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TOPIC HIGHLIGHT

#### 2016 Cirrhosis: Global view

# Outcomes of abdominal surgery in patients with liver cirrhosis

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### Abstract

Patients suffering from liver cirrhosis (LC) frequently require non-hepatic abdominal surgery, even before liver transplantation. LC is an important risk factor itself for surgery, due to the higher than average associated morbidity and mortality. This high surgical risk occurs because of the pathophysiology of liver disease itself and to the presence of contributing factors, such as coagulopathy, poor nutritional status, adaptive immune dysfunction, cirrhotic cardiomyopathy, and renal and pulmonary dysfunction, which all lead to poor outcomes. Careful evaluation of these factors and the degree of liver disease can help to reduce the development of complications both during and after abdominal surgery. In the emergency setting, with the presence of decompensated LC, alcoholic hepatitis, severe/advanced LC, and significant extrahepatic organ dysfunction conservative management is preferred. A multidisciplinary, individualized, and specialized approach can improve outcomes; preoperative optimization after risk stratification and careful management are mandatory before surgery. Laparoscopic techniques can also improve outcomes. We review the impact of LC on surgical outcome in non-hepatic abdominal surgeries required in this cirrhotic population before, during, and after surgery.

Key words: Liver cirrhosis; Outcomes; Coagulopathy; Nutritional status; Abdominal surgery; Adaptive



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immune dysfunction; Cirrhotic cardiomyopathy

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**Core tip:** The prevalence of chronic liver disease is increasing. Patients with liver cirrhosis may be more likely to need non-hepatic abdominal surgery than the non-cirrhotic population. The rising incidence of non-alcoholic fatty liver disease and non-alcoholic steatohepatitis may increase the prevalence of cirrhotic patients within the abdominal surgery population. The pathophysiological characteristics of the condition raise the surgical risk and the likelihood of poor prognosis. A review of the assessment and outcomes for nonhepatic abdominal procedures in these patients is essential since most current recommendations are based on observational studies.

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### INTRODUCTION

Mortality and morbidity rates are high in patients with liver cirrhosis (LC) requiring non-hepatic abdominal surgical procedures<sup>[1,2]</sup>. Despite the lack of reliable epidemiological data, the incidence of LC is expected to increase due to alcohol consumption and the potential high rates of hepatitis C virus (HCV) and hepatitis B virus (HBV) in coming years in European countries<sup>[3,4]</sup>. In addition, the obesity epidemic in Western countries, which is associated with metabolic syndrome, is expected to generate a large cohort of patients with non-alcoholic steatohepatitis (NASH) and non-alcoholic fatty liver disease (NAFLD) that may potentially change the etiological pattern of LC<sup>[5,6]</sup>.

LC patients may require surgery for abdominal wall hernia, gallstones<sup>[7]</sup>, peptic ulcer disease, biliary, small bowel, colo-rectal, and pancreatic diseases as well as liver procedures, such as transplantation. Surgery represents an additional source of stress for compromised liver function, and perioperative complications frequently appear in spite of significant advances in surgical and intensive care management<sup>[8]</sup>. Preoperative risk stratification is difficult due to the limited accuracy of the tools available to assess LC, and, in some cases, due to the absence of diagnosis. The mortality and morbidity risks are associated with the severity of the underlying LC<sup>[9,10]</sup>. This is one of the factors that explains the wide variation in outcomes recorded by different studies, in addition to patient characteristics, the surgical approach used, and the

levels of expertise of the surgeons, anesthetists, and intensive care unit (ICU) staff. Liver function is usually assessed by the Child-Turcott-Pugh (CTP) and Model for End-Stage Liver Disease (MELD) scores<sup>[11]</sup>. Early studies found 30 d mortality rates after surgery were 10% in CTP-A, 30% in CTP-B, and 76%-82% in CTP-C, figures that have not significantly changed in more recent assessments<sup>[12,13]</sup>. Despite the poor results, advances in the medical management of LC and life expectancy have increased the eligibility of these patients for abdominal surgery<sup>[14]</sup>. To be able to give definitive recommendations and indications for non-hepatic abdominal surgery in the cirrhotic population, it is important to identify the patients most likely to benefit from it. There is also a need to assess contemporary surgical techniques and the various scoring systems currently in use.

This review summarizes the outcomes of patients with LC undergoing non-hepatic abdominal surgery. Indexed articles in Medline of series of patients with LC who underwent non-hepatic abdominal surgery between 1950 and March 2014 were reviewed using the OVID interface. We aimed to select manuscripts addressing outcome based on the degree of LC assessed with MELD and/or CTP scores. Articles addressing the pathophysiology of cirrhotic patients and the clinical implications in non-hepatic abdominal surgery were selected based on their importance, their date of publication, and the citations of the manuscripts. As for articles describing the different types of surgery in LC patients, the most recent publications were selected in order to preserve comparability between contemporary surgical techniques. For these reasons, the present review does not follow the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA statement).

## PREOPERATIVE MANAGEMENT AND PATHOPHYSIOLOGICAL CONSIDERATIONS OF LC BEFORE SURGERY

The estimation of liver functional reserve and the identification of coexisting pathophysiological disorders, such as coagulopathy, malnutrition, cardiomyopathy, renal dysfunction, respiratory dysfunction, ascites, and hyponatremia, are key issues in the preoperative evaluation of LC patients scheduled for non-hepatic abdominal surgery. Despite the lack of evidence-based guidelines for the management of these patients, clinical and surgical teams are acquiring experience with LC patients undergoing non-hepatic abdominal surgery, and the body of knowledge of the pathophysiology of LC in the literature is increasing. Therefore, some general recommendations for the care of these patients are in order<sup>[15]</sup>.

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### Liver function

The occurrence of portal hypertension (PHT) in LC leads to variceal bleeding, ascites, spontaneous bacterial peritonitis (SBP), and hepatic encephalopathy. Patients with LC are at a higher risk of liver-related complications during surgery due to inappropriate response to surgical stress. Mortality is higher in nonhepatic abdominal surgery in the presence of PHT, which is also an indicator of illness progression<sup>[10,15]</sup>. The presence of high intraoperative PHT is associated with postoperative mortality due to rebleeding<sup>[16]</sup>. Preoperative evaluation of liver function by a physician experienced in managing liver disease in patients with confirmed or suspected LC is needed in order to stage the severity<sup>[17]</sup>. In clinical practice, severity should be assessed by means of both CTP and MELD scores, although MELD is more objective<sup>[18]</sup>. These scores should be assessed throughout the patient's admission because rapid deterioration indicates that surgery may not be safe. The value of indocyanine green plasma disappearance rate has been demonstrated in patients with LC undergoing cardiac surgery<sup>[19,20]</sup> and partial hepatectomy<sup>[21]</sup>, and it may be useful for assessing liver functional reserve in this scenario.

As a general rule, surgery is safe in CTP A patients without PHT and MELD < 10, but its pros and cons should be carefully considered in patients with severe  $LC^{[22,23]}$ . Surgery should be avoided if possible in the setting of acute viral hepatitis, alcoholic hepatitis, acute liver failure, acute renal failure, severe coagulopathy, hypoxemia, and/or cardiomyopathy<sup>[14]</sup>.

Regarding the etiology of LC, certain specific considerations need to be addressed before surgery. Due to the stress that surgery represents in patients with autoimmune hepatitis who are receiving or have recently received steroids, increasing the dose is a possible alternative. D-Penicillamine, the chelating agent used to treat Wilson disease, interferes with wound healing and should be reduced 2 wk before surgery and until the wound has healed completely. Alcohol consumption must be discontinued, especially in patients with a history of alcohol abuse, because of its associations with the risk of hepatotoxicity from drugs such as paracetamol, poor wound healing, bleeding, delirium, infections, and alcoholic hepatitis (the last of these being a clear contraindication for elective surgery in those patients)<sup>[24-26]</sup>. In addition, alcohol abstinence is very difficult, and the majority of patients have superimposed alcoholic hepatitis<sup>[26]</sup>.

