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Received: 2021.05.31 Accepted: 2021.09.06 Available online: 2021.10.04 Published: 2021.11.05			Single-Center Experience with Hepatic Artery Reconstruction During Living Donor Liver Transplantation: Microscope Versus Surgical Loupe		
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		kground: Aethods:	Hepatic artery reconstruction during living donor liver transplantation is a critical step. To perform this risky procedure, a microscope has been used. However, it takes a long time to complete the procedure and it has a long and steep learning curve. Recently, some transplant surgeons have performed the procedure using a surgical loupe. We conducted this study to compare the outcomes after hepatic artery reconstruction using a microscope versus using a surgical loupe. We retrospectively reviewed the outcomes of 300 patients at our institution from April 2014 to July 2020. From April 2014 to September 2017 (era 1), hepatic artery reconstruction was performed using a microscope by an experienced plastic surgeon. From September 2017 to the end date (era 2), it was performed using a loupe (x5.0) by an experienced transplantation surgeon. There was no difference in most perioperative outcomes between the 2 groups, including the major postoperative complications of hepatic artery complications (2/150 versus 2/150, <i>P</i> =0.343). There was a statistically significant difference between the 2 groups in anastomosis time (42.4±11.8 versus 24.2±7.8, <i>P</i> <0.001) and the entire operation time (436.6±83.9 versus 415.3±68.5, <i>P</i> =0.035). We suggest that when the surgeon is familiar with a loupe and vascular anastomosis, hepatic artery reconstruction using a surgical loupe is a safe and feasible method with a shorter operation time. <b>Anastomosis, Surgical • Hepatic Artery • Liver Transplantation • Microscopy</b>		
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# Background

For end-stage liver disease and unresectable hepatocellular carcinoma, liver transplantation (LT) has been accepted as the treatment of choice (HCC) [1]. Due to a scarcity of deceased donor grafts, the rate of living donor liver transplantation (LDLT) has grown in recent years [2]. However, due to the small size of the hepatic artery, arterial reconstruction in LDLT is technically more challenging than in deceased donor liver transplantation (DDLT). Hepatic artery complication is one of the most severe complications after LDLT [3] and is the most common vascular complication after LDLT, with estimated incidence rates ranging from 2.5% to 11% [4,5]. Several complications related to hepatic artery reconstruction, such as hepatic artery thrombosis (HAT), hepatic artery stenosis, pseudoaneurysm, and hepatic artery dissection, may lead to graft loss and mortality [6]. After transplantation, all collateral vessels are removed from the graft liver, and arterial blood flow is supplied only by the anastomosed hepatic artery. Therefore, hepatic artery reconstruction in LDLT is the most important procedure for the safety of the recipient.

One of the most significant advancements in hepatic artery reconstruction was the development of microsurgical techniques using a microscope, which has greatly reduced the incidence of HAT and significantly increased graft survival [7,8]. In the past, most centers used a surgical microscope during hepatic artery reconstruction due to the small diameter of the hepatic artery [8]. However, the use of a large surgical microscope has several disadvantages. Due to the deep operative field, the respiratory movement of the patient, and the size disparity between the donor and recipient hepatic artery, reconstructing the hepatic artery using a microscope takes a long time [9]. The hepatic artery reconstruction field is usually 10-15 cm deep from the abdominal wall, making it difficult to visualize the surgical field under a microscope. Furthermore, overcoming the learning curve of microvascular procedures and gaining necessary experience is time-consuming. For this reason, some surgeons suggested that magnification with a surgical loupe could produce comparable outcomes as using a microscope [10]. In the past, hepatic artery reconstruction was done under a microscope at our center, but for the reasons mentioned above, it is currently performed using a surgical loupe. In this retrospective study, we reviewed our experience with hepatic artery reconstruction in LDLT and compared the outcomes between using a microscope versus a surgical loupe.

