Liver transplantation: Advances and perioperative care

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ABSTRACT

Liver transplantation is one of the treatments for many-life threatening liver diseases. Numerous advances in liver transplant surgery, anaesthesia and perioperative care have allowed for an increasing number of these procedures. The purpose of this review is to consider some of the important advances in perioperative care of liver transplant patients such as pre-operative evaluation, intraoperative monitoring and management and early extubation. A PubMed and EMBASE search of terms "Anaesthesia" and "Liver Transplantation" were performed with filters of articles in "English", "Adult" and relevant recent publications of randomised control trial, editorial, systemic review and non-systemic review were selected and synthesized according to the author's personal and professional perspective in the field of liver transplantation and anaesthesia. The article outlines strategies in organ preservation, training and transplant database for further research.

Key words: Adult liver transplantation, anaesthesia, hepatopulmonary syndrome, hepatorenal syndrome, liver transplantation, organ preservation, recent advances, reperfusion injury

INTRODUCTION

Liver transplantation remains one of the treatments for many life-threatening liver diseases. The first human liver transplantation was performed in 1963 in Denver, Colorado.^[1] Around 350 liver transplants have been performed from 1995 to 2011 in India.^[2] Numerous advances in the perioperative management, like expertise in surgical techniques, better preoperative optimisation, intraoperative monitoring and management, changes in immunosuppression regime and advances in post-operative management not only increased the number of this procedure but also the outcome. One-year survival post-liver transplant has increased from 72% to 79% in 1998 to 85% to 90% in 2008. Ten-year survival has increased from 33% in 1998 to 53% in 2008 and 66% in 2010.^[3,4] Anaesthesia for liver transplantation^[5,6] and postoperative management of liver transplant recipients^[7] have been reviewed recently. The purpose of this review is to consider some of the important advances in perioperative care of liver transplant patients, in particular to cadaveric liver transplantation, as there has been a recent review on live related liver transplantation by one of the authors.^[8]

PRE-OPERATIVE EVALUATION FOR LIVER TRANSPLANTATION

Prognostic indices for a 1-year survival without liver transplantation, e.g. Child-Turcotte-Pugh or Revised Model for end-stage liver disease (MELD) scores, in patients with chronic liver disease determines the urgency for liver transplantation.^[9] A thorough assessment of the patient's comorbidities, both associated with and independent of liver failure, actuates the perioperative risks.^[10] A summary of the investigations considered in the transplant period is listed in Table 1.

Pre-transplant infections disease screening (dental, ophthalmic and otorhinolaryngological and urogenital)

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Table 1: Ir	vestigations for a liver transplant recipient
Blood investigations	Complete blood count, blood sugar
	Liver function tests, serum electrolytes and creatinine, prothrombin time, activated partial thromboplastin time and fibrinogen and factor levels, particularly factor V
	Arterial blood gas, blood grouping and cross-match
	Additional tests to detect aetiology of liver disease like anti-nuclear antibodies, ceruloplasmin, viral markers, iron studies, liver biopsy, α -fetoprotein
Radiology	X-ray chest
	Abdominal ultrasound with Doppler
	Computerized tomography and magnetic resonance imaging of the abdomen
Electrocardiography	12-lead ECG
Endoscopies	
Pulmonary function testing	Upper and lower GI endoscopies
Pre-transplant infectious disease screening	Blood and urine culture

of both donors and recipients is necessary to avoid postoperative infectious complications. A psychosocial assessment of the transplant recipient will help in identifying the negative factors affecting the outcome and to develop interventions to prevent it.

Cardiac evaluation

A transthoracic echocardiography (TTE) on all patients undergoing evaluation for liver transplantation assesses the structural and functional status of the heart.

Coronary artery disease

A 5–26% incidence of Coronary artery disease (CAD) exists in patients undergoing liver transplantation.^[11] Patients with risk factors for CAD, such as diabetes, non-alcoholic steato hepatitis (NASH), previous CAD and peripheral vascular disease need further evaluation to rule out CAD. Presence of three out of four cardiac risk factors, i.e. age more than 50 years, hypertension, elevated cholesterol and obesity, requires stress testing. Patients with two or more risk factors could undergo non-invasive computerized tomography to measure their calcium stores and a positive value (more than 100) should undergo stress testing.^[11]

Exercise stress tests are difficult to perform in patients with chronic liver disease due to the patient's limited functional capacity. Dobutamine stress myocardial perfusion imaging and dobutamine stress echocardiography can be done for detecting myocardial ischemia in liver disease patients.

