

What happened in 133 consecutive hepatic artery reconstruction in liver transplantation in 1 year?

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Background: The immediate challenges during microvascular reconstruction of hepatic artery (HAR) during liver transplantation (LT) can be many. Hence, in order to give a cross sectional view of these problems this study over a period of 1 year, showing our routine practice, was taken up.

Methods: From January 2015 to December 2015, a total of 133 LTs were performed in Kaohsiung Chang Gung Memorial Hospital, Taiwan. All hepatic artery (HA) reconstructions were performed by a microvascular surgeon under an operating microscope.

Results: In the 133 patients, one artery was anastomosed in 123 (92.5%) patients, two in 9 (6.8%) patients and three in 1 (0.7%) of the patient. Eleven (8.3%) arteries were less than 2 mm in size (1–1.9 mm). There were intimal dissections (IDs) involving either the donor or the recipient arteries of mild to severe nature in 9 (6.8%) patients. Immediately following graft arterial anastomosis, either there was no flow or an intraoperative hepatic artery thrombosis (HAT) was found in nine (7.1%—8 LDLT, 4.8%—1 DDLT) patients. Immediate re-do anastomosis was done in all of these patients who did well in the follow-up. The overall post-operative success rate was 99.2%. One patient (0.8%) developed postoperative HAT due to infection during follow up and died due to sepsis.

Conclusions: Small vessels or HA injury are the frequently encountered problems by a micro vascular surgeon. The other problems could be ID, need to do multiple reconstructions, immediate HAT and ability to re-do the HAR immediately.

Keywords: Hepatic artery (HA); thrombosis; liver transplantation (LT); microvascular surgery; reconstruction

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Introduction

One of the most important steps during liver transplantation (LT) is the hepatic artery reconstruction (HAR). The surgical reconstruction technique is the prime risk factor for hepatic artery thrombosis (HAT). The use of microsurgical technique has helped to overcome the risk of HAT in many series (1-4). The higher magnification $(10-15\times)$ of the

microscope helps in surveying accurately and identifying any intimal flaps or injury to the hepatic artery before the reconstruction. When the size of the vessel is less than 2 mm, reconstruction using loupes or a smaller magnification (<6x) can lead to problems and is risky. In addition, there may be a need to use alternate vessel for reconstruction when the recipient hepatic artery may not

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Table 1 Demography of Recipients and diagnostic indications for liver transplantation (n=133)

Variables	n (%)
Male	83 (62.4)
Female	50 (37.6)
Age, years (range)	52.7 (0.4–68.9)
<2 years	19 (14.3)
2–18 years	10 (7.5)
18–60 years	76 (57.1)
>60–68.9 years	28 (21.1)
Biliary atresia	30 (22.6)
HBV HCC	21 (15.8)
HBV cirrhosis	18 (13.5)
HCV cirrhosis	21 (15.8)
HCV HCC	12 (9.0)
Alcoholic cirrhosis	13 (9.8)
Alcoholic cirrhosis with HCC	7 (5.3)
HBV HCV HCC	2 (1.5)
HCC	2 (1.5)
Alagille syndrome	1 (0.8)
Fulminant hepatic failure	1 (0.8)
Cryptogenic cirrhosis	1 (0.8)
HBV acute liver failure	1 (0.8)
Primary biliary cirrhosis with HCV	1 (0.8)
Primary biliary cirrhosis	1 (0.8)
Wilson's disease	1 (0.8)
MELD/PELD–LDLT-mean (range)	7.1 (2.0–37.0)
MELD/PELD–DDLT-mean (range)	21.4 (7.0–32.0)

HBV, hepatitis B virus; HCC, hepatocellular carcinoma; HCV, hepatitis C virus; MELD, Model for End-Stage Liver Disease; PELD, pediatric end-stage liver disease; LDLT, living donor liver transplantation; DDLT, deceased donor liver transplantation.

be available for reconstruction. Hence, to highlight the technical problems that were encountered and how they were tackled, we decided to look at our experience in the microvascular reconstruction of the hepatic artery in patients undergoing LT, over a period of 1 year, which gives a cross-sectional view of our routine practice in LT program at our center.

