

Immunohistochemical evaluation for outflow reconstruction using opened round ligament in living donor right posterior sector graft liver transplantation: A case report

Yukihiro Sanada, Yasunaru Sakuma, Hideki Sasanuma, Atsushi Miki, Takumi Katano, Yuta Hirata, Noriki Okada, Naoya Yamada, Yoshiyuki Ihara, Taizen Urahashi, Naohiro Sata, Yoshikazu Yasuda, Koichi Mizuta

Yukihiro Sanada, Takumi Katano, Yuta Hirata, Noriki Okada, Naoya Yamada, Yoshiyuki Ihara, Taizen Urahashi, Koichi Mizuta, Department of Transplant Surgery, Jichi Medical University, Shimotsuke City, Tochigi 329-0498, Japan

Yasunaru Sakuma, Hideki Sasanuma, Atsushi Miki, Naohiro Sata, Yoshikazu Yasuda, Department of Surgery, Jichi Medical University, Shimotsuke City, Tochigi 329-0498, Japan

Author contributions: Sanada Y contributed to study design, acquisition of data, analysis and interpretation of data, and drafting of the manuscript; Sasanuma H, Miki A, Katano T, Hirata Y, Okada N, Yamada N, Ihara Y and Urahashi T contributed to acquisition of data, and analysis and interpretation of data; Sakuma Y, Sata N, Yasuda Y and Mizuta K contributed to analysis and interpretation of data, and critical revision of the manuscript for important intellectual content; all authors read and approved the final manuscript.

Institutional review board statement: This study was reviewed and approved by the Ethics Committees of Jichi Medical University (15-106).

Informed consent statement: The patient involved in this study gave her informed consent authorizing use and disclosure of her protected health information.

Conflict-of-interest statement: We declare that we have no conflict-of-interest.

Open-Access: This article is an open-access article which was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: <http://creativecommons.org/licenses/by-nc/4.0/>

Manuscript source: Unsolicited manuscript

Correspondence to: Yukihiro Sanada, MD, PhD, Department of Transplant Surgery, Jichi Medical University, 3311-1 Yakushiji, Shimotsuke City, Tochigi 329-0498, Japan. yuki371@jichi.ac.jp
Telephone: +81-285-587069
Fax: +81-285-587069

Received: April 2, 2016
Peer-review started: April 7, 2016
First decision: May 30, 2016
Revised: June 11, 2016
Accepted: July 6, 2016
Article in press: July 6, 2016
Published online: September 14, 2016

Abstract

Utilizing the opened round ligament as venous grafts during liver transplantation is useful but controversial, and there are no pathological analyses of this procedure. Herein, we describe the first reported case of a pathological analysis of an opened round ligament used as a venous patch graft in a living donor liver transplantation (LDLT). A 13-year-old female patient with biliary atresia underwent LDLT using a posterior segment graft from her mother. The graft had two hepatic veins (HVs), which included the right HV (RHV; 15 mm) and the inferior RHV (IRHV; 20 mm). The graft RHV and IRHV were formed into a single orifice using the donor's opened round ligament (60 mm × 20 mm) as a patch graft during bench surgery; it was then anastomosed end-to-side with the recipient inferior vena cava. The recipient had no post-transplant complications involving the HVs, but she died of septic shock with persistent cholangitis and jaundice 86 d after LDLT. The HV anastomotic site had no stenosis or thrombus on autopsy. On pathology, there was adequate patency and continuity between the reci-

patient's HV and the donor's opened round ligament. In addition, the stains for CD31 and CD34 on the inner membrane of the opened round ligament were positive. Hepatic venous reconstruction using the opened round ligament as a venous patch graft is effective in LDLT, as observed on pathology.

Key words: Opened round ligament; Venous patch graft; Hepatic venous reconstruction; Living donor liver transplantation; All-in-one venoplasty

© **The Author(s) 2016.** Published by Baishideng Publishing Group Inc. All rights reserved.

