

Acute cholecystitis: When to operate and how to do it safely

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I would like to thank the AAST and President Cioffi for the great honor to present this Master Surgeon Lecture. On this date, I must acknowledge that it is September 11, and we need to take a moment of silence for our fallen colleagues and countrymen on that day.

My topic today is, “Acute Cholecystitis: When to Operate and How to Do It Safely.” The obvious question is why did I select such a mundane topic? It is estimated that 30% to 49% of surgeons will produce a bile duct injury during their careers. This event is difficult both for the patient and the surgeon. The premise of my talk is that **nearly all bile duct injuries during cholecystectomy are avoidable**. Approximately 700,000 cholecystectomies are performed per year in the United States, with an estimated incidence of bile duct injury in **0.5%** (3,500 patients). When laparoscopic cholecystectomy was initially introduced, bile duct injury was four times more frequent than for open cholecystectomy. Current estimate is that the **incidence of bile duct injury** remains **twice** as frequent with **laparoscopic** versus **open** cholecystectomy.^{1–29} A population-based study from Sweden, reviewing 153,000 cholecystectomies from 1987 through 2002, showed a slight increase in the incidence of bile duct injury despite decades of experience with laparoscopic cholecystectomy (0.32–0.47%).²⁶ Similarly, the incidence of bile duct injury in Japan is unchanged from 1990 to 2009 (0.66–0.62%).²⁹ Thus, laparoscopic cholecystectomy is clearly an operation that we have not perfected, despite how often it is performed.

The goals in today’s talk are as follows:

- Discuss the timing of operation for cholecystitis
- Discuss factors that predict the difficult cholecystectomy
- Discuss the role of percutaneous cholecystostomy in the management of acute cholecystitis
- Discuss how to minimize the risk of bile duct or vascular injury during cholecystectomy
- Discuss techniques and tricks for the difficult cholecystectomy, both open and laparoscopic
- Discuss what to do once an injury has been recognized.

Timing of Operation for Acute Cholecystitis

Indications listed by SAGES [Society of American Gastrointestinal and Endoscopic Surgeons] for laparoscopic cholecystectomy include **symptomatic** cholelithiasis, **biliary dyskinesia**, **acute cholecystitis**, and **biliary pancreatitis**.²⁸ **Twenty percent** of cholecystectomies are performed for **acute cholecystitis**. The Tokyo guidelines for the diagnosis of acute cholecystitis are shown in Table 1.^{29–41} **Asymptomatic** gallstones are generally **not** considered an **indication** for laparoscopic cholecystectomy. The first **question** to address is whether cholecystectomy should be performed during the **index hospitalization** for acute cholecystitis **or** the patient treated with **antibiotics** and **discharged** for **delayed** cholecystectomy, usually **6 weeks to 12 weeks after** the hospitalization. A series using the national Medicare sample claims data on 29,818 patients older than 65 years hospitalized for acute cholecystitis from 1996 to 2005 demonstrated that 75% of patients underwent cholecystectomy during that admission.⁴² **Median** time to operation was **1 day**, with **conversion** from **laparoscopic** to **open** cholecystectomy in **29%** of patients. Percutaneous cholecystostomy was applied in only 0.5% of patients. Thus, 25% of patients did not undergo

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TABLE 1. Tokyo Guidelines (TG13) Diagnostic Criteria for Acute Cholecystitis^{29–41}

A. Local signs of inflammation, etc.
(1) Murphy's sign, (2) RUQ mass/pain/tenderness
B. Systemic signs of inflammation, etc.
(1) Fever, (2) elevated CRP, (3) elevated WBC count
C. Imaging findings
Imaging findings characteristic of acute cholecystitis
Suspected diagnosis: one item in A + one item in B
Definite diagnosis: one item in A + one item in B + C
Acute hepatitis, other acute abdominal diseases, and chronic cholecystitis should be excluded.
CRP, C-reactive protein; RUQ, right upper abdominal quadrant; WBC, white blood cell.

cholecystectomy at the initial admission. The lack of cholecystectomy resulted in 38% gallstone-related admissions during the next 2 years (occurred in only 4% of the patients who had undergone cholecystectomy). Thus, it was concluded that laparoscopic/open cholecystectomy for acute cholecystitis in elderly patients should be performed during initial hospitalization.

In a population-based study from Ontario, 25,397 adult patients admitted from 2004 to 2011 with the first episode of acute cholecystitis were reviewed.^{7,13} Median follow-up was 3.4 years. Fifty-nine percent of patients underwent cholecystectomy during the index admission; 41% (10,304 patients) were discharged without cholecystectomy. Of the patients discharged without cholecystectomy, the incidence of gallstone-related event after discharge was 14% at 6 weeks, 19% at 12 weeks, and 29% at 1 year. Importantly, of these events, 30% were for biliary tract obstruction or pancreatitis, significant complications of cholelithiasis. Interestingly, these events were more frequent in patients aged 18 years to 34 years. At 1 year, the incidence of recurrent biliary tract disease was 42% in patients 18 years to 34 years, 32% in patients age 50 years to 64 years, 27% in patients age 65 years to 79 years, and 24% in patients older than 80 years. The authors concluded that increased risk in younger patients with recurrent gallstone disease reinforced the value of early cholecystectomy.

The Cochrane review published in 2013 reviewed six trials with 488 patients.⁴³ Early cholecystectomy was defined as within 7 days of clinical presentation. Delayed cholecystectomy was defined as greater than 6 weeks. The authors concluded that there was no significant difference in the incidence of bile duct injury, similar rate of conversion from laparoscopic to open cholecystectomy, and obviously shorter stay in patients who underwent early cholecystectomy. However, this Cochrane review is underpowered to evaluate significant difference in bile duct injury. It is estimated to document a 50% difference (statistically significant, appropriately powered) in incidence of bile duct injury that 30,000 patients would need to be included. In addition, the authors concluded that “all trials were at high risk of bias and might have overestimated the benefits or underestimated the harms of either early laparoscopic cholecystectomy or delayed laparoscopic cholecystectomy. However, trials with high risk of bias indicate that early laparoscopic cholecystectomy during acute cholecystitis seems safe and may shorten total hospital stay.”

KEY CONCEPT: Cholecystectomy should be performed during the index hospitalization for acute cholecystitis, unless the patient is deemed a prohibitive operative risk.

The next issue to be addressed is at what time point during the initial hospitalization cholecystectomy should be performed. In an article presented at the AAST, using the American College of Surgeons' National Surgical Quality Improvement Program files from 2005 to 2010, emergency cholecystectomy for acute cholecystitis in 5,268 patients was evaluated.⁴⁴ The primary predictor variable was preoperative hospital length of stay, reported as 0, 1, 2, 3, or 4 to 7 days. In this study, 83% of the patients underwent cholecystectomy at Day 0 or 1. As shown in Table 2, morbidity and mortality increased significantly from Days 0 to 2 through Days 4 to 7. This was probably more a factor of the patient's comorbid disease than the operation itself. If we specifically look at the impact of early operation, the conversion rate significantly increased by 2 days (nearly doubled) and continued to increase daily. The operative time increased significantly with delay to cholecystectomy. Obviously, the length of stay was increased as the operation was delayed. The authors concluded that “patients hospitalized for 2 or more days preoperatively had longer operative times and were significantly more likely to undergo conversion to open cholecystectomy. Any delay in operation beyond the day of admission resulted in a significantly longer length of stay.”

A population-based study from the SALTS [Swiss Association of Laparoscopic and Thoracoscopic Surgery] reported 4,100 patients undergoing emergency laparoscopic cholecystectomy from 1995 to 2006.⁴⁵ They were grouped by day of admission defined as Days 0, 1, 2, 3, 4 or 5, or 6 or later. The median age in this study was 60 years. Conversion rate from laparoscopic to open cholecystectomy was 12% at Day 0 and increased to 28% at Day 6 or later. Postoperative complications increased from 5.7% to 13%, from Day 0 to Day 6. Need for reoperation tripled from Day 0 to Day 6, from 0.9% to 3%. Thus, the authors showed that delaying laparoscopic cholecystectomy for acute cholecystitis resulted in significantly higher conversion rates and complications. The authors stated that “this investigation provides compelling evidence that acute cholecystitis merits surgery within 48 hours of admission.”

In the study presented at the American Surgical Association recently, 35 centers from Germany and Slovenia reported a randomized prospective study evaluating early versus delayed cholecystectomy.⁴⁶ Early cholecystectomy was within 24 hours

TABLE 2. Analysis of the Timing of Cholecystectomy During Admission for Acute Cholecystitis⁴⁴

Time to cholecystectomy, d	0 d	1 day	2 d	3 d	4–7 d
Outcome variable, d					
30-d mortality, %	0.8	0.9	1.8*	2.0	5.3
30-d morbidity, %	6.0	7.6	12.7*	15.2	19.1
Conversion to open cholecystectomy, %	16.3	21.3	28.9*	30.9	37.0
Operative time, mean, min	82	87	89*	91	98
Total length of stay, median, d	1	3	4*	6	9

*Significantly different from Day 0.

TABLE 3. Tokyo Guidelines 2013 (TG13) Severity Grading for Acute Cholecystitis^{29–41}

Grade I (Mild) Acute Cholecystitis

Grade I is acute cholecystitis in a healthy patient with no organ dysfunction and mild inflammatory changes in the gallbladder, making cholecystectomy a safe and low-risk operative procedure.

