

Milestones in the Evolution of Hepatic Surgery

Henri Bismuth, M.D., F.A.C.S. (Hon)^{1*}, Rony Eshkenazy, M.D.², and Arie Arish, M.D.²

¹Hepatobiliary Institute, Paul Brousse Hospital, Paris, France, and ²Hepato-Biliary Surgery Service, Department of General Surgery, Rambam Health Care Campus, Haifa, Israel

ABSTRACT

This paper describes the rapid evolution of modern liver surgery, starting in the middle of the twentieth century. Claude Couinaud studied and described the segmental anatomy of the liver, Thomas Starzl performed the first liver transplantations, and Henri Bismuth introduced the concept of anatomical resections. Hepatic surgery has developed significantly since those early days. To date, innovative techniques are applied, using cutting-edge technologies: Intraoperative ultrasound, techniques of vascular exclusion of the liver, new devices for performing homeostasis and dissection, laparoscopy for resections, and new drugs that allow the resection of previously unresectable tumors. The next stage in liver surgery will probably be the implementation of a multidisciplinary holistic approach to the liver-diseased patient that will ensure the best and most efficient treatments in the future.

KEY WORDS: Surgery, liver, transplantation, transection

The adult liver, the largest organ in the body, accounts for 2%–3% of overall body weight. Richly vascularized, the liver has an inflow carried through the portal vein and the hepatic artery and an outflow draining through the main and accessory hepatic veins. The main portal vein drains the splanchnic territory and carries 70%–80% of the overall hepatic inflow. It divides into two branches, the right and left portal veins, which are

then subdivided into sectorial and segmental branches. Blood pressure in the main portal vein is usually low, with a portocaval gradient (i.e. portal vein pressure minus central venous pressure) of less than 5 mmHg. In chronic liver disease, especially cirrhosis, the portocaval gradient may increase to the point of portal hypertension.

In the seventeenth and eighteenth centuries,

Abbreviations: IOC, intraoperative cholangiography; IOUS, intraoperative ultrasound; PVE, portal vein embolization.

Citation: Bismuth H, Eshkenazy R, Arish A. Milestones in the evolution of hepatic surgery. RMMJ 2011;2(1):e0021. doi:10.5041/RMMJ.10021

Copyright: © 2011 Bismuth et al. This is an open-access article. All its content, *except where otherwise noted*, is distributed under the terms of the Creative Commons Attribution License (<http://creativecommons.org/licenses/by/3.0>), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Conflict of interest: No potential conflict of interest relevant to this article was reported.

* To whom correspondence should be addressed. E-mail: henri.bismuth@pbr.aphp.fr

liver resections were carried out on trauma victims with a range of injuries occurring as a result of military combat. The first liver resections in a non-emergency setting resulted in uncontrolled bleeding and death.

HISTORY OF MODERN LIVER SURGERY

The German surgeon Carl Johann August Langenbuch performed the first successful hepatic resection in 1888 (he was the first to perform cholecystectomy in 1882).¹ He resected a part of the left lobe of the liver after ligating the vascular pedicles. The pathologic examination of the specimen revealed normal liver.

The first steps leading to modern liver surgery began in the late 1950s. Scattered publications from the United States describing a limited series of liver resections that had met with some success were published. The technique of liver resection at that time was ill defined.^{2,3}

Concomitantly, in 1952 in France Lortat Jacob published a manuscript on his experiences performing anatomical liver resections: right hepatectomy after vascular control of the right liver.⁴ In 1956, Claude Couinaud,^{5,6} after studying casts of

the liver, published his book, *The liver – anatomical studies and surgical studies*. He showed that, according to the distribution of the portal blood, the liver contains four parts that are subdivided by the hepatic veins into eight segments. He coined the segment numbers as follows: one for the central segment and two through eight for the seven other segments in a clock-wise fashion. This segmental division of the liver is the basis of modern functional and surgical liver anatomy.

Despite the anatomical discoveries of the 1950s, their application in surgical practice was limited. No clinical methods existed that could detect the existence of small liver tumors that might have required segmental resections. Physicians made their diagnoses based on physical symptoms (pain or a palpable tumor), at which point it was usually already too late for surgery. Only large liver resections could be performed.

The introduction of ultrasound in the early 1980s^{7,8} into common clinical practice allowed the clinician to diagnose asymptomatic small liver tumors of 2 and 3 cm and paved the way to rapid development of liver surgery. In fact, modern liver surgery began when functional liver anatomy discovered 30 years earlier was applied, enabling

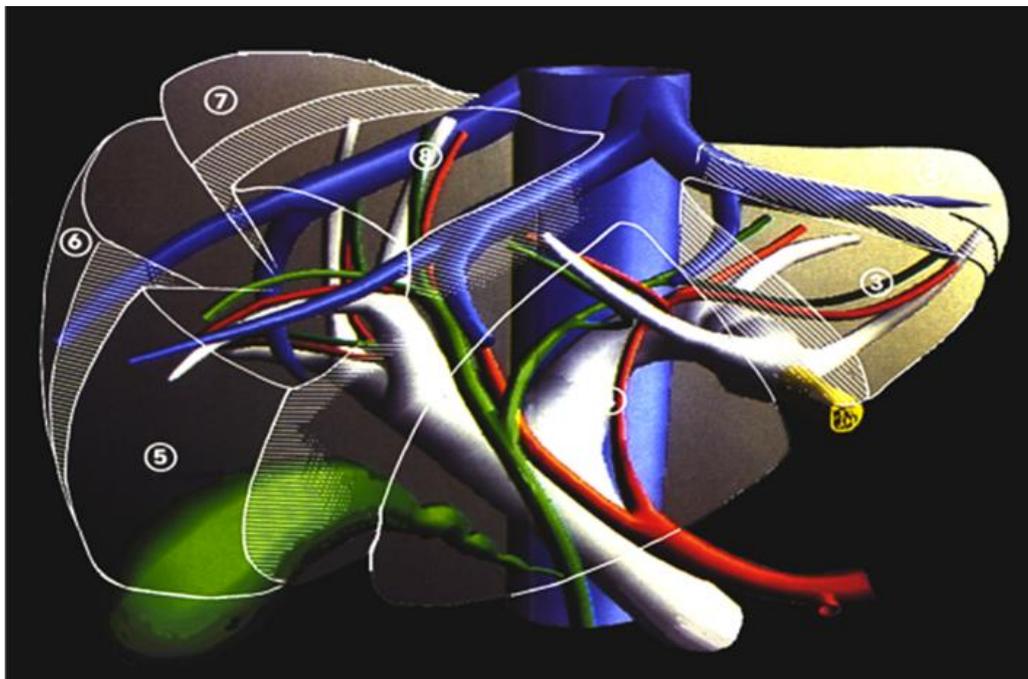


Figure 1. The transparent liver showing the main arteries (in red), portal vein (in white), hepatic veins (in blue), and main parts of the biliary system (in green). Liver segments are numbered, and the round ligament is designated in yellow.

segmental liver surgery.