Methods

From January 2015 to December 2015, a total of 133 LTs were performed in Kaohsiung Chang Gung Memorial Hospital, Taiwan. There were 113 living donor liver transplantation (LDLT) and 20 deceased donor liver transplantation (DDLT). Seventeen livers from the deceased donors were harvested at CGMH, Kaohsiung itself. The other 3 livers were harvested from the other hospitals. Twenty-nine cases were pediatric and 104 cases were adults. The male to female ratio was 83/50. The median age of the group was 52.7 years (range, 0.4–68.9 years). The indication for LT were biliary atresia (22.6%), hepatitis B virus (HBV) hepatocellular carcinoma (HCC) (15.8%), HBV cirrhosis (13.5%), hepatitis C virus (HCV) cirrhosis (15.8%), HCV HCC (9.0%), alcoholic cirrhosis (9.8%), alcoholic cirrhosis with HCC (5.3%) and other causes (8.3%). The median

Surgical technique

undergoing DDLT (Table 1).

All the hepatic artery reconstructions were completed using the posterior wall first, and a combined method for the anterior wall (5). The hepatic artery reconstructions were initiated following the completion of hepatic and portal vein reconstruction, once the graft was reperfused. All hepatic artery reconstructions were performed by a microvascular surgeon under an operating microscope with $6-15 \times$ magnification.

Model for End-Stage Liver Disease (MELD) score was 17 in patients undergoing LDLT and was 21 in those

The technique of reconstructing the arteries was described fully in our previous report (5). Briefly, the microvascular reconstructions were performed using an 8-0 prolene or a 9-0 non-absorbable nylon monofilament suture (Ethilon, Ethicon Inc., Somerville, NJ, USA) on a 9-0 gauge Micropoint needle. The vessels' posterior walls were reconstructed first using an interrupted suturing technique whereas the anterior wall was reconstructed using a continuous suturing and interrupted tying technique or a combined method. After that, the blood flow from the hepatic artery was immediately evaluated by a radiologist using a color Doppler ultrasound.

Postoperative care and follow-up

Cyclosporine-based immunosuppression was used for pediatric recipients, whereas, tacrolimus-based

Table 2 Details of the hepatic arterial size used in an astomoses in LDLT and DDLT $\ensuremath{\mathsf{DDLT}}$

Diameter of the hepatic arteries	Graft vessel, n (%)	Recipient vessel, n (%)
LDLT		
1–1.9 mm	11 (8.9)	10 (8.1)
2–3 mm	104 (84.6)	102 (82.9)
3.1–4 mm	8 (6.5)	11 (8.9)
DDLT		
2–3 mm	7 (33.3)	10 (47.6)
3.1–4 mm	14 (66.7)	11 (52.4)

LDLT, living donor liver transplantation; DDLT, deceased donor liver transplantation.

immunosuppression was employed for adults. Anticoagulation was administered immediately after the surgery. Similarly, after the transplantation, a Doppler ultrasound examination was performed to evaluate the blood flow in all the vessels daily during the first 2 weeks, every other day on the third week, and twice a week after that until discharge. A computed tomographic angiography (CTA) is performed to confirm any obstruction in the hepatic arteries if the Doppler examinations showed a poor or absence of flow in these vessels. The patient was immediately scheduled for surgery once an occlusion of hepatic artery was confirmed following the CTA, if done within 2 weeks after transplantation.

Statistics

Statistical analysis was performed using SPSS version 18.0 for windows. All the categorical data related to the patient's baseline characteristics were presented as frequencies and percentages. All normally distributed continuous data was presented as mean with standard deviation and range.

Results

In the 113 patients who underwent LDLT, right lobe of the liver was used in 63 (55.8%) patients, whereas left lobe was used in 39 (34.5%) and left lateral lobe was used in 11 (9.7%) patients. One artery was anastomosed in 104 (92.0%) patients, two in 8 (7.1%) patients and three in 1 (0.9%) of the patient. From these liver grafts, right hepatic artery was used for anastomosis in 63 (55.8%) patients, left hepatic artery was used for anastomosis in 41 (36.3%) patients, left hepatic artery with segment 4 artery was used in 8 (7.1%) patients and left hepatic artery, right gastro-epiploic artery (RGEA) and cystic artery were used in 1 (0.9%) patient. Most of these arteries (84.6%) were 2–3 mm in size, 7 (6.5%) were between 3.1-4 mm in size and 11 (8.9%) were less than 2 mm in size (1–1.9 mm) (*Tables 2,3*).