Core tip: Utilizing the opened round ligament as venous grafts during liver transplantation is useful but controversial, and there are no pathological analyses of this procedure. Herein, we describe the first reported case of pathological analysis of an opened round ligament used as a venous patch graft in living donor liver transplantation. The hepatic venous (HV) anastomotic site had no stenosis or thrombus on autopsy. On pathology, there was adequate patency and continuity between the recipient's HV and the donor's opened round ligament. In addition, the stains for CD31 and CD34 on the inner membrane of the opened round ligament were positive.

Sanada Y, Sakuma Y, Sasanuma H, Miki A, Katano T, Hirata Y, Okada N, Yamada N, Ihara Y, Urahashi T, Sata N, Yasuda Y, Mizuta K. Immunohistochemical evaluation for outflow reconstruction using opened round ligament in living donor right posterior sector graft liver transplantation: A case report. *World J Gastroenterol* 2016; 22(34): 7851-7856 Available from: URL: <http://www.wjgnet.com/1007-9327/full/v22/i34/7851.htm> DOI: <http://dx.doi.org/10.3748/wjg.v22.i34.7851>

INTRODUCTION

The use of the opened round ligament as a venous patch graft has become accepted in hepatopancreatobiliary surgery because of its easy availability^[1-3]. Recent studies have demonstrated the use of the opened round ligament as a venous patch graft in hepatic venous reconstruction during living donor liver transplantation (LDLT)^[4-7]. However, there is no consensus regarding the efficacy of using the opened round ligament as a venous patch graft. Although there are reports on the radiological patency of the opened round ligament after LDLT and reports on the pathological assessment of its epithelium at the time of transplant^[8,9], pathological analyses of long-term patency and continuity after LDLT have not been reported.

Herein, we describe the first reported case of a pathological analysis on an autopsy specimen of an opened round ligament used as a venous patch graft in hepatic venous reconstruction during LDLT. Approval

to conduct this study was obtained from the Ethics Committees of Jichi Medical University.

CASE REPORT

A 13-year-old female patient with biliary atresia was considered for LDLT because of decompensated liver cirrhosis with jaundice. Her body height and weight were 159.4 cm and 54.0 kg, respectively, and her standard liver volume was 1095 mL. The blood test results were as follows: white blood cells 9600/ μ L; hemoglobin 8.7 g/dL; hematocrit 26.0%; platelets 149000 / μ L; albumin 2.9 g/dL; creatinine 0.35 mg/dL; total bilirubin 31.6 mg/dL; aspartate aminotransferase 93 mU/mL; alanine aminotransferase 50 mU/mL; prothrombin time-international normalized ratio 1.40; and activated partial thromboplastin 58.3 s. The model for end-stage liver disease score was 23.

The patient underwent an ABO-compatible LDLT using her mother's posterior segment graft (390 g; graft volume to standard liver volume ratio: 35.6%). The graft had two hepatic veins (HVs), including the right hepatic vein (RHV; 15 mm) and the inferior RHV (IRHV; 20 mm) (Figure 1A). The graft RHV and IRHV were formed into a single orifice using the donor's opened round ligament (60 mm \times 20 mm) as a venous patch graft during bench surgery (Figure 1B); it was then anastomosed end-to-side with the recipient inferior vena cava (Figure 1C and D). Portal vein reconstruction using an interposition vein graft from the right saphenous vein of the donor was performed between the recipient portal vein and the graft posterior portal vein. Hepatic artery reconstruction using microsurgical technique was performed between the recipient right hepatic artery and the graft posterior hepatic artery. For biliary reconstruction, a Roux-en-Y hepaticojejunostomy was performed. Intraoperative color Doppler ultrasonography was performed to assess the blood flow velocity and pattern after vascular reconstruction, and the flow velocity and pattern were satisfactory. The length of the operation was 21 h and 8 min, and the bleeding volume was 7438 mL. Tacrolimus and methylprednisolone were used for standard post-operative immunosuppressive therapy.