The paper with the somewhat provocative title of “Surgical anatomy and anatomical surgery of the liver”⁹ published in 1982 was a turning-point in the practice of liver surgery. Navigating surgery on the basis of anatomy eliminated the use of “atypical” or “non-anatomical” resections of the liver which had resulted in bleeding, increased morbidity, and mortality sometimes as a result of liver necrosis. Liver resections based on anatomy gained popularity and evolved into bloodless surgery that removed independent segments or groups of two or more segments.

In 1984, intraoperative ultrasound (IOUS) was introduced into practice.^{10–14} The technique allowed the surgeon to understand liver vasculature and biliary duct anatomy and rendered the liver transparent (Figure 1). In fact, the introduction of IOUS into liver surgery had the same impact as the inclusion of intraoperative cholangiography (IOC) fifty years earlier in biliary surgery.¹⁵ By finding the liver’s lines of division with IOUS and

looking for a tumor, the surgeon could establish the relations among the portal elements, the hepatic veins, the hepatic parenchyma, and the tumor, and then decide what kind of anatomical resection needed to be carried out – “*Hepatectomie à la carte*” for each individual patient.

Control of bleeding during liver resection is a major challenge for the surgeon. It is particularly difficult in cirrhotic liver due to the fibrotic nature of the liver tissue. The indication, as well as the type of clamping, depends mainly on the size and location of the lesions to be resected, the quality of the liver parenchyma, the surgeon’s preferences, and unexpected operative events. In 1958 Lin introduced the finger fracture technique, which involves crushing of liver parenchyma by the surgeon’s fingers under inflow occlusion so as to isolate vessels and bile ducts for ligation.¹⁶ This technique was subsequently improved through the use of surgical instruments such as a small Kelly clamp for blunt dissection (clamp crushing or “Kellyclasia”).^{9,17,18}

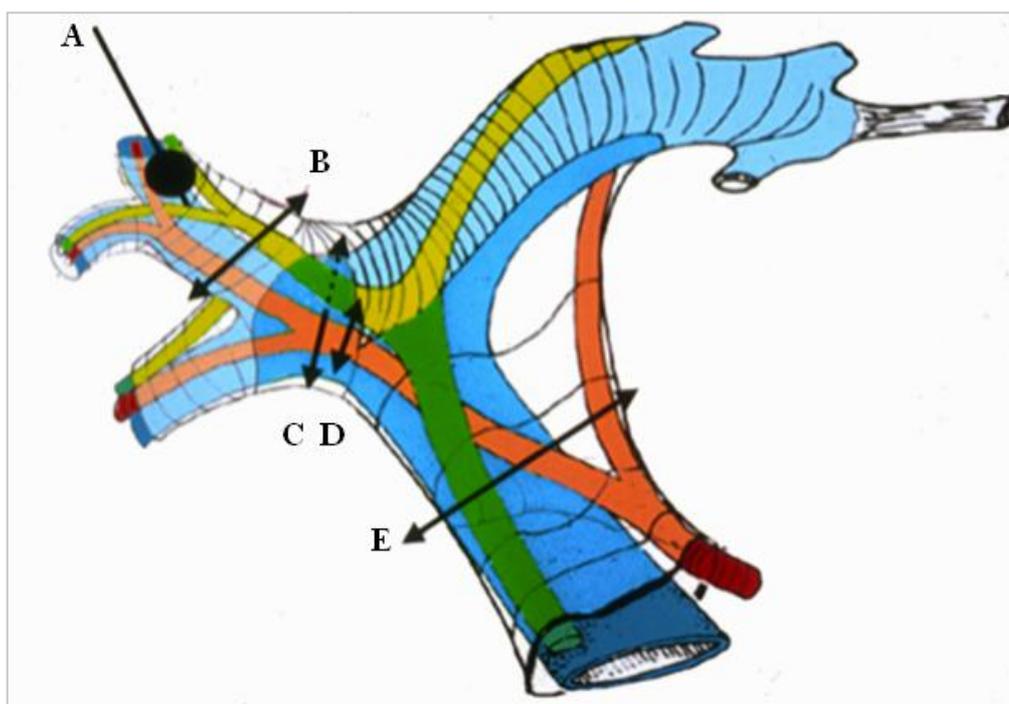


Figure 2. Glissonian pedicle elements: portal vein (in blue), hepatic artery (in red), and the bile ducts (in green). Hepatic inflow occlusion: A) Selective occlusion of segmental portal vein by a balloon introduced under ultrasonographic guidance. The arrows show the different sites of Glissonian clamping. B) Suprahilar clamping of one sector of the right liver; C and D) hilar selective clamping to the right liver; E) total pedicular clamping (Pringle maneuver).

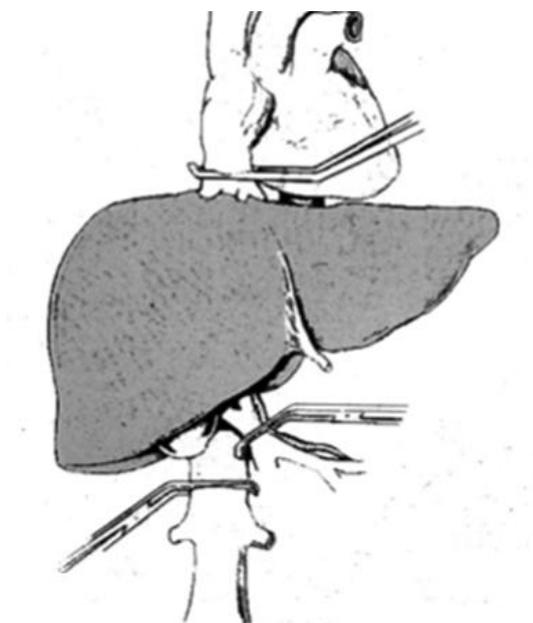


Figure 3. Total vascular exclusion of the liver by clamping the infrahepatic and suprahepatic inferior vena cava and the hepatoduodenal ligament.

