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**First quarter century of laparoscopic liver resection**

Morise Z *et al.* Quarter century of laparoscopic liver resection

Zenichi Morise, Go Wakabayashi

**Zenichi Morise,** Department of Surgery, Fujita Health University School of Medicine, Toyoake, Aichi 470-1192, Japan

**Go Wakabayashi,** Center for Advanced Treatment of HBP Diseases, Department of Surgery, Ageo Central General Hospital, Ageo, 362-8588 Saitama, Japan

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**Correspondence to: Zenichi Morise, MD, PhD, FACS, AGAF,** Department of Surgery, Fujita Health University School of Medicine, 1-98 Kutsukakecho Toyoake, Aichi 470-1192, Japan. zmorise@fujita-hu.ac.jp

**Telephone:** +81-562-939246

**Fax:** +81-562-935125

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**Abstract**

The beginnings of laparoscopic liver resection (LLR) were at the start of the 1990s, with the initial reports being published in 1991 and 1992. These were followed by reports of left lateral sectionectomy in 1996. In the years following, the procedures of LLR were expanded to hemi-hepatectomy, sectionectomy, segmentectomy and partial resection of posterosuperior segments, as well as the parenchymal preserving limited anatomical resection and modified anatomical (extended and/or combining limited) resection procedures. This expanded range of LLR procedures, mimicking the expansion of open liver resection in the past, was related to advances in both technology (instrumentation) and technical skill with conceptual changes. During this period of remarkable development, two international consensus conferences were held (2008 in Louisville, KY, United States, and 2014 in Morioka, Japan), providing up-to-date summarizations of the status and perspective of LLR. The advantages of LLR have become clear, and include reduced intraoperative bleeding, shorter hospital stay, and—especially for cirrhotic patients-lower incidence of complications (*e.g.,* postoperative ascites and liver failure). In this paper, we review and discuss the developments of LLR in operative procedures (extent and style of liver resections) during the first quarter century since its inception, from the aspect of relationships with technological/technical developments with conceptual changes.

**Key words:** Hepatectomy; Laparoscopic surgery; Liver cancer; History; Technology; Technique; Concept; Approach; Posture; Simulation

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**Core tip:** Laparoscopic liver resection (LLR) was introduced in early 1990s. Thereafter, LLR procedures have expanded to left lateral sectionectomy, hemi-hepatectomy, sectionectomy, segmentectomy and partial resection of posterosuperior segments, as well as parenchymal preserving limited and modified anatomical resection. This expansion is related to technological/technical developments with conceptual changes. During this period, two international consensus conferences summarized the up-to-date status and perspective of LLR. The current advantages of LLR include reduced intraoperative bleeding, shorter hospital stay, and lower incidence of complications. Here, we review and discuss the developments of LLR in operative procedures during the first quarter century since its inception.

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**INTRODUCTION**

Although laparoscopes were first introduced in the 1960s as diagnostic tools for urological and gynecological diseases, a new technology to create pneumoperitoneum and the development of the charged-coupled device (CCD) camera, which magnifies and projects laparoscopic images onto television monitors, led to the first laparoscopic cholecystectomy performed in the current style in 1987 by Phillipe Mouret of Lyon, France[1]. The procedure gained immediate acceptance according to its related clinical experiences of less pain and rapid recovery, in addition to the cosmetic advantages[2,3]. Since then, the field of laparoscopic surgery has expanded rapidly to include surgery for other abdominal organs and more complex and technically demanding abdominal surgery.

The beginnings of laparoscopic liver resection (LLR) were at the start of the 1990s, with the initial reports[4-6] published in 1991 and 1992. These were followed by the reports of left lateral sectionectomy (LLS)[7,8] in 1996. In the years following, the procedures of LLR were expanded to hemi-hepatectomy, sectionectomy, segmentectomy and partial resection of posterosuperior segments, as well as parenchymal preserving limited anatomical resection and modified anatomical (extended and/or combining limited) resection. This expanded range of LLR procedures, mimicking the expansion of open liver resection (OLR) in the past, was related to advances in both technology (instrumentation) and technical skill with conceptual changes (Table 1).