On post-operative day (POD) 8, portal vein thrombosis was detected by color Doppler ultrasonography, and the patient was given intravenous urokinase. On POD 26, acute cellular rejection was diagnosed by a graft liver biopsy, and she was treated with pulse steroids. On POD 40, cytomegalovirus viremia was detected by the C7-HRP test, and she underwent preemptive anti-cytomegalovirus therapy. On POD 48, portal vein balloon dilatation was performed by interventional radiology after the progressive narrowing of the portal vein anastomotic stricture to less than 2 mm was detected by color Doppler ultrasonography. During percutaneous transhepatic portography of the intrahepatic portal vein, bilhemia was detected, and

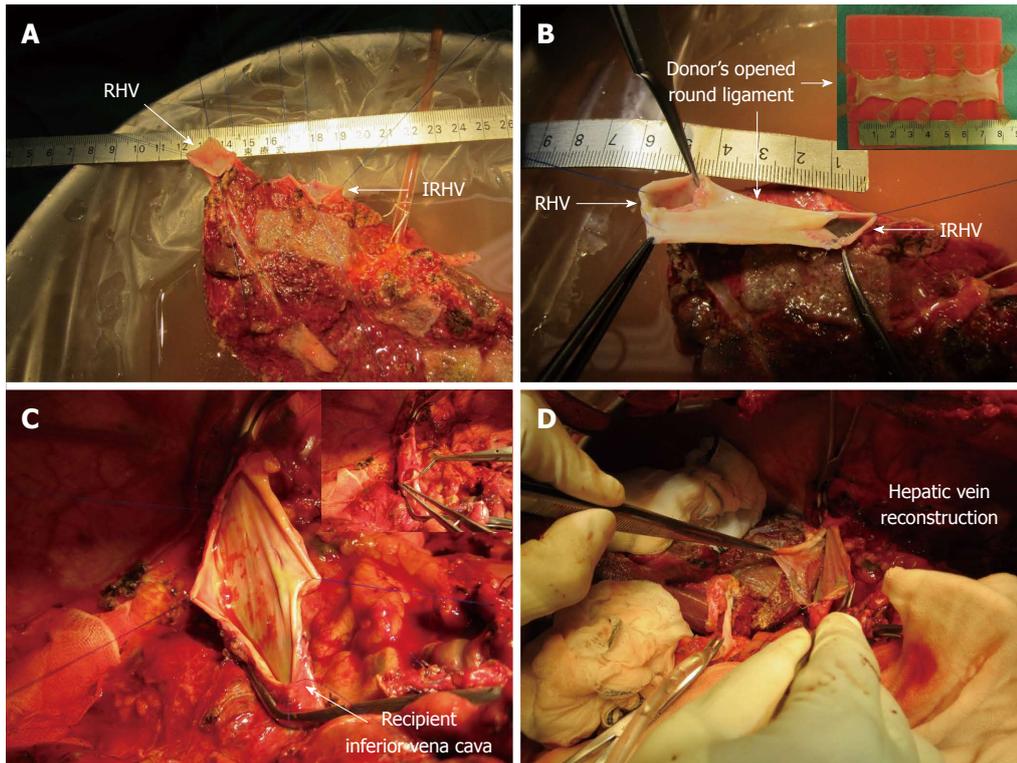


Figure 1 The graft had two hepatic veins. Including the right hepatic vein (RHV; 15 mm) and the inferior RHV (IRHV; 20 mm) (A). The graft RHV and IRHV were formed into a single orifice using the donor's opened round ligament (60 mm × 20 mm) as a venous patch graft during bench surgery (B). It was then anastomosed end-to-side with the recipient inferior vena cava (C and D).

thereafter, she suffered from persistent jaundice (total bilirubin 6.4 mg/dL). On POD 86, she developed a fever with liver dysfunction, and antibiotic treatment for acute cholangitis was administered. However, she died of septic shock with persistent cholangitis and bacteremia (with *Serratia marcescens*) on POD 88.