Grade II (Moderate) Acute Cholecystitis

Associated with any one of the following conditions:

1. Elevated white blood cell count ($>18,000/\mu\text{L}$)
2. Palpable tender mass in the right upper abdominal quadrant
3. Duration of complaints of >72 h
4. Marked local inflammation (gangrenous cholecystitis, pericholecystic abscess, hepatic abscess, biliary peritonitis, emphysematous cholecystitis)

Grade III (Severe) Acute Cholecystitis

Associated with dysfunction of any one of the following organs/systems:

- | | |
|-------------------------------|---|
| 1. Cardiovascular dysfunction | Hypotension requiring treatment with dopamine $\geq 5 \mu\text{g/kg/min}$ or any dose of norepinephrine |
| 2. Neurologic dysfunction | Decreased level of consciousness |
| 3. Respiratory dysfunction | $\text{Pao}_2/\text{FIO}_2$ ratio < 300 |
| 4. Renal dysfunction | Oliguria, creatinine $> 2.0 \text{ mg/dL}$ |
| 5. Hepatic dysfunction | Prothrombin time/international normalized ratio > 1.5 |
| 6. Hematologic dysfunction | Platelet count $< 100,000/\mu\text{L}$ |

of admission, and late cholecystectomy was defined as Days 7 to 45. Six hundred eighteen adult patients were randomized. Morbidity was significantly different, 12% in early cholecystectomy versus 34% in late cholecystectomy. They noted no difference in conversion rate, 10% versus 12%. Hospital length of stay was significantly increased in those who underwent delayed cholecystectomy. The authors concluded that “immediate laparoscopic cholecystectomy should be the therapy of choice for acute cholecystitis in operable patients.”

An interesting study by Catani showed correlation between duration of symptoms, rather than hospitalization, and length of operative time.⁴⁷ They reported a linear relationship between timing of surgery relative to duration of symptoms and operative time. There was an inflection point at 60 hours. At this point, each hour delay in cholecystectomy doubled the time added to the operation compared with operation earlier than 60 hours.

Another population-based study from Ontario, looked at 22,202 patients admitted with acute cholecystitis and undergoing cholecystectomy from 2004 to 2011.^{7,13} Early cholecystectomy was within 7 days of admission and compared with delayed cholecystectomy. The primary goal of the study was determination of the incidence of bile duct injury. They reported a doubling of the incidence of bile duct injury in delayed versus early cholecystectomy, 0.53% versus 0.28%, respectively ($p = 0.025$). The relative risk ratio with an advantage to early cholecystectomy was 0.53 (95% confidence interval, 0.31–0.90). As stated by the authors, this is the first study with significant power to detect a difference in bile duct injury, showing a clear advantage to early surgery for acute cholecystitis.

KEY CONCEPT: For acute cholecystitis, laparoscopic cholecystectomy should be performed on the day of admission or Day 1, unless there are clear contraindications.

I do think it is important that this not be performed at 2:00 AM or 3:00 AM, when the surgical team may be distracted by other issues or incoming patients. The patient admitted late at night or early in the morning should be on the operating room schedule as the first case, when the team is fresh and ready to deal with a difficult cholecystectomy.

Next, we need to discuss the Tokyo guidelines. These are important contributions generated by two dozen international experts on cholecystitis and biliary tract disease.^{29–41} An entire issue of the *Journal of Hepato-Biliary-Pancreatic Surgery* was devoted to this in 2007. These guidelines have been updated with other articles in 2013 and 2014 (Table 3). The Tokyo guidelines stratified acute cholecystitis into mild cholecystitis (Grade 1), moderate cholecystitis (Grade 2), and severe cholecystitis (Grade 3). *Mild cholecystitis (Grade 1)* is defined as cholecystitis in a healthy patient with no organ dysfunction and only mild inflammatory changes in the gallbladder. *Moderate cholecystitis (Grade 2)* has evidence of local inflammatory response or complaints for more than 72 hours. *Severe cholecystitis (Grade 3)* is acute cholecystitis accompanied by any evidence of organ dysfunction. As shown in the flow chart, defining the grade of acute cholecystitis determines management (Fig. 1). The patient with mild cholecystitis, that is, without complicating factors, should undergo early laparoscopic cholecystectomy. Severe, Grade 3 acute cholecystitis is best served by urgent gallbladder drainage, usually percutaneously. Less well defined is the ideal treatment for patients with moderate acute cholecystitis, where either percutaneous

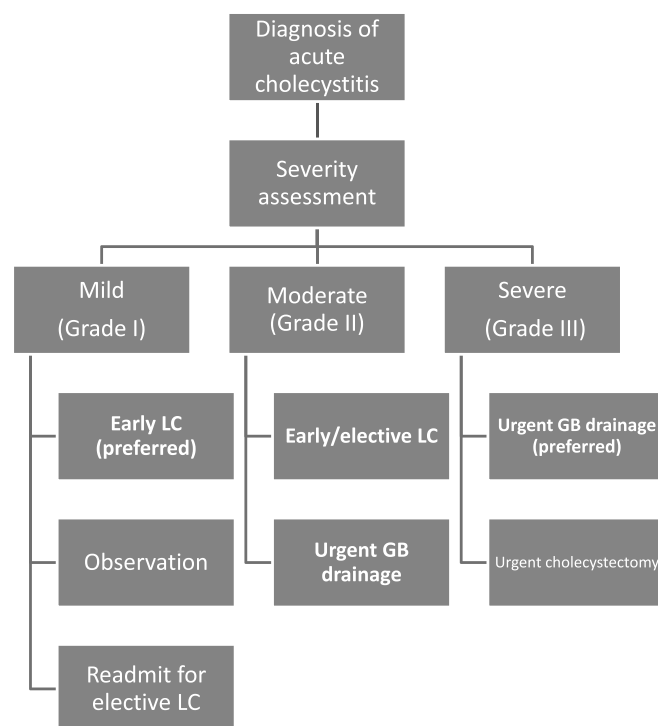


Figure 1. Tokyo guidelines for management of acute cholecystitis. (Source: Miura et al.³⁴ Reproduced with permission from John Wiley and Sons.)

drainage or laparoscopic cholecystectomy is appropriate based on a combination of factors. In patients with an elevated white blood cell count, palpable mass in the right upper quadrant, or signs of significant local inflammation, percutaneous drainage as an acute treatment followed by delayed cholecystectomy may be the safest option. The management of a patient who is classified as Grade 2 solely based on the duration of complaints for more than 72 hours is a more difficult decision. Often, cholecystectomy in such a patient is straightforward. At other times, acute inflammation and scarring are encountered, and the operation is difficult. This is an issue where we do not have a clear answer. Several authors have recommended that during the index hospitalization, unless there are clear reasons otherwise, any patient with acute cholecystitis should undergo operation, despite the duration of symptoms.^{43,48,49} However, they do concede that the surgeon must accept a longer and more difficult operation, and the skill set of the surgeon must be considered as well.

Antibiotics in Acute Cholecystitis

There is a relative paucity of high-quality studies examining the use of antibiotics in acute cholecystitis. Positive bile cultures, however, correlate with progression of cholecystitis to a more severe form,^{50–52} so the decision to begin antibiotics should be made shortly after the diagnosis has been established. According to the Tokyo guidelines, antibiotics are not necessary in patients with minimal abdominal pain and mild inflammatory findings.^{31,37} In these patients, who may be experiencing biliary colic as opposed to true acute cholecystitis, nonsteroidals may prevent progression to acute cholecystitis and may improve gallbladder function.⁵³ For the vast majority of patients, however, antibiotics should be started promptly. According to the Surgical Infection Society and Infectious Diseases Society of America guidelines,⁵⁴ mild cases of acute cholecystitis can be adequately treated with a first- (cefazolin), second- (cefuroxime), or third- (ceftriaxone) generation cephalosporin. Antibiotics should be discontinued 24 hours after cholecystectomy unless infection has spread outside the gallbladder wall.^{31,54,55} For complicated Grade II (pericholecystic abscess or perforated gallbladder) or Grade III cholecystitis, antibiotics should be continued until the patient is afebrile, has normalized white blood cell count, and is free of abdominal findings.³¹

For more severe cases or in those of advanced age or who are immunosuppressed, coverage should be broadened to include enterococci by using either an extended-spectrum penicillin or cephalosporin, a carbapenem, or a quinolone in combination with metronidazole. The Tokyo guidelines^{31,37} are similar except that they recommend a penicillin/ β -lactamase inhibitor in even mild (Grade I) cases because of the likelihood of β -lactamase production by intestinal organisms. Furthermore, these authors suggest that cultures of bile and the gallbladder wall “should be performed at all available opportunities, especially in severe cases” and that antibiotic coverage should be tailored depending on sensitivity results. Antibiotics should not be selected on the basis of biliary penetration because bile penetration by the antibiotic in the setting of obstruction (acute cholecystitis) essentially stops.⁵⁶

Percutaneous Cholecystostomy

The indications for percutaneous cholecystostomy are still not well defined.^{57–59} For the less common cases of Grade 3 acute cholecystitis, cholecystostomy insertion is recommended by the Tokyo guidelines.^{24,33,34} In addition, cholecystostomy is a safe option in patients with less severe cholecystitis who are considered poor surgical candidates or when a difficult dissection is encountered. Predictors of failure of antibiotic treatment alone and thus consideration for cholecystostomy tube include being older than 70 years, history of diabetes, and persistent leukocytosis of more than 15,000/ μ L at 48 hours.⁶⁰ Continued drainage must be established because aspiration alone is not as effective.⁶¹ Success rates of more than 80% are similar whether the procedure is performed for calculous or acalculous cholecystitis,^{62–65} and clinical improvement is generally seen within 72 hours.^{62,66,67} Mortality following the procedure is high (5–40%) but generally is related to the severity of the underlying disease process.^{62–65} As stated in a recent systematic review of percutaneous cholecystostomy, “there is no doubt that percutaneous cholecystostomy together with antibiotics can convert a septic cholecystitis into a non-septic condition.”⁶⁸ However, specific indications and criteria are still not well defined.^{58,69}