During this period of remarkable development, two international consensus conferences (ICCLLR) were held (2008 in Louisville, KY, United States[9] and 2014 in Morioka, Japan[10]), providing up-to-date summarizations of the status and perspective of LLR. The anxieties over LLR-specific complications, including gas-embolism, were eased by the cautious application of these procedures to and the long-term outcomes of selected patients for LLR, which were confirmed as similar to those for OLR. The advantages of LLR became clearly established, in particular, reduced intraoperative bleeding, shorter hospital stay, and—especially for cirrhotic patients—lower incidence of complications (*e.g.,* postoperative ascites and liver failure).

In this review of the developments of LLR in operative procedures (extent and style of liver resections) that have occurred during the first quarter century since its inception, we discuss the relationships of these advances in technological/technical aspects of LLR with conceptual changes.

**DEVELOPMENT OF LLR**

***Partial resection of anterolateral segments and LLS: The beginnings of LLR***

The initial reports of LLR by Reich *et al*[4], Katkhouda *et al*[5], Gagner *et al*[6] appeared in 1991 and 1992. These were followed by reports of LLS by Azagra *et al*[7] and Kaneko *et al*[8] in 1996. Although segment level Glissonian pedicles and thick hepatic veins should be divided in LLS, the lesions located in the anterolateral segments (segments 2, 3, 4b, 5, 6) are more accessible laparoscopically than those in the posterosuperior segments (1, 4a, 7, 8). Also, the relatively small transection plane of LLS lies in a caudal-to-cranial direction and is vertical when the patient is in supine position, making it easier to handle in the natural laparoscopic view and to access with ports below the costal-arch level. Therefore, LLS is a big partial resection of anterolateral segments in some aspects; indeed, the first development of the LLR procedure involved anterolateral partial resection to LLS. LLS is the most straightforward sectionectomy procedure, as in OLR, and the standardization of this procedure has emerged recently as a topic of considerable discussion[11,12].

***Hemi-hepatectomy and feasibility studies***

The first report of hemi-hepatectomy was in 1997 by Hüscher *et al*[13], just 1 year after the LLS reports. The transection plane of hemi-hepatectomy, like the one in LLS, lies in the caudal-to-cranial direction and is vertical in supine position, making it easier to handle *via* the laparoscopic approach. Hemi-hepatectomies are the second-most straightforward procedure, again as in OLR, after anterolateral partial resection and LLS[14,15]. However, stable transection maneuvers were required in this step of development, since the transection plane in hemi-hepatectomies is a large area. Advances in technologies and instrumentation contributed to the step[16,17]. In the early stage of LLR development, pre-transectional coagulation *via* coagulating energy devices proved important in reducing the possibility of intra-operative massive bleeding. Development of transection maneuvers that mimic open maneuvers, such as crash-clamp transection and Cavitron ultrasonic surgical aspirator (CUSA; or its equivalent) transection, was accomplished by adaptation of various energy devices (to achieve coagulation, sealing and shearing) and laparoscopic CUSA, accompanied by inflow control[17,18]. Differences exist between the right and left hemi-hepatectomy forms of the major hepatectomies, these specifically involve mobilization of the liver and handling of the caudate lobe and IVC. The mobilization procedure for the left liver is relatively straightforward, except for the dissection of the roots of the middle and left hepatic veins. Also, when it is performed without resection of the Spiegel lobe, there is no need for dissection of the IVC. On the other hand, the right hemi-hepatectomy is usually performed with resection of the para-caval caudate lobe and, therefore, necessitates dissection of the IVC and right adrenal gland. During mobilization of the right liver, handling of the heavy and large-volume right liver is also much more demanding, complicating the laparoscopic surgical procedure which occurs without the surgeon’s hands being present in the operative field. As such, the procedure of laparoscopic right hemi-hepatectomy has developed more slowly than that of left[13-15].