There were no post-transplant complications involving the HVs, and the radiological patency between the opened round ligament and the HV was confirmed on POD 82 (Figure 2). The HV anastomotic site had no stenosis or thrombus on an autopsy (Figure 3A and B). The patency and continuity between the donor's opened round ligament and the HV were adequate on pathological examination. In addition, the stains for CD31 and CD34 on the inner membrane of the opened round ligament were positive, a finding also observed in the graft HV (Figure 3C and D).

DISCUSSION

Outflow block in LDLT can potentially result in liver congestion, graft failure, and death^[10]. When a right liver graft is used for LDLT and the graft HV is anastomosed end-to-end with the recipient HV or end-to-side with the recipient inferior vena cava, the liver graft expands in all directions in the limited right subphrenic space during liver graft regeneration. Consequently, the HV anastomosis can be compressed and twisted in some circumstances^[11]. In addition,

multiple HV reconstructions are sometimes required in a right liver graft LDLT. Therefore, reports have indicated that it is important to simplify HV reconstruction and enlarge the graft HV orifice by an all-in-one sleeve patch graft venoplasty, which has shown excellent outcomes^[11-14].

The efficacy of various venous grafts used in HV reconstruction has been reported. These grafts include auto-venous grafts (from the inferior mesenteric vein, gonadal vein, external iliac vein, internal jugular vein, and saphenous vein)^[13,15,16], venous grafts from a living donor, native portal veins^[17], opened round ligaments^[4-9,12], cryopreserved homografts^[18-22], and polytetrafluoroethylene grafts^[23,24]. However, the harvest of a venous graft from a living donor or the recipient himself is not easy. In addition, the availability of cryopreserved homografts is limited, and the adverse effects of polytetrafluoroethylene grafts are unclear. Alternatively, the opened round ligament is easier to use, and less invasive than other venous grafts, but it is controversial in regard to its long-term patency. Recently, use of the opened round ligament as a venous patch graft in hepatic venous reconstruction during LDLT has been reported^[4-7]. Although there are reports on the radiological patency of the opened round ligament after LDLT and reports on the pathological assessment of its epithelium at the time of transplant^[8,9], there are no pathological analyses of patency and continuity. In this case, we

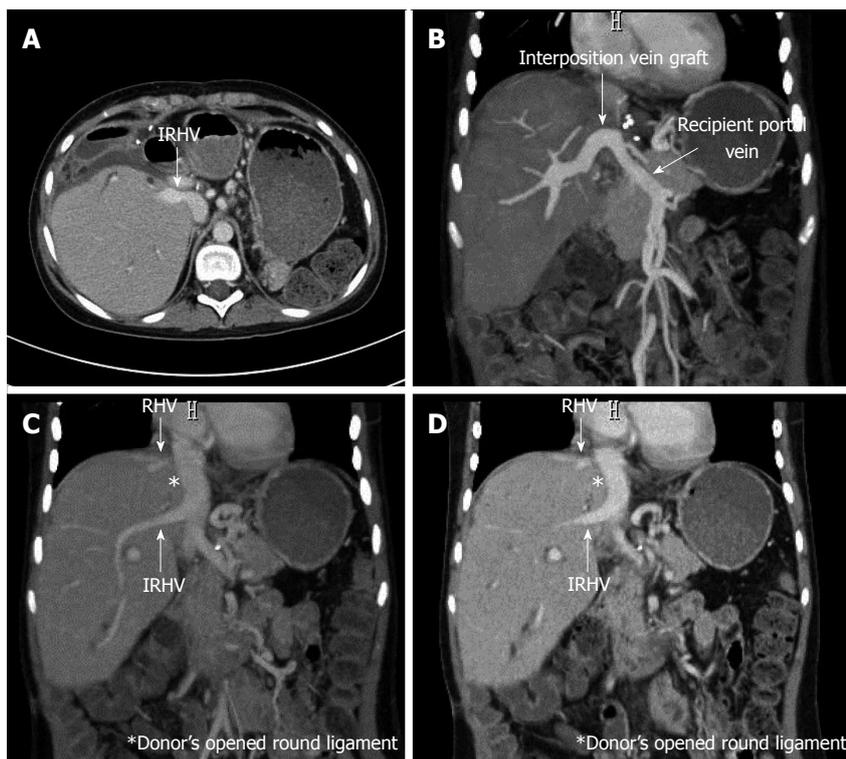


Figure 2 Abdominal enhanced computed tomography on post-operative day 82. The radiological patency of the donor's opened round ligament and the hepatic vein was confirmed (A, B, C, D). RHV: Right hepatic vein; IRHV: Inferior right hepatic vein.

