001) (Figure 1). As shown in Figure 2, tumor-free survivals were also comparable in the PHT-R and NPHT-R groups (*P* = 0.258).

Long-Term Outcomes in CTP Class A Patients

Considering only patients with CTP class A liver function (150 patients: 55 in the PHT-R group, 66 in the NPHT-R group, and 29 in the PHT-O group), patients in the PHT-R group had similar 1-, 3-, and 5-year survival rates compared with those in the NPHT-R group: 81.2% and 78.8%, 57% and 63.6%, and 28.7% and 38.8%, respectively (*P* = 0.336), which were significantly better than those treated with TACE or local ablation (67.7%, 22.8%, and 9.1%, respectively, *P* = 0.002) (Figure 3).

Univariate and Multivariate Analysis

Based on the univariate analysis, the best predictors of a poor prognosis for cirrhotic patients with HCC were age over 60 years, splenomegaly, albumin <35 g/L, multiple number of nodules, and serum AFP level over 200ng/mL (Table 5). Multivariate analysis of these 5 factors showed only AFP level and number of nodules were identified as an independent predicting

factors for survival (*P* = 0.048, 0.030, respectively) (Table 6). Although neither presence of PHT nor elevated serum bilirubin level was associated with overall survivals.

DISCUSSION

Hepatic resection for HCC has become a safe procedure in cirrhotic patients, with lower postoperative mortality, morbidity and improved long-term results.^{18–20} This progress has been mainly due to improved perioperative managements and better

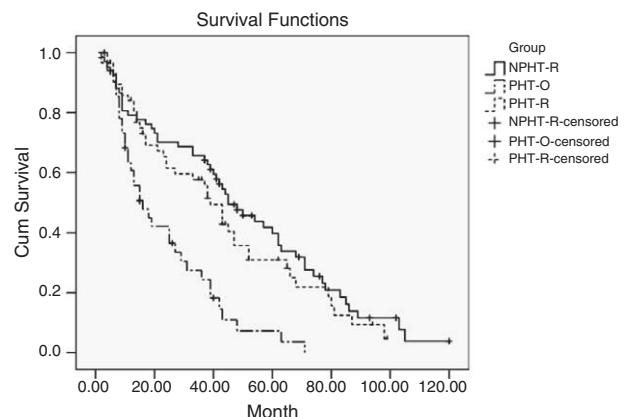


FIGURE 1. Overall survival curves of 167 cirrhotic patients undergoing liver resection for hepatocellular carcinoma with (PHT-R group) or without portal hypertension (NPHT-R group), and those with portal hypertension but performed other treatments (PHT-O group) (Kaplan–Meier method, long-rank test, *P* = 0.313 between PHT-R and NPHT-R groups; *P* < 0.001 between PHT-R and PHT-O groups).

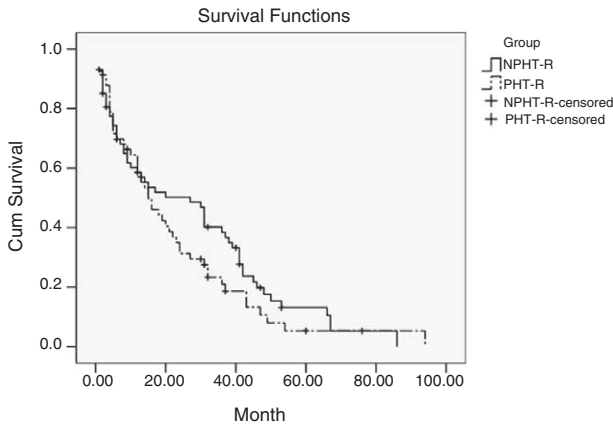


FIGURE 2. Tumor-free survivals of 125 cirrhotic patients undergoing liver resection for hepatocellular carcinoma with (PHT-R group) or without (NPHT-R group) portal hypertension (Kaplan–Meier method, long-rank test, $P=0.258$).

