Technical Considerations in Liver Transplantation

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Types

- OLTX: Orthotopic liver Tx; placed in the anatomically correct position
- ALTX: Auxiliary liver Tx; placement of donor liver in the presence of native liver (part or all).
 - Orthotopic: in correct position after partial removal of native organ.
 - Heterotopic: in other place.
- SLTX: Segmental liver Tx; placement of portion of donor-liver.
- Cadaveric (whole or split) & Living-donor
- Donor with Cardiac-Death (DCD)

Technique

- Hepatectomy is the most difficult part of the procedure (bleeding, adhesions, reperfusion coagulopathy, risk of bowel violation); veno-venous bypass can help with bleeding by decompressing portal pressure.
- After [hepatectomy + retrohepatic IVC removal], a cadaveric [liver graft + donor IVC] is placed with a subdiaphragmatic end-to-end IVC interposition.
- Portal vein anastomosis: end-to-end
- Hepatic artery anastomosis: end to end
- Biliary reconstruction: duct-to-duct or hepaticojejunostomy.

Hepatectomy

Technique Hepatectomy

- Dissection of hilium is most important part.
- Preserve as much length of hepatic artery
 & portal vein as possible.
- Recipient's Hepatic artery dissection:
 - starts at Rt & Lt branches, then
 - runs to the confluence,
 - then gastro-duodenal art,
 - finally to common hepatic artery;
 - avoid traction & intimal dissection

Technique Hepatectomy

- Dissection of cystic duct & CBD:
 - with preservation of surrounding tissue to prevent ischemia/necrosis.
- Portal vein dissection:
 - is done after Hepatic artery and bile duct division;
 - all soft tissue around is removed from liver hilium until pancreas head.

Technique Hepatectomy Precautions

- Avoid injury to Rt adrenal gland:
 - may cause massive bleed and need adrenalectomy.
- Avoid injury to Rt renal vein during IVC dissection.
- Avoid injury to Rt phrenic nerve:
 - may cause paralysis of Rt hemidiaphragm.

Anhepatic Phase

Veno-veno bypass:

- to decrease hemodynamic instability after clamp of IVC & PV,
- bypass is done before hepatectomy, from IVC & PV to SVC,
- heparin bonded cannulae and a motor.

Advantages of v-v bypass:

- 1. Avoids cardiovascular instability, specially in FHF & noncirrhotics (no porto-systemic collateral circulation),
- 2. Decompresses portal v. pressure, decreasing blood loss,
- 3. Avoids mesenteric stasis/bowel edema, which may cause bacterial translocation,
- 4. Protects renal function (no Renal vein stasis),
- 5. Facilitates a safer & longer anhepatic phase, for proper dissection/hemostasis.

- Potential Indications for Veno-Venous bypass:
 - Fulminant hepatic failure.
 - Severe retroperitoneal collateralization.
 - Poor pre-operative renal function.
 - Hypotension after "test-clamp" of IVC, despite adequate volume loading.
 - Intestinal or mesenteric edema.
 - Inexperience with the procedure.

- Disadvantages of v-v bypass:
 - Complications in 10-30%
 - Lymphocoele in inguinal/axillary incision
 - Seroma in cannulae insertion site
 - Hematoma
 - Wound infection
 - DVT
 - Nerve injury
 - Air embolus with death (cannulae removal)
- Possible alternative: Piggyback technique.

Implantation

Technique Implantation

- 1. IVC end-to-end anastomosis:
 - donor's IVC is interposed into the severed recipient's IVC (vs piggy-back into recipient's supra-hepatic veins "common tunnel")
- 2. Portal vein end-to-end anastomosis:
 - donor to recipient.
 - In PV thrombosis (0.6-64%), can do:
 - a) eversion thrombectomy,
 - b) interposition graft to SMV,
 - c) cavo-portal hemitransposition,
 - d) anastomosis to large venous collateral,
 - e) arterialization of Portal vein