Of the patients who undergo percutaneous cholecystostomy and those whose tubes are removed, the need for delayed cholecystectomy remains controversial, with reports ranging from 0% to 87%.^{58,69–74} de Mestral et al.⁶⁹ reported in their population based study that approximately 40% will have recurrent biliary tract disease within 1 year following cholecystostomy. In their review of 47 articles and 1,724 patients, Winbladh et al.⁶⁸ observed that more than 40% of patients eventually underwent cholecystectomy. A prospective randomized trial (the CHOCOLATE Trial) in the Netherlands is underway, comparing early cholecystectomy with percutaneous cholecystostomy.⁷⁵

Factors Predicting the Difficult Cholecystectomy

Conversion from laparoscopic to open cholecystectomy should not be viewed as a failure. With a difficult cholecystectomy, it is critical to operate under the premise that bile duct injury is never an acceptable outcome and thus, if necessary, conversion is the safest option. Preoperative factors predict the patient for whom difficult cholecystectomy or need for a conversion can be expected. These include male patients, age greater than 70 years, inflammation, duration of symptoms for the acute episode, chronicity and duration of symptoms with recurrent disease, an impacted stone, gallbladder wall thickness, pericholecystic fluid, elevated white blood cell count, previous upper abdominal surgery, repeated bouts of cholecystitis, or a contracted gallbladder on imaging.^{24,27,49,60,76–84}

Why Do Bile Duct Injuries Still Occur?

KEY CONCEPT: We would agree that we each want bile duct injury to be on the list of complications that we never have.

So, why do bile duct injuries still occur? Common factors include anatomic variation, acute inflammation, chronic scarring, misperception, and error traps. Misperception by the surgeon of what he or she is seeing in the operative field is a

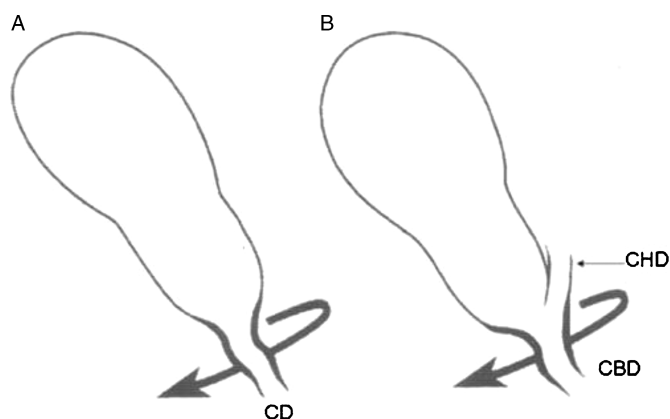


Figure 2. Infundibular view error trap. Dangerous anatomic variants of right posterior hepatic duct draining into the following: A, cystic duct; B, gall bladder neck; and C, common hepatic duct. (Source: Strasberg et al.¹⁹ Reproduced with permission from Elsevier, Inc.)

major factor in generating bile duct injury.^{6,8,17–22,80} In short, the surgeon sees what he or she believes and does not believe what he or she sees, and thus, the injury occurs.^{6,80} Along the same lines, Strasberg and colleagues^{17–22} discuss error traps. As noted by several authors, during the past two decades of laparoscopic cholecystectomy, the bile duct injuries seen may be less common but more severe.^{17–22,85} Strasberg and colleagues^{17–22} define an **error trap as an operative approach that works well in most circumstances but is prone to fail under certain circumstances**. Similar to the misperception issues, with an error trap, because the technique usually works, the surgeon develops confidence in it and fails to recognize when dangerous circumstances are present. The error traps that Strasberg and colleagues described are as follows:^{17–22}

1. The “infundibular view” error trap
2. Fundus down cholecystectomy in the face of severe inflammation
3. Failure to perceive the absence of an **aberrant right** hepatic duct on cholangiography (IOC). (I would add failure to

recognize an aberrant right hepatic duct or posterior right hepatic duct intraoperatively as well.)

4. Injury to the common bile duct in the case of a “parallel union” cystic duct.

The usual approach to the gallbladder is starting from the infundibulum and then working toward the fundus. It is taught that the taper between infundibulum and cystic duct identifies cystic duct. In a single view, this can be misleading, especially with any inflammation, and the common duct can be mistakenly divided, believing it is the cystic duct (“infundibular view error trap”) (Fig. 2). This produces the classic injury with resection of a portion of the common bile duct.

The error trap with an open, top-down cholecystectomy again is caused by what is normally safe, applied in a dangerous situation. Strasberg states that the worst injuries occur in those patients who undergo conversion from laparoscopic to open cholecystectomy, performed **top-down** because of marked inflammation and difficult dissection. This initially seems counterintuitive but will make sense as we explain it. The perceived, safe operative plane coming down the medial wall of the gallbladder is now obliterated by an inflammatory reaction, which incorporates the right-sided porta hepatis and the common bile duct. Thus, this injury is commonly associated with major biliary and vascular injury, at times requiring liver resection for the ischemic injury.

The variability of the right posterior hepatic duct includes drainage into the cystic duct, gallbladder neck, or common hepatic duct (Fig. 3).^{25,86,87} With the infundibular approach to the gallbladder, injury to such an aberrant posterior right hepatic duct is nearly unavoidable. However, with a top-down approach on the gallbladder, the aberrant right posterior hepatic duct can generally be seen and protected; leave a rim of infundibulum to protect the duct. In addition, this aberrant posterior right hepatic duct will often not be seen on an IOC because the cholangi catheter is introduced into the cystic duct below insertion of the aberrant duct.

If a posterior right hepatic duct is transected and not recognized, the clinical presentation is uncommon but classical. Generally, a clip is on the proximal duct, but the liver side of the duct is draining freely (Fig. 4). This case shows an IOC

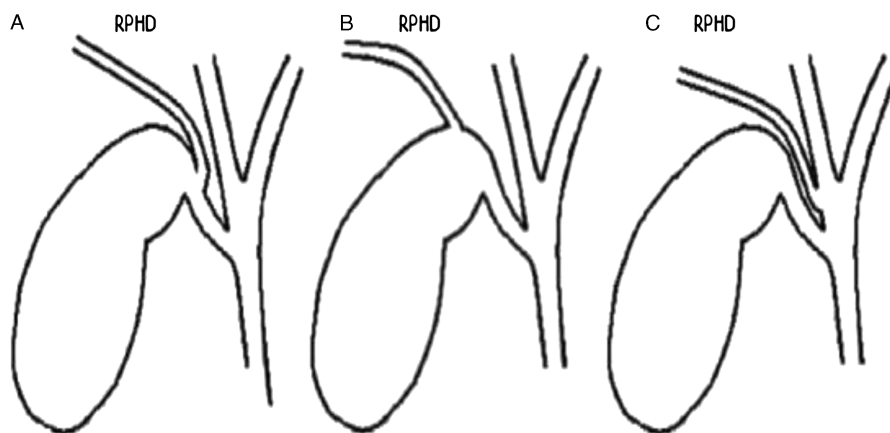


Figure 3. Common anomalies of the posterior right hepatic duct. (Source: Wojcicki et al.⁸⁶ Reproduced with permission from Baishideng Publishing Group Co.)

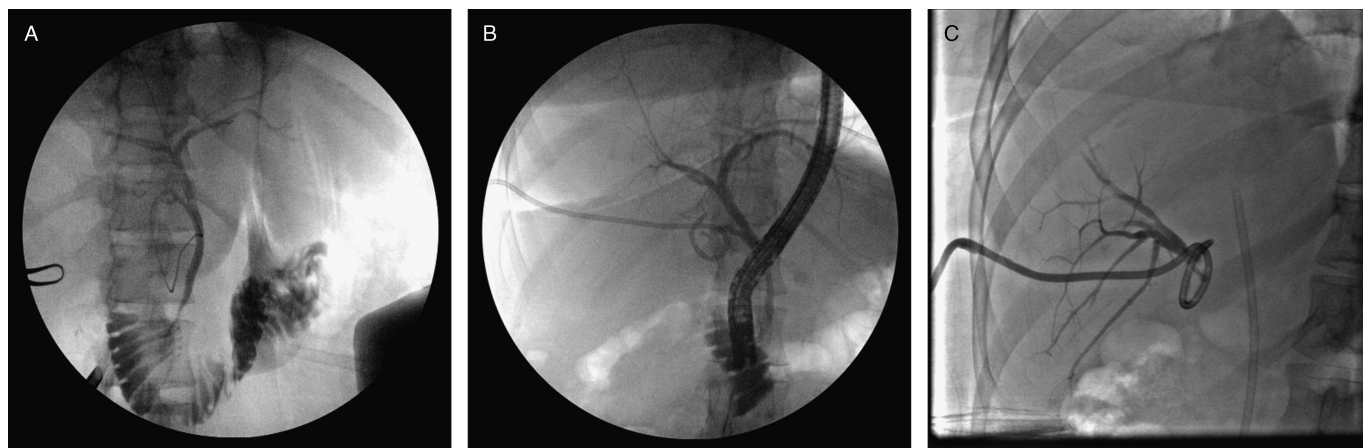


Figure 4. Studies from a patient with a transected posterior right hepatic duct. *A*, An IOC interpreted as normal. *B*, An ERCP a week postoperatively interpreted as normal. *C*, A sinogram showing filling of the posterior right lobe.

and endoscopic retrograde cholangiopancreatography (ERCP) a week later (for a bile leak), which are both interpreted as normal. Sometimes, what you do not see is as important as what you do see on these studies. Absent on both the IOC and ERCP is filling of the posterior right lobe. When contrast is injected through the drain as a sinogram, the transected right posterior sectoral duct fills (Fig. 4C). This requires either Roux-en-Y to the duct remnant or liver resection (as was performed in this case).

