

Intraperitoneal metastasis of hepatocellular carcinoma after spontaneous rupture: A case report

Min-Chang Hung, Hurng-Sheng Wu, Yueh-Tsung Lee, Chih-Hung Hsu, Dev-Aur Chou, Min-Ho Huang

Min-Chang Hung, Hurng-Sheng Wu, Yueh-Tsung Lee, Chih-Hung Hsu, Dev-Aur Chou, Min-Ho Huang, Department of Surgery, Chang Bing Show-Chwan Memorial Hospital, Changhua 505, Taiwan, China

Author contributions: Hung MC wrote the paper; Wu HS revised the paper; Lee YT, Hsu CH, Chou DA, and Huang MH performed the research.

Correspondence to: Min-Chang Hung, Department of surgery, Chang Bing Show-Chwan Memorial Hospital, No. 6, Lugang Rd., Lugang Township, Changhua 505, Taiwan, China. hmjohn@mail2000.com.tw

Telephone: +886-4-7813888-73120 Fax: +886-4-7073226

Received: December 17, 2007 Revised: May 19, 2008

Accepted: May 26, 2008

Published online: June 28, 2008

Abstract

Rupture of hepatocellular carcinoma (HCC) is a life-threatening complication. Peritoneal metastasis of HCC after spontaneous rupture was seldom noted. We report a case of intraperitoneal metastasis of HCC after spontaneous rupture. A previously asymptomatic 72-year-old man was admitted due to dull abdominal pain with abdominal fullness. He had a history of HCC rupture 10 mo ago and transarterial embolization was performed at that time. Abdominal computer tomography (CT) scan showed a huge peritoneal mass over the right upper quadrant area. Surgical resection was arranged and subsequent microscopic examination confirmed a diagnosis of moderately-differentiated HCC.

© 2008 The WJG Press. All rights reserved.

Key words: Hepatocellular carcinoma; Spontaneous rupture; Peritoneal metastasis

Peer reviewers: Fritz E von Weizsacker, Professor, Department of Medicine, Schlosspark Klinik, Humboldt University, Berlin 14059, Germany; Dr. Paolo Del Poggio, Hepatology Unit, Department of Internal Medicine, Trevisio Hospital, Piazza Ospedale 1, Trevisio Bg 24047, Italy

Hung MC, Wu HS, Lee YT, Hsu CH, Chou DA, Huang MH. Intraperitoneal metastasis hepatocellular carcinoma after spontaneous rupture: A case report. *World J Gastroenterol* 2008; 14(24): 3927-3931 Available from: URL: <http://www.wjgnet.com/1007-9327/14/3927.asp> DOI: <http://dx.doi.org/10.3748/wjg.14.3927>

INTRODUCTION

Hepatocellular carcinoma (HCC) is one of the most common malignancies worldwide. It is usually manifested in the 6th and 7th decades of life. Extrahepatic metastases are seen in 64% of patients with HCC. The most frequent sites of extrahepatic metastases are lung, abdominal lymph node and bone, but peritoneal dissemination is unusual^[1,2]. The incidence of spontaneous rupture of HCC is about 8%-26% in Asia^[3-5] and the mortality rate of HCC patients is 10%^[6]. However, peritoneal metastasis of HCC after spontaneous rupture is seldom noted. Here, we report a case of intraperitoneal metastasis of HCC after spontaneous rupture 10 mo ago, which was treated with transarterial embolization.

CASE REPORT

A previously asymptomatic 72-year-old man had a history of chronic hepatitis C-related liver cirrhosis without regular follow-up. Sudden nausea and vomiting with watery diarrhea were noted on January 2006. Then he was sent to Yun-Lin Branch of National Taiwan University Hospital for help. Abdominal computer tomography (CT) scan showed a huge HCC that was suspicious of rupture. Under the request of his family, he was transferred to our hospital and transarterial embolization was performed on January 31, 2006. After discharge, he was regularly followed up at our Gastrointestinal (GI) Outpatient Department (OPD). Dull abdominal pain over the right upper quadrant area, accompanied with fullness sensation, was noted in November 2006. Besides, he also had body weight loss of about ten kilograms in one year. So he visited our hospital again. Abdominal CT scan revealed a peritoneal mass in the right upper quadrant peritoneal area and hepatoma recurrence was considered (Figure 1). Transarterial embolization was arranged again, but failed. After consultation with the surgeon, he was admitted for surgical resection.