# **Material and Methods**

This study included 304 patients who received LDLT at our center between April 2014 and July 2020. We retrospectively reviewed 300 patients after excluding 4 patients who underwent a second LT. We reviewed patient demographics and perioperative findings such as age, sex, disease etiology, laboratory data, operative outcomes, and postoperative complications. We divided the patients into 2 groups depending upon whether we performed hepatic artery reconstruction using a microscope or a surgical loupe. From April 2014 to September 2017 (era 1), hepatic artery reconstruction was performed with a microscope by a plastic surgeon. From September 2017 to the end date (era 2), it was performed with a surgical loupe (×5.0) by an experienced transplantation surgeon. The study design was approved by our center's Institutional Review Board.

During era 1, hepatic artery reconstruction was performed with interrupted sutures using 9-0 monofilament nylon (Ethilon®, ETHICON Inc., a Johnson & Johnson Company, Somerville, NJ, USA) under a microscope by an experienced plastic surgeon. The plastic surgeon, who was well-trained in microsurgery using a microscope, participated in the operations during era 1. During era 2, the procedure was performed with continuous sutures using 8-0 monofilament polypropylene (PROLENE® ETHICON, Inc.) under surgical loupe (×5.0) by an experienced transplantation surgeon. It was usually performed in a continuous fashion with a parachute technique (Figures 1, 2), but sometimes in interrupted ways if the size of the hepatic artery was too small or there was a considerable size difference between the hepatic artery of the recipient and graft liver. This transplantation surgeon had sufficient experience in liver transplantation, including major hepatic vessels such as the portal vein and hepatic veins. Since we started using a surgical loupe for hepatic artery reconstruction, all patients have undergone the operation by surgical loupe without exception. During the era 1 and era 2 study periods, a single plastic surgeon and single transplantation surgeon performed the hepatic artery reconstruction, respectively.

After reconstruction of the vascular structure, we routinely performed intraoperative Doppler ultrasound to evaluate the vascular patency. If the ultrasound did not show the hepatic artery flow or if it appeared abnormal, we checked whether there was an anatomical problem or arterial spasm. If an arterial spasm was the cause of the hepatic artery flow abnormality, we waited until it was resolved. If a surgical factor such as thrombosis, angulation, kinking, or tension around the anastomosis was the cause of the hepatic artery flow problem, we performed a revision of the hepatic artery reconstruction.

We performed Doppler ultrasound on post-transplantation days 1, 3, 5, and 13, and a dynamic angio-computed tomography (CT) scan was performed on post-transplantation days 7 and 20 to check vascular patency. If we found vascular abnormalities on Doppler ultrasound or an abrupt elevation of liver enzymes, we used CT angiography to check for vascular complications.



Figure 1. (A-D) Illustration of parachute technique used in hepatic artery reconstruction using a loupe. (Adobe Sketch, V 4.7, Sketch B.V.)



Figure 2. Photograph of hepatic artery reconstruction with parachute technique. (A) After posterior wall suture. (B) Completion of continuous suture.

The immunosuppression regimen included tacrolimus, mycophenolate mofetil, and steroids. We did not use everolimus as an immunosuppression agent during the post-transplantation period. After evaluating bleeding risk, prostaglandin E1 was administered intravenously postoperatively and maintained for 7 days. We did not use heparin or acetylsalicylic acid routinely as prophylaxis to prevent HAT.