The most common configuration of the cystic duct joining the common duct is angular (75%). However, the parallel union occurs in 20%. Especially with any degree of inflammation, this fused cystic duct and common duct generate a situation where injury is more likely. Similarly, a spiral union between cystic duct and common duct can be misinterpreted.

Chronic scarring from recurrent or neglected bouts of cholecystitis is as dangerous as acute inflammation. This contracts all of the portal structures from the inflammatory response, thus obliterating the usual safe planes. This can be predicted based on preoperative history and imaging that shows a shrunken, contracted gallbladder. Cholecystectomy in these circumstances can be particularly difficult.

Cholecystectomy: How to Do It Safely

The essentials for safe laparoscopic cholecystectomy begin with a 30-degree or 45-degree high-definition laparoscope. Take full advantage of the angled scope, visualizing from different angles continuously as the operation proceeds. Hunter⁸⁸ describes many of these key principles nicely in his 1991 article. The assistant grasps the fundus cephalad and retracts this toward the patient's right shoulder. This reduces redundancy in the infundibulum and exposes the cystic duct. A second grasper retracts the infundibulum laterally to make the cystic duct perpendicular to the common bile duct and again separate the gallbladder from the common bile duct (Fig. 5). The key principles for safe laparoscopic cholecystectomy include the following:^{17–22,88}

- 30-degree or 45-degree high-definition laparoscope
- Cephalad traction on the dome of the gallbladder

- Lateral traction on the infundibulum
- Finding the gallbladder wall and staying on it
- Dissecting from above down to the neck
- Widely opening the hepatocystic triangle
- Moving the infundibulum back and forth (wave the flag), repeatedly looking at both sides of the gallbladder
- Critical view of safety
- Dividing the cystic duct as close to the gallbladder as possible
- Never dividing the cystic duct with any cauterizing instrument—if it turns out to be the common bile duct, the resulting ischemic injury will only lessen the chances for a good repair

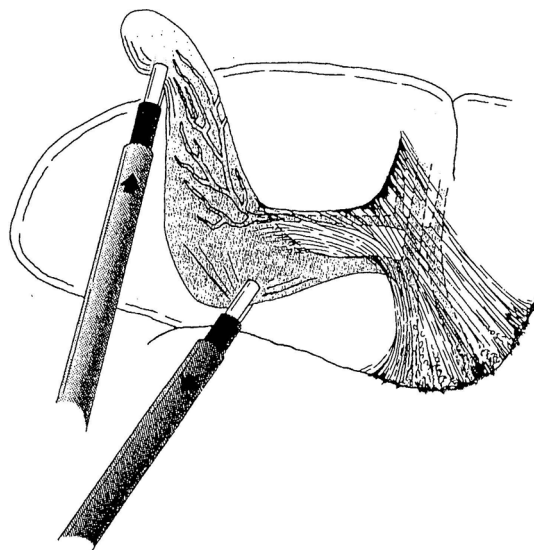


Figure 5. The assistant grasps the fundus cephalad and retracts this toward the patient's right shoulder. This reduces redundancy in the infundibulum and exposes the cystic duct. A second grasper retracts the infundibulum laterally to make the cystic duct perpendicular to the common bile duct, and again separate the gallbladder from the common bile duct. (Source: Hunter.⁸⁸ Reproduced with permission from Elsevier, Inc.)

KEY CONCEPT: Operative dissection technique versus method to identify anatomy.

Related but different principles include how we dissect the gallbladder and how we safely identify the anatomy. Dissection techniques include the *infundibular technique*, which is most commonly used; the *fundus first (top-down)*; and what we call the *semi-top-down technique*. The *infundibular technique* is how most of us have learned. As mentioned, this is a technique that works majority of the time but will fail in predictable circumstances, specifically anatomic variation or inflammation.

KEY CONCEPT: What is safest and best for an open procedure is safest and best for a laparoscopic procedure.

With infundibulum-first cholecystectomy, we violate this principle. Thus, it should not be a surprise at times that this generates problems. The fundus first (top-down) has been well described, mimicking what we do for open cholecystectomy.^{89–92} Certainly, with acute inflammation, this is the preferred approach. However, this can be awkward because of the floppiness of the gallbladder when it is fully detached from

the liver. Gently retracting the liver surface will generally stabilize this. On occasion, a liver retractor may be necessary.

The semi-top-down technique of laparoscopic cholecystectomy combines the advantages of both approaches and minimizes the disadvantages. Dissection is started higher on the gallbladder, above the infundibulum of the gallbladder (Fig. 6A–E). The peritoneum is scored circumferentially, lateral side first, coming across the peritoneum over the infundibulum of the gallbladder, then opening the peritoneum coming up the medial side of the gallbladder, being careful not to enter the cystic artery as you do so. Then, by rolling the gallbladder back and forth, the gallbladder can be largely detached from the liver, leaving only the fundus attached to again provide easy retraction. At this point and only at this point is the infundibulum and its junction with the cystic duct approached, thus generating a top-down approach to the cystic duct and cystic artery. When proceeding with the semi-top-down taking only tissues that you see through clearly, any structures that may be encountered such as an aberrant duct, right hepatic

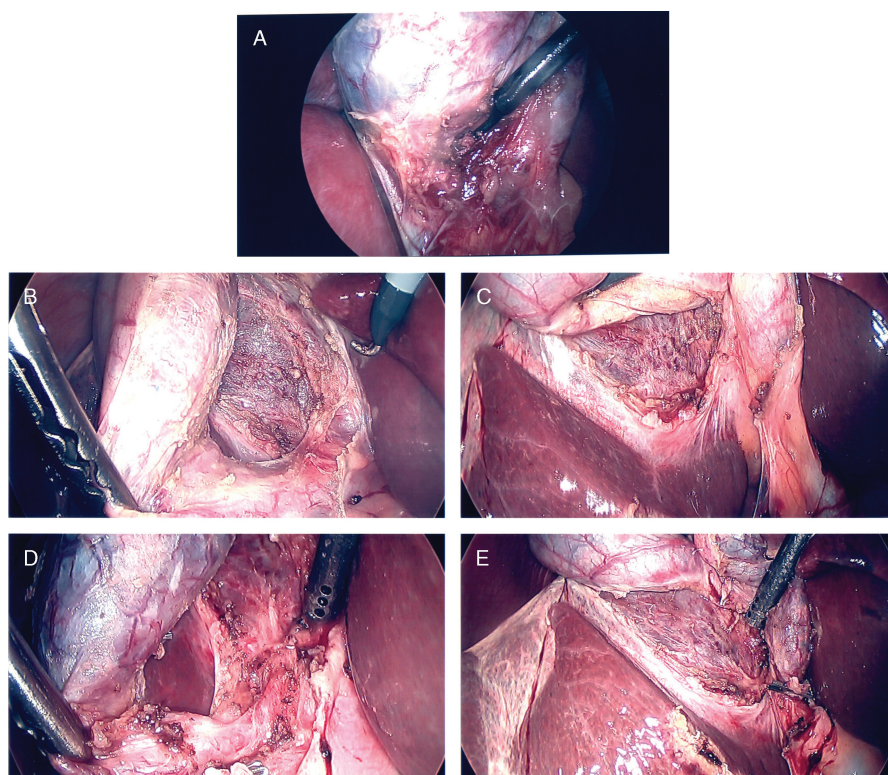


Figure 6. Technique of semi-top-down laparoscopic cholecystectomy. A, Dissection is started on the gallbladder, above the infundibulum of the gallbladder. The peritoneum is scored circumferentially, lateral side first, coming across the peritoneum over the infundibulum of the gallbladder, then opening the peritoneum coming up the medial side of the gallbladder, being careful not to enter the cystic artery as you do so. Dissection of the gallbladder off the liver is being completed here on the lateral aspect of the gallbladder. B and C, Then by rolling the gallbladder from one side to the other, the gallbladder can be fully detached from the liver, leaving only the fundus attached, to provide full exposure. D, At this point and only at this point is the infundibulum and its junction with the cystic duct approached, thus generating a top-down approach to the infundibulum, cystic duct, and cystic artery. When proceeding with the semi-top-down taking only tissues that you see through clearly, any structures that may be encountered such as an aberrant duct, right hepatic artery, or posterior cystic artery can be seen and avoided. E, An exaggerated critical view of safety has resulted. The cystic artery has been divided, and the cystic duct is clearly defined and ready for clipping and dividing.

artery, or posterior cystic artery can be seen and avoided. As you proceed with this dissection, often, the cystic artery widely separates from the gallbladder. At this point in the operation, what you have generated is an exaggerated critical view of safety. It is now clear which structures are cystic artery and cystic duct, having proceeded in essentially a top-down dissection.

KEY CONCEPT: The safest plane for dissection in a cholecystectomy, open or laparoscopically, is on the wall of the gallbladder. Dissection away from the wall of the gallbladder will lead to trouble.

Operative Tricks and Tips

Operating on an acutely inflamed gallbladder for acute cholecystitis or hydrops is challenging and difficult. When placing the laparoscope and seeing this, you must stop and ask the following questions. How sick is my patient? Will he or she tolerate an open cholecystectomy? Will he or she tolerate a long operation? How do I protect the structures in the porta hepatis? Maybe most critical, can I protect the structures in the porta hepatis? If it is clear that the patient is too ill or the anatomy is hazardous from the inflammation, then cholecystostomy is the appropriate option. If it is decided that cholecystectomy can be performed safely, then the gallbladder generally must be decompressed.