Surgical intervention was arranged on January 24, 2007. Operative methods were segmental hepatectomy (S6 and partial S5), excision of extrahepatic tumor, and cholecystectomy. The operation showed a huge tumor (12 cm × 8 cm × 6 cm) over the right upper quadrant area just below liver parenchyma (Figure 2) with its blood supplied from the omentum. Besides, two small mass lesions (3 cm × 2 cm and 2 cm × 1 cm) were found over

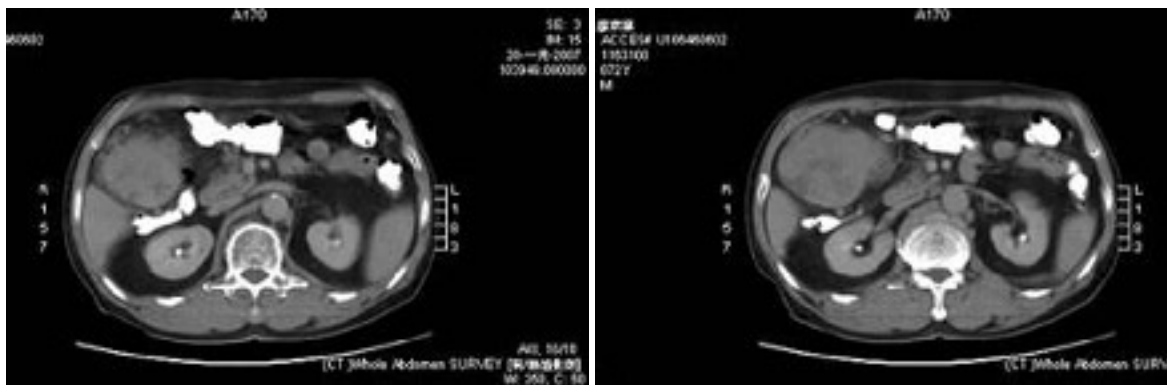


Figure 1 A well-defined mass about 10 cm in diameter in RUQ peritoneal cavity anterior to liver parenchyma.