Coronary angiography should be followed in patients with abnormal stress test results to rule out CAD. The American College of Cardiology/American Heart Association recommends coronary angiogram in patients more than 40 years of age as part of evaluation for transplantation if they have a known or suspected CAD or have stable angina.^[12] While there is no definite guideline as to when patients with CAD should undergo revascularization, there are case reports of combined liver transplant and cardiac surgery, particularly from high-volume centres.^[13] Liver transplant in patients with known CAD can be delayed for medical optimization and/or revascularization.^[14]

Cirrhotic cardiomyopathy

Cirrhotic cardiomyopathy characterized by abnormal or blunted cardiovascular response to stress, is present in as many as 50% of the patients undergoing liver transplant. These patients have inotropic and chronotropic incompetence, systolic and diastolic dysfunction, prolonged QT and electrical and mechanical dyssynchrony. Yet, these patients demonstrate a normal or increased myocardial contractility at rest.^[15]

Rapid haemodynamic changes during clamp release of the inferior vena cava increases filling pressures of the heart. There is an increased risk of congestive heart failure due to the impaired diastolic ventricular relaxation.

Pulmonary evaluation

Hepatopulmonary syndrome

In about 8–24% of patients with liver disease, Hepatopulmonary syndrome (HPS) exists. This is characterized by a decreased systemic arterial oxygenation (PaO_2 less than 80 mmHg or an alveolar arterial gradient of more than 15 mmHg on room air), pulmonary vascular dilation [contrast-enhanced echocardiography or abnormal uptake in the brain (more than 6%) with radioactive lung perfusion scanning] and liver disease.

Intravenous bubble contrast test can distinguish HPS from intracardiac shunts such as patent foramen ovale.^[16]

Portopulmonary hypertension

Portopulmonary hypertension (POPH) is characterized by increased mean pulmonary artery pressure (mPAP) greater than 25 mmHg at rest or greater than 30 mmHg with exercise, elevated pulmonary vascular resistance greater than 240 dyne/s/ cm⁻⁵ and normal or decreased pulmonary artery wedge pressure less than 15 mmHg. Among patients awaiting liver transplantation, the incidence of POPH is 2–10%.

A screening TTE uses tricuspid regurgitation to estimate right ventricular systolic pressure (RVSP). If the RVSP is more than 50 mmHg, these patients should undergo right heart catheterization to exclude other causes of pulmonary hypertension, such as high flow state and increased central volume and/or left ventricular dysfunction causing increased mPAP.^[16]

Hepatic hydrothorax

In cirrhotic patients, a pleural effusion greater than 500 mL without cardiac, pulmonary or pleural disease is termed hepatic hydrothorax.^[17]

Renal evaluation

Renal function is an important predictor of survival with chronic liver failure and liver transplantation.^[18]

Hepatorenal syndrome (incidence 18–40%) is a diagnosis of exclusion in patients with chronic liver insufficiency. It is characterized by liver disease, serum creatinine concentration more than 1.5 mg/dL (133 mmol/L) that is not reduced with the administration of albumin after discontinuation of diuretics for 2 days, absence of nephrotoxic drugs, shock and absence of findings suggestive of renal parenchymal disease (urinary protein greater than 500 mg/day, more than 50 red cells/high-power field or abnormal kidneys on ultrasonography).^[19]

INTRAOPERATIVE MONITORING AND MANAGEMENT

Haemodynamic monitoring

Haemodynamic monitoring is essential for successful liver transplantation.^[20] Additional to the standard cardiovascular monitors (electrocardiogram, pulseoximetry, invasive and non-invasive blood pressure), there is a need for cardiac output monitoring. There are numerous invasive and non-invasive cardiac output monitors available to monitor the haemodynamic change associated with liver transplantation.

Pulmonary artery catheter (PAC) is the gold standard used in haemodynamic monitoring during liver transplant to which other monitors have been compared.^[21] However, PAC has its own limitations, including poor real time updating, catheter migration and wedging. Furthermore, wedge pressure used to estimate left ventricular end diastolic pressure (LVEDP) as pre-load may be misleading in patient with liver disease. Table 2 shows the comparison of cardiac output monitors used in liver transplantation.^[22-29]

Monitoring of central venous oxygen saturation and mixed venous oxygen saturation during liver transplantation is of little value as they do not reflect changes in cardiac output nor oxygen supply demand mismatch during rapid haemodynamic disturbances.^[30,31]

Haemodynamic management

End-stage liver disease is often a high cardiac output state with decreased systemic vascular resistance and depleted intravascular volume. During the different stages of liver transplantation, i.e. pre-anhepatic phase, anhepatic phase and neohepatic phase, there are rapid fluid shifts due to blood loss, inferior vena cava clamping and reperfusion.