Importantly, performing a cholecystectomy on an acutely inflamed, hydroptic gallbladder involves a paradigm shift in operative strategy as compared with the straightforward cholecystectomy. Now, the strategy for the protection of the portal structures is to find and stay only on the wall of the gallbladder (at times submucosa) and know where not to be. The surgeon must know that attempts or persistence in obtaining the classical critical view of safety will lead to biliary or vascular injury. One of the difficulties in this operation is finding the wall and staying on the wall of the gallbladder. In my mind's eye, what I see when I encounter a hydroptic, acutely inflamed gallbladder is analogous to an onion—with multiple peels of inflammatory tissue. Carefully dissect through these layers to safely get onto the wall of the gallbladder, often the submucosa, and complete the dissection in this plane. Again, emphasizing the fact that the safest plane for dissection, open or laparoscopic, is on the wall of the gallbladder.

Partial Cholecystectomy

KEY CONCEPT: At times, the safest plane is viewing the anatomy from within the gallbladder itself.

Partial cholecystectomy has been documented by several authors as a safe and durable option in treating acute cholecystitis.^{93–100} Lateral, medial, and anterior walls of the gallbladder are excised using electrocautery. The densely adherent posterior wall is left on the liver. The mucosa is fully cauterized. As you proceed proximally, you are now within the infundibulum of the gallbladder and visualizing infundibulum and cystic duct from within the gallbladder. Be certain that all stones are extracted. The mucosa is then oversewn with a purse string suture, being certain not to get deeply enough such that portal structures are at risk. Another option in the setting of acute inflammation if the gallbladder can be safely taken off the liver but the infundibulum is markedly inflamed is amputation of the gallbladder at the infundibulum.¹⁰⁰ The anatomy

can again be identified from within the gallbladder; determine the junction of cystic duct and infundibulum. Dissection can often be continued in a safe plane, circumcising the inflamed peritoneum off the gallbladder wall and continuing the dissection. As applied earlier, oversewing the cystic duct from within may be the safest option in this setting. If you cannot safely close the cystic duct from within, in uncommon circumstances where it is not clear that a stitch can be placed safely, a drain is left.

As mentioned, identification methods and technique of dissection are related but different. We will discuss three methods to identify the anatomy during cholecystectomy: the critical view of safety, IOC, and intraoperative ultrasonography. The critical view of safety, espoused by Strasberg for two decades, has been confirmed in multiple studies to be an effective method.^{9,17,22,101–103}

KEY CONCEPT: There are three essential components of the critical view of safety as follows:

1. **At least one third of the gallbladder must be dissected from the liver bed**
2. **The Triangle of Calot must be widely cleared**
3. **Only cystic artery and cystic duct remain as the two structures between the gallbladder and the hepatic ligament**

In an interesting study, adequacy of the critical view of safety was reviewed in photos from 100 cases.¹⁰³ All three criteria were met in only half, with inadequate dissection of the gallbladder off the liver plate as the most common deficiency. Thus, in application of the critical view of safety, all three criteria are required to safely identify anatomy.

Intraoperative Cholangiography

IOC has also been applied as a method for the identification of structures.^{1,3–5,8,10,14,15,23,104} The purposes of IOC include the following: to prevent retained common bile duct stones, to define the biliary anatomy, and to prevent or identify



Figure 7. Computed tomography of a 61-year-old woman, hypotensive and tachycardic on presentation, which shows a liver abscess contiguous to marked cholecystitis. She was taken to the operation room for cholecystectomy and drainage of the liver abscess. The patient died of bleeding from the middle hepatic vein.

bile duct injury. For those who perform cholangiography selectively, which is our approach, the indications include history suggestive of common duct stones including pancreatitis or jaundice or any question of the biliary anatomy during cholecystectomy. Multiple studies have evaluated routine IOC as a means to make laparoscopic cholecystectomy safer;^{1,3–5,8,10,14,15,23,104} the data are conflicting. Several observational cohort studies suggest that routine use of IOC can reduce the risk of common bile duct injury by 50%.^{10,15} In a large meta-analysis by Ludwig et al.¹⁰ of more than 300,000 laparoscopic cholecystectomies, which included 405 major bile duct injuries, the incidence of major bile duct injury was 0.21% in the group where routine cholangiography was used as compared with 0.43% in the selective cholangiography group, representing a statistically significant reduction. Furthermore, 87% of the injuries were diagnosed at the time of surgery in the routine group, compared with only 45% in the selective group. Proponents of routine cholangiography cite this reduction in incidence, earlier recognition of the injury, and perhaps, more successful repair and outcomes as the major reasons to use cholangiography routinely.^{3,10,15} Opponents, however, claim that routine cholangiography is not cost-effective, adds unnecessary time to the operative procedure, and is not always effective at preventing or identifying injury.⁸⁰ A recent editorial in support of routine cholangiography asked, “why are we still debating?”³ In contrast, in a systematic review of IOC published recently, eight randomized trials with 1,715 patients were evaluated.¹ There were only two cases of bile duct injury, confirming that it was underpowered. The authors concluded that “there is no robust evidence to support or abandon the use of IOC to prevent retained stones or bile duct injury.” Another recent review of 92,392 Medicare patients with matched cohorts reported that 40% of patients underwent IOC and 60% did not. The authors concluded that, when confounders were controlled, “intraoperative cholangiography is not effective as a preventive strategy against common bile duct injury during cholecystectomy.”¹⁴

The IOC is dependent on correct interpretation by the surgeon, such as, the transected posterior right hepatic duct described previously. In addition, failure of IOC to prevent bile duct injuries is predictable and relates to (a) filling the CBD only to the bifurcation and not completely filling the liver and, perhaps more importantly, (b) the lack of experience of the general surgeon in reading cholangiograms, particularly the concept of what you do not see is often more important than what you do see. In contrast, bile duct injury found early on IOC leading to prompt diagnosis and treatment improves outcome from injury to the bile duct.

Intraoperative Ultrasonography

Laparoscopic ultrasonography (LUS) is an alternative to IOC for intraoperative assessment of biliary anatomy.^{5,28,105–111} LUS can delineate the common bile duct; cystic duct–common bile duct junction; hepatic artery; portal vein; anomalous anatomy, particularly vascular; and choledocholithiasis. A definite learning curve is associated with LUS, estimated to be 30 to 50 cases. Visualization of the distal common bile duct is more difficult with LUS, and IOC also has the advantage of

confirming free flow of bile (contrast) into the duodenum. Once proficiency with LUS is attained, it is less time consuming than IOC, without radiation exposure, and can be repeated during the operation. Biffl et al.¹⁰⁷ reported 842 cholecystectomies, with their practice initially split regarding routine LUS. They reported LUS to be associated with fewer bile duct complications (bile duct injury, retained stones, cystic duct leaks) than without LUS. In their meta-analysis assessing accuracy of LUS in the detection of choledocholithiasis, Aziz et al.¹⁰⁵ reported sensitivity of 0.87 and specificity of 1.00, nearly identical to IOC (sensitivity, 0.87; specificity, 0.99). Machi et al. have drawn similar conclusions.^{105–111} The SAGES guidelines²⁸ determined that the literature provided Level II, Grade B data for both LUS and IOC as means to delineate biliary anatomy and prevent bile duct injury. Other technologies to delineate biliary anatomy and avoid bile duct injury include passive infrared cholangiography, light cholangiography, near-infrared fluorescence cholangiography, and hyperspectral cholangiography.

KEY CONCEPT: Beware of the middle hepatic vein.

The middle hepatic vein bisects right and left lobes and normally runs within millimeters of the gallbladder fossa. In 20% of patients, a branch of the middle hepatic vein is essentially in the gallbladder plate.¹¹² Particularly when performing cholecystectomy for acute cholecystitis, drifting off the wall of the gallbladder may result in life-threatening hemorrhage with injury to the middle hepatic vein (Fig. 7).

What to Do When a Bile Duct Injury Occurs

If recognized intraoperatively, one must assess his or her ability to repair the injury. The best result comes from an early repair, and the first repair has the best outcome. Except in the most unusual of circumstances, avoid a duct-to-duct anastomosis; do a tension-free Roux-Y. If the surgeon is inexperienced with such a repair, leave the bile duct alone and simply place a drain immediately next to the duct and transfer the patient. The expertise of the surgeon dealing with this complication will impact long-term outcome. If the hepatic artery has also been injured, it is probably best not to repair the bile duct immediately, but wait several months until collaterals have

KEY CONCEPTS

- Perform the cholecystectomy during the index hospitalization for acute cholecystitis.
 - Perform the cholecystectomy within 24–48 h of admission.
 - Know the error traps; avoid them.
 - Semi-top-down technique
 - Critical view of safety
 - ±IOC
 - ±Intraoperative ultrasonography
 - The safest plane for dissection—open or laparoscopic—is on the wall of the gallbladder.
 - Sometimes, the safest plane is viewing things from within the gallbladder.
 - Avoid the use of cautery near the common bile duct or previously placed clips.
 - Know when cholecystostomy is the right operation—know when not to operate.
-

developed. The liver parenchyma can easily survive on the portal vein alone as approximately 70% to 75% of the parenchymal blood flow comes from the portal vein; however, the biliary system is heavily dependent on arterial blood flow.

If the injury is recognized after surgery, place a drain percutaneously and transfer the patient. The ideal treatment if a delayed repair is required is to place a percutaneous transhepatic cholangiocatheter (PTC) (which is difficult because of decompressed ducts) and an intra-abdominal drain (percutaneously if possible) to limit/drain the bile peritonitis. The common hepatic duct will scar down around the PTC and, the abdominal drain will cease draining bile. The abdominal drain can then be removed, and the bile duct can be repaired months later. Obviously, the PTC cannot be clamped but must remain connected to external drainage.