#### Coagulopathy

Coagulopathy is the major concern for surgeons, because both pro- and anti-coagulants present reductions in LC. Thrombocytopenia due to poor nutritional status, hypersplenism and/or bleeding from varices may exacerbate bleeding problems<sup>[27,28]</sup>. Myelosuppression due to hepatitis C, folate deficiency, and ethanol toxicity can also aggravate thrombopoietin deficiency<sup>[27]</sup>. However, primary homeostasis may not be defective in LC, and a low platelet count, if not severe, should not be automatically considered as an index of an increased risk of bleeding<sup>[29]</sup>. Thrombocytopenia of < 50000/µL for moderate risk and 100000/µL for high risk procedures must be corrected by platelet transfusion prior to surgery<sup>[30]</sup>. Infections, renal failure, malabsorption, and/or malnutrition can adversely affect coagulation in decompensated LC.

Conventional coagulation tests measure only part of the process of thrombin generation. The prothrombin time-derived international normalized ratio (PT-INR) is universally used to assess bleeding risk and prognosis in MELD score and to guide treatment of coagulation disturbances in clinical practice; however, it only measures the activity of procoagulants. Thromboelastography (TEG) provides a better assessment of the degree of coagulopathy and offers information allowing immediate transfusion therapy. It is useful before and during surgery for guiding transfusion therapy<sup>[31]</sup>. Serum fibrinogen, a key factor in fibrin clot formation, and protein C deficiency may also determine the risk of bleeding in LC<sup>[32]</sup>.

Coagulopathy is one of the factors of LC that can be modified preoperatively with intravenous vitamin K replenishment and cryoprecipitate transfusions in order to maintain a serum fibrinogen level < 100 mg/dL. Preoperative administration of fresh-frozen plasma to correct the INR should be avoided due to its ineffectiveness and its association with volume overload, exacerbation of PHT, risk of infections, and transfusionrelated acute lung injury<sup>[33,34]</sup>. Administration of desmopressin and tranexamic acid should be considered when platelet dysfunction and hyperfibrinolytic state are suspected<sup>[35]</sup>. This may help survival after surgery and convert CTP C patients into CTP B by improving the coagulopathy<sup>[36]</sup>.

The lack of improvement in PT-INR with the administration of vitamin K may reflect a poor hepatic reserve and outcome. In addition, plasma hemoglobin levels lower than 10 g/dL and higher intraoperative transfusion needs are independent prognostic factors for mortality in these patients<sup>[1,18]</sup>. These findings are linked with a major deterioration of coagulation and the association of perioperative blood product transfusion with the triggering of the inflammatory response<sup>[37]</sup>.

#### Immune dysfunction and nutritional status

Infections are an important cause of death during the postoperative period due to an innate and adaptive immune dysfunction in LC, which is associated with the development of liver dysfunction and an enhanced susceptibility to acute inflammatory processes<sup>[38,39]</sup>. This is also linked with nutritional status: between 50% and 90% of LC patients lack sufficient nutritional reserve and have a poor metabolic state, with inadequate inflammatory and immune responses to surgery<sup>[40,41]</sup>.

Malnutrition in patients undergoing abdominal surgery is associated with a greater risk of postoperative complications and longer hospitalization<sup>[42]</sup>. Although the evidence is controversial, preoperative nutrition support may play an important role in preventing complications<sup>[43]</sup>. Nutritional disorders are more severe with alcoholic cirrhosis than with non-alcoholic cirrhosis, which can explain the poorer prognosis of this subgroup of patients when undergoing surgery<sup>[44]</sup>. A combination of transferrin, prealbumin, and albumin seems an adequate approach for nutritional evaluation in LC<sup>[45]</sup>. Indeed, hypoalbuminemia has been associated with poor outcome in abdominal and colon cancer interventions in patients with advanced LC<sup>[8,46]</sup>.

### Cardiovascular and renal dysfunction

Preoperative evaluation of cardiovascular dysfunction in LC is crucial. Cardiovascular diseases are a common cause of mortality in LC because of the severity of liver injury, and inflammation is strongly associated with an increased cardiovascular risk and an atherogenic lipid profile<sup>[47]</sup>. This is especially relevant in NASH/NAFLD chronic liver disease because the etiological risk factors are the same for both LC and cardiovascular disease, even taking into account the expected rise in the incidence of NASH/NAFLD worldwide<sup>[48]</sup>. A hyperdynamic circulatory state with increased cardiac output in response to splanchnic arterial vasodilatation is an inherent characteristic of LC, leading to heart failure with the progression of liver disease. Cirrhotic cardiomyopathy develops a variety of progressive clinical manifestations and is characterized by diastolic dysfunction and impaired inotropic and chronotropic competence, leading to a suboptimal ventricular contractile response during stressful conditions<sup>[49,50]</sup>.

The dynamic assessment of preoperative cardiac function with dobutamine stress echocardiography may play a role in the indication of abdominal surgery and postoperative management in the setting of LC. Cirrhotic cardiomyopathy may also be involved in the pathogenesis of hepatorenal syndrome (HRS) or the development of acute kidney injury (AKI). AKI can lead to a positive fluid balance, resulting in vital organ edema related to cardiac output performance<sup>[51,52]</sup>. LC may cause renal dysfunction and HRS; which occurs in conjunction with microcirculatory dysfunction in other organs, including the heart and the peripheral vascular bed<sup>[53]</sup>. Assessment of preoperative renal function is of paramount importance due to the high influence of AKI on postoperative survival in LC patients<sup>[54,55]</sup>. Thus, the identification of high creatinine levels as an independent predictor of complications and mortality in LC patients undergoing surgery was not surprising<sup>[56]</sup>.

### Ascites, respiratory dysfunction, and hyponatremia

Ascites and fluid overload may cause or aggravate pulmonary function due to atelectasis and pulmonary edema. In advanced LC, hepatopulmonary syndrome, portopulmonary hypertension, and hepatic hydrothorax are typical pulmonary complications. The endexpiratory lung volume may fall, leading to impairment in the mechanics of the respiratory system, lung, and chest wall as well as gas-exchange. Thus, initial use of moderate Positive End Expiratory Pressure is advisable to improve oxygenation and compliance without causing adverse effects in the respiratory function<sup>[57-59]</sup>.

Hemodynamic changes, such as systemic vasodilatation and volume retention in the splanchnic bed secondary to PHT, reduce preload and the associated retention of water and sodium in order to compensate for the low effective circulatory volume. These mechanisms explain the development of ascites and the impairment of the kidneys in the elimination of solute-free water. Ultimately, this also explains the development of hyponatremia, which is associated with increased morbidity and mortality in patients with  $LC^{[60]}$ . Sodium < 130 mmol/L in non-hepatic abdominal surgery and ascites in general surgery are markers of poor prognosis in LC patients<sup>[1,8,56]</sup>.

Preoperative portal decompression by transjugular intrahepatic portosystemic shunt (TIPS) seems to be a safe procedure for reducing the risk of bleeding in the presence of varices and the risk of ascites even in decompensated LC. Although there is not enough evidence to support its routine use before nonhepatic surgery, it could be considered in patients with significant ascites, extensive abdominal varices, or both. Reducing portal pressure seems to be helpful in decreasing bleeding during surgery<sup>[61-63]</sup>.

If possible, these complications need to be screened in candidates for surgery, especially in those with a medical history consistent with LC-related complications, and then medically optimized prior to surgery. Despite their implications for the perioperative course of LC patients undergoing any type of surgery, they have not been adequately addressed in non-hepatic abdominal surgery, especially when compared with other major surgeries, such as cardiac surgery<sup>[2]</sup>. In Table 1 we summarize recommendations for preoperative evaluation in patients with LC scheduled for non-hepatic abdominal surgery.

### INTRAOPERATIVE MANAGEMENT

Patient management during surgery is crucial for outcomes in LC. A team-based approach involving hepatologists, surgeons, and anesthetists with experience in treating LC patients is required, ideally at a specialist center<sup>[1,8,9]</sup>.