#### **Statistical Analysis**

The mean and standard deviation are presented for all numerical data. To compare the continuous variables based on their distribution, the t test was performed. To compare the categorical variables the chi-squared test or Fisher's exact test was used after the assumptions were checked. All analyses were

Variable	Microscope (n=150)	Loupe (n=150)	p-value
Recipient age (year)	52.6±9.5	55.3±10.1	0.016
Recipient sex, Male (n, %)	98 (65.3%)	114 (76.0%)	0.042
BMI (kg/m²)	24.8±4.1	24.5±3.7	0.419
Original disease (n, %)			0.163
HBV	90 (60.0%)	78 (52.0%)	
HCV	11 (7.3%)	6 (4.0%)	
Alcohol	28 (18.7%)	41 (27.3%)	
Others	21 (14%)	25 (16.7%)	
Accompanying HCC (n, %)	75 (50%)	73 (48.7%)	0.817
Pre-transplantation TACE (n, %)	35 (23.3%)	33 (22.0%)	0.783
Pretransplant hepatectomy (n, %)	17 (11.3%)	12 (8.0%)	0.329
Major hepatectomy (n, %)	8 (5.3%)	5 (3.3%)	0.395
CTP score	8.3±2.8	8.1±2.8	0.937
MELD score	16.7±10.2	16.2±9.5	0.71
ABO-I LDLT (n, %)	23 (15.3%)	32 (21.3%)	0.179
Graft type: right lobe (n, %)	149 (99.3%)	136 (90.6%)	0.001
GRWR (%)	1.2±0.2	1.1±0.2	0.47

Table 1. Demographic characteristics and preoperative factors of the recipients and donors.

BMI – body mass index; HBV – hepatitis B virus; HCV – hepatitis C virus; HCC – hepatocellular carcinoma; TACE – transarterial chemoembolization; CTP score - Child-Turcotte-Pugh score; MELD score – model for end-stage liver disease score; LDLT – living donor liver transplantation; GRWR – graft-to-recipient weight ratio.

performed using IBM<sup>®</sup> SPSS<sup>®</sup> Statistics. A *P* value of less than 0.05 was taken to indicate statistical significance.

## **Results**

## Patient and Perioperative Characteristics

A total of 300 patients who underwent LDLT during the study period were enrolled. The difference between the 2 groups was not statistically significant, except for recipient age, recipient sex, and graft type (right lobe versus left lobe). The mean age of the recipients was 53.99±9.89 (median: 55, 7~77) years and the mean body mass index (BMI) of the recipients was 24.68±3.92. The most common cause of the original liver disease was viral hepatitis-related liver cirrhosis (n=185, 61.6%), followed by alcohol-related liver cirrhosis (n=69, 32.0%). A total of 148 patients were accompanied by HCC (49.3%). The mean Child-Turcotte-Pugh (CTP) and model for end-stage liver disease (MELD) scores were 8.26±2.871 and 16.54±9.886, respectively. A total of 55 patients (18.3%) underwent ABOincompatible LDLT. There was no statistically significant difference between the 2 groups in pre-transplantation transcatheter arterial chemoembolization (TACE) history or pre-transplantation

hepatectomy history, which is considered to be a factor that can induce HAT (**Table 1**).

## **Operative Variables and Surgical Outcomes**

Compared to the microscope group, the loupe group showed advantages in anastomosis time ( $42.47\pm11.83$  versus  $24.27\pm7.83$ , mean $\pm$ SD, *P*<0.001) and the entire operation time ( $436.66\pm83.91$  versus  $415.35\pm68.55$ , mean $\pm$ SD, *P*=0.035). However, there was no statistically significant difference in ascites, recipient liver weight, estimated blood loss, packed red blood cell (PRC) transfusions, graft fatty change, or hepatic artery size (**Table 2**).

There was no statistically significant difference in postoperative complications such as postoperative bleeding, hepatic artery complications, biliary strictures, and biliary leaks. There was a total of 15 re-operations, all of which were due to postoperative bleeding. The length of hospital stay, length of ICU stay, and in-hospital mortality showed no statistically significant differences (**Table 3**).