In many centers, ultrasonic dissection using the Cavitron Ultrasonic Surgical Aspirator (CUSA; Tyco Healthcare, Mansfield, MA, USA) has become the standard technique for liver parenchyma dissection. With this technology, the liver tissue is fragmented with ultrasonic energy and aspirated, thus exposing small vascular and ductal structures that can be ligated or clipped with titanium hemoclips.¹⁹ The water jet dissector employs a pressurized jet of water instead of ultrasonic energy, in order to fragment the liver parenchyma tissue and expose the vascular and ductal structures.²⁰ However, this technique has not become as popular as ultrasonic dissection.

To reduce blood loss during liver resection, intermittent inflow occlusion by clamping of the portal triad (Pringle maneuver) is frequently used.^{21,22} However, there is a limit to how long the Pringle maneuver can be applied. Prolonged inflow occlusion (over 120 minutes) may have deleterious effects on liver functions.²³ Hepatic inflow occlusion can be directed to one side or to a segment by clamping the Glissonian pedicle at the hilum or inside the liver parenchyma (Figure 2).^{9,24–29}

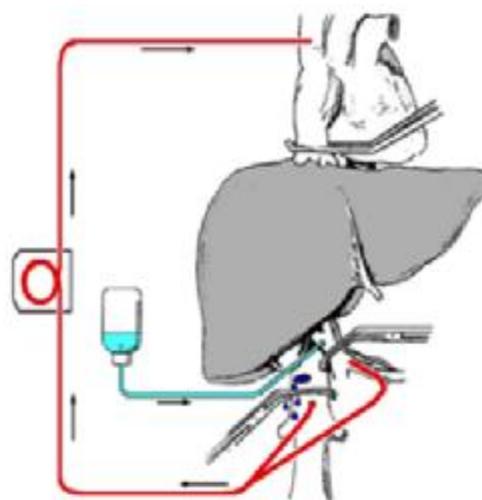


Figure 4. Total vascular exclusion of the liver with hypothermia as described by Fortner et al.⁴⁰ The liver is excluded (as in Figure 3). Veno-venous bypass of the liver is performed (red lines), and hypothermic solution is infused into the portal vein (in blue).

The majority of liver resections can be done with no clamping at all.^{30,31} In some patients total hepatic vascular isolation is needed. This isolation can be partial (meaning occlusion of the inflow and only one hepatic vein^{32,33}) or total (meaning occlusion of the inflow). Outflow occlusion is obtained by occluding the vena cava above and below the hepatic veins^{34–37} (Figure 3). In cases of a small tumor adherent to a hepatic vein, isolation of the corresponding liver by clamping the inflow and the hepatic vein can facilitate and render surgery safe if the lesion is resected with the adherent vein. When a large tumor is found to have entered the vena cava, this technique enables bloodless resection of the involved vena cava and safe reconstruction of its continuity.^{38,39} Total vascular exclusion (TVE) of the liver can be applied safely for as long as 60 minutes. This can be extended to 8 hours by using hypothermia (Figure 4), as is done in liver transplantation. The liver resection can be done *in situ in vivo* as first described by Fortner et al.,^{40–42} *ex situ in vivo* as described by Hannoun et al.,^{42–45} or *ex situ ex vivo* as described by Pichlmayr et al.⁴⁶ (Figure 5, next page). This innovative approach to liver resection has a high rate of complication and even mortality.

Another concept in liver surgery that guides

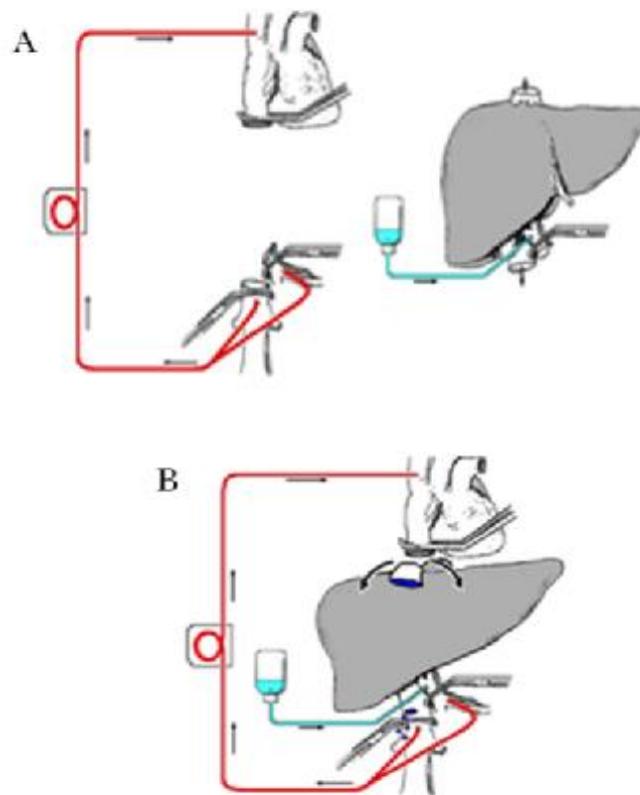


Figure 5. Total vascular exclusion for complex liver resections. A) The *ex-situ ex-vivo* technique described by Pichlmayr et al.⁴⁶ B) The *ex-situ in-vivo* technique described by Hannoun et al.⁴³

surgeons toward safe surgery is that “the liver is parenchyma tissue with blood vessels inside”. The rationale behind this concept is to dissect this parenchyma tissue while ensuring hemostasis. The concept was popularized following the development of new devices that enabled such dissection, e.g. the harmonic scalpel,^{47–49} LigaSure,^{50–52} tissue link,⁵³ radiofrequency,^{54,55} and the “Habib sealer”.^{56,57} These devices allow the parenchyma to be cut without having to clamp the pedicle. Nevertheless, the clamp crushing technique is still widely used.^{58–61}

The main indication for liver resection today is liver metastasis resulting from advanced cancers of the colon and rectum. Apart from liver surgery for trauma or hepatocellular carcinoma in cirrhotic patients where the mortality is high,⁶² the overall operative mortality in liver resections is be-

tween 0% and 2%.^{63,64} Trained liver surgery teams can achieve less than 1% mortality. This is a great advance in comparison to the mortality in liver surgery in early reports, which reached a mortality rate as high as 20%.⁶⁵

LIVER SURGERY – A NEW SPECIALTY

In medicine and surgery, the use of special technologies and knowledge has facilitated the development of subspecialties. Cardiac surgery, for example, emerged as a specialty of surgery because of the use of special machines and instruments. This was not the case for liver surgery.