The anastomosis patterns of the vessels between the liver graft and the recipient vessel, including the two or three vessel anastomoses are shown in Table 3. The commonest pattern was the right hepatic artery of the graft was anastomosed to either the left hepatic artery of the recipient in 27 (23.9%) patients or the right hepatic artery of the recipient in 24 (21.2%) patients. The second common pattern was the left hepatic artery was anastomosed to either the recipient's right hepatic artery (16.8%) or the left hepatic artery (10.6%). Two cases (1.8%) needed radial artery graft (RAG) interposition graft and three cases (2.7%) needed reconstruction with RGEA. In the recipient, 8.1% (6) arteries used were smaller than 2 mm (1–1.9 mm). Majority (82.9%) were 2-3 mm in size (Table 2). The color Doppler evaluation done by a radiologist, immediately after the anastomosis, showed a mean Vmax 47±15 cm/s and a mean resistance index (RI) 0.65±0.14. The average time taken for anastomoses was 20-25 minutes.

In the 20 patients who underwent DDLT, whole liver was used as a graft in 9 (45.0%) patients, split right lobe in 5 (25.0%) patients and split left lobe was used in another 5 (25.0%) patients. Reduced liver graft was used in 1 (5.0%) patient. One artery was used for anastomosis in 19 (95.0%) patients and, two arteries were used in 1 (5.0%) patient. From these liver grafts, the proper hepatic artery was the most common used for anastomosis in 6 (30.0%) patients, followed by the common hepatic artery in 5 (25.0%) and LHA in 4 (20.0%) patients. Both the common hepatic artery and the splenic artery were used in 1 (5.0%) patients. The pattern of the arterial anastomoses is shown in Table 3. Most of the donor arteries were more than 3 mm (66.7%) in size. Most of the recipient arteries were also more than 3 mm (52.4%) (Table 2). The RGEA had to be used in one patient (5.0%). The colour Doppler evaluation done by a radiologist, immediately after the anastomosis, showed a mean Vmax 57.3±29 cm/s, mean RI 0.59±0.09. The average time taken for anastomoses was 15-20 minutes.

There were intimal dissections (IDs) involving either the donor (7) or the recipient (8) arteries of mild to severe nature in 9 (6.8%) patients. In 12 of them (9%), there was injury to the artery without ID. Immediately following graft arterial anastomosis, either there was poor flow (Vmax <0.20 m/s)

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Table 3 Type of arterial anastomoses in patients of LDLT (n=113) and DDLT (n=20)

Liver transplantation	Anastomosed arteries	n (%)
LDLT		
Single arterial anastomosis	RHA-LHA	27 (23.9)
	RHA-RHA	24 (21.2)
	LHA-RHA	19 (16.8)
	LHA-LHA	12 (10.6)
	RHA-A4	6 (5.3)
	LHA-LGA	3 (2.7)
	RHA-GEA	3 (2.7)
	LHA-A4	4 (3.5)
	LHA-PHA	2 (1.8)
	LHA-RAG-PHA	1 (0.9)
	RHA-LGA	1 (0.9)
	RHA-RAG-LHA	1 (0.9)
Two arterial anastomosis	LHA-RHA, A4-LHA	2 (1.8)
	LHA-LHA, A4-A4	2 (1.8)
	LHA-LGA, A4-RHA	1 (0.9)
	LHA-LHA, A4-RHA	1 (0.9)
	LHA-A4, A4-LHA	1 (0.9)
	LHA-RHA, A4-A4	1 (0.9)
Three arterial anastomosis	LHA-RGA, RGA-LHA cystic-A4	.,1 (0.9)
DDLT		
One arterial anastomosis	PHA-RHA	4 (20.0)
	CHA-RHA	3 (15.0)
	LHA-RHA	3 (15.0)
	PHA-LHA	2 (10.0)
	CHA-PHA	2 (10.0)
	RHA-A4 LHA-PHA	1 (5.0) 1 (5.0)
	RHA-RHA	1 (5.0)
	SA-RHA	1 (5.0)
	GDA-GEA	1 (5.0)
Two arterial anastomosis	SA-SA, CHA-CHA	1 (5.0)