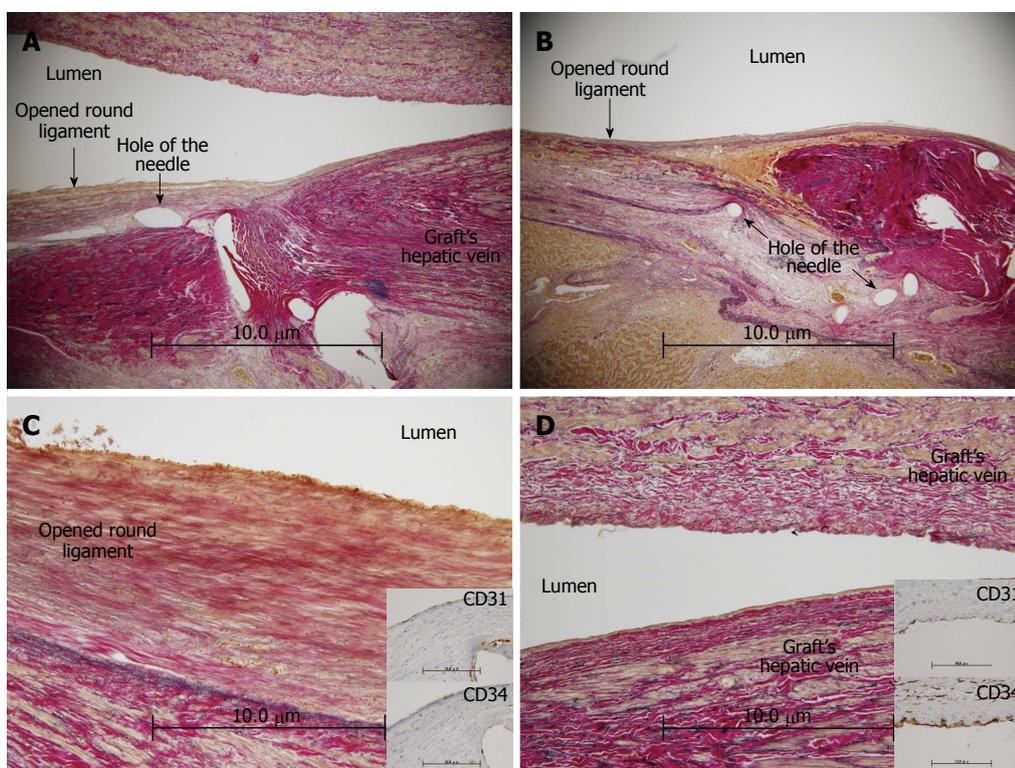


Figure 3 Pathology of the hepatic venous anastomotic site on autopsy. No stenosis or thrombus was identified (A and B). On pathology, there was adequate patency and continuity between the recipient's hepatic vein and the donor's opened round ligament. In addition, the stains for CD31 and CD34 on the inner membrane of the opened round ligament were positive, a finding also observed in the graft hepatic vein (C and D).

conducted a pathological autopsy evaluation of a donor's opened round ligament used as a venous patch graft in LDLT. This is the first report that demonstrates the pathological efficacy of the opened round ligament after surgery. The HV anastomotic site had no stenosis or thrombus (Figure 3A and B). On pathology, there was adequate patency and continuity between the recipient HV and the donor's opened round ligament. In addition, the stains for CD31 and CD34 on the inner membrane of the opened round ligament were positive, a finding also observed in the graft HV (Figure 3C and D). Therefore, we believe that the pathological patency and the existence of venous endothelial cells support the efficacy of using the opened round ligament as a venous patch graft.