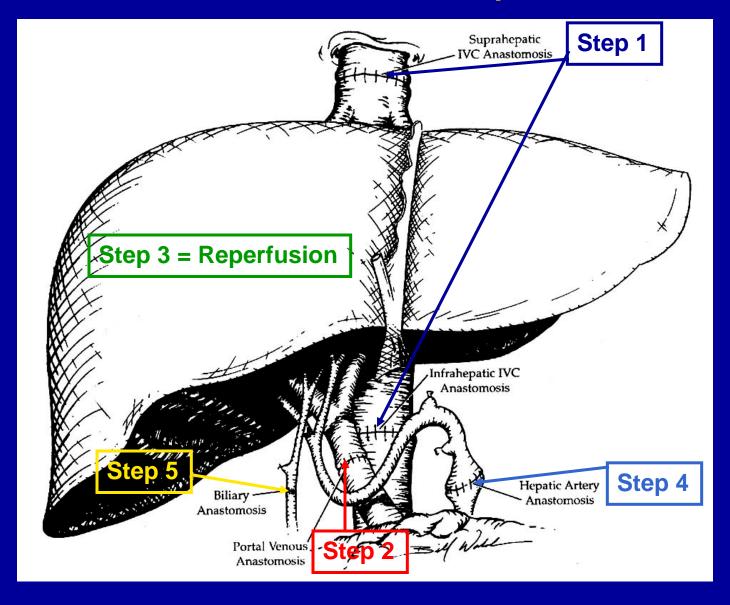
Technique Implantation

- 3. Reperfusion of liver with recipient's blood;
 - unstable time with risk of arrhythmia, hypotension, pulmonary edema, hyperkalemia;
 - liver is pre-perfused with Lactate Ringer or albumin to decrease risk.
- 4. End-to-end hepatic artery anastomosis.
 - In celiac axis stenosis: use a "donor iliac artery conduit graft" from infrarenal or supra-celiac aorta, to the donor's HA.
 - Do not use artificial grafts (high risk for thrombosis or infection).

Technique Implantation

- 5. Biliary anastomosis ("Achilles tendon" of LTX):
 - a) Choledochocholedochostomy: frequently over T-tube,
 - b) Roux-en-Y Choledochojejunostomy

Standard Technique



- Piggy-back technique can help to avoid venovenous bypass.
 - Dissection of recipient's caudate & Rt lobe from retrohepatic IVC to expose Rt, middle & Lt suprahepatic veins.
 - Clamp at IVC junction, then cut of all 3 veins leaving as much afferent hepatic veins as possible.
 - All 3 afferent veins are interconnected to make a single hepatic vein tunnel cuff which opens into the recipient's IVC.
 - Tunnel cuff will later be connected to the donor's efferent IVC (afferent donor's IVC will be ligated).

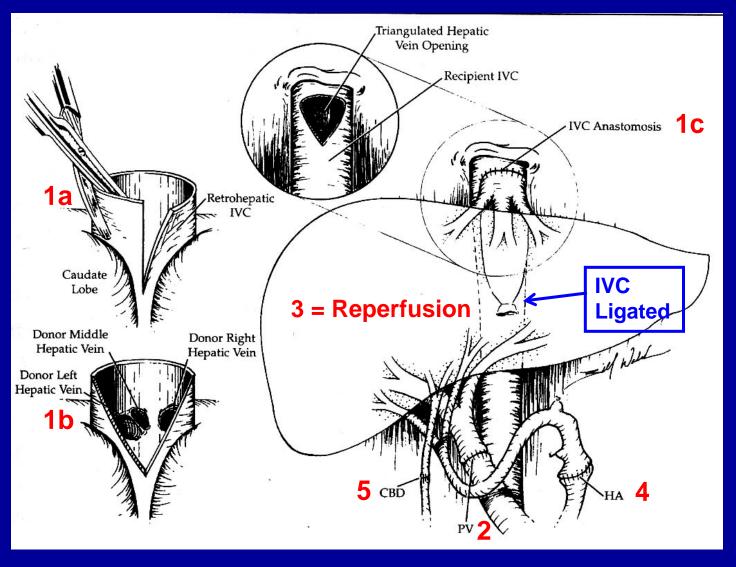
Advantages of "Piggy-back" technique:

- No need for lower IVC anastomosis
- Less risk of adrenal gland injury
- Less risk of renal v. injury
- Less bleeding
- Shorter anhepatic phase
- Less hemodynamic instability; v-v bypass needed only if portal vein clamp causes instability
- In FHF & non-cirrhotics, temporary porto-caval shunt may be needed to decompress the bowel (no porto-systemic collaterals)

Disadvantages of "Piggy-back" technique:

Compression of IVC causing thrombosis (Budd-Chiari like)

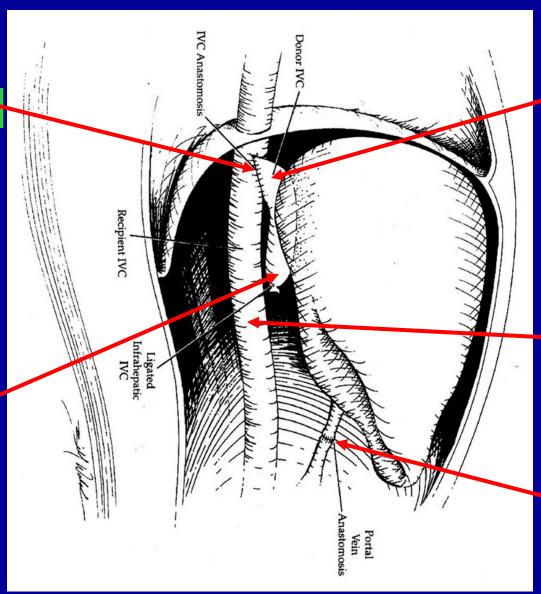
Piggyback Technique #2



Piggyback Technique

IVC Anastomosis

Ligated Donor Infrahepatic IVC



Suprahepatic Donor IVC

Recipient IVC

Portal Vein Anastomosis

Indications for Choledochojejunostomy

- Donor-recipient duct size discrepancy
- Diseased recipient bile duct
 - Primary sclerosing cholangitis
 - Secondary biliary cirrhosis
 - Choledocholithiasis
- Biliary duct malignancy
- Poor blood supply to recipient's bile duct.
- Inability to pass bile-probe through ampulla.

Post-Procedure

Intraoperative Signs of Hepatic Function

- Restoration of hemodynamic stability
- Adequate urine output
- Acid-base stabilization
- Normalization of coagulation
- Normalization of body temperature
- Maintenance of glycemia
- Adequate bile production
- Good texture & color of liver.

Early Post-operative Signs of Hepatic Function in ICU

- Hemodynamic stability
- Awakening from anesthesia
- Clearance of lactate
- Resolution of hypoglycemia
- Normalization of coagulation/ Factor V
- Resolution of elevated aminotransferases
- Good quantity of golden-brown bile.

Complications

Primary Nonfunction:

- Occurs in 4-10% LTX.
- Features: hepatocyte necrosis, without vascular complication
 - hepatic encephalopathy,
 - coagulopathy,
 - minimal bile output,
 - renal & multisystem failure,
 - persistent hypothermia,
 - hemodynamic instability,
 - high lactate & liver enzymes.
- Those with hemodynamic instability or multiorgan failure need urgent retransplantation.

Primary Nonfunction:

- Donor risk factors:
 - prolonged cold ischemia,
 - unstable donor,
 - high steatosis,
 - older age,
 - hypernatremia,
 - non-heartbeating donor

Criteria for 1A Status Primary Non-Function

- Non function in first 7 days, and:
- AST > 3000 IU and at least one:
 - INR > /= 2.5
 - Arterial pH </= 7.3 or Venous pH </= 7.25, and/or Lactate >/= 4 mMol/L
- Anhepathic Phase