During this and the next step of development, the hand-assisted (HALS) procedure and hybrid (laparoscopic-assisted) procedure helped to reduce the technical difficulty of LLR in pure laparoscopic setting[19-22]. Also during this step of development, an encouraging feasibility study of LLR—including left hemi-hepatectomy, LLS, segementectomy and partial resection of segments 3, 4, 5, 6—was reported by Cherqui *et al*[23] in 2000. This report concluded, “Laparoscopic resections are feasible and safe in selected patients with left-sided and right-peripheral lesions requiring limited resection.”

***Left medial, right anterior and posterior sectionectomies***

In the summary paper from the first ICCLLR[9], LLR was divided into the following three categories: I, small wedge resections; II, resections of the left lateral section or anterior segments (4b, 5, 6); III, hemi-hepatectomies, trisectionectomies and resections of posterior segments (4a, 7, 8). Category III was referred to as “major LLR”. The section on major LLR in this summary paper concluded, “Major LLR have been performed with safety and efficacy equaling OLR in highly specialized centers.” Also, in the section on hepatocellular carcinoma (HCC) treatment and LLR in this summary paper, anatomic segmental resection was recommended, instead of non-anatomical partial resection, due to the related lower rates of local recurrence. Especially for those patients with HCC and chronic liver diseases (CLDs), laparoscopic left medial, right anterior and right posterior sectionectomies were recommended as the next-step procedures after hemi-hepatectomy, in order to accomplish the preservation of residual liver function and to maintain oncological efficacy equal to that of OLR[24].

The transection planes in sectionectomies are larger in area and more difficult to handle than those in hemi-hepatectomies. Also, hilar dissection with individual vessel preparation for processing territorial vessels cannot be performed in this level. Although the Glissonian approach has been employed for hemi-hepatectomy[25] alongside hilar dissection with individual vessel preparation, the importance of the Glissonian approach is greater in sectionectomies and in more limited anatomical resections. Both extrahepatic[26] and intrahepatic[27] laparoscopic Glissonian approaches have been reported and employed widely, as in OLR, for this step in the development of LLR.

 On the other hand, handling of the transection plane-especially the border between the anterior and posterior sections—is one of the key obstacles for right anterior and posterior sectionectomies[28]. Since the liver is located in the subphrenic rib cage, in OLR, surgeons open the subphrenic cage with a large subcostal incision and lifting-up of the costal arch; after which, the surgeon dissects the retro-peritoneal attachments and physically picks-up the liver with his/her left hand in order to manipulate the intact organ (Figure 1A). However, in LLR, there are no instruments as good as the surgeon’s left hand and, moreover, no anterior space available without abdominal wall incision. Therefore, laparoscopic right anterior and posterior sectionectomies are technically demanding to obtain a fine surgical field that will ensure hemostasis and an appropriate surgical margin in handling the transection plane beneath the large and heavy right liver in the small subphrenic rib cage.

Postural changes have been employed to conquer this obstacle. Semi-prone[29,30] and left lateral[31] position LLR were reported as capable of allowing for acquirement of fine surgical view and manipulation for sectionectomies in the right liver. Also, a paper on lateral position posterior sectionectomy published by our group[31] described the new concept of “caudal approach in LLR” (Figure 1B); in this approach, the laparoscopic specific view and manipulation access is made from the caudal direction, using ports entering below the costal-arch level and going into the subphrenic rib cage. The summary paper of the second ICCLLR[10], explains this concept as follows: “The caudal approach, which relies on visual magnification, offers improved exposure around the right adrenal gland and the vena cava and greatly facilitates identification of the Laennec’s capsule and the Glissonian pedicle at the hilar plate.”