In conclusion, using the opened round ligament as a venous patch graft is easy, less invasive, and had been shown to be pathologically effective. The accumulation of further cases and the long-term observation of this case are needed to confirm our findings.

ACKNOWLEDGMENTS

We thank Professor Akira Tanaka (Department of Pathology, Jichi Medical University) for helpful advices regarding pathological evaluations.

COMMENTS

Case characteristic

A 13-year-old female patient with biliary atresia underwent living donor liver transplantation using a posterior segment graft from her mother, and she died of septic shock with persistent cholangitis and jaundice 86 d after living donor liver transplantation.

Clinical diagnosis

The patient was diagnosed septic shock with persistent cholangitis and jaundice 86 d after living donor liver transplantation.

Differential diagnosis

No-obstructive afferent loop syndrome.

Laboratory diagnosis

Biliary stasis.

Imaging diagnosis

The radiological patency between the opened round ligament and the hepatic vein was confirmed on POD 82.

Pathological diagnosis

On pathology, there was adequate patency and continuity between the recipient's hepatic vein and the donor's opened round ligament. In addition, the stains for CD31 and CD34 on the inner membrane of the opened round ligament were positive, a finding also observed in the graft hepatic vein.

Treatment

The antibiotic treatment for acute cholangitis was administered.

Related reports

The efficacy of various venous grafts used in hepatic vein reconstruction

has been reported. These grafts include auto-venous grafts (from the inferior mesenteric vein, gonadal vein, external iliac vein, internal jugular vein, and saphenous vein), venous grafts from a living donor, native portal veins, opened round ligaments, cryopreserved homografts, and polytetrafluoroethylene grafts.

Term explanation

Opened round ligament and all-in-one venoplasty.

Experiences and lessons

Hepatic venous reconstruction using the opened round ligament as a venous patch graft is effective in living donor liver transplantation, as observed on pathology.

Peer-review

The manuscript provides anecdotal support for the use of opened round ligament in hepatic vein reconstruction.