Variable	Microscope (n=150)	Loupe (n=150)	p-value
Operation time (min)	436.6±83.9	415.3±68.5	0.035
Anastomosis time (min)	42.4±11.8	24.2±7.8	<0.001
Ascites (cc)	1623.1±2794.1	1230.8±2429.5	0.196
Recipient liver weight (g)	1053.9±360.7	1145.6±616.6	0.117
Estimated blood loss (cc)	3770.6±3952.6	4014.3±3784.1	0.587
PRC transfusion (pack)	8.6±8.7	8.2±7.7	0.728
Graft fatty change (%)	3.7±6.2	4.4±6.1	0.829
Hepatic artery size (mm)	1.9±0.8	2.1±0.3	0.25
Complications			
Postoperative bleeding (n, %)	10 (6.7%)	5 (3.3%)	0.185
Hepatic artery complication (n, %)	2 (1.3%)	2 (1.3%)	1.0
Biliary stricture (n, %)	4 (2.7%)	1 (0.7%)	0.176
Biliary leak (n, %)	18 (12.0%)	13 (8.7%)	0.343
Re-operation (n, %)	10 (6.7%)	5 (3.3%)	0.185
Length of ICU stay	7.12±4.293	6.87±3.110	0.567
Length of hospital stay (days)	25.70±12.469	27.02±17.939	0.46
In-hospital mortality (n, %)	8 (5.3%)	8 (5.3%)	1.0

#### Table 2. Comparison of intraoperative findings and postoperative outcomes.

PRC - packed red cells, ICU - Intensive Care Unit.

#### Table 3. Comparison of postoperative outcomes.

Variable	Microscope (n=150)	Loupe (n=150)	p-value
Complications			
Postoperative bleeding (n, %)	10 (6.7%)	5 (3.3%)	0.185
Hepatic artery complication (n, %)	2 (1.3%)	2 (1.3%)	1.0
Biliary stricture (n, %)	4 (2.7%)	1 (0.7%)	0.176
Biliary leak (n, %)	18 (12.0%)	13 (8.7%)	0.343
Re-operation (n, %)	10 (6.7%)	5 (3.3%)	0.185
Length of ICU stay	7.12 <u>+</u> 4.293	6.87±3.110	0.567
Length of hospital stay (days)	25.70±12.469	27.02 <u>+</u> 17.939	0.46
In-hospital mortality (n, %)	8 (5.3%)	8 (5.3%)	1.0

ICU - Intensive Care Unit.

## Discussion

Although LT is the most effective treatment for end-stage liver disease and HCC, it is still considered a high-risk operation with significant morbidity and mortality. LDLT has been established as an alternative to overcome the graft shortage in Asia, where the supply of deceased donors remains below 5 per 1 million population per year [2,11]. However, the surgical procedures for arterial reconstruction in LDLT are more complicated than in DDLT and can cause hepatic arterial complications such as HAT. The introduction of surgical microscope in the 1990s was one of the most significant improvements in hepatic artery reconstruction [8]. Microsurgical techniques overcame the risk of HAT, with an incidence reduced to 1.7% [12].

Although the use of microsurgical techniques in LDLT minimized the problems associated with a high incidence of HAT, it has several disadvantages. First of all, the surgical microscope is bulky

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and needs to be protected with a sterile plastic bag, which requires additional work. It also requires tricky maneuvering and has difficulties focusing in the abdominal cavity [13]. It offers an operative view at a particular prefixed angle, but time is required to change this setting. Compared to microsurgery on the head, neck, and extremities, it is difficult to make precise observations due to the constant movement of the liver graft in accordance with ventilation. Therefore, ventilation needs to be withheld sometimes during hepatic artery reconstruction using a surgical microscope. For these reasons, some surgeons prefer suturing the artery under a surgical loupe, with more convenient conditions and comparable reported surgical outcomes [14,15].

Postoperative results, including postoperative complications, were comparable between the 2 groups at our center, and the results using a surgical loupe were superior to those of a microscope in terms of operation time and anastomosis time. Other factors such as previous treatment history like TACE or hepatectomy or the size of the hepatic artery can also affect the postoperative outcomes including HAT, but there was no statistically significant difference between the 2 groups. The difference in the incidence by graft type was mainly due to the chronological nature of the 2 groups. As the experience of our center increased, the use of the left lobe became more frequent. The difference in the ratio between the 2 groups may have caused selection bias, but there are several reports that suggest similar postoperative outcomes including HAT between LDLT using the left and right lobes [16-18].