RHA, right hepatic artery; LHA, left hepatic artery; A4, artery to segment 4; LGA, left gastric artery; RGA, right gastric artery; GEA, gastro epiploic artery; RAG, radial artery graft; RPHA, right posterior hepatic artery; PHA, proper hepatic artery; CHA, common hepatic artery; SA, splenic artery; GDA, gastroduodenal artery; LDLT, living donor liver transplantation; DDLT, deceased donor liver transplantation. or an intra-operative HAT was found in 9 (7.1%—8 LDLT, 4.8%—1 DDLT) patients. Immediate re-do anastomosis was done in all of these patients (*Table 4*).

Three patients (2.7%) of the LDLT group developed bile leak in the immediate postoperative period needing re-exploration and repair. Two (1.5%) patients needed an endoscopic retrograde biliary drainage (ERBD) and stenting for the biliary stricture. All these patients were followed up after surgery. At the time of writing this manuscript, the minimum duration of follow-up was 28 months and the maximum duration of follow-up was up to 36 months. There were three deaths (2.3%). One patient who underwent DDLT died immediately following the surgery because of diffuse intracerebral bleeding. One of the patients had an equivocal finding of HAT 19 days after LDLT, but was showing normal liver functions, hence was treated conservatively. Visualization of hepatic artery was noted 2 months after LDLT. He later developed hepatic abscess which was drained but ultimately died of sepsis at the end of 3 months. Since he had equivocal finding of the hepatic artery during follow-up and since the radiologist felt that visualized hepatic artery could be due collaterals, this could be a case of post-operative HAT treated conservatively. The third patient (LDLT group) developed in intrahepatic haematoma and herpes zoster. He also had an episode of gastrointestinal bleeding and finally succumbed to sepsis at the end of 10 weeks.

Discussion

This study was mainly taken up to look at the challenges during microvascular reconstruction of the hepatic artery in LT and also to see how we could overcome the same technically. It has been shown by various other studies in the literature that HAT following LT can be reduced from 25% without a microscope to less than 4% with a microscope (7-9). However, the microvascular surgeons may have to face some problems during the reconstruction. As it can be seen in *Figure 1*, we had problems like small arteries, need to do multiple reconstructions, hepatic arterial injury, ID, immediate HAT, need of re-doing the anastomosis, need to use RGEA and interposition RAG grafts.

Liver grafts vessel or recipient vessel less than 2 mm diameter is considered to be a risk factor for HAT (10). Narrow diameter of the hepatic arteries complicates the reconstructive procedures both in adult and pediatric LDLTs (10). In our study, we had 21 (18.6%) instances,