DISCLOSURE

The authors declare no conflicts of interest.

REFERENCES

1. Ford JA, Soop M, Du J, Loveday BP, Rodgers M. Systematic review of intraoperative cholangiography (IOC) in cholecystectomy. *Brit J Surg*. 2012;99:160–167.
2. Akyurek N, Salman B, Irkorucu O, Tascilar O, Yuksel O, Sare M, Tatlicioglu E. Laparoscopic cholecystectomy in patients with previous abdominal surgery. *JSLs*. 2005;9:178–183.
3. Ausania F, Holmes LR, Ausania F, Iype S, Ricci P, White SA. Intraoperative cholangiography in the laparoscopic cholecystectomy era: why are we still debating? *Surg Endosc*. 2012;26:1193–1200.
4. Buddingh KT, Weersma RK, Savenije RA, van Dam GM, Nieuwenhuijs VB. Lower rate of major bile duct injury and increased intraoperative management of common bile duct stones after implementation of routine intraoperative cholangiography. *J Am Coll Surg*. 2011;213:267–274.
5. Buddingh KT, Nieuwenhuijs VB, van Buuren L, Hulscher JB, de Jong JS, van Dam GM. Intraoperative assessment of biliary anatomy for prevention of bile duct injury: a review of current and future patient safety interventions. *Surg Endosc*. 2011;25:2449–2461.
6. Davidson AM, Pappas TN, Murray EA, Hilleren DJ, Johnson RD, Baker ME, Newman GE, Cotton PB, Meyers WC. Mechanisms of major biliary injury during laparoscopic cholecystectomy. *Ann Surg*. 1992;215:196–202.
7. deMestral C, Rotstein OD, Laupacis A, Hoch JS, Zagorski B, Nathens AB. A population-based analysis of the clinical course of 10,304 patients with acute cholecystitis, discharged without cholecystectomy. *J Trauma Acute Care Surg*. 2013;24:460–465.
8. Eikermann M, Siegel R, Broeders I, Dziri C, Fingerhut A, Gutt C, Jaschinski T, Nassar A, Paganini AM, Pieper D, et al.; European Association for Endoscopic Surgery. Prevention and treatment of bile duct injuries during laparoscopic cholecystectomy: the clinical practice guidelines of the European Association for Endoscopic Surgery. *Surg Endosc*. 2012;26(11):3003–39.
9. Heistermann HP, Tobusch A, Palmes D. Prevention of bile duct injuries after laparoscopic cholecystectomy: the critical view of safety [in German]. *Zentralbl Chir*. 2006;131:460–465.
10. Ludwig K, Bernhardt J, Steffen H, Lorenz D. Contribution of intraoperative cholangiography to incidence and outcome of common bile duct injuries during laparoscopic cholecystectomy. *Surg Endosc*. 2002;16:1098–1104.
11. Ingraham AM, Cohen ME, Ko CY, Hall BL. A current profile and assessment of North American cholecystectomy: results from the American College of Surgeons National Surgical Quality Improvement Program. *J Am Coll Surg*. 2010;211:176–186.
12. Murphy MM, Ng S-C, Simons JP, Csikesz NG, Shah SA, Tseng JF. Predictors of major complications after laparoscopic cholecystectomy: surgeon, hospital or patient? *J Am Coll Surg*. 2010;211:73–80.
13. deMestral C, Rotstein OD, Laupacis A, Hoch JS, Zagorski B, Alali AS, Nathens AB. Comparative operative outcomes of early and delayed cholecystectomy for acute cholecystitis. *Ann Surg*. 2014;259:10–15.
14. Sheffield KM, Riall TS, Han Y, Kuo YF, Townsend CM Jr, Goodwin JS. Association between cholecystectomy with vs without intraoperative cholangiography and risk of common duct injury. *JAMA*. 2013;310:812–820.
15. Massarweh NN, Flum DR. Role of intraoperative cholangiography in avoiding bile duct injury. *J Am Coll Surg*. 2007;204:656–664.
16. Strasberg SM. Avoidance of bile duct injury during laparoscopic cholecystectomy. *J Hepatobiliary Pancreat Surg*. 2002;9:543–547.
17. Strasberg SM. Error traps and vasculo-biliary injury in laparoscopic and open cholecystectomy. *J Hepatobiliary Pancreat Surg*. 2008;15:284–292.
18. Strasberg SM, Brunt LM. Rationale and use of the critical view of safety in laparoscopic cholecystectomy. *J Am Coll Surg*. 2010;211:132–138.
19. Strasberg SM, Eagon CJ, Drebin JA. The “hidden cystic duct” syndrome and the infundibular technique of laparoscopic cholecystectomy—the danger of the false infundibulum. *J Am Coll Surg*. 2000;191:661–667.
20. Strasberg SM, Gouma DJ. ‘Extreme’ vasculobiliary injuries: association with fundus-down cholecystectomy in severely inflamed gallbladders. *HPB (Oxford)*. 2012;14:1–8.
21. Strasberg SM, Helton WS. An analytical review of vasculobiliary injury in laparoscopic and open cholecystectomy. *HPB (Oxford)*. 2011;13:1–14.
22. Strasberg SM, Hertl M, Soper NJ. An analysis of the problem of biliary injury during laparoscopic cholecystectomy. *J Am Coll Surg*. 1995;180:101–125.
23. Tomqvist B, Strömberg C, Persson G, Nilsson M. Effect of intraoperative cholangiography and early detection of bile duct injury on survival after cholecystectomy: population based cohort study. *Br Med J*. 2012;345:e6457.
24. Wolf AS, Nijse BA, Sokal SM, Chang Y, Berger DL. Surgical outcome of open cholecystectomy in the laparoscopic era. *Am J Surg*. 2009;197:781–784.
25. Wu YV, Linehan DC. Bile duct injuries in the era of laparoscopic cholecystectomies. *Surg Clin North Am*. 2010;90:787–802.
26. Waage A, Nilsson M. Iatrogenic bile duct injury: a population based study of 152,776 cholecystectomies in the Swedish inpatient registry. *Arch Surg*. 2006;141:1207–1213.
27. Visser BC, Parks RW, Garden OJ. Open cholecystectomy in the laparoscopic era. *Am J Surg*. 2008;195:108–114.
28. Overby DW, Apelgren KN, Richardson W, Fanelli R; Society of American Gastrointestinal and Endoscopic Surgeons. SAGES guidelines for the clinical application of laparoscopic biliary tract surgery. *Surg Endosc*. 2010;24:2368–2386.
29. Yamashita Y, Takada T, Strasberg SM, Pitt HA, Gouma DJ, Garden OJ, Büchler MW, Gomi H, Dervenis C, Windsor JA, et al.; Tokyo Guideline Revision Committee; TG13 surgical management of acute cholecystitis. *J Hepatobiliary Pancreat Sci*. 2013;20:89–96.
30. Kimura Y, Takada T, Kawarada Y, Nimura Y, Hirata K, Sekimoto M, Yoshida M, Mayumi T, Wada K, et al. Definitions, pathophysiology, and epidemiology of acute cholangitis and cholecystitis: Tokyo guidelines. *J Hepatobiliary Pancreat Surg*. 2007;14:15–26.
31. Gomi H, Solomkin JS, Takada T, Strasberg SM, Pitt HA, Yoshida M, Kusachi S, Mayumi T, Miura F, Kiriyaama S, et al.; Tokyo Guideline Revision Committee. TG13 antimicrobial therapy for acute cholangitis and cholecystitis. *J Hepatobiliary Pancreat Sci*. 2013;20:60–70.
32. Hirota M, Takada T, Kawarada Y, Nimura Y, Miura F, Hirata K, Mayumi T, Yoshida M, Strasberg S, Pitt H. Diagnostic criteria and severity assessment of acute cholecystitis: Tokyo guidelines. *J Hepatobiliary Pancreat Surg*. 2007;14:78–82.
33. Yamashita Y, Takada T, Kawarada Y, Nimura Y, Hirota M, Miura F, Mayumi T, Yoshida M, Strasberg S, Pitt HA, et al. Surgical treatment of patients with acute cholecystitis: Tokyo guidelines. *J Hepatobiliary Pancreat Surg*. 2007;14:91–97.
34. Miura F, Takada T, Strasberg SM, Solomkin JS, Pitt HA, Gouma DJ, Garden OJ, Büchler MW, Yoshida M, Mayumi T, et al.; Tokyo Guidelines Revision Committee. TG13 flowchart for the management of acute cholangitis and cholecystitis. *J Hepatobiliary Pancreat Sci*. 2013;20:47–54.
35. Yasuda H, Takada T, Kawarada Y, Nimura Y, Hirata K, Kimura Y, Wada K, Miura F, Hirota M, Mayumi T, et al. Unusual cases of acute