### Anesthesiology

Intraoperative management depends more on the anesthesiologist than on the surgeon, because the use of anesthesia tends to cause hepatic decompensation in patients with  $LC^{[64]}$ . Fluid management can be difficult since crystalloids (*e.g.*, Hartmann's solution or



#### Table 1 Recommendations for evaluation before non-hepatic abdominal surgery in liver cirrhotic patients

Elements to evaluate		Recommended tests	Recommended action
Homeostasis	Thrombo-cytopenia	Platelet number and function by	Preoperative transfusion if:
		means of thromboelastography	Platelet > $50000/\mu L \rightarrow moderate risk procedures$
		017	Platelet > $10000/\mu L \rightarrow$ high risk procedures
			Consider desmopressin (300 µg intranasal) if uremia or altered platelet function in
			thromboelastography
	Coagulopathy	PT-INR; thromboelastography.	IV replenishment of vitamin K ( $\ge 10$ mg OD during 3 d)
		Serum fibrinogen;	Cryoprecipitate if serum fibrinogen $\ge 100 \text{ mg/dL}$
		Thromboelastography	
		C	onsider Tranexamic acid (10 mg/kg TD during 2-7 d)
Liver function	PTH	Abdominal US	Consider the less invasive surgical treatment or avoid surgery if severe PTH
			Consider TIPS
	Ascites	Diagnostic ascitic tap; check	Discard SBP
		diuretics response	Antibiotic prophylaxis or treatment.
			Sodium restriction and diuretics (careful monitoring of renal function avoiding
			hyponatremia)
			Large volume of paracentesis for uncontrolled ascites
	Esophageal varices	Upper endoscopy; Abdominal	Consider prophylactic treatment (i.e., $\beta$ -blockers, variceal banding) based of risk of
		US	bleeding
Immune	Malnutrition,	White blood cells count;	Optimize protein and caloric intake (higher requirements than normal individuals)
function and	hypoalbuminemia	Nutritional biomarkers:	Vitamin B1 in alcoholics
nutritional		Albumin, Pre-albumin,	Administer antibiotic prophylaxis if suspected concurrent infections
status		transferrin; muscle wasting	(Other than SBP)
	Glucose intolerance	Laboratory testing	Insulin infusion
cardiac	Cardiomyopathy	Dobutamine	Consider the less invasive surgical treatment or avoid surgery if severe cardiac
function		stress	dysfunction
		echocardiography	Consider close invasive monitoring and hemodynamic strategy in order to
			preserve normal cardiac function and avoid organ hypoperfusion
			(especially liver and kidney)
			Consider $\beta$ -blockers in perioperative period
Renal function	Renal dysfunction;	Serum creatinine;	Avoid dehydration if possible before surgery
	Hepatorenal	Glomerular filtration rate;	Avoid positive fluid balance during perioperative course
	syndrome	Evaluate normal Blood Pressure	(if hemodynamics allow that)
		and cardiac performance	
Pulmonary	Hydrothorax;	Chest-X ray;	Optimize pulmonary function:
function	HPS;	Electrocardiogram and	Discard high arterial pulmonary pressure
	PPH	echocardiography;	Discard pleural effusion/thoracentesis if necessary
		Spirometry	If HPS/PPH evaluate appropriate therapy (i.e., IV epoprostenol, sildenafil)
CNS	HE	Clinical assessment;	Use of lactulose despite absence of HE if medical past history or PTH
		Ammonia serum levels	Treat or avoid potential triggers of HE (i.e., diuretics, infections, constipations,
			CNS depressants, azotemia, uremia, hyponatremia)

PT-INR: Prothrombin time-derived international normalized ratio; IV: Intravenous; PTH: Portal hypertension; US: Ultrasound; TIPS: Transjugular intrahepatic portosystemic shunt; SBP: Spontaneous bacterial peritonitis; HPS: Hepatopulmonary syndrome; PPH: Portopulmonary hypertension; CNS: Central nervous system; HE: Hepatic encephalopathy.

saline) may worsen ascites and peripheral edema and have little effect over intravascular volumes<sup>[65]</sup>. Indeed, intravascular volume may be depleted, even in the setting of extravascular volume overload<sup>[14]</sup>. Thus, it seems more appropriate to provide fluid support in the form of a volume expander if necessary, prioritizing fluid restriction. Blood products should be used routinely, and TEG can achieve optimal monitoring of coagulopathy. The increased physiological demand that surgery represents in patients with identified cirrhotic cardiomyopathy can lead to heart failure. Minimization of large fluctuations in preload and afterload are recommended in order to improve outcomes. The avoidance of heart failure during the perioperative period has important prognostic implications<sup>[66]</sup>. It is important to note that most of these patients are currently receiving treatment with  $\beta$ -blockers<sup>[67]</sup>, which

may ultimately reduce the cardiac response under stressful conditions. Thus, a closer monitoring of cardiac performance and the prompt use of inotropes and vasoconstrictors are mandatory in order to avoid low cardiac output and intraoperative hypotension.

Perioperative antibiotic prophylaxis should be given in the presence of ascites to prevent postoperative SBP or bacteremia secondary to SBP, which may occur during the procedure. The most frequent microbiological causes, such as gram-negative bacteria, must be covered<sup>[68]</sup>. Although quinolones are the most frequently used antibiotic for SBP prophylaxis, an individualized approach is needed due to the emergence of multidrug-resistant agents<sup>[69]</sup>.

Pain and sedative management regimens need to be minimized and adapted according to the degree of liver disease and the alterations to the metabolism Table 2 Modifications in operative laparoscopic techniques in non-hepatic abdominal surgery in liver cirrhosis who underwent cholecystectomy and hernia repair

Ref.	Modified technique	Objective and advantage	
Laparoscopic cholecystectomy			
Friel <i>et al</i> <sup>[77]</sup> , 1999	Use of open technique using Hassan's trocar	Prevent inadvertent puncture of umbilical varix	
Shiff <i>et al</i> <sup>[78]</sup> , 2005	Placement of the trocar in the right paramedian position		
Clark <i>et al</i> <sup>[79]</sup> , 2001	Use of additional ports	Facilitate laparoscopic technique and prevent	
	Performance of retrograde cholecystectomy	complications in cases of severe gallbladder	
Clark <i>et al</i> <sup>[79]</sup> , 2001	Modified subtotal cholecystectomy	inflammation	
Palanivelu et al <sup>[80]</sup> , 2006			
Friel <i>et al</i> <sup>[77]</sup> , 1999	Mechanical compression from introduced surgical sponges	Facilitate haemostasis	
	( <i>i.e.</i> , oxidized cellulose)		
	Application of ultrasonic energy via harmonic scalpel		
	Use of argon beam coagulator through an operative port		
Laparoscopic hernia repair			
Belli <i>et al</i> <sup>[81]</sup> , 2006	Minimally invasive and tension-free laparoscopic technique	Prevent inadvertent puncture of collateral veins	
		Prevent recurrence rates and wound infections	
McAlister et al <sup>[82]</sup> , 2003	Dual mesh prosthesis: fixation of mesh in a preperitoneal space	Prevent recurrence rates and mesh migration	
	Sterile fashion of mesh insertion	Prevent wound infections	

caused by these anesthetic drugs. Portal blood flow is reduced as a result of PHT, and anesthetic agents may reduce hepatic blood flow by 30%-50% as a result of impaired autoregulation. In addition, myocardial depression and vasodilatation are frequent in most anesthetic agents<sup>[70]</sup>. Agents with minimal hepatic metabolism (< 0.2%), such as isoflurane, desflurane, and sevoflurane together with nitrous oxide, are an appropriate choice for the operating room, along with propofol as a narcotic agent<sup>[71,72]</sup>. These agents also cause fewer disturbances in hepatic arterial blood flow than others<sup>[73]</sup>. Atracurium and cisatracurium are preferred as muscle relaxants since they are metabolized independently from the liver; vecuronium and rocuronium, on the other hand, are metabolized exclusively by the liver and must be avoided<sup>[74,75]</sup>.

Some authors argue that epidural anesthesia should be avoided in LC patients due to the complications derived from the coagulopathy<sup>[8]</sup>. However, epidural analgesia has many benefits, not only in terms of pain control during and after surgery, but also in terms of reducing pulmonary, cardiovascular, thromboembolic, and gastrointestinal complications and enhancing the recovery of gut function after abdominal surgery<sup>[76]</sup>. In our opinion, epidural anesthesia may be safe in the absence of abnormal coagulation and/or platelet count, especially if evaluated preoperatively by means of TEG.

#### Surgical considerations

Surgery should ideally be elective, due to the higher than average risk associated with emergency surgery in cirrhotic patients<sup>[1,7-11]</sup>. Careful tissue handling and meticulous surgical technique are vital to prevent major bleeding during surgery. Morbidity and mortality rates vary greatly depending on the severity of the cirrhosis, the presence of PHT, complications of LC, and the nature of the surgical procedure.