		Table 2: Haemodynamic mon	itors in liver transplant	
Monitors	Physical principle	Advantage	Disadvantage	References
Flotrac™	Pulse contour analysis	No need for calibration	Not useful in high cardiac output states and during major haemodynamic shifts	[22,23]
LIDCO™	Pulse contour analysis	Comparable to pulmonary	Needs calibration with lithium	[23,24]
		artery catheter in liver transplant patients for cardiac output	Requires frequent calibration during blood loss and fluid boluses	
PICCO™	Pulse contour analysis	Provides ITBV, GEDV and ELW	Needs external calibration and frequent calibration	[25,26]
			Results not reproducible	
USCOM™	Ultrasound, flow, doppler	Comparable to pulmonary artery catheter in liver transplant patients for cardiac output	Does not measure pulmonary artery pressure, interobserver variability and poor-quality images	[27,28]
TEE	Ultrasound, flow	Direct visualization of heart, rapid real-time changes	Cost, expertise training, difficult to measure PA pressure, rupture of oesophageal varices and bleeding	[29]

TEE – Transthoracic echocardiography, ITBV – Intrathoracic blood volume, GEDV – Global end diastolic volume, ELW – Extravascular lung water

Proactive attempts to lower the central venous pressure (CVP) either by phlebotomy or avoiding plasma transfusion during the pre-anhepatic phase have shown to reduce red cell transfusions,^[32] but with variable outcomes in post-operative renal status.^[33,34] Cywinsky *et al.* in their retrospective review of data question the reduced graft dysfunction due to lower CVP in neohepatic phase.^[35]

Patients with liver disease have been shown to have lower vasopressin levels. Vasopressin is often added intraoperatively to maintain the systemic vascular resistance.^[36] Vasopressin reduces portal blood flow by selective splanchnic vasoconstriction and hence may be useful in reducing the intraoperative blood loss.^[37] There is no evidence to date on the vasopressor of choice in maintaining the blood pressure during liver transplantation. Orthotopic Epinephrine; norepinephrine and dopamine have all been used.^[38] Methylene blue at a dose of 0.5 mg/kg body weight over 10 min can be used as a rescue to treat hypotension due to vasopressor-resistant vasoplegic shock.^[39] Phenylephrine is often used to tide over acute hypotensive episodes; however, bear in mind that it increases the pulmonary vascular resistance.

Pulmonary vasodilators [intravenous prostacyclin (epoprostenol) and inhaled iloprost] have shown improvement in POPH, but the optimal management strategy has not yet been defined due to lack of randomized control trials.^[40]

Fluid management

Liver transplant surgery is associated with massive fluid shifts both from the perspective of intravascular volume depletion and large surgical blood loss. Albumin can be used in liver transplants as the patients are often hypoalbuminaemic and hypovolaemic;^[41] however, the cost factor may factor in the decision of its use. The use of hydroxyl ethyl starches (modern low-molecular weight) and Gelatins in liver transplant has not been supported by convincing evidence. Furthermore, these may affect the coagulation profile and possesses a risk to increased renal injury.^[42]

The selection of various crystalloids during OLT should be based on their pH, electrolyte composition, osmolarity and metabolism. There is no ideal crystalloid solution and all have limitations; normal saline (0.9% NS) causes hyperchloremic acidosis while the lactate in Ringers Lactate (RL) requires liver metabolism for its elimination. Further, RL is a hypotonic solution and may increase the intracellular fluid. Plasmalyte has a pH near normal, electrolyte and osmolarity similar to plasma and acetate, which is metabolised extrahepatically to bicarbonate, but it is proinflammatory and potentially cardiotoxic.^[43]

Haemostasis and liver disease

Patients for liver transplant present with a wide range of coagulation profiles ranging from normal coagulation (hepatocellular carcinoma) to gross coagulopathy (fulminant hepatic failure).