***Segmentectomies and partial resections of segments 7, 8 and 1***

Although LLS, segmentectomies for segments 5 and 6, and left medial sectionectomy (segment 4 segmentectomy) had been performed in the earlier stage of LLR development, segments 7, 8 and 1 remained unresolved challenge areas for segmentectomy and even for partial resection[32,33].

In LLR, and distinctive from OLR, more sectionectomies or right hepatectomies have been performed than segmentectomies or partial resections as treatment of tumors involving segments 7 and 8[9,34-36]. This trend can be explained by the fact that the straightforward transection plane of the liver, from caudal edge to the diaphragm in right hepatectomy or posterior sectionectomy, is more easily handled in LLR. In the laparoscope view from the caudal direction, the transection planes of segment 7 and 8 segmentectomies or partial resections are located in the deep small subphrenic space behind the liver, with segments 5 and 6 acting as physical obstacles to the lesions. Since surgeons need to create a precisely curved or angulated transection plane in the space, the parenchymal preserving segmentectomies or partial resections of the area are technically more difficult than performance of a posterior sectionectomy or right hepatectomy.

Adequate functional reserve of the liver after resection is as important as oncological efficacy, especially in impaired livers, as encountered in CLD patients with HCC[24]. An important consideration for LLR of this area, therefore, is how to obtain good and stable access that allow for sufficient and safe handling of the liver and tumors, so that a well-visualized transection plane can be acquired. To this end, intercostal (transdiaphragmatic) ports with total mobilization of the liver from the retroperitoneum have been applied to facilitate the direct lateral approach into the rib cage (in the abdominal cavity) and to segment 7 (Figure 1C)[37-39]. In addition, the thoracoscopic approach was employed for lesions in segment 8 (Figure 1D), with direct exposure of the tumor into the pleural cavity being achieved by incision on the diaphragm adjacent to the tumor[40,41]. Endoscopes have been placed in the abdominal cavity for the lateral approach using intercostal ports (Figure 1C), and in the pleural cavity for the thoracoscopic approach (Figure 1D).

On the other hand, postural changes, such as semi-prone positioning for tumors located in segment 7[28-30], have also been applied to solve the same problem. Although segment 7 is located in the bottom of the abdominal cavity when the patient is in supine position, that same area is located almost on the top of the abdominal cavity when the patient is in semi-prone position. Adapting those postural changes allows for the weight of the liver itself to facilitate its own mobilization, ultimately providing a good and stable surgical space above the liver. Ikeda *et al*[42] applied semi-prone position LLR with the use of intercostal ports to treat tumors in the anterosuperior and posterior segments.

There are still only a few reports, all with small numbers of cases, for laparoscopic isolated resection of the caudate lobe[33,43,44]. Although further experiences are needed for segment 1 LLR, especially for the total isolated resection of caudate lobe (Spiegel lobe, caudate process and paracaval portion), the fine laparoscopic caudal view to the vena cava area and behind the hilar plate, particularly from the left side with the incision on the gastro-hepatic ligament, could facilitate LLR for this area[44].

**FUTURE PERSPECTIVES OF LLR**

During the development of the LLR procedures, disadvantages of LLR have also been recognized. The lack of three-dimensional (3D) view was overcome by the development of the 3D laparoscope[45]. However, the lack of overview in the operative field (despite the local fine magnified view) combined with the lack of tactile sensation easily leads to disorientation on the perspective of the organs, tumors and the intrahepatic structures during LLR. Therefore, intraoperative laparoscopic ultrasonography and preoperative simulation/intraoperative navigation using reconstruction of preoperative imaging scans and the intraoperative implementation of near-infrared fluorescence scans with indocyanine green have become more important and are continued to be developed[46,47]. Based on the development of the imaging techniques, parenchymal preserving limited anatomical resection and modified (extended and/or combining limited) anatomical resection are advocated[48-51]. Robotic-assisted LLR holds the promise of facilitating a more precise surgery in certain situations, such as bile duct reconstruction[47,52,53].