patient evaluation, but the definition of selection criteria is still controversial. According to the guidelines published by EASL and AASLD in 2001 and 2011, surgical resection was only recommended in noncirrhotic patients or in patients who have cirrhosis but still have well-preserved liver function, normal bilirubin, and hepatic vein pressure gradient <10 mm Hg.^{3,4} One reason for narrowing the indication for resection was high incidence of postoperative liver decompensation and poor long-term survivals. This evaluation was mainly based on the 1996 and 1999 studies by the Barcelona group.^{21,22} The 2 studies containing a small series of 29 and 74 CTP class A cirrhotic patients, respectively, proved that increased portal pressure was the most powerful independent factor for postoperative liver decompensation in multivariate analysis. With respect to long-term outcomes, PHT and serum bilirubin levels were revealed to be the independent prognostic factors for overall survival after resection. Choi et al²³ also reported that liver-related

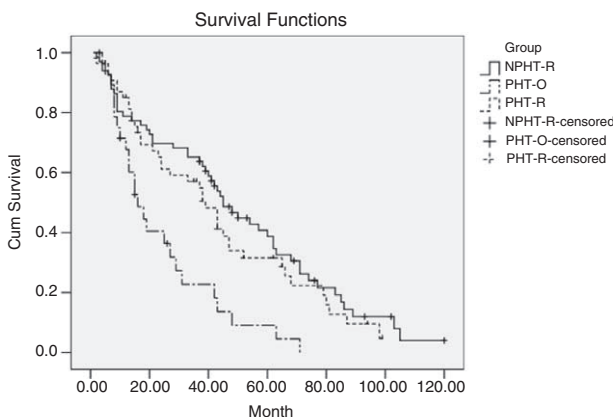


FIGURE 3. Overall survival curves of 150 CTP class A cirrhotic patients undergoing liver resection for hepatocellular carcinoma with (PHT-R group) and without portal hypertension (NPHT-R group), and those with portal hypertension but performed other treatment (PHT-O group) (Kaplan–Meier method, long-rank test, $P=0.336$ between PHT-R and NPHT-R groups; $P=0.002$ between PHT-R and PHT-O groups; $P<0.001$ between NPHT-R and PHT-O groups).

TABLE 5. Univariate Analysis of Predictive Factors for Overall Survival After Hepatectomy for Hepatocellular Carcinoma in Cirrhotic Patients With (PHT-R Group) or Without (NPHT-R Group) Portal Hypertension ($n=125$)

Variables	No.	P
Age >60 y (yes/no)	20/105	0.045
Gender (male/female)	116/9	0.340
HBsAg (positive/negative)	116/9	0.716
Platelet $>10^5/mm^3$ (yes/no)	73/52	0.711
Splenomegaly (yes/no)	76/49	0.030
Esophageal varices (yes/no)	27/98	0.763
Portal hypertension (yes/no)	58/67	0.313
Albumin >35 g/L (yes/no)	106/19	0.042
Total bilirubin >20.5 μ mol/L (yes/no)	34/91	0.660
Blood transfusion (yes/no)	31/94	0.418
Tumor diameter >5 cm (yes/no)	75/50	0.937
Number of nodules (single/multiple)	102/23	0.014
CTP classification (A/B)	121/4	0.882
α -Fetoprotein (ng/mL) >200 (yes/no)	58/67	0.044
MELD classification score p)	19/106	0.778
(Kaplan–Meier method, long >9 (yes/no)		
Extent of hepatectomy <2 segments (yes/no)	85/40	0.211

CTP = Child–Turcotte–Pugh, HBsAg = hepatitis B surface antigen, MELD = model for end-stage liver disease.

complications were significantly higher in the PHT group and the 5-year overall survival rate was significantly higher in the non-PHT group. But this evaluation is not widely accepted, and whether the existence of PHT has been considered a prognostic factor remains unclear.^{24,25}

In 2009, Cucchetti et al⁸ published a retrospective analysis consisting of 241 cirrhotic patients divided into 2 groups according to the presence ($n=89$) or absence ($n=152$) of PHT at the time of surgery. The study demonstrated that patients with PHT experienced worse preoperative liver function and survival in comparison to those without PHT; after one-to-one matching by propensity score, patients with ($n=78$) and without PHT ($n=78$) had the same preoperative characteristics and showed the same intraoperative course, postoperative occurrence of liver failure, morbidity, and 5-year survival rates. The only predictors of postoperative liver failure were MELD score and extent of hepatectomy. Several other studies have also concluded that the short- and long-term outcomes were acceptable in patients with PHT, if patients had a similar preoperative liver function.^{5–7} Therefore, the optimal treatment for HCC in cirrhotic patients with PHT is still controversial.