When it began, liver surgery used nothing but the instruments of general surgery. Today, with

the exploitation of special technology and instrumentation such as operative ultrasound, ultrasonic dissectors, argon coagulators, cryotherapy, radiofrequency, and extracorporeal circulation, liver surgery has become a subspecialty of general surgery, or more precisely, a *hyper-specialty*.

One step on the way to liver surgery becoming a specialty was the advent of liver transplantation. When liver surgeons began performing transplantation, liver resection was a step in the procedure. Surgery on the graft itself or on the recipient, which may have meant heterotopic liver transplantation and reducing the size of the graft,^{66,67} operating on the living donor,⁶⁸ performing a domino transplantation,^{69,70} or a split liver procedure,⁷¹⁻⁷⁴ were all done by liver surgeons doing liver transplantation.

Liver surgery is now a specialty, as one of the three branches of gastro-intestinal (GI) surgery. These are upper GI, colorectal, and hepato-bilio-pancreatic surgery. Moreover, it is not a narrow field of surgery. Today's liver surgeons need to master a variety of techniques and tactics: open liver surgery, laparoscopic surgery, endoscopic surgery, and percutaneous surgeries (minimal access surgery).

The two branches, liver surgery and liver transplantation, overlap:

- Techniques of liver transplantation in liver surgery: External bypass in extensive resections, cooling (topical, hypothermia, *ex-situ* resections).
- Techniques of liver surgery in liver transplantation: Reduced size graft, split liver, living donor.

Today's liver surgeon needs to be familiar with anatomy, echography, vascular surgery, microsurgery, immunology, hepatology, and oncology.

THE PRESENT

Improving the technique of liver resection and increasing the number of patients eligible for liver resection are the main objectives because the only way to cure a patient, in cases of liver tumors, is to offer him or her the chance of a liver resection.

There are two main ways to render a patient eligible for liver resection: by changing the tumor

size through chemotherapy or changing the remaining liver size by performing portal vein embolization.⁷⁵⁻⁸⁰

In cases of multiple, bilateral, unresectable colorectal liver metastases, the strategy must be multimodal, starting with neoadjuvant chemotherapy. In patients with a normal liver, portal vein embolization (PVE) is indicated when the future liver remnant volume is predicted to be <30%.⁸¹⁻⁹⁵ PVE may also be useful in patients who have evidence of chemotherapy-related liver injury. If there is a positive response to chemotherapy, then surgery should follow. Limited resection of the colorectal liver metastases in one lobe can be performed as the first stage combined with radio frequency ablation of other lesions and PVE of the contralateral portal vein. Alternatively, ligation and alcoholization of the portal vein can be performed intraoperatively. PVE, ligation, and/or alcoholization produce, on the one hand, essential atrophy of the implicated lobe and, on the other hand, significant hypertrophy of the contralateral lobe. This strategy enhances the volume of the future liver remnant. Studies have shown an approximately 10%–25% increase in the size of the liver remnant after PVE.⁸¹⁻⁹⁵ The hypertrophy is complete by the third week. In a second stage, a major resection of the contralateral liver can be carried out safely.⁹⁶⁻⁹⁸ Usually, patients undergoing these procedures receive postoperative chemotherapy. With this approach, currently there are no limits to the resection other than those imposed by the volume and function of the liver remnant.

Is laparoscopic liver surgery an abdominal approach or a technical improvement? It is too early to compare laparoscopic and open surgery since there are much fewer laparoscopic liver resections reported^{99,100} than open liver resections. Many thousands of open liver resections are performed every year. Laparoscopic liver surgery is feasible and safe for certain indications, e.g. left lateral segmentectomy for benign or malignant disease.¹⁰¹⁻¹⁰⁶ Harvesting of the left lateral segment for living related transplantation is done by laparoscopy by some expert surgeons (experts in liver surgery, liver transplantation, and laparoscopy) with excellent results.¹⁰⁷

There are some reports on robotic liver resections, though these are still preliminary descriptions.¹⁰⁸

THE FUTURE

The future of liver surgery and new developments in the field are moving in a multidisciplinary direction. Today, oncologists, hepatologists, surgeons, endoscopists, radiologists, pathologists, and researchers are all members of the team treating patients. Their joint efforts are intended to give patients the best and most efficient treatment. In the coming years, the positive results of this cooperation should lead to new and successful treatment procedures.