Table 4 Dei	tails of re-do	arterial anast	tomoses	, anastomoses using RGEA	and RAG		
Case No.	Age/sex	Primary disease	Graft type	Final anastomosis graft vessel-recipient vessel	Complication and management	Vmax (cm/s)/RI	Result
Re-do							
1174	43/M	HCV	LDLT	RHA-RHA	No flow, intima injury over the recipient HA-trim back	32/0.76	Doing well
1179	57/M	HBV	LDLT	RHA-LHA	HAT after anastomosis & re-do	50/0.47	Doing well
1199	55/M	НВV, НСV	LDLT	RHA-LHA	acute HAT due to donor HA injury, trim back	46/0.5	Doing well
1206	63/F	HCV	LDLT	RHA-A4	Intima injury over the donor artery, trim back	46/0.68	Doing well
1228	54/F	НВV, НСС	LDLT	RHA-RHA	HAT after anastomosis due to mild id, trim back	23/0.68	Doing well
1260	44/F	HCV	LDLT	LHA-LHA, A4-A4	donor LHA injury, trim back, HAT after HA anastomosis over LHA and A4, re-do	70/0.72	Doing well
1349	52/F	НВV, НСС	DDLT	CHA-RHA	HAT due to recipient HA injury over branch, re-do	64/0.68	Doing well
Using RGE/	٩						
1183	60/M	НСС	LDLT	RHA-GEA	HA blocked due to previous TAE, hence use of GEA	54/0.60	Died, 3 months later due to sepsis
1237	64/M	НВV, НСС	LDLT	RHA-GEA	initial use RHA, HAT due HA injury and recipient severe ID- immediate re-do using GEA	31/0.55	Doing well
1254	38/M	НВV, НСС	LDLT	RHA-GEA	donor mild id and thick wall, recipient HA severe ID-using GEA, HAT-twice, immediate re-do-thrice	29/0.33	Doing well
19		BA	DDLT	GDA-GEA	Re-transplantation, no suitable recipient hepatic arteries, hence GEA	92/0.49	Doing well
Using RAG							
1219	0.4/M	BA	LDLT	LHA-RAG-PHA	Donor LHA ligated incidentally; repair first; recipient RHA segmental narrow-excision; repair using RAG	51/0.62	Doing well
1220	43/M	HBV	LDLT	RHA-RAG-LHA	donor HA injury, recipients HA mild id, trim back, inadequate length-hence RAG interposition graft	32/0.68	Doing well
RAG, radial hepatic arte deceased d	artery graft sry; HAT, he lonor liver tr	; RGEA, righ patic artery th ansplantation	t gastro hrombo n; RI, re	epiploic artery; HBV, hepasis; CHA, common hepati sis; CHA, common hepati sistance index.	tititis B virus; HCC, hepatocellular carcinoma; HCV, hepatitis C c artery; BA, biliary atresia; GDA, gastro-duodenal artery; LDL	virus; RHA, right he 7, living donor liver t	patic artery; LHA, left ransplantation; DDLT,

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Figure 1 Bar chart showing the number of problematic cases. HAR, hepatic artery reconstruction; GEA, gastro epiploic artery.

in the LDLT group, of either the donor vessels or the recipient vessels were less than 2 mm in size. In the past, liver graft with a vessel less than 2 mm was considered as a contraindication for transplantation (2,6,7). However, there was no statistical significance as comparison of result between the group with diameter less than 2 mm and the group with the diameter more than 2 mm. There was vessel size discrepancy in some of these cases which was overcome by cutting the smaller vessel obliquely and resorting to end to side anastomosis in one of the instances where a three vessel anastomosis was necessary. We also reconstructed all of the arteries with the pattern of obliquely cutting to prevent kinking or twist. Other methods like fish-moth method or funnelization method were not used in the present study.

In our previous study, we had noted that the incidence of multiple graft arteries even in right lobe graft could be 4.7% and was much higher in left lobe grafts (11). However, selective reconstruction of a single hepatic artery is sufficient, even in the presence of multiple arteries (12-14). Multiple reconstructions are usually associated with increase in the operation time, the duration of hepatic arterial reconstruction and blood loss. All these factors may increase the occurrence of a biliary stricture (15,16). Hence, many of the transplant surgeons avoid using a graft with multiple small vessels. But, still, we had to do multiple reconstructions in 9 (6.8%) cases, because both arteries were all significant in terms of angiosome in the graft.

The cause of hepatic arterial injury or ID could be because of excessive pulling during the dissection which may cause severe vasospasm and may also cause shedding of the intima. Another major cause is transarterial embolization (TAE) or chemoembolization to treat HCC patients. These procedures or the agents used in these procedures can damage the intima. The ID could be higher in patients undergoing TAE for HCC and has been classified as mild, moderate and severe (5). In mild and moderate ID, the HA can be used after trimming back but if ID is severe an alternative vessel needs to be used. Injury or ID can predispose to HAT. We had 9 (6.8%) cases of ID and 13 (9.8%) cases of hepatic artery injury in this study which had to be managed.