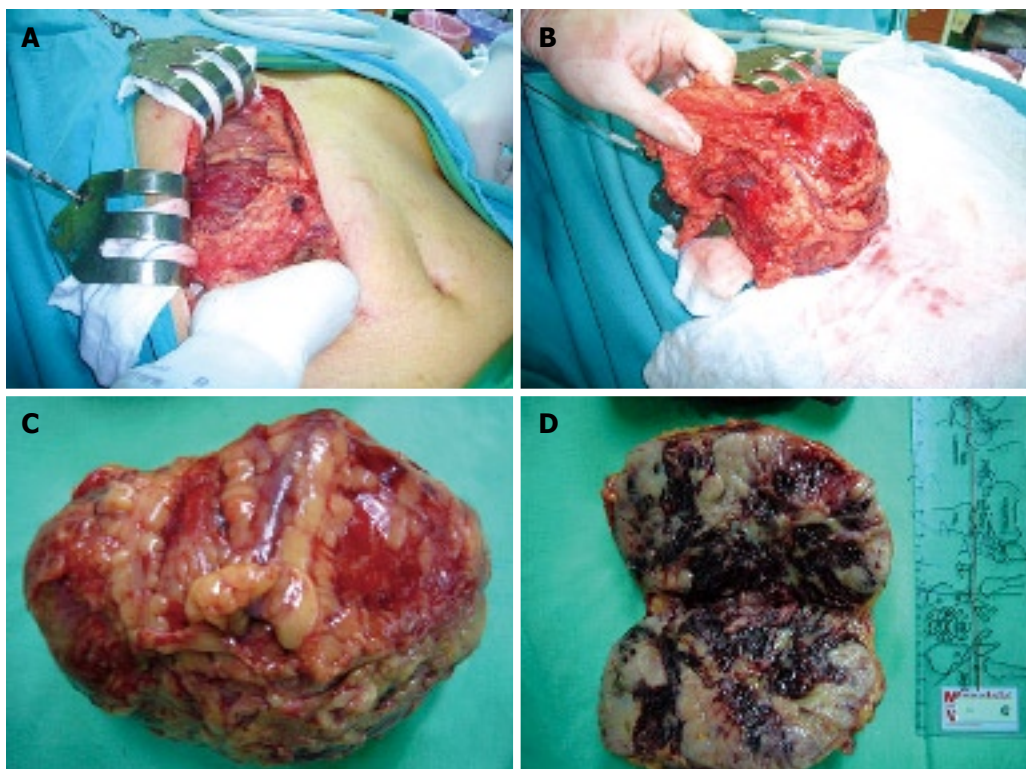


Figure 2 A huge tumor (12 cm × 8 cm × 6 cm) over the right upper quadrant area just below liver (A), blood supply of tumor from the omentum (B), and intraperitoneal tumor (C, D).

S5 and S6, respectively (Figure 3). Microscopy showed that the huge extrahepatic tumor and two intrahepatic lesions were moderately differentiated.

DISCUSSION

Intraperitoneal metastasis of non-ruptured HCC is rare, but the risk of such a metastasis of ruptured HCC increases^[7]. The mechanism of spontaneous rupture of HCC is not exactly clear. Hypotheses include rapid expansion of the tumor and central necrosis, venous hypertension caused by venous obstruction due to direct tumor invasion, mild trauma or compression by the diaphragm associated with respiratory movement, coagulopathies such as thrombocytopenia and disturbed prothrombin synthesis, and vascular injury leading to hemorrhage and subsequent rupture^[4,8-10]. Recently, Zhu *et al* postulated that the poor function of macrophage phagocytosis could result in cumulation of immune

complex and deposition on vascular wall. Then vascular wall could become stiff and weak due to the proliferated fragment elastin and damaged collagen, which would make blood vessels more prone to splitting and result in hemorrhage and rupture of HCC^[11,12]. Large tumor size, peripheral location and protruding contour are all associated with an increased risk for rupture of HCC^[4,13,14].

The diagnosis of HCC rupture is based on blood-stained ascites plus imaging studies and symptoms^[2]. The most common symptom is sudden onset of abdominal pain (66%-100%)^[6,15-17]. Yeh *et al* also found that the presence of sudden-onset abdominal pain is the only independent indicator of ruptured HCC^[18]. Abdominal ultrasonography and computed tomography improve the rate of preoperative diagnosis^[13,14,19,20]. Peripheral location, protruding contour, discontinuity of hepatic surface, surrounding hematoma and elevated ascitic CT number are helpful signs for the diagnosis of ruptured HCC. In addition, enucleation sign on helical CT could

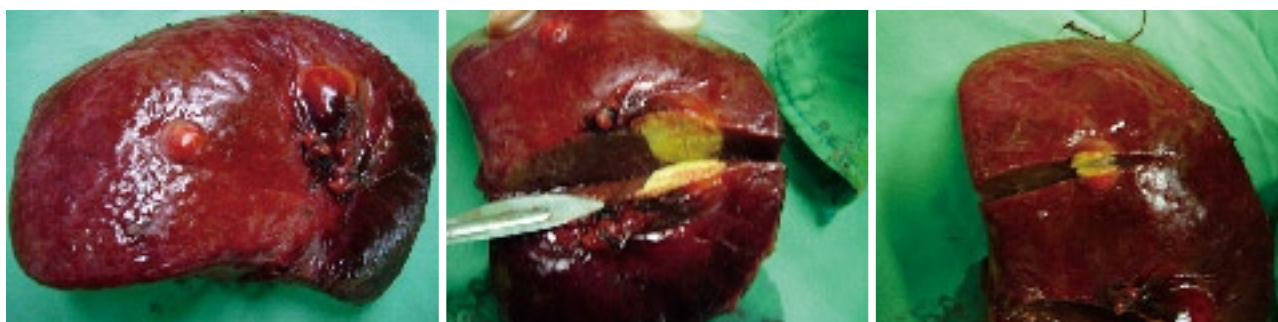


Figure 3 Two small mass lesions (3 cm × 2 cm and 2 cm × 1 cm) over S5 and S6, respectively.

