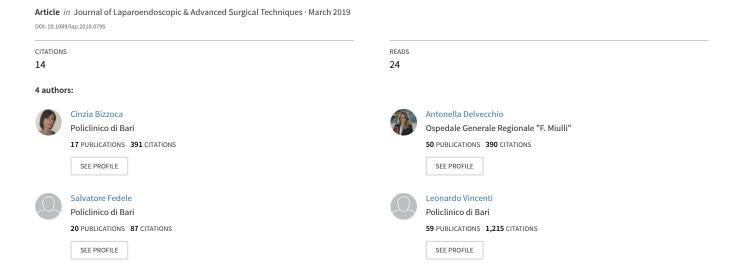
# Simultaneous Colon and Liver Laparoscopic Resection for Colorectal Cancer with Synchronous Liver Metastases: A Single Center Experience



Volume 00, Number 00, 2019 © Mary Ann Liebert, Inc. DOI: 10.1089/lap.2018.0795

# Simultaneous Colon and Liver Laparoscopic Resection for Colorectal Cancer with Synchronous Liver Metastases: A Single Center Experience

Cinzia Bizzoca, MD, Antonella Delvecchio, MD, Salvatore Fedele, MD, and Leonardo Vincenti, MD

### **Abstract**

**Background:** The one-stage approach for colorectal cancer (CRC) with synchronous liver metastases (SLM) has demonstrated advantages, when feasible, in terms of oncological radicality and reduction in sanitary costs. The simultaneous laparoscopic approach to both colon cancer and liver metastases joins the advantages of mininvasiveness to the one-stage approach.

**Methods:** During the period from February 2011 to July 2017, a single surgeon performed 17 laparoscopic colorectal operations with simultaneous liver resection for CRC with SLM. Colorectal procedures included 9 rectal resections, 6 left colectomies, and 2 right colectomies. Associated hepatic resections included 1 left hepatectomy, 1 right posterior sectionectomy, 2 segmentectomies, and 13 wedge resections. We analyzed retrospectively the patient's short-term outcome and operative and oncologic results.

**Results:** There was no conversion to open surgery. Six patients (35%) had minor complications (Clavien–Dindo grade I–II), whereas only 2 patients (12%) had major complications (Clavien–Dindo grade III–IV) and no mortality occurred. The median time of discharge was 8.6 (range 5–36) days. We obtained 94% of R0 resection margin on the liver specimen and 100% of negative distal and circumferential margin in case of rectal resection. An average of 20 lymphnodes were retrieved in the colorectal specimen.

**Conclusions:** Simultaneous mini-invasive colorectal and liver resection is a challenging but feasible procedure. The advantages of treating primary cancer and metastases in the same recovery justify the morbidity rate, especially because the most of the complications are minor and no cases of mortality occurred. Further experience is needed to better understand how to reduce the morbidity rate.

**Keywords:** simultaneous resection, colorectal resection, synchronous liver metastases, laparoscopic surgery, colorectal cancer

## Introduction

COLORECTAL CANCER (CRC) is the third most common cancer worldwide and the fourth cause of death for malignancy. At diagnosis, about 15%–25% of patients have synchronous liver metastases (SLM) and only a minority of patients has a resectable liver disease. The one-stage approach for CRC with resectable SLM has demonstrated advantages, when feasible, in terms of oncological radicality and reduction in costs, without increasing cumulative morbidity and mortality. The decision to perform a simultaneous or staged resection depends on multiple variables, such as complexity of hepatic and colorectal resection, patient's

performance status and underlying comorbidities, besides symptoms, location, and extent of disease. Surgery is still the only curative treatment for liver metastases, but nowadays an important role is played by adjuvant chemotherapy, which has improved the response and survival rates for these patients. In this scenario, a faster recovery of the patient after surgery is paramount, to favor a prompt start of chemotherapy. Laparoscopy has become a standard practice for the treatment of both colon and rectal cancer, due to the well-known advantages in terms of shorter hospital stay and faster recovery of the patient if compared to the open approach, with similar oncological outcome. All 11-17 On the contrary, the mini-invasive approach for the treatment of liver metastases

2 BIZZOCA ET AL.

has gained acceptance only in the last few years. The difficulties encountered in the development of this kind of surgery are mainly related to a steep learning curve, especially in case of major hepatectomies and posterior segments. <sup>18,19</sup> Moreover, major concerns regard aspects such as risk of bleeding, fear of gas embolism, and oncological safety. <sup>3,18,20</sup> Laparoscopic liver resections (LLR) are challenging procedures, but the association with several advantages, including decreased morbidity rate (such as wound and pulmonary complications), less need for blood transfusion, earlier recovery of the patient, and facilitation of iterative hepatic surgery, has been demonstrated. <sup>4,18,20,21</sup> Furthermore, LLR have similar on cological outcome if compared with open surgery. <sup>3,20,22–25</sup> Nevertheless, the laparoscopic synchronous approach to both CRC and liver metastases is still uncommon, as demonstrated by the limited number of patients included in most of the series published in literature. <sup>4,26–43</sup>

This retrospective study has the goal to analyze the short-term and oncological outcome of simultaneous laparoscopic resection of CRC and liver metastases on a series of 17 patients.