The incidence of gall stones and hernia is higher in LC than in the non-cirrhotic population due to the increased intra-abdominal pressure. Cholecystectomy and hernia repair surgery are the most frequently described surgeries in LC, and modified laparoscopic techniques have been proposed in these cases in order to minimize morbidity (Table 2)[77-82]. The use of laparoscopy allows non-exposure of viscera and restricts electrolytic and protein losses<sup>[79]</sup>. From the technical point of view, the use of coagulation instruments like the harmonic scalpel causes less postoperative pain and lower morbidity<sup>[83,84]</sup>. The use of laparoscopic general surgery is also supported by the lower morbidity rates, lower bleeding complications, shorter operating time and hospital stay (despite the higher conversion rates to open surgery during the procedure, especially in the case of emergency surgery), and fewer bleeding-related complications compared to noncirrhotic patients<sup>[77,85,86]</sup>. The laparoscopic approach also helps to reduce morbidities such as surgical site infection and hemorrhage<sup>[87]</sup>. A wide range of interventions can be performed laparoscopically in LC patients: splenectomies, colectomies, Nissen fundoplication, Heller's myotomy, gastric bypass, radical nephrectomy<sup>[88]</sup>, appendectomies<sup>[89]</sup>, suture closure, and placement of an omental patch for treatment of a perforated gastric ulcer<sup>[90]</sup>.

Although, in general, outcomes are poorer in LC patients than in the non-LC population in terms of morbidity and mortality, mortality rates are quite similar in certain surgeries thanks to the technical advances and experience gained in recent years (Table 3)<sup>[10,77-79,81-87,89,91-101]</sup>. One example is abdominal wall surgery. Elective repair of umbilical hernia in patients with LC and ascites is indicated due to the poor outcomes with conservative management<sup>[102-104]</sup>; and, even in advanced LC, inguinal hernia repair obtains similar outcomes to those recorded in the non-LC

Type of surgery	Morbidities	Mortality in LC population	Mortality in non-LC population			
Cholecystectomy <sup>[77-79,83-86,91-97]</sup>						
Laparotomy	30%-35%	1%-7.7%	0.5%-1%			
Laparoscopy	13%-33%	<1%	<1%			
Colorectal surgery <sup>[10,98]</sup>	43%	14%-29% (20.9%-35.8% if ES)	5%			
Radical gastric surgery <sup>[99]</sup>	56% (53.3% CTP A, 67.7% CTP B)	-	80% (5-yr)			
Appendectomy <sup>[87,89]</sup>						
Laparotomy	5%	9%	0.7%			
Laparoscopy	<1%	<1%	<1%			
Pancreatic surgery <sup>[100]</sup>	69% (67% CTP A, 100% CTP B)	9% (3% CTP A, 100% CTP B)	-			
Abdominal wall surgery <sup>[81,82,101-110]</sup>						
Umbilical hernia	7%-20%	< 1%-5.5%	<1%			
Inguinal hernia	6.3%-10.9%	< 1%-2.7%	<1%			

Table 3 Outcomes of liver cirrhosis who underwent non-hepatic abdominal surgery based on type of surgery

LC: Liver cirrhosis; ES: Emergency surgery; CTP: Child-Turcotte-Pugh; ICU: Intensive care unit.

population and improves quality of life<sup>[81,82,100-110]</sup>.

### **POSTOPERATIVE MANAGEMENT**

Postoperative management should ideally be performed in the ICU, at least during the first 24 h, especially in CTP B and C. It is important to monitor for the complications that may be expected depending on the degree of LC and type of surgery. As discussed above, postoperative management should be based on the pathophysiological characteristics of LC. Physicians should focus on fluid management, vigilance, and prophylaxis of infections (especially surgical site infections); opioids and sedatives should be used cautiously. The inadequate administration of normal saline can lead to the development of ascites. Salt restriction with both intravenous fluid and oral intake is mandatory in order to prevent the postoperative occurrence of AKI, ascites, and/or hepatic encephalopathy<sup>[111,112]</sup>. Early introduction of lactulose should prevent encephalopathy. In the presence of fast decompensation of liver function, physicians should first rule out infection<sup>[113]</sup>. The dosage of analgesics for pain control should be based on the degree of liver dysfunction because the drug metabolism of simple analgesics, such as paracetamol, non-steroidal, and opioid analgesics, may be impaired. If opioids are necessary, fentanyl is preferred because it does not produce active metabolites; however, it may accumulate in the fat tissue for several days and is cleared through the liver<sup>[114]</sup>.

### CONCLUSION

Patients with LC undergoing non-hepatic abdominal surgery are at an increased risk of poor outcome. Emergency surgery should be avoided if possible, and conservative management is preferred in the presence of decompensated LC, alcoholic hepatitis, severe/advanced LC, and significant extrahepatic organ dysfunction. A multidisciplinary team approach involving surgeons, anesthesists, intensivists, and gastroenterologist/

hepatologists together with specialized hospital staff with experience in the perioperative management of those patients can improve outcomes. Since there are no formal guidelines and few randomized controlled trials have been performed, the pathophysiological characteristics of LC mean that an individualized approach to the care of the patients is essential. Preoperative optimization after risk stratification is mandatory before surgery. Laparoscopic techniques may improve outcomes in those patients, and recent advances in cholecystectomy, abdominal wall surgery, and appendectomy have reduced mortality to levels similar to those found in the non-LC population. Future prospective randomized studies are needed to assess the effect of preoperative TIPS, to compare elective surgery vs conservative management, and to compare preoperative assessment with MELD and CTP scores for specific surgical procedures. These studies should also assess the efficacy of new approaches, especially before surgery, in order to establish formal guidelines for the management of patients with LC undergoing nonhepatic abdominal surgery.

### REFERENCES

- Neeff H, Mariaskin D, Spangenberg HC, Hopt UT, Makowiec F. Perioperative mortality after non-hepatic general surgery in patients with liver cirrhosis: an analysis of 138 operations in the 2000s using Child and MELD scores. *J Gastrointest Surg* 2011; 15: 1-11 [PMID: 21061184 DOI: 10.1007/s11605-010-1366-9]
- 2 Lopez-Delgado JC, Esteve F, Javierre C, Ventura JL, Mañez R, Farrero E, Torrado H, Rodríguez-Castro D, Carrio ML. Influence of cirrhosis in cardiac surgery outcomes. *World J Hepatol* 2015; 7: 753-760 [PMID: 25914775 DOI: 10.4254/wjh.v7.i5.753]
- 3 Hatzakis A, Wait S, Bruix J, Buti M, Carballo M, Cavaleri M, Colombo M, Delarocque-Astagneau E, Dusheiko G, Esmat G, Esteban R, Goldberg D, Gore C, Lok AS, Manns M, Marcellin P, Papatheodoridis G, Peterle A, Prati D, Piorkowsky N, Rizzetto M, Roudot-Thoraval F, Soriano V, Thomas HC, Thursz M, Valla D, van Damme P, Veldhuijzen IK, Wedemeyer H, Wiessing L, Zanetti AR, Janssen HL. The state of hepatitis B and C in Europe: report from the hepatitis B and C summit conference\*. *J Viral Hepat* 2011; 18 Suppl 1: 1-16 [PMID: 21824223 DOI: 10.1111/ j.1365-2893.2011.01499.x]
- 4 Harris RJ, Thomas B, Griffiths J, Costella A, Chapman R,

Ramsay M, De Angelis D, Harris HE. Increased uptake and new therapies are needed to avert rising hepatitis C-related end stage liver disease in England: modelling the predicted impact of treatment under different scenarios. *J Hepatol* 2014; **61**: 530-537 [PMID: 24824282 DOI: 10.1016/j.jhep.2014.05.008]