 No AST requirement for recipient of segmental grafts of LDLT

Small-for-Size Syndrome

- Partial liver graft unable to meet functional demands of recipient: poor early graft function in absence of ischemia.
- Prevention: in cirrhotic "Graft-Weight to Body-Weight Ratio" (GWBWR) must be =/> 0.85%
- Manifestations:
 - Poor bile production
 - Prolonged cholestasis
 - Significant ascites
 - Coagulopathy

- Small-for-Size Syndrome
- Biochemical profile:
 - Elevated Direct (& total) bili
 - Mild/moderate elevation of ALT & AST
 - Prolonged PT
- Histologic Features:
 - Cholestasis with "bile plugs"
 - Areas of regeneration & ischemia with patchy necrosis.
- Prognosis: 50% of recipients will die of sepsis within 4-6 weeks.

- Small-for-Size Syndrome
- Recipient Factors Predictive of poor-outcome/ SFSS
 - Graft total mass
 - Poor metabolic & physical recipient condition
 - Advanced chronic liver disease & severe portal hypertension
 - Impaired venous inflow and/or outflow.

Hepatic Artery Stenosis

- Caliber decrease of > 50%;
- Occurs in 5%.
 - Asymptomatic or elevated liver enzymes.
 - U/S doppler: resistive index < 0.5 with increase in focal peak velocity.
- Therapy:
 - early: surgical repair;
 - weeks later: angioplasty (90% success)

Hepatic Artery Thrombosis

- Pathogenesis: intimal dissection due to vigorous manipulation or clamping.
 - In children may be asymptomatic (rare in adults).
 - Manifestations:
 - acute, massive necrosis,
 - intrahepatic duct necrosis with central biloma,
 - multiple strictures, or
 - intermittent bacteremia.

Hepatic Artery Thrombosis

- HAT may be segmental;
 - Lt HAT is usually benign.
 - Rt HAT causes biliary strictures.
- DX: angiography.
- Treatment:
 - urgent revascularization;
 - may need retransplantation (biliary sepsis, intraabdominal infection)

Criteria for 1A Status HAT with Primary Non-Function

- Non function in first 7 days, and:
- AST > 3000 IU and at least one:
 - INR > /= 2.5
 - Arterial pH </= 7.3 or Venous pH </= 7.25, and/or Lactate >/= 4 mMol/L
- Anhepathic Phase
- No AST requirement for recipient of segmental grafts of LDLT
- List with MELD of 40 if HAT does not cause PNF or if between 7-14 days post-op.

Portal Vein Stenosis

- Manifestations: post-LTX ascites or elevated enzymes.
 - U/S doppler or CT angiography are diagnostic.
 - Confirmed by SMA angiography (late films).
- Treatment: Without therapy may evolve to thrombosis.
 - a) early: surgical,
 - b) late presentation: angioplasty or stenting.

Vascular Complications

Portal Vein Thrombosis

- Rare.
- May cause graft dysfunction or massive ascites.
- Due to:
 - kinked or redundant vein, or
 - anastomotic stricture/twist, or
 - poor mesentery flow due to "steal syndrome" from venous collaterals.
- Treatment: revascularization, +/- ligation of large collaterals.

Vascular Complications

Hepatic Outflow Obstruction

- More common after "piggy-back" technique.
- 2.5-6% have iatrogenic Budd-Chiari.
- High morbidity and mortality.
- Due to "rotation" of liver graft or to anastomotic stricture.
- Suprahepatic cava stenosis can cause:
 - liver dysfunction,
 - ascites, or
 - impaired renal function.
- Side-to-side cavo-cavoplasty may be protective.
- **DX**: cavagram with measurement of pressure gradient
- **Treatment**: angioplasty, stent, or surgical repair

- Incidence: 15-20%.
- Mortality: 10%.
- Presentation: nonspecific; elevation of bili, alk. phosph & GGT.

Biliary stenosis:

- Is the most common complication;
- Due to imperfect anastomosis or ischemia.
- May cause abnormal enzymes or recurrent cholangitis.
- Treatment:
 - Dilatation + stent (ERCP or PTC).
 - May need creation of, or revision of choledochojejunostomy.