REFERENCES

- 1 **Yamanaka N**, Yasui C, Yamanaka J, Tanaka T, Ando T, Kuroda N, Okamoto E. Recycled use of reopened umbilical vein for venous reconstruction in hepatopancreatobiliary surgery. *J Am Coll Surg* 2000; **190**: 497-501 [PMID: 10757391 DOI: 10.1016/S1072-7515(99)00288-4]
- 2 **Watanabe M**, Yamazaki K, Tsuchiya M, Otsuka Y, Tamura A, Shimokawa K, Kaneko H, Teramoto T. Use of an opened umbilical vein patch for the reconstruction of the injured biliary tract. *J Hepatobiliary Pancreat Surg* 2007; **14**: 270-275 [PMID: 17520202 DOI: 10.1007/s00534-006-1183-4]
- 3 **Gunasekaran G**, Mosna LC, Savino JA. Portomesenteric reconstruction using an umbilical vein patch during pancreaticoduodenectomy (Whipple procedure). *J Am Coll Surg* 2013; **217**: e9-e11 [PMID: 23870231 DOI: 10.1016/j.jamcollsurg.2013.04.034]
- 4 **Ikegami T**, Shimada M, Imura S, Morine Y, Kanemura H, Mori H, Arakawa Y, Hanaoka J. Beneficial use of the round ligament as a patch graft for vena cava reconstruction. *J Hepatobiliary Pancreat Surg* 2008; **15**: 581-584 [PMID: 18987926 DOI: 10.1007/s00534-007-1319-1]
- 5 **Sakamoto S**, Kasahara M, Shigeta T, Fukuda A, Tanaka H, Matsuno N. Feasibility of using the graft's umbilical vein as a patch graft for hepatic vein reconstruction in pediatric living donor liver transplantation. *Transpl Int* 2010; **23**: 436-437 [PMID: 19737372 DOI: 10.1111/j.1432-2277.2009.00953.x]
- 6 **Kamei H**, Onishi Y, Ogawa K, Uemoto S, Ogura Y. Living donor liver transplantation using a right liver graft with additional vein reconstructions for patient with situs inversus. *Am J Transplant* 2014; **14**: 1453-1458 [PMID: 24725262 DOI: 10.1111/ajt.12692]
- 7 **Toshima T**, Ikegami T, Matsumoto Y, Yoshiya S, Harimoto N, Yamashita Y, Yoshizumi T, Ikeda T, Shirabe K, Maehara Y. One-step venous reconstruction using the donor's round ligament in right-lobe living-donor liver transplantation. *Surg Today* 2015; **45**: 522-525 [PMID: 25231939 DOI: 10.1007/s00595-014-1004-3]
- 8 **Mergental H**, Gouw AS, Slooff MJ, de Jong KP. Venous outflow reconstruction with surgically reopened obliterated umbilical vein in domino liver transplantation. *Liver Transpl* 2007; **13**: 769-772 [PMID: 17457890 DOI: 10.1002/lt.21067]
- 9 **Ikegami T**, Wang H, Imai D, Bekki Y, Yoshizumi T, Yamashita Y, Toshima T, Soejima Y, Shirabe K, Maehara Y. Pathological analysis of opened round ligaments as venous patch grafts in living donor liver transplantation. *Liver Transpl* 2013; **19**: 1245-1251 [PMID: 23894128 DOI: 10.1002/lt.23716]
- 10 **Lee S**, Park K, Hwang S, Lee Y, Choi D, Kim K, Koh K, Han S, Choi K, Hwang K, Makuuchi M, Sugawara Y, Min P. Congestion of right liver graft in living donor liver transplantation. *Transplantation* 2001; **71**: 812-814 [PMID: 11330547]
- 11 **Lee SG**. Techniques of reconstruction of hepatic veins in living-donor liver transplantation, especially for right hepatic vein and major short hepatic veins of right-lobe graft. *J Hepatobiliary Pancreat Surg* 2006; **13**: 131-138 [PMID: 16547674 DOI: 10.1007/

s00534-005-1019-7]