REFERENCES

1. Langenbuch C. Ein Fall von Resektion eines linksseitigen Schnurlappens der Leber. *Berl Klin Wochenschr* 1888;25:37. German
2. Quattlebaum JK, Quattlebaum JK Jr. Technic of hepatic lobectomy. *Ann Surg* 1959;149:648–51. [doi:10.1097/0000658-195905000-00005](https://doi.org/10.1097/0000658-195905000-00005)
3. Fineberg C, Goldburgh WP, Templeton JY 3rd. Right hepatic lobectomy for primary carcinoma of the liver. *Ann Surg* 1956;144:881–92. [doi:10.1097/0000658-195611000-00013](https://doi.org/10.1097/0000658-195611000-00013)
4. Lortat-Jacob JL, Robert HG. Well defined technic for right hepatectomy. *Presse Med* 1952;60:549–51. French
5. Couinaud C. Lobes et segments hépatiques. *Presse Med* 1954;62:709. French
6. Couinaud C. *Le foie*. Paris: Masson; 1957. French
7. Stellamor K. [About the advantages of the sonographic diagnostic in the field of the radiologist (author's transl)]. *Rontgenblätter* 1976;29:544–52. German
8. Gosink BB. Evaluation of hepatic neoplasms. *AJR AM J Roentgenol* 1980;134:621.
9. Bismuth H. Surgical anatomy and anatomical surgery of the liver. *World J Surg* 1982;6:3–9. [doi:10.1007/BF01656368](https://doi.org/10.1007/BF01656368)
10. Makuuchi M, Hasegawa H, Yamazaki S. Intraoperative ultrasonic examination for hepatectomy. *Ultrasound Med Biol* 1983;Suppl 2:493–7.
11. Makuuchi M, Hasegawa H, Yamazaki S. Development on segmentectomy and subsegmentectomy of the liver due to introduction of ultrasonography. *Nippon Geka Gakkai Zasshi* 1983;84:913–7. Japanese
12. Bismuth H, Castaing D. Peroperative echography in hepatobiliary surgery. *Ann Gastroenterol Hepatol (Paris)* 1984;20:221–3. French
13. Bismuth H, Castaing D, Kunstlinger F. Peroperative echography in hepatobiliary surgery. *Presse Med* 1984;13:1819–22. French
14. Bismuth H, Houssin D, Castaing D. Major and minor segmentectomies “régées” in liver surgery. *World J Surg* 1982;6:10–24. [doi:10.1007/BF01656369](https://doi.org/10.1007/BF01656369)
15. Neuhaus W. [Intraoperative manometry and roentgenography of biliary system and its significance for surgery of the bile ducts]. *Langenbecks Arch Klin Chir Ver Dtsch Z Chir* 1953;275:395–403. [doi:10.1007/BF01438628](https://doi.org/10.1007/BF01438628)
16. Lin TY, Tsu K, Mien C, Chen C. Study on lobectomy of the liver. *J Formosa Med Assoc* 1958;57:742–9.
17. Lin TY. A simplified technique for hepatic resection: the crush method. *Ann Surg* 1974;180:285–90. [doi:10.1097/0000658-197409000-00005](https://doi.org/10.1097/0000658-197409000-00005)
18. Lin TY. Results in 107 hepatic lobectomies with a preliminary report on the use of a clamp to reduce blood loss. *Ann Surg* 1973;177:413–21. [doi:10.1097/0000658-197304000-00006](https://doi.org/10.1097/0000658-197304000-00006)
19. Pamecha V, Gurusamy KS, Sharma D, Davidson BR. Techniques for liver parenchymal transection: a meta-analysis of randomized controlled trials. *HPB (Oxford)* 2009;11:275–81.
20. Rau HG, Wichmann MW, Schinkel S, et al. [Surgical techniques in hepatic resections: Ultrasonic aspirator versus Jet-Cutter. A prospective randomized clinical trial]. *Zentralbl Chir* 2001;126:586–90. German [doi:10.1055/s-2001-16573](https://doi.org/10.1055/s-2001-16573)
21. Pringle JHV. Notes on the arrest of hepatic hemorrhage due to trauma. *Ann Surg* 1908;48:541–9. [doi:10.1097/0000658190810000-00005](https://doi.org/10.1097/0000658190810000-00005)
22. Man K, Fan ST, Ng IO, Lo CM, Liu CL, Wong J. Prospective evaluation of Pringle maneuver in hepatectomy for liver tumors by a randomized study. *Ann Surg* 1997;226:704–13. [doi:10.1097/0000658-199712000-00007](https://doi.org/10.1097/0000658-199712000-00007)
23. Man K, Fan ST, Ng IO, et al. Tolerance of the liver to intermittent pringle maneuver in hepatectomy for liver tumors. *Arch Surg* 1999;134:533–9. [doi:10.1001/archsurg.134.5.533](https://doi.org/10.1001/archsurg.134.5.533)
24. Lazorthes F, Chiotasso P, Chevreau P, Materre JP,

- Roques J. Hepatectomy with initial suprahepatic control of intrahepatic portal pedicles. *Surgery* 1993;113:103–8.
25. Launois B, Jamieson GG. The posterior intrahepatic approach for hepatectomy or removal of segments of the liver. *Surg Gynecol Obstet* 1992;174:155–8.
 26. Figueras J, Lopez-Ben S, Llado L, et al. Hilar dissection versus the “glissonian” approach and stapling of the pedicle for major hepatectomies: a prospective, randomized trial. *Ann Surg* 2003;238:111–9. [doi:10.1097/00006582003070000015](https://doi.org/10.1097/00006582003070000015)
 27. Shimamura Y, Gunven P, Takenaka Y, et al. Selective portal branch occlusion by balloon catheter during liver resection. *Surgery* 1986;100:938–41.
 28. Castaing D, Garden OJ, Bismuth H. Segmental liver resection using ultrasound-guided selective portal venous occlusion. *Ann Surg* 1989;210:20–3. [doi:10.1097/0000658-198907000-00003](https://doi.org/10.1097/0000658-198907000-00003)
 29. Bismuth H, Houssin D, Castaing D. Major and minor segmentectomies “reglees” in liver surgery. *World J Surg* 1982;6:10–24. [doi:10.1007/BF01656369](https://doi.org/10.1007/BF01656369)
 30. Capussotti L, Muratore A, Ferrero A, Massucco P, Ribero D, Polastri R. Randomized clinical trial of liver resection with and without hepatic pedicle clamping. *Br J Surg* 2006;93:685–9. [doi:10.1002/bjs.5301](https://doi.org/10.1002/bjs.5301)
 31. Nuzzo G, Giuliani F, Giovannini I, Vellone M, De Cosmo G, Capelli G. Liver resections with or without pedicle clamping. *Am J Surg* 2001;181:238–46. [doi:10.1016/S0002-9610\(01\)00555-4](https://doi.org/10.1016/S0002-9610(01)00555-4)
 32. Smyrniotis VE, Kostopanagiotou GG, Contis JC, et al. Selective hepatic vascular exclusion versus Pringle maneuver in major liver resections: prospective study. *World J Surg* 2003;27:765–9. [doi:10.1007/s00268-003-6978-8](https://doi.org/10.1007/s00268-003-6978-8)
 33. Cherqui D, Malassagne B, Colau PI, Brunetti F, Rotman N, Fagniez PL. Hepatic vascular exclusion with preservation of the caval flow for liver resections. *Ann Surg* 1999;230:24–30. [doi:10.1097/0000658-199907000-00004](https://doi.org/10.1097/0000658-199907000-00004)
 34. Huguet C, Gallot D, Offenstadt G, Coloigner M. [Total vascular exclusion of the liver in extensive hepatic exeresis. Value and limits]. *Nouv Presse Med* 1976;5:1189–92. French
 35. Bismuth H, Castaing D, Garden OJ. Major hepatic resection under total vascular exclusion. *Ann Surg* 1989;210:13–19. [doi:10.1097/0000658-198907000-00002](https://doi.org/10.1097/0000658-198907000-00002)
 36. Offenstadt G, Huguet C, Gallot D, Bloch P. Hemodynamic monitoring during complete vascular exclusion for extensive hepatectomy. *Surg Gynecol Obstet* 1978;146:709–13.
 37. Huguet C, Nordlinger B, Galopin JJ, Bloch P, Gallot D. Normothermic hepatic vascular exclusion for extensive hepatectomy. *Surg Gynecol Obstet* 1978;147:689–93.
 38. Maeba T, Okano K, Mori S, et al. Retrohepatic vena cava replacement of hepatic malignancies without using total hepatic vascular exclusion or extracorporeal bypass. *Hepatogastroenterology* 2001;48:1455–60.
 39. Ohwada S, Kawashima Y, Ogawa T, et al. Extended hepatectomy with ePTFE graft vena caval replacement and hepatic vein reconstruction: a case report. *Hepatogastroenterology* 1999;46:1151–5.
 40. Fortner JG, Shiu MH, Kinne DW, et al. Major hepatic resection using vascular isolation and hypothermic perfusion. *Ann Surg* 1974;180:644–652. [doi:10.1097/0000658-197410000-00030](https://doi.org/10.1097/0000658-197410000-00030)
 41. Azoulay D, Eshkenazy R, Andreani P, et al. In situ hypothermic perfusion of the liver versus standard total vascular exclusion for complex liver resection. *Ann Surg* 2005;241:277–85. [doi:10.1097/01.sla.0000152017.62778.2f](https://doi.org/10.1097/01.sla.0000152017.62778.2f)
 42. Delrivière L, Hannoun L. In situ and ex situ procedures for complex major liver resections requiring prolonged hepatic vascular exclusion in normal and diseased livers. *J Am Coll Surg* 1995;181:272–6.
 43. Hannoun L, Balladur P, Delva E, et al. “Ex situ-in vivo” surgery of the liver: a new technique in liver surgery. Principles and preliminary results. *Gastroenterol Clin Biol* 1991;15:758–61. French
 44. Hannoun L, Borie D, Balladur P, et al. [Ex situ-in vivo hepatic resection. Technique and initial results]. *Chirurgie* 1992;118:292–6; discussion 296–7. French.
 45. Hannoun L, Panis Y, Balladur P, et al. Ex-situ in vivo liver surgery. *Lancet* 1991;337:1616–7. [doi:10.1016/0140-6736\(91\)93321-Y](https://doi.org/10.1016/0140-6736(91)93321-Y)
 46. Pichlmayr R, Grosse H, Hauss J, et al. Technique and preliminary results of extracorporeal liver surgery (bench procedure) and of surgery of the in situ perfused liver. *Br J Surg* 1990;77:21–6. [doi:10.1002/bjs.1800770107](https://doi.org/10.1002/bjs.1800770107)