According to the guidelines published by EASL and AASLD, HCC patients with PHT were recommended for other

TABLE 6. Multivariate Analysis

Variables	Hazard Ratio	95% CI	P
Tumor number (multiple)	1.805	1.059–3.076	0.030
α -Fetoprotein >200 ng/mL	1.506	1.004–2.259	0.048

CI = confidence interval.

treatments. Liver transplantation is recognized as the first choice for cirrhotic patients with HCC within Milan criteria, allowed removing the diseased liver together with the tumor itself, but the current scarcity of organs and extremely high cost restricted only a limited number of candidates to be transplanted at specialized centers.²⁶ TACE and local ablation are recommended for patients with HCC who were not appropriate for resection or transplantation according to the guidelines.^{3,4} However, the long-term outcomes were not satisfactory enough and there are numerous reports demonstrating that hepatic resection was superior with overall and tumor-free survivals than TACE or thermal ablations for HCC,^{3,27–32} but whether the result is the same in patients with PHT is still unknown.

As reported by Capussotti et al,⁵ operative mortality related to liver function or portal pressure (liver failure, portal vein thrombosis, esophageal bleeding) was significantly higher in patients with PHT. However, recent advances in surgical techniques and perioperative care for patients with cirrhosis have reduced the number of cirrhosis-related complications and deaths. The occurrence of perioperative morbidity was similar in patients who underwent hepatic resection with or without PHT, as was postoperative liver dysfunction. It seems that patients in the PHT-R group had higher rate of postoperative intra-abdominal hemorrhage requiring reoperation compared with those in the NPHT-R group (5.2% vs 1.5%), though the difference was not significant ($P = 0.244$). But one must bear in mind that patients with PHT usually had prolonged prothrombin time and severe thrombocytopenia, which may be responsible for higher rate of postoperative hemorrhage. For patients in the PHT-R group in this study, there was only 1 case of perioperative death due to liver failure, 8 (13.8%) experienced and 5 (8.6%) died of variceal bleeding during follow-up time, the majority of the deaths were associated with the recurrence of HCC. This was echoed in the study by Ishizawa et al,⁶ who argued that the benefits of hepatic resection in patients with HCC and PHT might outweigh the risks of cirrhosis-related mortality after resection. Moreover, neither PHT nor preoperative bilirubin level was confirmed as an independent predicting factor for survival by multivariate analysis. Our results indicate that liver resection can be performed safely in PHT patients with satisfactory and acceptable 5-year overall survival rate, justifying the surgical indications for HCC with PHT if the liver function was still well compensated.

Patients with PHT who were performed hepatic resection in this study had similar 5-year survivals compared with those reported by Capussotti et al⁵ (28.1% and 28.9%, respectively), but were obviously lower than those in the reports by Ishizawa et al⁶ (56%) and Cucchetti et al⁸ (56.5%). One explanation is that patients in this study had significant larger tumor size. The result was as expected that patients with a more advanced stage of cancer were going to have worse long-term survival. Moreover, most of the patients in our study were HBV related. It was significantly different from the previous published literatures, in which the majority of the patients had HCV as cirrhosis etiology.^{5–8} Patients with HBV or HCV may have distinct clinical profiles and tumor burdens, which have implications for long-term survival.

This study has several limitations. It is a retrospective study with limited number of patients in a single center. A randomized clinical trial with larger number of patients would provide stronger evidence to get a conclusion.

In conclusion, PHT should not be considered as an absolute contraindication to liver resection for HCC in cirrhotic patients with compensated liver functions. Multivariate analysis

identifying serum AFP level and number of tumors can be used as independent predicting factors for survival. This study provided important evidence for surgeons that selected HCC patients with PHT could benefit from hepatic resection as compared with TACE or thermal ablation.

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