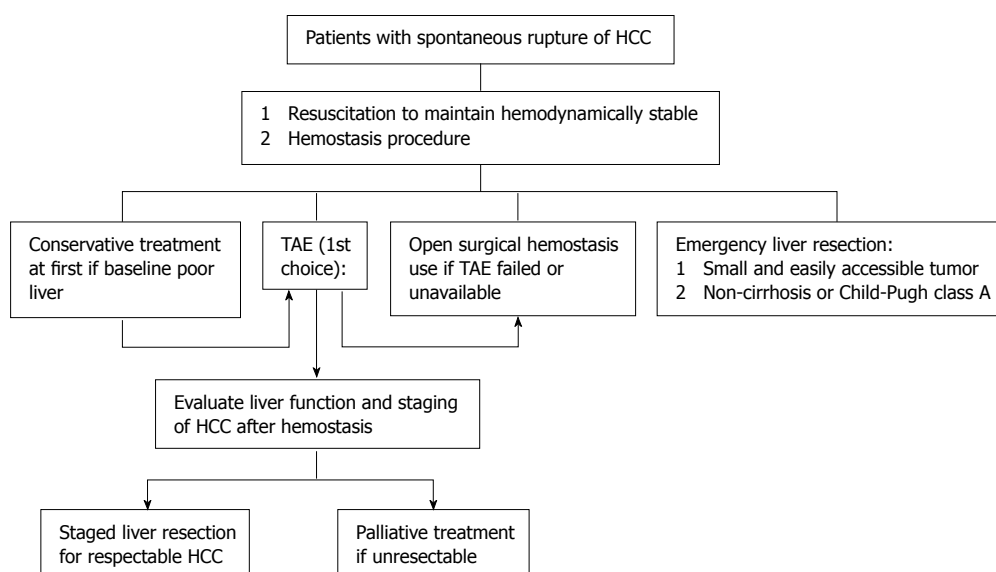


Figure 4 Logarithm about how to approach to the patient with spontaneously ruptured HCC.

be more specific^[14,20,21].

Treatment of ruptured HCC is primarily aimed at controlling hemorrhage and preserving the functional liver parenchyma as possible. Open surgical method was the mainstay of treatment during 1960s-1980s. It was reported that various surgical procedures, including perihepatic packing, suture plication of bleeding tumors, injection of alcohol, hepatic artery ligation (HAL), and liver resection, are effective against hemostasis^[10,22-26]. Besides transarterial embolization (TAE) and transarterial chemoembolization (TACE) for palliative treatment in patients with unresectable HCC, TAE is also gradually used for hemostasis in spontaneous rupture of HCC. In addition, Ng *et al*^[27] also reported that ruptured HCC could be treated with radiofrequency ablation as a salvage procedure. To our knowledge, no prospective randomized controlled trials have found the best method for hemostasis. There is evidence that TACE is the preferred method to arrest tumor bleeding^[13,28-34]. Besides, two-stage hepatectomy is advisable because it can prolong the survival of selected patients^[35-40]. Figure 4 is the logarithm about how to approach the patients with spontaneous ruptured HCC^[35-40].

Spontaneous rupture of HCC with intraperitoneal hemorrhage is a life-threatening complication with a high mortality rate. Prognosis is associated with poor liver reserve, advanced disease and severity of hemor-

rhage^[41]. The median survival time is around 4-5 mo after HCC rupture, and only few patients can have a long-term survival^[42,43]. Thus, implanted metastases usually do not become clinically apparent. Ong *et al*^[44] reported the first case of peritoneal metastasis after HCC rupture in 1996 and then only sporadic case reports have been published. Most reported cases of peritoneal metastases are documented 8 mo after rupture^[7,41,45-47], but Ryu *et al*^[48] and Lin *et al*^[49] reported that peritoneal metastasis can be found 4 and 3 mo respectively after rupture episode. A single metastatic tumor is the most common presentation. Resection is the treatment of choice for peritoneal metastasis if possible and might offer long-term survival benefits^[7,44-48].