- 12 **Chen CL**, Yap AQ, Concejero AM, Liu CY. All-in-one sleeve patch graft venoplasty for multiple hepatic vein reconstruction in living donor liver transplantation. *HPB (Oxford)* 2012; **14**: 274-278 [PMID: 22404267 DOI: 10.1111/j.1477-2574.2012.00442.x]
- 13 **Soejima Y**, Ueda N, Fukuhara T, Yoshizumi T, Ikegami T, Yamashita Y, Sugimachi K, Taketomi A, Maehara Y. One-step venous reconstruction for a right lobe graft with multiple venous orifices in living donor liver transplantation. *Liver Transpl* 2008; **14**: 706-708 [PMID: 18433050 DOI: 10.1002/lt.21401]
- 14 **Sugawara Y**, Makuuchi M, Sano K, Imamura H, Kaneko J, Ohkubo T, Matsui Y, Kokudo N. Vein reconstruction in modified right liver graft for living donor liver transplantation. *Ann Surg* 2003; **237**: 180-185 [PMID: 12560775 DOI: 10.1097/01.SLA.0000048444.40498.AD]
- 15 **Hwang S**, Lee SG, Park KM, Kim KH, Ahn CS, Moon DB, Ha TY. Quilt venoplasty using recipient saphenous vein graft for reconstruction of multiple short hepatic veins in right liver grafts. *Liver Transpl* 2005; **11**: 104-107 [PMID: 15690544 DOI: 10.1002/lt.20314]
- 16 **Ikegami T**, Shirabe K, Yoshiya S, Soejima Y, Yoshizumi T, Uchiyama H, Toshima T, Motomura T, Maehara Y. One-step reconstruction of the right inferior hepatic veins using auto-venous grafts in living-donor liver transplantation. *Surg Today* 2013; **43**: 769-776 [PMID: 23247889 DOI: 10.1007/s00595-012-0449-5]
- 17 **Mori A**, Kaido T, Ogura Y, Ogawa K, Hata K, Yagi S, Yoshizawa A, Isoda H, Shibata T, Uemoto S. Standard hepatic vein reconstruction with patch plasty using the native portal vein in adult living donor liver transplantation. *Liver Transpl* 2012; **18**: 602-607 [PMID: 22253117 DOI: 10.1002/lt.23387]
- 18 **Sugawara Y**, Makuuchi M, Akamatsu N, Kishi Y, Niiya T, Kaneko J, Imamura H, Kokudo N. Refinement of venous reconstruction using cryopreserved veins in right liver grafts. *Liver Transpl* 2004; **10**: 541-547 [PMID: 15048798 DOI: 10.1002/lt.20129]
- 19 **Hwang S**, Lee SG, Ahn CS, Park KM, Kim KH, Moon DB, Ha TY. Cryopreserved iliac artery is indispensable interposition graft material for middle hepatic vein reconstruction of right liver grafts. *Liver Transpl* 2005; **11**: 644-649 [PMID: 15915499 DOI: 10.1002/lt.20430]
- 20 **Kilic M**, Aydin U, Sozbilen M, Ozer I, Tamsel S, Demirpolat G, Atay Y, Alper M, Zeytinlu M. Comparison between allogenic and autologous vascular conduits in the drainage of anterior sector in right living donor liver transplantation. *Transpl Int* 2007; **20**: 697-701 [PMID: 17511830 DOI: 10.1111/j.1432-2277.2007.00499.x]
- 21 **Yaprak O**, Balci NC, Dayangac M, Demirbas T, Guler N, Ulusoy L, Tokat Y, Yuzyer Y. Cryopreserved aortic quilt plasty for one-step reconstruction of multiple hepatic venous drainage in right lobe living donor liver transplantation. *Transplant Proc* 2011; **43**: 2817-2819 [PMID: 21911171 DOI: 10.1016/j.transproceed.2011.06.064]
- 22 **Wang CC**, Lopez-Valdes S, Lin TL, Yap A, Yong CC, Li WF, Wang SH, Lin CC, Liu YW, Lin TS, Concejero AM, Eng HL, Henry D, Cheng YF, Jawan B, Chen CL. Outcomes of long storage times for cryopreserved vascular grafts in outflow reconstruction in living donor liver transplantation. *Liver Transpl* 2014; **20**: 173-181 [PMID: 24382821]
- 23 **Hwang S**, Jung DH, Ha TY, Ahn CS, Moon DB, Kim KH, Song GW, Park GC, Jung SW, Yoon SY, Namgoong JM, Park CS, Park YH, Park HW, Lee HJ, Lee SG. Usability of ringed polytetrafluoroethylene grafts for middle hepatic vein reconstruction during living donor liver transplantation. *Liver Transpl* 2012; **18**: 955-965 [PMID: 22511404]
- 24 **Thorat A**, Jeng LB, Yang HR, Li PC, Li ML, Yeh CC, Chen TH, Hsu SC, Poon KS. Outflow reconstruction for right liver allograft with multiple hepatic veins: "V-plasty" of hepatic veins to form a common outflow channel versus 2 or more hepatic vein-to-inferior vena cava anastomoses in limited retrohepatic space. *Liver Transpl* 2016; **22**: 192-200 [PMID: 26390259 DOI: 10.1002/lt.24342]

P- Reviewer: Boucek CD **S- Editor:** Yu J **L- Editor:** A
E- Editor: Ma S





Published by **Baishideng Publishing Group Inc**

8226 Regency Drive, Pleasanton, CA 94588, USA

Telephone: +1-925-223-8242

Fax: +1-925-223-8243

E-mail: bpgooffice@wjgnet.com

Help Desk: <http://www.wjgnet.com/esps/helpdesk.aspx>

<http://www.wjgnet.com>



ISSN 1007-9327



9 771007 932045