47. Gertsch P, Pelloni A, Guerra A, Krpo A. Initial experience with the harmonic scalpel in liver surgery. *Hepatogastroenterology* 2000;47:763-6.
48. Schmidbauer S, Hallfeldt KK, Sitzmann G, Kantelhardt T, Trupka A. Experience with ultrasound scissors and blades (UltraCision) in open and laparoscopic liver resection. *Ann Surg* 2002;235:27-30. [doi:10.1097/0000658-200201000-00004](https://doi.org/10.1097/0000658-200201000-00004)
49. Aldrighetti L, Pulitanò C, Arru M, Catena M, Finazzi R, Ferla G. "Technological" approach versus clamp crushing technique for hepatic parenchymal transection: a comparative study. *J Gastrointest Surg* 2006;10:974-9. [doi:10.1016/j.gassur.2006.02.002](https://doi.org/10.1016/j.gassur.2006.02.002)
50. Saiura A, Yamamoto J, Koga R, Seki M, Yamaguchi T. Liver transection using the LigaSure sealing system. *HPB (Oxford)* 2008;10:239-43.
51. Ikeda M, Hasegawa K, Sano K, et al. The vessel sealing system (LigaSure) in hepatic resection: a randomized controlled Trial. *Ann Surg* 2009;250:199-203. [doi:10.1097/SLA.0b013e3181a334f9](https://doi.org/10.1097/SLA.0b013e3181a334f9)
52. Nanashima A, Tobinaga S, Abo T, Nonaka T, Sawai T, Nagayasu T. Usefulness of the combination procedure of crash clamping and vessel sealing for hepatic resection. *J Surg Oncol* 2010;102:179-83. [doi:10.1002/jso.21575](https://doi.org/10.1002/jso.21575)
53. Poon RT, Fan ST, Wong J. Liver resection using a saline-linked radiofrequency dissecting sealer for transection of the liver. *J Am Coll Surg* 2005;200:308-13. [doi:10.1016/j.jamcollsurg.2004.10.008](https://doi.org/10.1016/j.jamcollsurg.2004.10.008)
54. Stella M, Percivale A, Pasqualini M, et al. Radiofrequency-assisted liver resection. *J Gastrointest Surg* 2003;7:797-801. [doi:10.1016/S1091-255X\(03\)00137-9](https://doi.org/10.1016/S1091-255X(03)00137-9)
55. Poon RT, Ng KK, Lam CM, et al. Learning curve for radiofrequency ablation of liver tumors: prospective analysis of initial 100 patients in a tertiary institution. *Ann Surg* 2004;239:441-9. [doi:10.1097/01.sla.0000118565.21298.0a](https://doi.org/10.1097/01.sla.0000118565.21298.0a)
56. Weber JC, Navarra G, Jiao LR, Nicholls JP, Jensen SL, Habib NA. New technique for liver resection using heat coagulative necrosis. *Ann Surg* 2002;236:560-3. [doi:10.1097/0000658200211000-00004](https://doi.org/10.1097/0000658200211000-00004)
57. Ferko A, Lesko M, Subrt Z, et al. A modified radiofrequency-assisted approach to right hemihepatectomy. *Eur J Surg Oncol* 2006;32:1209-11. [doi:10.1016/j.ejso.2006.07.013](https://doi.org/10.1016/j.ejso.2006.07.013)
58. Jarnagin WR, Gonen M, Fong Y, et al. Improvement in perioperative outcome after hepatic resection: analysis of 1,803 consecutive cases over the past decade. *Ann Surg* 2002;236:397-406. [doi:10.1097/0000658-200210000-00001](https://doi.org/10.1097/0000658-200210000-00001)
59. Imamura H, Seyama Y, Kokudo N, et al. One thousand fifty-six hepatectomies without mortality in 8 years. *Arch Surg* 2003;138:1198-206. [doi:10.1001/archsurg.138.11.1198](https://doi.org/10.1001/archsurg.138.11.1198)
60. Sun HC, Qin LX, Lu L, et al. Randomized clinical trial of the effects of abdominal drainage after elective hepatectomy using the crushing clamp method. *Br J Surg* 2006;93:422-66. [doi:10.1002/bjs.5260](https://doi.org/10.1002/bjs.5260)
61. Lesurtel M, Selzner M, Petrowsky H, McCormack L, Clavien PA. How should transection of the liver be performed? A prospective randomized study in 100 consecutive patients: comparing four different transection strategies. *Ann Surg* 2005;242:814-22. [doi:10.1097/01.sla.0000189121.35617.d7](https://doi.org/10.1097/01.sla.0000189121.35617.d7)
62. Mullin EJ, Metcalfe MS, Maddern GJ. How much liver resection is too much? *Am J Surg* 2005;190:87-97. [doi:10.1016/j.amjsurg.2005.01.043](https://doi.org/10.1016/j.amjsurg.2005.01.043)
63. Asiyanbola B, Chang D, Gleisner AL, et al. Operative mortality after hepatic resection: are literature-based rates broadly applicable? *J Gastrointest Surg* 2008;12:842-51. [doi:10.1007/s11605-008-0494-y](https://doi.org/10.1007/s11605-008-0494-y)
64. Dimick JB, Cowan JA Jr, Knol JA, Upchurch GR Jr. Hepatic resection in the United States, indications, outcomes, and hospital procedural volumes from a nationally representative database. *Arch Surg* 2003;138:185-91. [doi:10.1001/archsurg.138.2.185](https://doi.org/10.1001/archsurg.138.2.185)
65. Almersjö O, Bengmark S, Hafström L. Liver resection for cancer. *Acta Chir Scand* 1976;142:139-44.
66. Houssin D, Franco D, Berthelot P, Bismuth H. Heterotopic liver transplantation in end-stage HBsAg-positive cirrhosis. *Lancet* 1980;1:990-3. [doi:10.1016/S0140-6736\(80\)91435-X](https://doi.org/10.1016/S0140-6736(80)91435-X)
67. Bismuth H, Houssin D. Reduced-sized orthotopic liver graft in hepatic transplantation in children. *Surgery* 1984;95:367-70.
68. Brölsch CE, Stevens LH, Whittington PF. The use of reduced-size liver transplants in children, including split livers and living related liver transplants. *Eur J Pediatr Surg* 1991;1:166-71. [doi:10.1055/s-2008-1042480](https://doi.org/10.1055/s-2008-1042480)
69. Azoulay S, Samuel D, Castaing D, et al. Domino liver transplants for metabolic disorders: experience