Peritoneal metastasis after spontaneous rupture of HCC is rare. This is our first experience with such a patient. Our patient developed peritoneal metastasis, which was documented 10 mo after spontaneous HCC rupture. The time from rupture to documentation of peritoneal metastasis is similar to other case reports. Because few cases of peritoneal metastasis after ruptured HCC have been reported, the association between metastatic tumor and viral infection (HBV or HCV), AFP level, or age is lacking. Lin *et al*^[49] found that most reported cases are males, but the impact of gender is still unclear. The disease-free time is around 7-45 mo according to previous case reports^[40-42,44]. Our patient had no peritoneal

recurrence until December, 2007 and is now regularly followed up at OPD.

REFERENCES

- Katyal S, Oliver JH 3rd, Peterson MS, Ferris JV, Carr BS, Baron RL. Extrahepatic metastases of hepatocellular carcinoma. *Radiology* 2000; **216**: 698-703
- Nakashima T, Okuda K, Kojiro M, Jimi A, Yamaguchi R, Sakamoto K, Ikari T. Pathology of hepatocellular carcinoma in Japan. 232 Consecutive cases autopsied in ten years. *Cancer* 1983; **51**: 863-877
- Ong GB, Taw JL. Spontaneous rupture of hepatocellular carcinoma. *Br Med J* 1972; **4**: 146-149
- Chen CY, Lin XZ, Shin JS, Lin CY, Leow TC, Chen CY, Chang TT. Spontaneous rupture of hepatocellular carcinoma. A review of 141 Taiwanese cases and comparison with nonrupture cases. *J Clin Gastroenterol* 1995; **21**: 238-242
- Goel AK, Sinha S, Kumar A, Chattopadhyay TK. Spontaneous hemoperitoneum due to rupture of hepatocellular carcinoma. *Trop Gastroenterol* 1993; **14**: 152-155
- Miyamoto M, Sudo T, Kuyama T. Spontaneous rupture of hepatocellular carcinoma: a review of 172 Japanese cases. *Am J Gastroenterol* 1991; **86**: 67-71
- Sonoda T, Kanematsu T, Takenaka K, Sugimachi K. Ruptured hepatocellular carcinoma evokes risk of implanted metastases. *J Surg Oncol* 1989; **41**: 183-186
- Zhu LX, Wang GS, Fan ST. Spontaneous rupture of hepatocellular carcinoma. *Br J Surg* 1996; **83**: 602-607
- Tanaka T, Yamanaka N, Oriyama T, Furukawa K, Okamoto E. Factors regulating tumor pressure in hepatocellular carcinoma and implications for tumor spread. *Hepatology* 1997; **26**: 283-287
- Chearanai O, Plengvanit U, Asavanich C, Damrongsak D, Sindhvananda K, Boonyapisit S. Spontaneous rupture of primary hepatoma: report of 63 cases with particular reference to the pathogenesis and rationale treatment by hepatic artery ligation. *Cancer* 1983; **51**: 1532-1536
- Zhu LX, Geng XP, Fan SD. [Mechanism of spontaneous rupture of hepatocellular carcinoma.] *Zhonghua Waike Zazhi* 2004; **42**: 1036-1039
- Zhu LX, Geng XP, Fan SD. [Ultrastructure study on patients with spontaneous rupture of hepatocellular carcinoma] *Zhonghua Waike Zazhi* 2006; **44**: 161-164
- Kanematsu M, Imaeda T, Yamawaki Y, Seki M, Goto H, Sone Y, Iinuma G, Mochizuki R, Doi H. Rupture of hepatocellular carcinoma: predictive value of CT findings. *AJR Am J Roentgenol* 1992; **158**: 1247-1250
- Choi BG, Park SH, Byun JY, Jung SE, Choi KH, Han JY. The findings of ruptured hepatocellular carcinoma on helical CT. *Br J Radiol* 2001; **74**: 142-146
- Chen TZ, Wu JC, Chan CY, Sheng WY, Yen FS, Chiang JH, Chau GY, Lui WY, Lee SD. Ruptured hepatocellular carcinoma: treatment strategy and prognostic factor analysis. *Zhonghua Yixue Zazhi (Taipei)* 1996; **57**: 322-328
- Xu HS, Yan JB. Conservative management of spontaneous ruptured hepatocellular carcinoma. *Am Surg* 1994; **60**: 629-633
- Leung KL, Lau WY, Lai PB, Yiu RY, Meng WC, Leow CK. Spontaneous rupture of hepatocellular carcinoma: conservative management and selective intervention. *Arch Surg* 1999; **134**: 1103-1107
- Yeh CN, Lee WC, Jeng LB, Chen MF, Yu MC. Spontaneous tumour rupture and prognosis in patients with hepatocellular carcinoma. *Br J Surg* 2002; **89**: 1125-1129