On the other hand, there are specific advantages in LLR, besides those advantages common to all laparoscopic surgeries. For one, improved direct exposure with magnification could be obtained under the laparoscopic specific view to the liver inside the rib cage. This allows clearer access to the surgical field without the destruction of the surrounding environments, such as collateral vessels in patients with HCC and liver cirrhosis, and without inducing compression damage on the liver parenchyma[54,55]. Pneumoperitoneum pressure during laparoscopic surgery could reduce the amount of bleeding from the hepatic vein concomitantly with inflow control[54]. This creates a very dry surgical field, with clear visualization of the detailed internal structures of the liver. After the second ICCLLR in Morioka (2014), two important studies using propensity score analysis on about 5,000 patients’ data were published, and both showed the short-term benefits of LLR without deteriorating long-term results, compared with open procedure[56,57].

In the context of these advantages, several endeavors have now been attempted with the aim of increasing the adoption rate of LLR to clinical practice. To help ensure the safe and consistent extended application of the procedure, studies aimed at determining the learning curve of LLR were published[58,59] and a difficulty scoring system[60] (*i.e*. calculated according to tumor condition, resection style and liver condition) for the appropriate selection of the patient according to the surgeon’s skill set was proposed in the second ICCLLR[10]. Randomized clinical trials are underway and two have been completed[61,62], and registries have been started in several nations and areas[63,64]. It is likely that LLR will become a more standardized procedure with wider application in the second quarter century based upon the experiences in the first quarter century.

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**Table 1 Development of laparoscopic liver resection over the first 25 years**

|  |  |
| --- | --- |
| **Year** | **Procedure[Ref]  Related developments****(technological, technical, conceptual)** |
| 1991 | 1st report of LLR[4-6] (partial resection in AL) |
| 1996 | LLS[7,8] |
| 1997 | Hemi-hepatectomy[13-15] energy devices (coagulating, sealing, shearing)CUSAHALS[19,20] and hybrid[21,22]inflow control[17,18] |
| 2000s-2010s | Sectionectomy (right posterior, right anterior, left medial)Glissonian approach(extra-[26], intra-hepatic[27])caudal approach[0,31]postural change[29-31]Segmentectomy and partial resection of segments 7, 8, 1postural change[29-31]caudal approach[10,31]lateral approach[37-39] (intercostal port)tracoscopic approach[40,41]Limited anatomical resection and modified anatomical (extended and/or combining limited) resection[48-51] Simulation and navigation[46,47] |

3D: Three-dimensional; AL: Anterolateral segments; CUSA: Cavitron ultrasonic surgical aspirator; HALS: Hand-assisted laparoscopic surgery; Hybrid: Laparoscopic-assisted LLR; LLR: Laparoscopic liver resection; LLS: Left lateral sectionectomy; Ref: Reference number in the References section.









**Figure 1 Schema of open liver resection (A), laparoscopic liver resection (regular caudal approach, B), laparoscopic liver resection (lateral approach, C) and thracoscopic liver resection (D).** Red arrows indicate the directions of view and manipulation in each approach. A: In the open approach, the subcostal cage containing the liver is opened with a large subcostal incision and instruments are used to lift the costal arch, after which the liver is dissected and mobilized (lifted) from the retroperitoneum; B: In the regular laparoscopic caudal approach, the laparoscope and forceps are placed into the subcostal cage from the caudal direction, and the surgery is performed with minimal alteration and destruction of the associated structures; C: In the laparoscopic lateral approach, the intercostal (transdiaphragmatic) ports combined with total mobilization of the liver from the retroperitoneumcan allow the direct lateral approach into the cage and to the posterosuperior tumors; D: Thoracoscopic approach is employed for lesions in segment 8, with direct exposure of the tumor into the pleural cavity upon incision on the diaphragm adjacent to the tumor, with the endoscope placed in the pleural cavity.