Some consider the coagulopathy of liver failure to be a balanced coagulopathy having an equal decrease in pro- and anti-coagulant factors.^[44] The new state of coagulation lies in a narrow balance and hence can be tipped to either side by the complications of liver disease such as renal failure or infection resulting in bleeding, thrombosis or both.^[44]

Coagulation monitors

Prothrombin time (PT) and activated partial thromboplastin time (aPTT) have been found to have a limited role in OLT as they measure only the procoagulant pathway without considerations for platelets function or fibrinolysis.^[45] Thromboelastogram (TEG), rotational thromboelastometry (ROTEM) and Sonoclot measure the viscoelastic properties of blood during all stages of thrombus formation, tests stability, firmness of the clot and fibrinolysis. Hence, they provide a detailed assessment of both pro- and anticoagulant status of the blood.^[46] Use of TEG and ROTEM has been found to reduce the transfusion requirements in OLT,^[47] but it is important that these systems are well validated and supported by a coagulation laboratory.^[48]

Blood component management

Pre-anhepatic phase is associated with blood loss due to dissection of the native liver, particularly in patients with end-stage liver disease. Haemodilution and hypothermia, resulting in altered coagulation status, add on to the pre-existing anaemia and coagulopathy. Aggressive correction of the coagulation status may lead to hypervolaemia and increasing blood loss in addition to the effect on ionized hypocalcaemia. The aim at this stage is to avoid large volume of transfusion and dilutional coagulopathy.^[5] During the anhepatic phase, there is ionized hypocalcaemia, failure of clearance of thromboplastin and a rapid rise in tissuetype plasminogen activator with increase in plasmin activity and a hyperfibrinolytic state in addition to the pre-existing coagulopathy.^[5] Antifibrinolytics are used in the liver transplantation to prevent the hyperfibrinolytic state that may occur during the anhepatic and neohepatic phases due to increased tissue plasminogen activator in the absence of plasminogen activator inhibitor and alpha 2 antiplasmin. Use of antifibrinolytics has been questioned by the occurrence thromboembolic complications,^[49] but a recent metaanalysis fails to reveal any such association.^[50] Antifibrinolytics are used if there is evidence of fibrinolysis and there are no contraindications for its use, such as Budd-Chiari syndrome or hypercoagulable states. In Donation after Cardiac Death (DCD) donors where tissue plasminogen activator (tPA) has been given to prevent biliary complications, the use of antifibrinolytics may be concerning.^[50]

The neohepatic phase is associated with a multifactorial coagulopathy of hyperfibrinolysis, dilutional coagulopathy, heparin-like effect, platelet dysfunction, hypothermia and hypocalcaemia.^[5]

There is no evidence on which to base cut-off values for haemoglobin or coagulation parameters need to be corrected prior to blood product transfusion. However, there is evidence to support the association between red cell, fresh frozen plasma or platelet transfusion and an increased morbidity and mortality,^[51] but considerations should be made in interpreting this data as massive transfusion is a result of either massive blood loss or coagulopathy.

Intraoperative blood salvage and retransfusion of the autologous blood is a cost-effective method to reduce the need for allogenic red blood cells in OLT provided there are no contraindications to cell savage^[52] and the centre is a high turnover (with a minimum of 80 cases of cell savage per year).^[53]

There is no evidence to support the routine use of recombinant factor VIIa in liver transplantation. However, there are several case reports where it has been used in massive bleeding.^[54] Analysis of randomized controlled trials for the safety of off -label use of factor VIIa found an increased incidence of arterial thrombosis.^[55]

Neuro monitoring

Patients with liver disease undergoing transplantation are at risk of a wide range of neurological complications, including cerebral oedema, encephalopathy, seizures, hypoxia and central pontine myelinolysis.^[56] Conventional intracranial pressure (ICP) monitoring in liver transplant patients is complicated by bleeding and infection without any survival advantage.^[57] In acute liver failure patients, the use of transcranial Doppler, which measures cerebral perfusion pressures, has proven advantages. However, its use in the perioperative period has yet to be defined.^[58]

The inhalational anaesthetic requirement is less for patients with liver disease compared with healthy patients.^[59] Anaesthetic requirements vary along with their MELD scores in different phases of liver transplantation.^[60] Bispectral index (BIS) monitoring has been used in the peritransplant period to monitor consciousness and hence can be used intraoperatively to titrate the anaesthetic requirement.^[61] The routine use of BIS monitoring in OLT is of limited value because of its cost and minimal clinical utility in decision making.^[62]

Neurological management

Patients with elevated ICP should be placed with head elevated by 30°. Osmotic diuresis with IV mannitol is effective in reducing cerebral oedema, but runs the risk of fluid overload and pulmonary oedema in patients with hepatorenal syndrome.