One of the unique features of this study was noticing the HAT intraoperatively, immediately after the anastomosis using the color Doppler and correcting the same immediately by re-doing the anastomosis. There were 7 (5.3%) such instances including the one in the DDLT. This immediate HAT was either due to vessel wall injury or the hepatic arterial ID. There were no instances of late (post-transplantation) re-do anastomosis in this study. As a result, all these patients had a good postoperative outcome. A late (>24 hours) re-do anastomosis could be associated with less desirable outcome (17). In this study we had to do immediate re-do anastomosis not only for HAT but also when we found poor flow (Vmax <0.20 m/s) or intimal injury following the anastomosis which would have led to HAT. The total number of re-do was in 9 (6.8%) cases. We also performed Doppler examination in all the vessels daily during the first 2 weeks, every other day on the third week, and twice a week after that until discharge. These could detect compromised circulation and re-do reconstruction earlier to prevent re-transplantation.

Gastric vessels can be used for hepatic arterial revascularization with good results (18). RGEA and left gastric artery (LGA) can be used in patients with ID of the recipient artery or those who develop HAT during thrombectomy and revascularization (18). The gastroepiploic artery has been used in coronary artery bypass graft surgery by the cardiac surgeons (19). The splenic artery has also been used in hepatic arterial revascularization but with the slight disadvantage of splenic infarction (20). In this study, we had to use RGEA in 4 (3%) cases. In two of the cases it was used as a re-do procedure to overcome HAT. In the other two cases it was used primarily as a substitute since recipient hepatic artery was blocked in one case due to previous TAE for HCC and in the other case the hepatic artery was not available due retransplantation in a case of biliary atresia.

RGEA is our first choice as an alternative conduit for hepatic arterial reconstruction. However, if ID involves common hepatic artery or celiac axis, they are not suitable for the use. If RGEA is not available we choose the other alternative like RAG as interposition graft. When RGEA and LGA are anomalous or extremely short, RAG can be used as it offers a considerable length, appropriate diameter and excellent long-term patency (21). Ileocolic artery along with 17-cm RAG has been used as a secondary conduit for HA alternative in an adult LDLT (22). We had to use RAG in two cases (1.8%) of LDLT. In the first case, recipient vessel showed a narrow segment which had to be excised which resulted in shortening and hence RAG had to be used. In the second case, the recipient vessel showed ID and had to be trimmed which resulted in inadequate length and hence RAG had to be used. Radial artery is more similar as hepatic artery in term of diameter and resistance to compression. The venous graft is more vulnerable to be compressed by tissue. The radial artery was the first choice if interposition graft was needed.

The incidence of post-operative HAT in our study was 0.8% (one case of LDLT) which is one of the lowest. This patient had a doubtful diagnosis of HAT 19 days post LDLT and hence was managed conservatively and eventually had good flow from collaterals 2 months later.

There are arguments favoring arterial reconstruction using either a microscope or a loupe but not showing a statistical significance with respect to HAT (6,23-27); but there are no randomized trials. Our view is that loupes can give only a fixed working distance, fixed magnification, smaller field of view, limited Illumination, can cause weight burden on head and nose and can result in a spinal compression in a surgeon. On the other hand, microscope can give a changeable working distance, changeable magnification, large field of view, motorized focus and zoom, xenon Illumination, image/video recording and ergonomic device adjustment which are all advantageous if the technique is mastered. The higher magnification of the microscope helps in surveying accurately and identifying any intimal flaps or injury to the hepatic artery accurately before the reconstruction itself. When the size of the vessel is less than 2 mm, reconstruction using loupes or a smaller magnification (<6×) can lead to problems and is risky. In contrast, there was no contraindication using vessels less than 2 mm under microscope. We strongly recommend the use of microscope for all the hepatic artery reconstruction.

However, the causes for HAT can range from a simple technical factor to more complicated donor and recipient size, graft volume, graft vascular resistance, excessive portal pressure or flow etc. And now recently hypercoagulable states identified by thromboelastography have been added to the list of parameters (28,29). This study which gives a cross-sectional view represents probably the technical peak of the microvascular surgeon when the HAT could be at its lowest. Similar reports with no HAT have been reported (30,31).

To conclude, even with the technical advances in microsurgical reconstruction of hepatic artery, still small vessels or hepatic artery injury are the frequently encountered problems by a micro vascular surgeon. The other problems could be ID, need to do multiple reconstructions, immediate HAT and ability to re-do the same immediately. The micro vascular surgeon should have the capacity to use alternate conduits like RGEA or interposition grafts like RAG. With experience it is possible to bring down the HAT to the lowest possible level.

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Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

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