Although hepatic artery reconstruction in a continuous fashion is less time-consuming, there is a concern that it can cause hepatic artery stenosis. However, if there is no hepatic artery dissection and the size of the hepatic artery is not too small (<2 mm), the continuous fashion showed results similar to those of the interrupted fashion [19], consistent with the results in our center. Since we start using a loupe for hepatic artery reconstruction, we usually do reconstruction using continuous fashion. However, when the size of hepatic artery from the graft liver was small, especially when the size discrepancy between the graft and recipient hepatic artery is considerable, we choose an interrupted fashion. Most of the complications related to the hepatic artery are due to surgical factors such as kinking or the wrong axis, but these complications are thought to be irrelevant to whether the method is continuous or interrupted.

In this study, the postoperative outcomes between the 2 groups were not significantly different. However, there were 2 cases of postoperative complications related to the hepatic artery in each group. In our center, re-operation was done when HAT occurred within 1 week after surgery, and interventions were performed when HAT occurred after 1 week. The first case was hepatic artery occlusion, which was detected on postoperative day (POD) 14 by a routine follow-up CT. We inserted a stent via radiologic intervention and the hepatic artery was successfully recanalized. The second case was hepatic artery bleeding with a pseudoaneurysm on POD 9. We tried embolization with intervention, but it failed. We performed a re-operation, and the bleeding was successfully controlled. The third case was HAT on POD 13. We performed an intervention, but it failed. The patient was in poor condition and eventually died. The fourth case was HAT, which occurred on POD 3 but the condition of the graft artery was so poor that it was not possible to perform a re-operation or intervention. Fortunately, despite the poor condition of the hepatic artery, the patient's condition and LFT were relatively stable, and the hepatic artery was partially recanalized after 6 months using acetylsalicylic acid.

Because hepatic artery complications can cause serious problems in LDLT, hepatic artery reconstruction is one of the most important procedures. Since research reported that hepatic artery anastomosis using a microscope dramatically lowers hepatic artery-related complications [20], the microscope method has been regarded as a standard technique in most LDLT centers. There have been several studies comparing use of the microscope and loupe for microvascular anastomosis, but it is also argued that the difference in the manner of anastomosis whether microscope or loupe is used is insignificant, since both can use the same method of magnification [21]. However, most centers still prefer the microscope method over the loupe method, despite the disadvantage of the microscope method, which requires additional training [22]. In our study, the loupe method had comparable postoperative results to the microscope method and had shorter anastomosis time and operation time. This may be another advantage of the loupe method, as previous studies reported that a prolonged operation time can increase postoperative complications [23,24]. In addition, reduced operating time can minimize surgeon fatigue and enhance the surgeon's focus on the operation.

This study had some limitations. First, this was a single-center retrospective study, so our results may not be applicable to other centers and some significant biases may have affected the selection of the controls. Due to the retrospective design, data on some operative factors such as the status of the hepatic artery intima or the number of hepatic artery revisions were not collected properly. However, in this study, the microscope method and the loupe method were each performed by a single surgeon who was familiar with the technique, so it seems that the surgeon-related factors were controlled.

# Conclusions

This study shows that there was no significant difference in most perioperative outcomes between the microscope group

and the loupe group. Therefore, we suggest that when the surgeon is familiar with a loupe and vascular anastomosis, hepatic artery reconstruction using a surgical loupe will be a safe and feasible method with a shorter operation time.

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#### **Declaration of Figures' Authenticity**

All figures submitted have been created by the authors, who confirm that the images are original with no duplication and have not been previously published in whole or in part.

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