In the setting of intracranial hypertension refractory to mannitol therapy, therapeutic hypernatraemia (serum sodium 145–155 mmol/L) may be used. Moderate hypothermia (32°–34°C) may also prevent or control elevated ICP. Induction of a barbiturate coma should be considered if osmotic therapy and hypothermia fail.^[63] Mechanical hyperventilation to reduce pCO_2 , except for temporary reduction in raised ICPs, is no longer considered as an optimum intervention.^[64]

ORGAN PRESERVATION

The aim of organ preservation is to maintain the viability of the donor graft and to allow for the restoration of normal function of the organ.

Cold storage and perfusate

Static cold storage (SCS) is the standard method of liver preservation. It is based on the principle that hypothermia reduces the metabolism and enzymatic activity and slows down cell death, but during the process it causes cellular oedema. Cellular oedema is prevented by addition of various substances to the preservative fluid. Euro Collins, University of Wisconsin (UW), Histidine-Tryptophan-Ketoglutarate (HTK), Marshall's hypertonic citrate (HOC) and Celsior are few of the perfusates being used in liver preservation.^[65,66] UW and HTK are the most common perfusate fluids used in liver transplantation. UW being expensive with increased viscosity and a high potassium/low sodium ratio (120 mM/L, 25 mM/L) has been its major disadvantages. HTK is inexpensive, less viscous and has a low potassium/high sodium ratio (9 and 15 mM/L) and has found a larger acceptance in liver transplantation.^[67] HTK contains low potassium and therefore does not need to be flushed before reperfusion.^[66]

Feng *et al.* in their metaanalysis of studies comparing UW and HTK did not find any major differences in outcome in liver transplants from live donor or from heart beating donors, but found that HTK is better at preventing biliary complications and more bile production in patients with marginal non-heart beating donors. This is probably due to better flushing as a result of low viscosity.^[68]

Machine perfusion

Hypothermic machine perfusion (HMP) combines the advantage of protecting the organ with hypothermia and continuous perfusion to restoration of energy and oxygenation.^[69] Although extensively used in renal and in lung transplantation, the evidence for HMP in liver transplantation is based mainly on animal data. There is a paucity of publications on the benefits in human subjects. Guarrera *et al.*^[70] show that in 20 liver transplant patients there was a benefit of HMP over SCS.

The use of normothermic machine perfusion (NMP) permits continuous machine perfusion while avoiding the complications of hypothermia, such as cold preservation injury and ischaemic rewarming during reperfusion, and provides nutrition at physiological temperatures.^[71] However, there are no human studies till date. The advantage of HMP and NMP has to be tested in large randomized human clinical trials before being put into practice.

Ischemia reperfusion injury

During the periods of ischemia and reperfusion of the liver, there is a microcirculatory failure, activation of Kupffer cells and production of reactive oxygen species, inflammatory responses and apoptosis of the hepatocytes.^[72]

Ischemia reperfusion injury (IRI) is associated with primary graft dysfunction and delayed graft function. Strategies of prevention of IRI include ischaemic preconditioning (IP) and pharmacological management. A recent review on the effect of ischaemic preconditioning on liver transplant (live and deceased donor) shows evidence of IP on decreasing the IRI in liver transplant patients. But, most of the studies use small sample size and the periods of IP are not uniform.^[73]

Some of the pharmacological manipulations aiming at reduction in IRI include antioxidants, free radical scavengers and preconditioning.

N-acetyl cysteine (NAC) is being used in liver transplantation patients to prevent renal failure and IRI of the new liver. NAC, in addition to its direct antioxidant property, replenishes glutathione and acts as a free radical scavenger.^[74] Systemic review of the use of NAC in liver transplantation for the purpose of preventing IRI of the graft liver is inconclusive.^[75] A recent RCT with use of NAC for preventing hepatorenal IRI did not show any improvement.^[76]

Inhalational anaesthetics, especially sevoflurane, have been shown to offer protection against IRI in the myocardium in cardiac patients.^[77] There is some evidence to suggest that sevoflurane possess the same properties in liver surgery patients by reduced transaminase levels.^[78] Infusion of remifentanil before induction of ischaemia has been shown to reduce ischaemic injury in rat liver model, and its clinical use has yet to be determined.^[79]

FAST TRACKING AND EARLY EXTUBATION

Early extubation in selected patients has shown to improve early graft function and reduce duration of stay in the Intensive Care Unit and nosocomial infections. However, selection of patients for early extubation depends on duration of the surgery, amount of blood and products transfused, patients' pre-operative status (MELD score), ischaemia time and status of the graft.^[80,81] A safe operating room extubation after liver transplantation (SORELT) prediction rule may be used to select patients for early extubation, but requires validation^[82] [Table 3].