- 5 Wong RJ, Ahmed A. Obesity and non-alcoholic fatty liver disease: Disparate associations among Asian populations. *World J Hepatol* 2014; 6: 263-273 [PMID: 24868320 DOI: 10.4254/wjh.v6.i5.263]
- 6 Corey KE, Kaplan LM. Obesity and liver disease: the epidemic of the twenty-first century. *Clin Liver Dis* 2014; 18: 1-18 [PMID: 24274861 DOI: 10.1016/j.cld.2013.09.019]
- 7 del Olmo JA, Flor-Lorente B, Flor-Civera B, Rodriguez F, Serra MA, Escudero A, Lledó S, Rodrigo JM. Risk factors for nonhepatic surgery in patients with cirrhosis. *World J Surg* 2003; 27: 647-652 [PMID: 12732995 DOI: 10.1007/s00268-003-6794-1]
- 8 Telem DA, Schiano T, Goldstone R, Han DK, Buch KE, Chin EH, Nguyen SQ, Divino CM. Factors that predict outcome of abdominal operations in patients with advanced cirrhosis. *Clin Gastroenterol Hepatol* 2010; 8: 451-47, quiz e58 [PMID: 20036761 DOI: 10.1016/j.cgh.2009.12.015]
- 9 Teh SH, Nagorney DM, Stevens SR, Offord KP, Therneau TM, Plevak DJ, Talwalkar JA, Kim WR, Kamath PS. Risk factors for mortality after surgery in patients with cirrhosis. *Gastroenterology* 2007; 132: 1261-1269 [PMID: 17408652 DOI: 10.1053/ j.gastro.2007.01.040]
- 10 Csikesz NG, Nguyen LN, Tseng JF, Shah SA. Nationwide volume and mortality after elective surgery in cirrhotic patients. *J Am Coll* Surg 2009; 208: 96-103 [PMID: 19228510 DOI: 10.1016/j.jamcoll surg.2008.09.006]
- 11 Rai R, Nagral S, Nagral A. Surgery in a patient with liver disease. J Clin Exp Hepatol 2012; 2: 238-246 [PMID: 25755440 DOI: 10.1016/j.jceh.2012.05.003]
- 12 Garrison RN, Cryer HM, Howard DA, Polk HC. Clarification of risk factors for abdominal operations in patients with hepatic cirrhosis. *Ann Surg* 1984; 199: 648-655 [PMID: 6732310]
- 13 Mansour A, Watson W, Shayani V, Pickleman J. Abdominal operations in patients with cirrhosis: still a major surgical challenge. *Surgery* 1997; 122: 730-735; discussion 735-736 [PMID: 9347849 DOI: 10.1016/S0039-6060(97)90080-5]
- 14 Friedman LS. Surgery in the patient with liver disease. *Trans Am Clin Climatol Assoc* 2010; **121**: 192-204; discussion 205 [PMID: 20697561]
- 15 Møller S, Henriksen JH, Bendtsen F. Extrahepatic complications to cirrhosis and portal hypertension: haemodynamic and homeostatic aspects. *World J Gastroenterol* 2014; 20: 15499-15517 [PMID: 25400435 DOI: 10.3748/wjg.v20.i42.15499]
- 16 Sun YW, Chen W, Luo M, Hua R, Liu W, Huo YM, Wu ZY, Cao H. Evaluation of surgical procedure selection based on intraoperative free portal pressure measurement in patients with portal hypertension. *Hepatobiliary Pancreat Dis Int* 2010; 9: 269-274 [PMID: 20525554]
- 17 O'Leary JG, Friedman LS. Predicting surgical risk in patients with cirrhosis: from art to science. *Gastroenterology* 2007; 132: 1609-1611 [PMID: 17428482 DOI: 10.1053/j.gastro.2007.03.016]
- 18 Befeler AS, Palmer DE, Hoffman M, Longo W, Solomon H, Di Bisceglie AM. The safety of intra-abdominal surgery in patients with cirrhosis: model for end-stage liver disease score is superior to Child-Turcotte-Pugh classification in predicting outcome. *Arch Surg* 2005; 140: 650-654; discussion 655 [PMID: 16027329 DOI: 10.1001/archsurg.140.7.650]
- 19 Weis F, Kilger E, Beiras-Fernandez A, Hinske CL, Nassau K, Adnan L, Vicol C, Kur F, Möhnle P. Indocyanine green clearance as an outcome prediction tool in cardiac surgery: a prospective study. *J Crit Care* 2014; **29**: 224-229 [PMID: 24332990 DOI: 10.1016/j.jerc.2013.10.023]
- 20 Sander M, Spies CD, Berger K, Schröder T, Grubitzsch H, Wernecke KD, von Heymann C. Perioperative indocyanine green clearance is predictive for prolonged intensive care unit stay after coronary artery bypass grafting--an observational study. *Crit Care* 2009; 13: R149 [PMID: 19747406 DOI: 10.1186/cc8045]

- 21 Imamura H, Sano K, Sugawara Y, Kokudo N, Makuuchi M. Assessment of hepatic reserve for indication of hepatic resection: decision tree incorporating indocyanine green test. *J Hepatobiliary Pancreat Surg* 2005; **12**: 16-22 [PMID: 15754094 DOI: 10.1007/ s00534-004-0965-9]
- 22 de Goede B, Klitsie PJ, Lange JF, Metselaar HJ, Kazemier G. Morbidity and mortality related to non-hepatic surgery in patients with liver cirrhosis: a systematic review. *Best Pract Res Clin Gastroenterol* 2012; 26: 47-59 [PMID: 22482525 DOI: 10.1016/ j.bpg.2012.01.010]
- 23 Hanje AJ, Patel T. Preoperative evaluation of patients with liver disease. *Nat Clin Pract Gastroenterol Hepatol* 2007; 4: 266-276 [PMID: 17476209 DOI: 10.1038/ncpgasthep0794]
- 24 **Peacock EE Jr**. Control of wound healing and scar formation in surgical patients. *Arch Surg* 1981; **116**: 1325-1329 [PMID: 7283705 DOI: 10.1001/archsurg.1981.01380220069011]
- 25 Strassburg CP, Manns MP. Treatment of autoimmune hepatitis. Semin Liver Dis 2009; 29: 273-285 [PMID: 19676000 DOI: 10.1055/s-0029-1233534]
- 26 Papastergiou V, Burroughs AK, Tsochatzis EA. Prognosis and treatment of patients with acute alcoholic hepatitis. *Expert Rev Gastroenterol Hepatol* 2014; 8: 471-486 [PMID: 24716632 DOI: 10.1586/17474124.2014.903800]
- Kleinegris MC, Bos MH, Roest M, Henskens Y, Ten Cate-Hoek A, Van Deursen C, Spronk HM, Reitsma PH, De Groot PG, Ten Cate H, Koek G. Cirrhosis patients have a coagulopathy that is associated with decreased clot formation capacity. *J Thromb Haemost* 2014; 12: 1647-1657 [PMID: 25142532 DOI: 10.1111/jth.12706]
- 28 Witters P, Freson K, Verslype C, Peerlinck K, Hoylaerts M, Nevens F, Van Geet C, Cassiman D. Review article: blood platelet number and function in chronic liver disease and cirrhosis. *Aliment Pharmacol Ther* 2008; 27: 1017-1029 [PMID: 18331464 DOI: 10.1111/j.1365-2036.2008.03674.x]
- 29 Violi F, Basili S, Raparelli V, Chowdary P, Gatt A, Burroughs AK. Patients with liver cirrhosis suffer from primary haemostatic defects? Fact or fiction? *J Hepatol* 2011; 55: 1415-1427 [PMID: 21718668 DOI: 10.1016/j.jhep.2011.06.008]
- 30 Samama CM, Djoudi R, Lecompte T, Nathan N, Schved JF; French Health Products Safety Agency (AFSSAPS) Expert Group. Perioperative platelet transfusion. Recommendations of the French Health Products Safety Agency (AFSSAPS) 2003. *Minerva Anestesiol* 2006; 72: 447-452 [PMID: 16682914]
- 31 Tripodi A, Caldwell SH, Hoffman M, Trotter JF, Sanyal AJ. Review article: the prothrombin time test as a measure of bleeding risk and prognosis in liver disease. *Aliment Pharmacol Ther* 2007; 26: 141-148 [PMID: 17593061 DOI: 10.1111/ j.1365-2036.2007.03369.x]
- 32 Tripodi A, Primignani M, Lemma L, Chantarangkul V, Dell'Era A, Iannuzzi F, Aghemo A, Mannucci PM. Detection of the imbalance of procoagulant versus anticoagulant factors in cirrhosis by a simple laboratory method. *Hepatology* 2010; **52**: 249-255 [PMID: 20578143 DOI: 10.1002/hep.23653]
- Amarapurkar PD, Amarapurkar DN. Management of coagulopathy in patients with decompensated liver cirrhosis. *Int J Hepatol* 2011; 2011: 695470 [PMID: 22164337 DOI: 10.4061/2011/695470]
- 34 Youssef WI, Salazar F, Dasarathy S, Beddow T, Mullen KD. Role of fresh frozen plasma infusion in correction of coagulopathy of chronic liver disease: a dual phase study. *Am J Gastroenterol* 2003; 98: 1391-1394 [PMID: 12818286 DOI: 10.1111/ j.1572-0241.2003.07467.x]
- 35 Shah NL, Intagliata NM, Northup PG, Argo CK, Caldwell SH. Procoagulant therapeutics in liver disease: a critique and clinical rationale. *Nat Rev Gastroenterol Hepatol* 2014; 11: 675-682 [PMID: 25023035 DOI: 10.1038/nrgastro.2014.121]
- 36 D'Albuquerque LA, de Miranda MP, Genzini T, Copstein JL, de Oliveira e Silva A. Laparoscopic cholecystectomy in cirrhotic patients. Surg Laparosc Endosc 1995; 5: 272-276 [PMID: 7551278]
- 37 **Cata JP**, Wang H, Gottumukkala V, Reuben J, Sessler DI. Inflammatory response, immunosuppression, and cancer recurrence