- cholecystitis and cholangitis: Tokyo guidelines. *J Hepatobiliary Pancreat Surg.* 2007;14:98–113.
36. Yokoe M, Akada T, Strasberg SM, Solomkin JS, Mayumi T, Gomi H, Pitt HA, Gouma DJ, Garden OJ, Büchler MW, et al.; Tokyo Guidelines Revision Committee. New diagnostic criteria and severity assessment of acute cholecystitis in revised Tokyo guidelines. *J Hepatobiliary Pancreat Sci.* 2012;19:578–585.
37. Yoshida M, Takada T, Kawarada Y, Tanaka A, Nimura Y, Gomi H, Hirota M, Miura F, Wada K, Mayumi T, Solomkin JS, et al. Antimicrobial therapy for acute cholecystitis: Tokyo guidelines. *J Hepatobiliary Pancreat Surg.* 2007;14:83–90.
38. Takada T, Kawarada Y, Nimura Y, Yoshida M, Mayumi T, Sekimoto M, Miura F, Wada K, Hirota M, Yamashita Y, et al. Background: Tokyo guidelines for the management of acute cholecystitis and cholangitis. *J Hepatobiliary Pancreat Surg.* 2007;14:1–10.
39. Tsuyuguchi T, Itoi T, Takada T, Strasberg SM, Pitt HA, Kim MH, Supe AN, Mayumi T, Yoshida M, Miura F, et al.; Tokyo Guideline Revision Committee. TG13 indications and techniques for gallbladder drainage in acute cholecystitis. *J Hepatobiliary Pancreat Sci.* 2013;20:81–88.
40. Mayumi T, Takada T, Kawarada Y, Nimura Y, Yoshida M, Sekimoto M, Miura F, Wada K, Hirota M, Yamashita Y, Nagino M, et al. Result of the Tokyo consensus meeting Tokyo guidelines. *J Hepatobiliary Pancreat Surg.* 2007;14:114–121.
41. Fujii Y, Ohuchida J, Chijiwa K, Yano K, Imamura N, Nagano M, Hiyoshi M, Otani K, Kai M, Kondo K. Verification of Tokyo Guidelines for diagnosis and management of acute cholangitis. *J Hepatobiliary Pancreat Sci.* 2012;19:487–491.
42. Riall TS, Zhang D, Townsend CM Jr, Kuo YF, Goodwin JS. Failure to perform cholecystectomy for acute cholecystitis in elderly patients is associated with increased morbidity, mortality and cost. *J Am Coll Surg.* 2010;210:668–679.
43. Gurusamy KS, Davidson C, Gluud C, Davidson BR. Early versus delayed laparoscopic cholecystectomy for people with acute cholecystitis. *Cochrane Database Syst Rev.* 2013;6:CD005440.
44. Brooks KR, Scarborough JE, Vaslef SN, Shapiro ML. No need to wait: an analysis of the timing of cholecystectomy during admission for acute cholecystitis using the American College of Surgeons National Surgical Quality Improvement Program database. *J Trauma Acute Care Surg.* 2013;74:167–174.
45. Banz V, Gsponer T, Candinas D, Guller U. Population-based analysis of 4113 patients with acute cholecystitis: defining the optimal time-point for laparoscopic cholecystectomy. *Ann Surg.* 2011;254:964–970.
46. Gutt CN, Encke J, Königer J, Harnoss JC, Weigand K, Kipfmüller K, Schunter O, Götze T, Golling MT, Menges M, Klar E, et al. Acute cholecystitis: early vs late cholecystectomy, a multicenter randomized trial (ACDC Study). *Ann Surg.* 2013;258:385–393.
47. Catani M, Modini C. Laparoscopic cholecystectomy in acute cholecystitis: a proposal of a safe and effective technique. *Hepatogastroenterology.* 2007;54:2186–2191.
48. Farooq T, Buchanan G, Manda V, Kennedy R, Ockrim J. Is early laparoscopic cholecystectomy safe after the “safe period”? *J Laparoendosc Adv Surg Tech A.* 2009;19:471–474.
49. To KB, Cherry-Bukowiec JR, Englesbe MJ, Terjmanian MN, Shijie C, Campbell DA Jr, Napolitano LM. Emergent versus elective cholecystectomy: conversion rates and outcomes. *Surg Inf.* 2013;14:512–519.
50. Pitt HA, Postier RG, Cameron JL. Consequences of preoperative cholangitis and its treatment on the outcome of operation for choledocholithiasis. *Surgery.* 1983;94:447–452.
51. Maluenda F, Csendes A, Burdiles P, Diaz J. Bacteriological study of choledochal bile in patients with common duct stones, with or without acute suppurative cholangitis. *Hepatogastroenterology.* 1989;36:132–135.
52. Jaafar G, Persson G, Svenblad B, Sandblom G. Outcomes of antibiotic prophylaxis in acute cholecystectomy in a population based gallstone surgery registry. *Br J Surg.* 2014;101:69–73.
53. Goldman G, Kahn PJ, Alon R, Wiznitzer T. Biliary colic treatment and acute cholecystitis prevention by prostaglandin inhibitor. *Dig Dis Sci.* 1989;34:809–811.
54. Solomkin JS, Mazuski JE, Bradley JS, Rodvold KA, Goldstein EJ, Baron EJ, O'Neill PJ, Chow AW, Dellinger EP, Eachempati SR, et al. Diagnosis and management of complicated intra-abdominal infection in adults and children: guidelines by the Surgical Infection Society and Infectious Disease Societies of America. *Clin Infect Dis.* 2010;50:133–164.
55. Regimbeau JM, Fuks D, Pautrat K, Mauvais F, Haccart V, Msika S, Mathonnet M, Scotté M, Paquet JC, Vons C, et al.; FRENCH Study Group. Effect of postoperative antibiotic administration on postoperative infection following cholecystectomy for acute cholecystitis. *JAMA.* 2014;312:145–154.
56. Van den Hazel SJ, Speelman P, Tytgat GN, Dankert J, van Leeuwen DJ. Role of antibiotics in the treatment and prevention of acute and recurrent cholangitis. *Clin Infect Dis.* 1994;19:279–286.
57. Wang C-H, Chou HC, Liu KL, Lien WC, Wang HP, Wu YM. Long-term outcome of patients with acute cholecystitis receiving antibiotic treatment: a retrospective cohort study. *World J Surg.* 2014;38(2):347–354.
58. Gurusamy KS, Rossi M, Davidson BR. Percutaneous cholecystostomy for high risk patients with acute calculous cholecystitis. *Cochrane Database Syst Rev.* 2013;8:CD007088.
59. Joseph T, Unver K, Hwang GL, Rosenberg J, Sze DY, Hashimi S, Kothary N, Louie JD, Kuo WT, Hofmann LV, et al. Percutaneous cholecystostomy for acute cholecystitis: ten year experience. *J Vasc Interv Radiol.* 2012;23:83–88.
60. Barak O, Elazary R, Appelbaum L, Rivkind A, Almog G. Conservative treatment for acute cholecystitis: clinical and radiographic predictors of failure. *Isr Med Assoc J.* 2009;11:739–743.
61. Ito K, Fujita N, Noda Y, Kobayashi G, Kimura K, Sugawara T, Horaguchi J. Percutaneous cholecystostomy versus gallbladder aspiration for acute cholecystitis: a prospective randomized controlled trial. *AJR Am J Roentgenol.* 2004;183:193–196.
62. Berber E, Engle KL, String A, Garland AM, Chang G, Macho J, Pearl JM, Siperstein AE. Selective use of tube cholecystostomy with interval laparoscopic cholecystectomy in acute cholecystitis. *Arch Surg.* 2000;135:341–346.
63. Byrne MF, Suhocki P, Mitchell RM, Pappas TN, Stiffler HL, Jowell PS, Branch MS, Baillie J. Percutaneous cholecystostomy in patients with acute cholecystitis: experience of 45 patients at a US referral center. *J Am Coll Surg.* 2003;297:206–211.
64. Griniatsos J, Petrou A, Pappas P, et al. Percutaneous cholecystostomy without interval cholecystectomy as definitive treatment of acute cholecystitis in elderly and critically ill patients. *South Med J.* 2008;101:586–590.
65. Spira RM, Petrou A, Pappas P, Revenas K, Karavokyros I, Michail OP, Tsigis C, Giannopoulos A, Felekouras E. Percutaneous transhepatic cholecystostomy and delayed laparoscopic cholecystectomy in critically ill patients with acute calculous cholecystitis. *Am J Surg.* 2002;183:62–66.
66. Granlund A, Karlson BM, Elvin A, Rasmussen I. Ultrasound-guided percutaneous cholecystostomy in high risk surgical patients. *Langenbecks Arch Surg.* 2001;386:212–217.
67. Davis CA, Landercasper J, Gundersen LH, Lambert PJ. Effective use of percutaneous cholecystostomy in high risk surgical patients: techniques, tube management and results. *Arch Surg.* 1999;134:727–731.
68. Winblad A, Gullstrand P, Svanvik J, Sandström P. Systematic review of cholecystostomy as a treatment option in acute cholecystitis. *HPB (Oxford).* 2009;11:183–193.
69. de Mestral C, Gomez D, Haas B, Zagorski B, Rotstein OD, Nathens AB. Cholecystostomy: a bridge to hospital discharge but not delayed cholecystectomy. *J Trauma Acute Care Surg.* 2013;74:175–180.
70. Li M, Li N, Ji W, Quan Z, Wan X, Wu X, Li J. Percutaneous cholecystostomy is a definitive treatment for acute cholecystitis in elderly high-risk patients. *Am Surg.* 2013;79:524–527.
71. McKay A, Abulfaraj M, Lipschitz J. Short and long term outcomes following percutaneous cholecystostomy for acute cholecystitis in high-risk patients. *Surg Endosc.* 2012;26:1343–1351.
72. Hatzidakis AA, Prassopoulos P, Petinarakis I, Sanidas E, Chrysos E, Chalkiadakis G, Tsiftsis D, Gourtsoyannis NC. Acute cholecystitis in high-risk patients: percutaneous cholecystostomy vs conservative treatment. *Eur Radiol.* 2002;12:1778–1784.
73. Abi-Haidar Y, Sanchez V, Williams SA, Itani KM. Revisiting percutaneous cholecystostomy for acute cholecystitis based on a 10-year experience. *Arch Surg.* 2012;147:416–422.