OUTCOMES OF LIVER TRANSPLANTATION

Graft dysfunction is one of the dreaded complications of liver transplantation. Various factors predict the occurrence of graft dysfunction^[83] [Table 4]. A simple model of donor age and pre-operative MELD (D-MELD) defined a subgroup of donor–recipient matches with significantly poorer short- and long-term outcomes (D-MELD>1600).^[84]

COST-EFFECTIVENESS IN PERIOPERATIVE CARE FOR LIVER TRANSPLANTATION

The perioperative management of liver transplant is not uniform across the institutions and countries. Cost of the individual procedures plays a major role in the decision making during the perioperative care. Recently, a cost-effective analysis of liver transplantation perioperative care modalities in lowand medium-income countries has been published,

Table 3:	Safe operating room extubation after liver transplantation score
Major criteria	≥7 packed red blood cells intraoperatively
	End of surgery lactate ≥3.4 m mol/L
Minor criteria	Patient not at home at the time before surgery
	Duration of surgery ≥5 h
	Vasoactive drug infusions (dopamine >5 mcg/kg/min or nor epinephrine >0.05 mcg/kg/min)

Consider extubation if patient has less than two major, less than three minor or one major and two minor

Table reproduced with permission from Skurzak S, Stratta C, Schellino MM, Fop F, Andruetto P, Gallo M, *et al.* Extubation score in the operating room after liver transplantation. Acta Anaesthesiol Scand 2010 Sep"1; 54(8):970-978.

suggesting the best way to maximize cost-effectiveness in a resource-limited environment^[85] [Table 5].

CENTRAL DATABASE, RESEARCH AND TRAINING

At present, the number of centres performing liver transplantation in India is limited, but the number is steadily increasing.

An electronic central database of organ transplants (including liver), like the United Network for Organ Sharing (UNOS) database of the United States, has been established by the Indian Society of Organ Transplantation to collect the information, to help in long-term follow-up and in research studies that can improve future patient care.^[2] Simulation and porcine model can be used as a teaching tool for training in management of liver transplantation.^[86,87]

CONCLUSION

Liver transplantation has evolved over the years in terms of pre-operative preparation, perioperative monitoring and management, organ perfusion and

Table 4: Predictors of graft dysfunction			
Donor factors	Procurement and operative-related factors	Recipient factors	
Donor age	Prolonged cold ischaemia time	Reduced renal function	
Macrovesicular steatosis	Length of the anhepatic phase >100 min	Greater serum bilirubin	
High inotropic drug use		Dialysis before transplant	
Hypernatraemia		Underweight (body mass index <18.5 kg/m ²)	
Prolonged ICU stay		2 /	
Craft weight/reginight weight ratios <0.00/			

Graft weight/recipient weight ratios <0.8% Donors after cardiac death

Evidence	Perioperative modalities
Good evidence (highly recommended)	Pre-operative doppler echocardiography
	Fluid warmers
	Red cell salvage (Cell-saver)
	Tranexamic acid
	Early extubation protocol
Conflicting or low level of evidence (optional)	Myocardial perfusion imaging for routine screening
	Veno venous bypass
	PA catheter
	Thromboelastography
	Human albumin solution
	Haemostatic drugs other than tranexamic acid (EACA and rFVIIa)
No good evidence (no recommendation)	Continuous cardiac output
	TEE
	SvO ₂
	Anaesthetic drugs
	Number of anaesthetic personnel

Rando K, Niemann CU, Taura P, Klinck J. Optimizing cost-effectiveness in perioperative care for liver transplantation: A model for low-to-medium income countries. Liver Transpl 2011 Aug 11. doi: 10.1002/lt.22405. [Epub ahead of print] fast tracking. Studies are needed in the areas of haemodynamic monitoring, blood and coagulation management and fast tracking to prove the efficacy of one technique over the other. Training anaesthesia residents with simulation and animal models will prepare them to face the challenge of managing these complicated patients. Creating a national database will help in future research.

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KERALA STATE CHAPTER

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D. Application / Nomination

The applications / nominations with reprint of publications and other supporting documents should reach the Co-ordinator on or before 1st August, 2012 (both email and surface mail).

Dr. Thomas Koshy

Co-ordinator, KPR Endowment, Professor in Cardiac Anesthesiology,

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