after perioperative blood transfusions. *Br J Anaesth* 2013; **110**: 690-701 [PMID: 23599512 DOI: 10.1093/bja/aet068]

- 38 Bonnel AR, Bunchorntavakul C, Reddy KR. Immune dysfunction and infections in patients with cirrhosis. *Clin Gastroenterol Hepatol* 2011; 9: 727-738 [PMID: 21397731 DOI: 10.1016/j.cgh.2011.02.031]
- 39 Sipeki N, Antal-Szalmas P, Lakatos PL, Papp M. Immune dysfunction in cirrhosis. *World J Gastroenterol* 2014; 20: 2564-2577 [PMID: 24627592 DOI: 10.3748/wjg.v20.i10.2564]
- 40 McGuinness J, Bouchier-Hayes D, Redmond JM. Understanding the inflammatory response to cardiac surgery. *Surgeon* 2008; 6: 162-171 [PMID: 18581753 DOI: 10.1016/S1479-666X(08)80113-8]
- 41 Eghtesad S, Poustchi H, Malekzadeh R. Malnutrition in liver cirrhosis: the influence of protein and sodium. *Middle East J Dig Dis* 2013; 5: 65-75 [PMID: 24829672]
- 42 Leide da Silva Nunes F, Calado Ferreira Pinheiro Gadelha P, Damasceno de Souza Costa M, Carolina Ribeiro de Amorim AC, Bezerra da Silva Mda G. Nutritional status and its impact on time and relocation in postoperative complications of abdominal patients undergoing surgery. *Nutr Hosp* 2014; **30**: 629-635 [PMID: 25238841 DOI: 10.3305/nh.2014.30.3.7628]
- 43 Burden S, Todd C, Hill J, Lal S. Pre-operative nutrition support in patients undergoing gastrointestinal surgery. *Cochrane Database Syst Rev* 2012; 11: CD008879 [PMID: 23152265 DOI: 10.1002/14651858.CD008879.pub2]
- 44 Roongpisuthipong C, Sobhonslidsuk A, Nantiruj K, Songchitsomboon S. Nutritional assessment in various stages of liver cirrhosis. *Nutrition* 2001; 17: 761-765 [PMID: 11527674 DOI: 10.1016/S0899-9007(01)00626-8]
- 45 Fuhrman MP, Charney P, Mueller CM. Hepatic proteins and nutrition assessment. J Am Diet Assoc 2004; 104: 1258-1264 [PMID: 15281044 DOI: 10.1016/j.jada.2004.05.213]
- 46 Lai CC, You JF, Yeh CY, Chen JS, Tang R, Wang JY, Chin CC. Low preoperative serum albumin in colon cancer: a risk factor for poor outcome. *Int J Colorectal Dis* 2011; 26: 473-481 [PMID: 21190025 DOI: 10.1007/s00384-010-1113-4]
- 47 Alkhouri N, Tamimi TA, Yerian L, Lopez R, Zein NN, Feldstein AE. The inflamed liver and atherosclerosis: a link between histologic severity of nonalcoholic fatty liver disease and increased cardiovascular risk. *Dig Dis Sci* 2010; 55: 2644-2650 [PMID: 19960252 DOI: 10.1007/s10620-009-1075-y]
- 48 Vernon G, Baranova A, Younossi ZM. Systematic review: the epidemiology and natural history of non-alcoholic fatty liver disease and non-alcoholic steatohepatitis in adults. *Aliment Pharmacol Ther* 2011; 34: 274-285 [PMID: 21623852 DOI: 10.1111/j.1365-2036.2011.04724.x]
- 49 Zardi EM, Abbate A, Zardi DM, Dobrina A, Margiotta D, Van Tassell BW, Afeltra A, Sanyal AJ. Cirrhotic cardiomyopathy. *J Am Coll Cardiol* 2010; 56: 539-549 [PMID: 20688208 DOI: 10.1016/ j.jacc.2009.12.075]
- 50 Theocharidou E, Krag A, Bendtsen F, Møller S, Burroughs AK. Cardiac dysfunction in cirrhosis - does adrenal function play a role? A hypothesis. *Liver Int* 2012; 32: 1327-1332 [PMID: 22292920 DOI: 10.1111/j.1478-3231.2011.02751.x]
- 51 Bagshaw SM, Bellomo R. The influence of volume management on outcome. *Curr Opin Crit Care* 2007; 13: 541-548 [PMID: 17762233 DOI: 10.1097/MCC.0b013e3282e2a978]
- 52 Testani JM, Khera AV, St John Sutton MG, Keane MG, Wiegers SE, Shannon RP, Kirkpatrick JN. Effect of right ventricular function and venous congestion on cardiorenal interactions during the treatment of decompensated heart failure. *Am J Cardiol* 2010; 105: 511-516 [PMID: 20152246 DOI: 10.1016/j.amjcard.2009.10.020]
- 53 Carl DE, Sanyal A. The management of hepatorenal syndrome. *Minerva Gastroenterol Dietol* 2009; 55: 207-226 [PMID: 19305378]
- 54 Lopez-Delgado JC, Esteve F, Torrado H, Rodríguez-Castro D, Carrio ML, Farrero E, Javierre C, Ventura JL, Manez R. Influence of acute kidney injury on short- and long-term outcomes in patients undergoing cardiac surgery: risk factors and prognostic value of a modified RIFLE classification. *Crit Care* 2013; **17**: R293 [PMID: 24330769 DOI: 10.1186/cc13159]
- 55 Lopez-Delgado JC, Esteve F, Javierre C, Torrado H, Carrio ML,

Rodríguez-Castro D, Farrero E, Lluís Ventura J, Manez R. Predictors of long-term mortality in patients with cirrhosis undergoing cardiac surgery. *J Cardiovasc Surg* (Torino) 2015; **56**: 647-654 [PMID: 24670881]