- Corr P, Chan M, Lau WY, Metreweli C. The role of hepatic arterial embolization in the management of ruptured hepatocellular carcinoma. *Clin Radiol* 1993; **48**: 163-165
- Pombo F, Arrojo L, Perez-Fontan J. Haemoperitoneum secondary to spontaneous rupture of hepatocellular carcinoma: CT diagnosis. *Clin Radiol* 1991; **43**: 321-322
- Ishihara M, Kobayashi H, Ichikawa T, Cho K, Gemma K, Kumazaki T. The value of emergency CT studies in spontaneous rupture of hepatocellular carcinoma. Analysis for tumor protrusion and hemorrhagic ascites. *Nippon Ika Daigaku Zasshi* 1997; **64**: 532-537
- Lai EC, Wu KM, Choi TK, Fan ST, Wong J. Spontaneous ruptured hepatocellular carcinoma. An appraisal of surgical treatment. *Ann Surg* 1989; **210**: 24-28
- Chen MF, Hwang TL, Jeng LB, Jan YY, Wang CS. Clinical experience with hepatic resection for ruptured hepatocellular carcinoma. *Hepatogastroenterology* 1995; **42**: 166-168
- Descottes B, Lachachi F, Valleix D, Durand-Fontanier S, Sodji M, Pech de Laclause B, Maissonnette F. [Ruptured hepatocarcinoma. Report of 22 cases] *Chirurgie* 1999; **124**: 618-625
- Chiappa A, Zbar A, Audisio RA, Paties C, Bertani E, Staudacher C. Emergency liver resection for ruptured hepatocellular carcinoma complicating cirrhosis. *Hepatogastroenterology* 1999; **46**: 1145-1150
- Vergara V, Muratore A, Bouzari H, Polastri R, Ferrero A, Galatola G, Capussotti L. Spontaneous rupture of hepatocellular carcinoma: surgical resection and long-term survival. *Eur J Surg Oncol* 2000; **26**: 770-772
- Ng KK, Lam CM, Poon RT, Law WL, Seto CL, Fan ST. Radiofrequency ablation as a salvage procedure for ruptured hepatocellular carcinoma. *Hepatogastroenterology* 2003; **50**: 1641-1643
- Sato Y, Fujiwara K, Furui S, Ogata I, Oka Y, Hayashi S, Ohta Y, Iio M, Oka H. Benefit of transcatheter arterial embolization for ruptured hepatocellular carcinoma complicating liver cirrhosis. *Gastroenterology* 1985; **89**: 157-159
- Le Neel JC, De Cervens T, Comy M, Dupas B, Letessier E, Mirallie E. [Ruptured hepatocarcinoma. Report of 20 cases and review of the literature] *Chirurgie* 1994; **120**: 380-384
- Ngan H, Tso WK, Lai CL, Fan ST. The role of hepatic arterial embolization in the treatment of spontaneous rupture of hepatocellular carcinoma. *Clin Radiol* 1998; **53**: 338-341
- Yang Y, Cheng H, Xu A, Chen D, Wang Y, Yao X, Chen H, Wu M. [Transarterial embolization for hemorrhage due to spontaneous rupture in hepatocellular carcinoma] *Zhonghua Zhongliu Zazhi* 2002; **24**: 285-287
- Fujii M, Miyake H, Takamura K, Tashiro S. [Management of spontaneous ruptured hepatocellular carcinoma] *Nippon Geka Gakkai Zasshi* 2004; **105**: 292-295
- Buczkowski AK, Kim PT, Ho SG, Schaeffer DF, Lee SI, Owen DA, Weiss AH, Chung SW, Scudamore CH. Multidisciplinary management of ruptured hepatocellular carcinoma. *J Gastrointest Surg* 2006; **10**: 379-386
- Ribeiro MA Jr, Fonseca AZ, Chaib E, D'Ippolito G, Carnevale FC, Rodrigues JJ, Saad WA. An unusual approach to the spontaneous rupture of hepatocellular carcinoma. *Hepatogastroenterology* 2007; **54**: 1235-1238
- Hirai K, Kawazoe Y, Yamashita K, Kumagai M, Nagata K, Kawaguchi S, Abe M, Tanikawa K. Transcatheter arterial embolization for spontaneous rupture of hepatocellular carcinoma. *Am J Gastroenterol* 1986; **81**: 275-279
- Chen MF, Jan YY, Lee TY. Transcatheter hepatic arterial embolization followed by hepatic resection for the spontaneous rupture of hepatocellular carcinoma. *Cancer* 1986; **58**: 332-335
- Shuto T, Hirohashi K, Kubo S, Tanaka H, Hamba H, Kubota D, Kinoshita H. Delayed hepatic resection for ruptured hepatocellular carcinoma. *Surgery* 1998; **124**: 33-37
- 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
- Liu CL, Fan ST, Lo CM, Tso WK, Poon RT, Lam CM, Wong J. Management of spontaneous rupture of hepatocellular