74. Cherg N, Witkowski ET, Sneider EB, Wiseman JT, Lewis J, Litwin DE, Santry HP, Cahan M, Shah SA. Use of cholecystostomy tubes in the management of patients with primary diagnosis of acute cholecystitis. *J Am Coll Surg*. 2012;214:196–201.
75. Kortram K, van Ramshorst B, Bollen TL, Besselink MG, Gouma DJ, Karsten T, Kruij PM, Nieuwenhuijzen GA, Kelder JC, Tromp E, et al. Acute cholecystitis in high risk surgical patients: percutaneous cholecystostomy versus laparoscopic cholecystectomy (CHOCOLATE Trial): study protocol for a randomized controlled trial. *Trials*. 2012;13:7.
76. Lipman JM, Claridge JA, Haridas M, Martin MD, Yao DC, Grimes KL, Malangoni MA. Preoperative findings predict conversion from laparoscopic to open cholecystectomy. *Surgery*. 2007;142:556–565.
77. Brodsky A, Matter I, Sabo E, Cohen A, Abrahamson J, Eldar S. Laparoscopic cholecystectomy for acute cholecystitis: can need for conversion and the probability of complications be predicted? A prospective study. *Surg Endosc*. 2000;14:755–760.
78. Zhu B, Zhang Z, Wang Y, Gong K, Lu Y, Zhang N. A comparison of laparoscopic cholecystectomy for acute cholecystitis both within and beyond 72 h of symptom onset during the emergency admission: how golden is “golden”? *World J Surg*. 2012;36:2654–2658.
79. Wevers KP, vanWestreenen HL, Patijn GA. Laparoscopic cholecystectomy in acute cholecystitis: C-reactive protein level combined with age predicts conversion. *Surg Laparosc Endosc Percutan Tech*. 2013;23:163–166.
80. Way LW, Stewart L, Gantert W, Liu K, Lee CM, Whang K, Hunter JG. Causes and of laparoscopic bile duct injuries. *Ann Surg*. 2003;237:460–469.
81. Rosen M, Brody F, Ponsky J. Predictive factors for conversion of laparoscopic cholecystectomy. *Am J Surg*. 2002;184:254–258.
82. Livingston EH, Rege RV. A nationwide study of conversion from laparoscopic to open cholecystectomy. *Am J Surg*. 2004;188:205–211.
83. Lim KR, Ibrahim S, Tan NC, Lim SH, Tay KH. Risk factors for conversion to open surgery in patients with acute cholecystitis undergoing interval laparoscopic cholecystectomy. *Ann Acad Med Singapore*. 2007;36:631–635.
84. Kanaan SA, Murayama KM, Merriam LT, Dawes LG, Prystowsky JB, Rege RV, Joehl RJ. Risk factors for conversion of laparoscopic to open cholecystectomy. *J Surg Res*. 2002;106:20–24.
85. Chuang KI, Corley D, Postlethwaite DA, Merchant M, Harris HW. Does increased experience with laparoscopic cholecystectomy yield more complex bile duct injuries? *Am J Surg*. 2012;203:480–487.
86. Wojcicki M, Patkowski W, Chmurowicz T, Bialek A, Wiechowska-Kozłowska A, Stankiewicz R, Milkiewicz P, Krawczyk M. Isolated right posterior bile duct injury following cholecystectomy: report of two cases. *World J Gastroenterol*. 2013;19:6118–6121.
87. Babel N, Sakpal SV, Paragi P, Wellen J, Feldman S, Chamberlain RS. Iatrogenic bile duct injury associated with anomalies of the right hepatic sectoral ducts: a misunderstood and underappreciated problem. *HPB Surg*. 2009;2009:153269.
88. Hunter J. Avoidance of bile duct injury during laparoscopic cholecystectomy. *Am J Surg*. 1991;162:71–76.
89. Tuveri M, Calò PG, Medas F, Tuveri A, Nicolosi A. Limits and advantages of fundus-first laparoscopic cholecystectomy: lessons learned. *J Laparoendosc Adv Surg Tech A*. 2008;18:69–75.
90. Kelly MD. Laparoscopic retrograde (fundus first) cholecystectomy. *BMC Surg*. 2009;9:19–27.
91. Mahmud S, Masaud M, Canna K, Nassar AH. Fundus-first laparoscopic cholecystectomy. *Surg Endosc*. 2002;16:581–584.
92. Neri V, Ambrosi A, Fersini A, Tartaglia N, Valentino TP. Antegrade dissection in laparoscopic cholecystectomy. *JSLs*. 2007;11:225–228.
93. Bornman PC, Terblanche J. Subtotal cholecystectomy: for the difficult gallbladder in portal hypertension and cholecystitis. *Surgery*. 1985;98:1–6.
94. Michalowski K, Bornman PC, Krige JE, Gallagher PJ, Terblanche J. Laparoscopic subtotal cholecystectomy in patients with complicated acute cholecystitis or fibrosis. *Br J Surg*. 1998;85:9904–9906.
95. Nakajima J, Sasaki A, Obuchi T, Baba S, Nitta H, Wakabayashi G. Laparoscopic subtotal cholecystectomy for severe cholecystitis. *Surg Today*. 2009;39:870–875.
96. Sinha I, Smith ML, Safranek P, Dehn T, Booth M. Laparoscopic subtotal cholecystectomy without cystic duct ligation. *Br J Surg*. 2007;94:1527–1529.
97. Sharp CF, Garza RZ, Mangram AJ, Dunn EL. Partial cholecystectomy in the setting of severe inflammation is an acceptable consideration with few long-term sequelae. *Am Surg*. 2009;75:249–252.
98. Soleimani M, Mehrabi A, Mood ZA, Fonouni H, Kashfi A, Büchler MW, Schmidt J. Partial cholecystectomy as a safe and viable option in the emergency treatment of complex acute cholecystitis: a case series and review of the literature. *Am Surg*. 2007;5:498–507.
99. Henneman D, da Costa DW, Vrouwenraets BC, van Wagenveld BA, Lagarde SM. Laparoscopic partial cholecystectomy for the difficult gallbladder: a systematic review. *Surg Endosc*. 2013;27:351–358.
100. Hubert C, Annet L, van Beers BE, Gigot JF. The “inside approach of the gallbladder” is an alternative to the classic Calot’s triangle dissection for a safe operation in severe cholecystitis. *Surg Endosc*. 2010;24:2626–2632.
101. Yegiyants S, Collins JC. Operative strategies can reduce the incidence of major bile duct injury in laparoscopic cholecystectomy. *Am Surg*. 2008;74:985–987.
102. Averginos C, Kelgiorgi D, Touloumis Z, Baltatzi L, Derveniz C. One thousand laparoscopic cholecystectomies in a single surgical unit using the critical view of safety technique. *J Gastrointest Surg*. 2009;13:498–503.
103. Lam T, Usatoff V, Chan STF. Are we getting the critical view? A prospective study of photographic documentation during laparoscopic cholecystectomy. *HPB (Oxford)*. 2014;16(9):859–863.
104. Sanjay P, Kulli C, Polignano FM, Tait IS. Optimal surgical technique, use of intraoperative cholangiography and management of acute gallbladder disease: the results of a nationwide survey in UK and Ireland. *Ann R Coll Surg Engl*. 2010;92(4):302–6.
105. Aziz O, Ashrafian H, Jones C, Harling L, Kumar S, Garas G, Holme T, Darzi A, Zacharakis E, Athanasiou T. Laparoscopic ultrasonography versus intra-operative cholangiogram for the detection of common bile duct stones during laparoscopic cholecystectomy: a meta-analysis of diagnostic accuracy. *Int J Surg*. 2014;12:712–719.
106. Falcone RA Jr, Fegelman EJ, Nussbaum MS, Brown DL, Bebbe TM, Merhar GL, Johannigman JA, Luchette FA, Davis K Jr, Hurst JM. A prospective comparison of laparoscopic ultrasound vs intraoperative cholangiogram during laparoscopic cholecystectomy. *Surg Endosc*. 1999;13:784–788.
107. Biffi WL, Moore EE, Offner PJ, Franciose RJ, Burch JM. Routine intraoperative laparoscopic ultrasonography with selective cholangiography reduces bile duct complications during laparoscopic cholecystectomy. *J Am Coll Surg*. 2001;193:272–280.
108. Machi J, Johnson JO, Deziel DJ, Soper NJ, Berber E, Siperstein A, Hata M, Patel A, Singh K, Arregui ME. The routine use of laparoscopic ultrasound decreases bile duct injury: a multicenter study. *Surg Endosc*. 2009;23:384–388.
109. Tranter SE, Thompson MH. A prospective single-blinded controlled study comparing laparoscopic ultrasound of the common bile duct with operative cholangiography. *Surg Endosc*. 2003;17:216–219.
110. Perry KA, Myers JA, Deziel DJ. Laparoscopic ultrasound as the primary method for bile duct imaging during cholecystectomy. *Surg Endosc*. 2008;22:208–213.
111. Machi J, Oishi AJ, Tajiri T, Murayama KM, Furumoto NL, Oishi RH. Routine laparoscopic ultrasound can significantly reduce the need for selective intraoperative cholangiography during cholecystectomy. *Surg Endosc*. 2007;21:270–274.
112. Ball CG, MacLean AR, Kirkpatrick AW, Bathe OF, Sutherland F, Debru E, Dixon E. Hepatic vein injury during laparoscopic cholecystectomy: the unappreciated proximity of the middle hepatic vein to the gall bladder bed. *J Gastrointest Surg*. 2006;10:1151–1155.