- rience with familial amyloidotic polyneuropathy. *J Am Coll Surg* 1999;189:584–93. doi:10.1016/S1072-7515(99)00208-2
70. Figueras J, Parés D, Munar-Ques M, et al. Experience with domino or sequential liver transplantation in patients with familial amyloidotic polyneuropathy. *Transplant Proc* 2002;34:307–8. doi:10.1016/S0041-1345(01)02776-2
71. Bismuth H, Samuel D, Castaing D, et al. Orthotopic liver transplantation in fulminant and subfulminant hepatitis. The Paul Brousse experience. *Ann Surg* 1995;222:109–19. doi:10.1097/0000658-199508000-00002
72. Bismuth H, Morino M, Castaing D, et al. Emergency orthotopic liver transplantation in two patients using one donor liver. *Br J Surg* 1989;76:722–4. doi:10.1002/bjs.1800760723
73. Azoulay D, Astarcioglu I, Bismuth H, et al. Split-liver transplantation. The Paul Brousse policy. *Ann Surg* 1996;224:737–46; discussion 746–8. doi:10.1097/0000658-199612000-00009
74. Bismuth H, Azoulay D, Samuel D, et al. Auxiliary partial orthotopic liver transplantation for fulminant hepatitis. The Paul Brousse experience. *Ann Surg* 1996;224:712–24; discussion 724–6. doi:10.1097/0000658-199612000-00007
75. Adam R, Avisar E, Ariche A, et al. Five-year survival following hepatic resection after neoadjuvant therapy for nonresectable colorectal. *Ann Surg Oncol* 2001;8:347–53. doi:10.1007/s10434-001-0347-3
76. Bismuth E, Adam R, Lévi F, et al. Resection of nonresectable liver metastases from colorectal cancer after neoadjuvant chemotherapy. *Ann Surg* 1996;224:509–20. doi:10.1097/0000658-199610000-00009
77. Adam R, Delvart V, Pascal G, et al. Rescue surgery for unresectable colorectal liver metastases downstaged by chemotherapy: a model to predict long term survival. *Ann Surg* 2004;240:644–57; discussion 657–8.
78. Folprecht G, Gruenberger T, Bechstein WO, et al. Tumour response and secondary resectability of colorectal liver metastases following neoadjuvant chemotherapy with cetuximab: the CELIM randomised phase 2 trial. *Lancet Oncol* 2010;11:38–47. doi:10.1016/S1470-2045(09)70330-4
79. Min BS, Kim NK, Ahn JB, et al. Cetuximab in combination with 5-fluorouracil, leucovorin and irinotecan as a neoadjuvant chemotherapy in patients with initially unresectable colorectal liver metastases. *Onkologie* 2007;30:637–43. doi:10.1159/000109957
80. Pozzo C, Barone C, Kemeny NE. Advances in neoadjuvant therapy for colorectal cancer with liver metastases. *Cancer Treat Rev* 2008;34:293–301. doi:10.1016/j.ctrv.2008.01.004
81. Kawasaki S, Makuuchi M, Kakazu T, et al. Resection for multiple metastatic liver tumors after portal embolization. *Surgery* 1994;115:674–7.
82. Madoff D, Abdalla E, Vauthey J. Portal vein embolization in preparation for major hepatic resection: evolution of a new standard of care. *J Vasc Interv Radiol* 2005;16:779–90.
83. Elias D, Santoro R, Ouellet J, Osmak L, de Baere T, Roche A. Simultaneous percutaneous right portal vein embolization and left liver tumor radiofrequency ablation prior to a major right hepatic resection for bilateral colorectal metastases. *Hepatogastroenterology* 2004;51:1788–91.
84. Hemming A, Reed A, Howard R, et al. Preoperative portal vein embolization for extended hepatectomy. *Ann Surg* 2003;237:686–91. doi:10.1097/0000658-200305000-00011
85. Elias D, Ouellet J, De Baere T, Lasser P, Roche A. Preoperative selective portal vein embolization before hepatectomy for liver metastases: long-term results and impact on survival. *Surgery* 2002;131:294–9. doi:10.1067/msy.2002.120234
86. Azoulay D, Castaing D, Smail A, et al. Resection of nonresectable liver metastases from colorectal cancer after percutaneous portal vein embolization. *Ann Surg* 2000;231:480–6. doi:10.1097/0000658-200004000-00005
87. Shimada H, Tanaka K, Masui H, et al. Results of surgical treatment for multiple (>or=5 nodules) bi-lobar hepatic metastases from colorectal cancer. *Langenbecks Arch Surg* 2004;389:114–21. doi:10.1007/s00423-003-0447-6
88. Akasu T, Moriya Y, Takayama T. A pilot study of multimodality therapy for initially unresectable liver metastases from colorectal carcinoma: hepatic resection after hepatic arterial infusion chemotherapy and portal embolization. *Jpn J Clin Oncol* 1997;27:331–5. doi:10.1093/jjco/27.5.331
89. Broering D, Hillert C, Krupski G, et al. Portal vein embolization vs. portal vein ligation for induction of hypertrophy of the future liver remnant. *J Gastrointest Surg* 2002;6:905–13. doi:10.1016/S1091-255X(02)00122-1