- carcinoma: single-center experience. *J Clin Oncol* 2001; **19**: 3725-3732
- 40 **Mizuno S**, Yamagiwa K, Ogawa T, Tabata M, Yokoi H, Isaji S, Uemoto S. Are the results of surgical treatment of hepatocellular carcinoma poor if the tumor has spontaneously ruptured? *Scand J Gastroenterol* 2004; **39**: 567-570
- 41 **Yunoki Y**, Takeuchi H, Makino Y, Murakami I, Yasui Y, Tanakaya K, Kawaguchi K, Konaga E. Intraperitoneal seeding of ruptured hepatocellular carcinoma: case report. *Abdom Imaging* 1999; **24**: 398-400
- 42 **Tan FL**, Tan YM, Chung AY, Cheow PC, Chow PK, Ooi LL. Factors affecting early mortality in spontaneous rupture of hepatocellular carcinoma. *ANZ J Surg* 2006; **76**: 448-452
- 43 **Al-Mashat FM**, Sibiany AM, Kashgari RH, Maimani AA, Al-Radi AO, Balawy IA, Ahmad JE. Spontaneous rupture of hepatocellular carcinoma. *Saudi Med J* 2002; **23**: 866-870
- 44 **Ong GB**, Chu EP, Yu FY, Lee TC. Spontaneous rupture of hepatocellular carcinoma. *Br J Surg* 1965; **52**: 123-129
- 45 **Shirabe K**, Kitamura M, Tsutsui S, Maeda T, Matsumata T, Sugimachi K. A long-term survivor of ruptured hepatocellular carcinoma after hepatic resection. *J Gastroenterol Hepatol* 1995; **10**: 351-354
- 46 **Kosaka A**, Hayakawa H, Kusagawa M, Takahashi H, Okamura K, Mizumoto R, Katsuta K. Successful surgical treatment for implanted intraperitoneal metastases of ruptured small hepatocellular carcinoma: report of a case. *Surg Today* 1999; **29**: 453-457
- 47 **Kaido T**, Arai S, Shiota M, Imamura M. Repeated resection for extrahepatic recurrences after hepatectomy for ruptured hepatocellular carcinoma. *J Hepatobiliary Pancreat Surg* 2004; **11**: 149-152
- 48 **Ryu JK**, Lee SB, Kim KH, Yoh KT. Surgical treatment in a patient with multiple implanted intraperitoneal metastases after resection of ruptured large hepatocellular carcinoma. *Hepatogastroenterology* 2004; **51**: 239-242
- 49 **Lin CC**, Chen CH, Tsang YM, Jan IS, Sheu JC. Diffuse intraperitoneal metastasis after spontaneous rupture of hepatocellular carcinoma. *J Formos Med Assoc* 2006; **105**: 577-582

S- Editor Li DL L- Editor Wang XL E- Editor Ma WH