- Ampullary dysfunction:
- Due to cut of hepatic branch of vagus?
- Causes Donor+Recipient Bile Duct dilation.
- Treatment:
 - sphyncterotomy or conversion to choledochojejunostomy.

Multiple Intrahepatic Strictures:

- May be due to:
 - HA thrombosis/stenosis (more frequent in non-heartbeating donors),
 - Preservation damage,
 - Immune injury (lymphocytotoxic (+) cross-match),
 - Recurrent PSC,
 - CMV/ cryptosporidium/ microspora infection.
 - Treatment:
 - antibiotic prophylaxis;
 - may need retransplantation.

Bile leak:

- In c-c anastomosis:
 - usually at T-tube site & self-contained.
- In c-j anastomosis:
 - may be lethal due to infection that makes repair difficult (friable & edematous)
 - high risk of mycotic rupture of hepatic artery anastomosis.

Roux-en-Y loop complications:

- a) Bleeding at jejunojejunostomy:
 - self limited in half; need surgical hemostasis in other half.
- b) Internal mesentery hernia at j-j anastomosis;
 - Causes unexplained abdominal distention and vomiting, due to small bowel volvulus.
 - May be lethal due to bowel necrosis.
 - DX: CT scan showing "closed loop obstruction".
 - Treatment: surgical closure of mesentery defect.

Live-Donor Adult Liver Tx

- 5% of transplants in USA. (learning curve = 20 cases)
- Only 30-45% of potential donors donate;
- Aborted hepatectomy in 5% of donors
- Donor relation:
 - 30% offspring,20% sibling,
 - 20% parent,20% unrelated,
 - 10% other relative/unknown.
- Donor age: 50% > 50 years old.
- Donor risk: (14 death, 1 vegetative state, 2 LT/ 6-7000 live-donors)
 - 0.4% mortality,
 - 0.4-0.6% catastrophic complication, &
 - 35% morbidity
- Patient survival: equal to cadaver-donor.

Live-Donor Adult Liver Tx Disease-Specific Considerations

• HCC:

Must fulfill Milan Criteria

HCV:

 Is acceptable indication, but appropriate timing needs further investigation (not too early).

FHF:

Acceptable indication for emergency transplantation.

- Complete history & physical with "ideal & actual body weight".
- No smoking.
- Stop BCPs before surgery.
- Laboratory:
 - CBC, CMP,
 - serologic testing,
 comprehensive coagulation profile,
 - markers of liver disease,
 other tests as indicated by Hx & PE.
- Psychosocial evaluation.

- Radiology:
 - liver volume & vascular anatomy;
 - biliary anatomy pre-op or intra-op.
- Pre-op liver Bx is controversial. Do it if:
 - abnormal enzymes, or
 - steatosis by imaging, or
 - -BMI > 30,
 - donor genetically related to patient with AIH,
 PSC, or PBC

- Donor age-limit of 60 is considered appropriate.
- BMI > 30 may increase risk to donor, but is not absolute contraindication.
- Volumetric imaging analysis may overestimate liver volume by 10%.
- Calculated donor-remnant should be at least 30% of original liver volume & with complete venous drainage.
- Graft-liver-volume to recipient-body-weight ratio (GWBWR) should be =/> 0.8%
- GWBWR 0.85% or larger in portal-HTN/Child B or C to avoid SFSS.
- Without cirrhosis even 0.6% could be used in selected patients with normal liver function.
- Overall, 1-year survival is < 50% when GWBWR is < 0.8%)
- Later: Avoid excessive portal v. inflow and assure excellent hepatic vein outflow to prevent "graft flooding".

- ABO compatibility is recommended.
- ABO incompatible only in:
 - a) infants,
 - b) child < 1y/o without isoagglutinins,
 - c) emergency situation where no deceased-donor available
- Lab contraindications: HIV, HCV, HBsAg(+), anti-HBc(+)
- Thromboembolism & stress ulcer prophylaxis recommended.
- Autologous blood storage should be offered.