90. Takayama T, Makuuchi M. Preoperative portal vein embolization: is it useful? *J Hepatobiliary Pancreat Surg* 2004;11:17–20. [doi:10.1007/s00534-002-0800-0](https://doi.org/10.1007/s00534-002-0800-0)
91. Abdalla EK, Barnett CC, Doherty D, Curley SA, Vauthey JN. Extended hepatectomy in patients with hepatobiliary malignancies with and without preoperative portal vein embolization. *Arch Surg* 2002;137:675–80. [doi:10.1001/archsurg.137.6675](https://doi.org/10.1001/archsurg.137.6675)
92. Wakabayashi H, Ishimura K, Okano K, et al. Application of preoperative portal vein embolization before major hepatic resection in patients with normal or abnormal parenchyma. *Surgery* 2002;131:26–33. [doi:10.1067/msy.2002.118259](https://doi.org/10.1067/msy.2002.118259)
93. Adam R, Lucidi V, Bismuth H. Hepatic colorectal metastases: methods of improving resectability. *Surg Clin North Am* 2004;84:659–71. [doi:10.1016/j.suc.2003.12.005](https://doi.org/10.1016/j.suc.2003.12.005)
94. Goere D, Farges O, Leporrier J, Sauvanet A, Vilgrain V, Belghiti J. Chemotherapy does not impair hypertrophy of the left liver after right portal vein obstruction. *J Gastrointest Surg* 2006;10:365–70. [doi:10.1016/j.gassur.2005.09.001](https://doi.org/10.1016/j.gassur.2005.09.001)
95. Beal I, Anthony S, Papadopoulou A, et al. Portal vein embolization prior to hepatic resection for colorectal liver metastases and the effects of periprocedure chemotherapy. *Br J Radiol* 2006;79:473–8. [doi:10.1259/bjr/29855825](https://doi.org/10.1259/bjr/29855825)
96. Jaeck D, Bachellier P, Nakano H, et al. One or two-stage hepatectomy combined with portal vein embolization for initially nonresectable colorectal liver metastases. *Am J Surg* 2003;185:221–9. [doi:10.1016/S0002-9610\(02\)01373-9](https://doi.org/10.1016/S0002-9610(02)01373-9)
97. Adam R, Laurent A, Azoulay D, Castaing D, Bismuth H. Two stage hepatectomy: a planned strategy to treat unresectable liver tumors. *Ann Surg* 2000;232:777–85. [doi:10.1097/0000658-200012000-00006](https://doi.org/10.1097/0000658-200012000-00006)
98. Jaeck D, Oussoultzoglou E, Rosso E, Greget M, Weber JC, Bachellier P. A two-stage hepatectomy procedure combined with portal vein embolization to achieve curative resection for initially unresectable multiple and bilobar colorectal liver metastases. *Ann Surg* 2004;240:1037–51. [doi:10.1097/01.sla.0000145965.86383.89](https://doi.org/10.1097/01.sla.0000145965.86383.89)
99. Nguyen KT, Laurent A, Dagher I, et al. Minimally invasive liver resection for metastatic colorectal cancer: a multi-institutional, international report of safety, feasibility, and early outcomes. *Ann Surg* 2009;250:842–8. [doi:10.1097/SLA.0b013e3181bc789c](https://doi.org/10.1097/SLA.0b013e3181bc789c)
100. Nguyen KT, Gamblin TC, Geller DA. World review of laparoscopic liver resection – 2,804 patients. *Ann Surg* 2009;250:831–41. [doi:10.1097/SLA.0b013e3181b0c4df](https://doi.org/10.1097/SLA.0b013e3181b0c4df)
101. Cherqui D, Husson E, Hammoud R, et al. Laparoscopic liver resections: a feasibility study in 30 patients. *Ann Surg* 2000;232:753–762. [doi:10.1097/0000658-200012000-00004](https://doi.org/10.1097/0000658-200012000-00004)
102. Cherqui D. Laparoscopic liver resection. *Br J Surg* 2003;90:644–6. [doi:10.1002/bjs.4197](https://doi.org/10.1002/bjs.4197)
103. Vibert E, Perniceni T, Levard H, Denet C, Shahri NK, Gayet B. Laparoscopic liver resection. *Br J Surg* 2006;93:67–72. [doi:10.1002/bjs.5150](https://doi.org/10.1002/bjs.5150)
104. Gayet B, Cavaliere D, Vibert E, et al. Totally laparoscopic right hepatectomy. *Am J Surg* 2007;194:685–9. [doi:10.1016/j.amjsurg.2006.11.044](https://doi.org/10.1016/j.amjsurg.2006.11.044)
105. Gumbs AA, Gayet B, Gagner M. Laparoscopic liver resection: when to use the laparoscopic stapler device. *HPB (Oxford)* 2008;10:296–303.
106. Bryant R, Laurent A, Tayar C, Cherqui D. Laparoscopic liver resection-understanding its role in current practice: the Henri Mondor Hospital experience. *Ann Surg* 2009;250:103–11. [doi:10.1097/SLA.0b013e3181ad6660](https://doi.org/10.1097/SLA.0b013e3181ad6660)
107. Soubrane O, Cherqui D, Scatton O, et al. Laparoscopic left lateral sectionectomy in living donors: safety and reproducibility of the technique in a single center. *Ann Surg* 2006;244:815–20. [doi:10.1097/01.sla.0000218059.31231.b6](https://doi.org/10.1097/01.sla.0000218059.31231.b6)
108. Idrees K, Bartlett DL. Robotic liver surgery. *Surg Clin North Am* 2010;90:761–74. [doi:10.1016/j.suc.2010.04.020](https://doi.org/10.1016/j.suc.2010.04.020)