- 56 Ziser A, Plevak DJ, Wiesner RH, Rakela J, Offord KP, Brown DL. Morbidity and mortality in cirrhotic patients undergoing anesthesia and surgery. *Anesthesiology* 1999; 90: 42-53 [PMID: 9915311]
- 57 Feltracco P, Carollo C, Barbieri S, Pettenuzzo T, Ori C. Early respiratory complications after liver transplantation. *World J Gastroenterol* 2013; 19: 9271-9281 [PMID: 24409054 DOI: 10.3748/ wjg.v19.i48.9271]
- 58 Hemprich U, Papadakos PJ, Lachmann B. Respiratory failure and hypoxemia in the cirrhotic patient including hepatopulmonary syndrome. *Curr Opin Anaesthesiol* 2010; 23: 133-138 [PMID: 20019600 DOI: 10.1097/ACO.0b013e328335f024]
- 59 Huffmyer JL, Nemergut EC. Respiratory dysfunction and pulmonary disease in cirrhosis and other hepatic disorders. *Respir Care* 2007; 52: 1030-1036 [PMID: 17650360]
- 60 John S, Thuluvath PJ. Hyponatremia in cirrhosis: pathophysiology and management. *World J Gastroenterol* 2015; 21: 3197-3205 [PMID: 25805925 DOI: 10.3748/wjg.v21.i11.3197]
- 61 Azoulay D, Buabse F, Damiano I, Smail A, Ichai P, Dannaoui M, Castaing D, Bismuth H. Neoadjuvant transjugular intrahepatic portosystemic shunt: a solution for extrahepatic abdominal operation in cirrhotic patients with severe portal hypertension. J Am Coll Surg 2001; 193: 46-51 [PMID: 11442253 DOI: 10.1016/ S1072-7515(01)00911-5]
- 62 Gil A, Martínez-Regueira F, Hernández-Lizoain JL, Pardo F, Olea JM, Bastarrika G, Cienfuegos JA, Bilbao JI. The role of transjugular intrahepatic portosystemic shunt prior to abdominal tumoral surgery in cirrhotic patients with portal hypertension. *Eur J Surg Oncol* 2004; **30**: 46-52 [PMID: 14736522 DOI: 10.1016/ j.ejso.2003.10.014]
- 63 Vinet E, Perreault P, Bouchard L, Bernard D, Wassef R, Richard C, Létourneau R, Pomier-Layrargues G. Transjugular intrahepatic portosystemic shunt before abdominal surgery in cirrhotic patients: a retrospective, comparative study. *Can J Gastroenterol* 2006; 20: 401-404 [PMID: 16779457]
- 64 Kiamanesh D, Rumley J, Moitra VK. Monitoring and managing hepatic disease in anaesthesia. *Br J Anaesth* 2013; 111 Suppl 1: i50-i61 [PMID: 24335399 DOI: 10.1093/bja/aet378]
- 65 Friedman LS. The risk of surgery in patients with liver disease. *Hepatology* 1999; 29: 1617-1623 [PMID: 10347099 DOI: 10.1002/ hep.510290639]
- 66 Rahman S, Mallett SV. Cirrhotic cardiomyopathy: Implications for the perioperative management of liver transplant patients. *World J Hepatol* 2015; 7: 507-520 [PMID: 25848474 DOI: 10.4254/wjh. v7.i3.507]
- 67 Abid S, Ali S, Baig MA, Waheed AA. Is it time to replace propranolol with carvedilol for portal hypertension? *World J Gastrointest Endosc* 2015; 7: 532-539 [PMID: 25992192 DOI: 10.4253/wjge.v7.i5.532]
- 68 Biecker E. Diagnosis and therapy of ascites in liver cirrhosis. World J Gastroenterol 2011; 17: 1237-1248 [PMID: 21455322 DOI: 10.3748/wjg.v17.i10.1237]
- 69 de Mattos AA, Costabeber AM, Lionço LC, Tovo CV. Multiresistant bacteria in spontaneous bacterial peritonitis: a new step in management? *World J Gastroenterol* 2014; 20: 14079-14086 [PMID: 25339797 DOI: 10.3748/wjg.v20.i39.14079]
- 70 Gelman S. General anesthesia and hepatic circulation. Can J Physiol Pharmacol 1987; 65: 1762-1779 [PMID: 3319112 DOI: 10.1139/y87-276]
- Hoetzel A, Ryan H, Schmidt R. Anesthetic considerations for the patient with liver disease. *Curr Opin Anaesthesiol* 2012; 25: 340-347 [PMID: 22450699 DOI: 10.1097/ACO.0b013e3283532b02]
- 72 Mcclain RL, Ramakrishna H, Iii SA, Cartwright JA, Phar LG, Pai SL, Rodrigues ES, Martin AK, Shine TS. Anesthetic pharmacology and perioperative considerations for the end stage liver disease patient. *Curr Clin Pharmacol* 2015; 10: 35-46 [PMID: 24521189 DOI: 10.2174/1574884709666140212110036]
- 73 Mandell MS, Durham J, Kumpe D, Trotter JF, Everson GT,



#### Lopez-Delgado JC et al. Cirrhosis influence in non-hepatic abdominal surgery

Niemann CU. The effects of desflurane and propofol on portosystemic pressure in patients with portal hypertension. *Anesth Analg* 2003; **97**: 1573-1577 [PMID: 14633521 DOI: 10.1213/01. ANE.0000090741.63156.1B]

- 74 Lebrault C, Berger JL, D'Hollander AA, Gomeni R, Henzel D, Duvaldestin P. Pharmacokinetics and pharmacodynamics of vecuronium (ORG NC 45) in patients with cirrhosis. *Anesthesiology* 1985; 62: 601-605 [PMID: 2859815]
- 75 Khuenl-Brady KS, Sparr H. Clinical pharmacokinetics of rocuronium bromide. *Clin Pharmacokinet* 1996; **31**: 174-183 [PMID: 8877248 DOI: 10.2165/00003088-199631030-00002]
- 76 Nimmo SM, Harrington LS. What is the role of epidural analgesia in abdominal surgery? *Contin Educ Anaesth Crit Care Pain* 2014; 14: 224-229 [DOI: 10.1093/bjaceaccp/mkt062]
- 77 Friel CM, Stack J, Forse A, Babineau TJ. Laparoscopic cholecystectomy in patients with hepatic cirrhosis: a five-year experience. *J Gastrointest Surg* 1999; 3: 286-291 [PMID: 10481121 DOI: 10.1016/S1091-255X(99)80070-5]
- 78 Schiff J, Misra M, Rendon G, Rothschild J, Schwaitzberg S. Laparoscopic cholecystectomy in cirrhotic patients. *Surg Endosc* 2005; 19: 1278-1281 [PMID: 16021366 DOI: 10.1007/s00464-004-8823-z]
- 79 Clark JR, Wills VL, Hunt DR. Cirrhosis and laparoscopic cholecystectomy. Surg Laparosc Endosc Percutan Tech 2001; 11: 165-169 [PMID: 11444745]
- 80 Palanivelu C, Rajan PS, Jani K, Shetty AR, Sendhilkumar K, Senthilnathan P, Parthasarthi R. Laparoscopic cholecystectomy in cirrhotic patients: the role of subtotal cholecystectomy and its variants. J Am Coll Surg 2006; 203: 145-151 [PMID: 16864026 DOI: 10.1016/j.jamcollsurg.2006.04.019]
- 81 Belli G, D'Agostino A, Fantini C, Cioffi L, Belli A, Russolillo N, Langella S. Laparoscopic incisional and umbilical hernia repair in cirrhotic patients. *Surg Laparosc Endosc Percutan Tech* 2006; 16: 330-333 [PMID: 17057574 DOI: 10.1097/01. sle.0000213745.15773.c1]
- 82 McAlister V. Management of umbilical hernia in patients with advanced liver disease. *Liver Transpl* 2003; 9: 623-625 [PMID: 12783406 DOI: 10.1053/jlts.2003.50121]
- 83 Kandil T, El Nakeeb A, El Hefnawy E. Comparative study between clipless laparoscopic cholecystectomy by harmonic scalpel versus conventional method: a prospective randomized study. J Gastrointest Surg 2010; 14: 323-328 [PMID: 19882194 DOI: 10.1007/s11605-009-1039-8]
- 84 Bessa SS, Abdel-Razek AH, Sharaan MA, Bassiouni AE, El-Khishen MA, El-Kayal el-SA. Laparoscopic cholecystectomy in cirrhotics: a prospective randomized study comparing the conventional diathermy and the harmonic scalpel for gallbladder dissection. J Laparoendosc Adv Surg Tech A 2011; 21: 1-5 [PMID: 21166564 DOI: 10.1089/lap.2010.0255]
- 85 Ji W, Li LT, Wang ZM, Quan ZF, Chen XR, Li JS. A randomized controlled trial of laparoscopic versus open cholecystectomy in patients with cirrhotic portal hypertension. *World J Gastroenterol* 2005; 11: 2513-2517 [PMID: 15832428 DOI: 10.3748/wjg.v11. i16.2513]
- Puggioni A, Wong LL. A metaanalysis of laparoscopic cholecystectomy in patients with cirrhosis. *J Am Coll Surg* 2003; 197: 921-926 [PMID: 14644279 DOI: 10.1016/j.jamcollsurg.2003.08.0 11]
- 87 Poulsen TL, Thulstrup AM, Sørensen HT, Vilstrup H. Appendicectomy and perioperative mortality in patients with liver cirrhosis. *Br J Surg* 2000; 87: 1664-1665 [PMID: 11122181 DOI: 10.1046/j.1365-2168.2000.01599.x]
- 88 Cobb WS, Heniford BT, Burns JM, Carbonell AM, Matthews BD, Kercher KW. Cirrhosis is not a contraindication to laparoscopic surgery. *Surg Endosc* 2005; 19: 418-423 [PMID: 15624057 DOI: 10.1007/s00464-004-8722-3]
- 89 Tsugawa K, Koyanagi N, Hashizume M, Tomikawa M, Ayukawa K, Akahoshi K, Sugimachi K. A comparison of an open and laparoscopic appendectomy for patients with liver cirrhosis. Surg Laparosc Endosc Percutan Tech 2001; 11: 189-194 [PMID:

11444750]

- 90 Gentileschi P, Rossi P, Manzelli A, Lirosi F, Susanna F, Stolfi VM, Spina C, Gaspari AL. Laparoscopic suture repair of a perforated gastric ulcer in a severely cirrhotic patient with portal hypertension: first case report. JSLS 2003; 7: 377-382 [PMID: 14626407]
- 91 El-Awadi S, El-Nakeeb A, Youssef T, Fikry A, Abd El-Hamed TM, Ghazy H, Foda E, Farid M. Laparoscopic versus open cholecystectomy in cirrhotic patients: a prospective randomized study. *Int J Surg* 2009; 7: 66-69 [PMID: 19028148 DOI: 10.1016/ j.ijsu.2008.10.013]
- 92 Hamad MA, Thabet M, Badawy A, Mourad F, Abdel-Salam M, Abdel-Rahman Mel-T, Hafez MZ, Sherif T. Laparoscopic versus open cholecystectomy in patients with liver cirrhosis: a prospective, randomized study. *J Laparoendosc Adv Surg Tech A* 2010; 20: 405-409 [PMID: 20518688 DOI: 10.1089/lap.2009.0476]
- 93 Fernandes NF, Schwesinger WH, Hilsenbeck SG, Gross GW, Bay MK, Sirinek KR, Schenker S. Laparoscopic cholecystectomy and cirrhosis: a case-control study of outcomes. *Liver Transpl* 2000; 6: 340-344 [PMID: 10827236 DOI: 10.1053/lv.2000.6353]
- 94 Yeh CN, Chen MF, Jan YY. Laparoscopic cholecystectomy in 226 cirrhotic patients. Experience of a single center in Taiwan. Surg Endosc 2002; 16: 1583-1587 [PMID: 12085147 DOI: 10.1007/ s00464-002-9026-0]
- 95 Curro G, Baccarani U, Adani G, Cucinotta E. Laparoscopic cholecystectomy in patients with mild cirrhosis and symptomatic cholelithiasis. *Transplant Proc* 2007; **39**: 1471-1473 [PMID: 17580164 DOI: 10.1016/j.transproceed.2007.01.086]
- 96 Cucinotta E, Lazzara S, Melita G. Laparoscopic cholecystectomy in cirrhotic patients. *Surg Endosc* 2003; 17: 1958-1960 [PMID: 14577023 DOI: 10.1007/s00464-002-8852-4]
- 97 Morino M, Cavuoti G, Miglietta C, Giraudo G, Simone P. Laparoscopic cholecystectomy in cirrhosis: contraindication or privileged indication? *Surg Laparosc Endosc Percutan Tech* 2000; 10: 360-363 [PMID: 11147909]
- 98 Nguyen GC, Correia AJ, Thuluvath PJ. The impact of cirrhosis and portal hypertension on mortality following colorectal surgery: a nationwide, population-based study. *Dis Colon Rectum* 2009; 52: 1367-1374 [PMID: 19617746 DOI: 10.1007/ DCR.0b013e3181a80dca]
- 99 Jeong SH, Ahn HS, Yoo MW, Cho JJ, Lee HJ, Kim HH, Lee KU, Yang HK. Increased morbidity rates in patients with heart disease or chronic liver disease following radical gastric surgery. *J Surg Oncol* 2010; 101: 200-204 [PMID: 20063368 DOI: 10.1002/ jso.21467]
- 100 Warnick P, Mai I, Klein F, Andreou A, Bahra M, Neuhaus P, Glanemann M. Safety of pancreatic surgery in patients with simultaneous liver cirrhosis: a single center experience. *Pancreatology* 2011; **11**: 24-29 [PMID: 21336005 DOI: 10.1159/000323961]
- 101 Gray SH, Vick CC, Graham LA, Finan KR, Neumayer LA, Hawn MT. Umbilical herniorrhapy in cirrhosis: improved outcomes with elective repair. J Gastrointest Surg 2008; 12: 675-681 [PMID: 18270782 DOI: 10.1007/s11605-008-0496-9]
- 102 Eker HH, van Ramshorst GH, de Goede B, Tilanus HW, Metselaar HJ, de Man RA, Lange JF, Kazemier G. A prospective study on elective umbilical hernia repair in patients with liver cirrhosis and ascites. *Surgery* 2011; 150: 542-546 [PMID: 21621237 DOI: 10.1016/j.surg.2011.02.026]
- 103 Carbonell AM, Wolfe LG, DeMaria EJ. Poor outcomes in cirrhosis-associated hernia repair: a nationwide cohort study of 32,033 patients. *Hernia* 2005; 9: 353-357 [PMID: 16132187 DOI: 10.1007/s10029-005-0022-x]
- 104 Marsman HA, Heisterkamp J, Halm JA, Tilanus HW, Metselaar HJ, Kazemier G. Management in patients with liver cirrhosis and an umbilical hernia. *Surgery* 2007; 142: 372-375 [PMID: 17723889 DOI: 10.1016/j.surg.2007.05.006]
- 105 McKay A, Dixon E, Bathe O, Sutherland F. Umbilical hernia repair in the presence of cirrhosis and ascites: results of a survey and review of the literature. *Hernia* 2009; 13: 461-468 [PMID: 19652907 DOI: 10.1007/s10029-009-0535-9]
- 106 Ammar SA. Management of complicated umbilical hernias in

cirrhotic patients using permanent mesh: randomized clinical trial. *Hernia* 2010; **14**: 35-38 [PMID: 19727551 DOI: 10.1007/ s10029-009-0556-4]

- 107 Oh HK, Kim H, Ryoo S, Choe EK, Park KJ. Inguinal hernia repair in patients with cirrhosis is not associated with increased risk of complications and recurrence. *World J Surg* 2011; 35: 1229-1233; discussion 1234 [PMID: 21365342 DOI: 10.1007/ s00268-011-1007-9]
- 108 Patti R, Almasio PL, Buscemi S, Famà F, Craxì A, Di Vita G. Inguinal hernioplasty improves the quality of life in patients with cirrhosis. *Am J Surg* 2008; **196**: 373-378 [PMID: 18639226 DOI: 10.1016/j.amjsurg.2008.02.007]
- 109 Hansen JB, Thulstrup AM, Vilstup H, Sørensen HT. Danish nationwide cohort study of postoperative death in patients with liver cirrhosis undergoing hernia repair. *Br J Surg* 2002; **89**: 805-806 [PMID: 12027997 DOI: 10.1046/j.1365-2168.2002.02114.x]
- 110 Belghiti J, Durand F. Abdominal wall hernias in the setting of

cirrhosis. Semin Liver Dis 1997; **17**: 219-226 [PMID: 9308126 DOI: 10.1055/s-2007-1007199]

- 111 Chatzizacharias NA, Bradley JA, Harper S, Butler A, Jah A, Huguet E, Praseedom RK, Allison M, Gibbs P. Successful surgical management of ruptured umbilical hernias in cirrhotic patients. *World J Gastroenterol* 2015; 21: 3109-3113 [PMID: 25780312 DOI: 10.3748/wjg.v21.i10.3109]
- 112 Senousy BE, Draganov PV. Evaluation and management of patients with refractory ascites. *World J Gastroenterol* 2009; 15: 67-80 [PMID: 19115470 DOI: 10.3748/wjg.15.67]
- 113 Nusrat S, Khan MS, Fazili J, Madhoun MF. Cirrhosis and its complications: evidence based treatment. *World J Gastroenterol* 2014; 20: 5442-5460 [PMID: 24833875 DOI: 10.3748/wjg.v20. i18.5442]
- 114 Ojeda A, Moreno LA. Pain management in patients with liver cirrhosis. *Gastroenterol Hepatol* 2014; 37: 35-45 [PMID: 24309482 DOI: 10.1016/j.gastrohep.2013.05.007]

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