### **Materials and Methods**

From February 2011 to July 2017, a single surgeon L.V. performed 17 simultaneous resections of CRC with SLM. Eleven patients were males, 6 patients were females Eliminate (M/F 1,8) and the median age was 65 (range 46–88) years. Patients were preoperatively staged according to the TNM classification following colonoscopy, three-phases computed tomography of chest-abdomen-pelvis, and pelvic and/or abdominal magnetic resonance if further imaging was requested. The metastatic disease was a single liver nodule for 10 patients (59%), whereas in 7 cases (41%), multiple metastases were detected, with an average diameter of 28.8 (range 10–69) mm. Localization and characteristics of primary tumor and liver metastases at preoperative assessment are summarized in Table 1.

In agreement with the multidisciplinary team, none of the patients underwent chemotherapy or radiotherapy previous to surgery, because they suffered from severe stenosis and/or bleeding. All patients had a preserved hepatic function (Child-Pugh A), and there was no evidence of other metastases at preoperative CT scan, except in 2 cases of pulmonary nodules. Five patients (29%) had been previously operated. Preoperative assessment included evaluation of the surgical risk (ASA score) and the Eastern Cooperative Oncology Group performance status (ECOG). ASA IV patients were excluded, whereas the age itself was not considered as a contraindication if an ECOG 0-1 performance status was assigned. 44

Colorectal procedures included 2 right colectomies, 6 left colectomies, and 9 rectal resections, 5 with partial mesorectal excision (PME), 3 with total mesorectal excision (TME) and 1 Miles operation.

Hepatic resections included 1 left hepatectomy, 1 right posterior sectionectomy, 2 segmentectomies and 14 wedge resections (Table 2). We retrospectively analyzed the patient short-term outcome as primary endpoint, while operative and oncologic results were evaluated as secondary endpoints. Thirty-day mortality and morbidity rate were evaluated, according to the Clavien–Dindo classification. 45 Moreover, we

TABLE 1. PATIENTS' DEMOGRAPHICS AND PREOPERATIVE ASSESSMENT

Variable	Patient, $n=17$
Patients	
Age [mean $\pm$ SD, range (years)]	$65.2 \pm 12.5 \ (46-88)$
Gender, M/F	11/6
BMI [mean $\pm$ SD, range (kg/m <sup>2</sup> )]	$24.6 \pm 5.1 (41-19)$
Symptoms, Y/N	17/0
ASA score (I/II/III)	5/7/5
ECOG (0/1)	9/8
CHT, Y/N	0/17
RT, Y/N	0/19
Colorectal tumor	
Location	
Right colon	2
Left colon	6
Rectosigmoid junction	5 1 3
Upper rectum	1
Middle rectum	
Lower rectum	0
cTNM	0/2/1/14
T 1/2/3/4	0/2/1/14
N 0/1/2	3/4/10
Liver metastases	
N. hepatic nodules <sup>a</sup>	4.0
Single	10
Multiple	7
Location	10
Unilobar	12
Bilobar	5
Anterolateral	23
S 2/3/4b/5/6 Posterior	5/5/6/5/2 5
S 4a/7/8	1/1/3
S 44//8 Size (cm)	1/1/3
Size (ciii) ≤2	17
≥ ∠ >2	11
	11

<sup>a</sup>N. nodules 28.

ASA, American Society of Anesthesiologist physical status score; BMI, body mass index; CHT, chemotherapy; cTMN, clinical TNM stage; ECOG, Eastern Cooperative Oncology Group performance status; RT, radiotherapy; S, segment; SD, standard deviation.

analyzed postoperative transfusion rate and time of discharge. Operative results included median operative time, estimated blood loss (EBL), need for intraoperative transfusion and conversion rate. Finally, we analyzed oncological results in terms of number of harvested lymph nodes, rate of negative distal and circumferential resection margin for the primary cancer, and rate of R0 resection for the secondary tumor. Liver resections were defined as R0 if the neoplasm was microscopically more than 1 mm from resection margin. 46

# Operative technique

Trocars were positioned for the standard colorectal resection (3 accesses, 4 trocars in 1 case of rectal resection), but in 11 cases (65%) an additional trocar was introduced to allow the hepatic resection. Colorectal procedures were conducted using an ultrasound, radiofrequency or combined energy device, depending on the availability of surgical instruments during the years (Table 2). The usual technique provided a lateral-to-medial approach for left colon and

TABLE 2. SURGICAL TECHNIQUE AND OPERATIVE RESULTS

Variable	Data
No. of trocars  3 4 5	6 10 1
Colorectal resection Right colectomy Left colectomy Proctectomy Miles	2 6 8 1
Anastomosis Intracorporeal Extracorporeal No	15 1 1
Hepatic resection Wedge resection Segmentectomy Right sectionectomy Left hepatectomy	13 2 1 1
Pringle maneuver Yes No	0 17
Hemostatics Yes No	5 12
Device Electrocoagulation Ultrasound Radiofrequency Combined energy	4 3 2 8
Conversion to open Yes No	0 17
Intraoperative transfusion Yes No	2 15
Operative time (t) $t \le 120'$ $120' < t \le 180'$ $180' < t \le 240'$ t > 240'	4 7 4 2

rectum and a medial-to-lateral resection for right colectomy. Specimens were extracted through a 5 cm incision protected by an impermeable bag. The elective laparotomy was a Pfannenstiel incision, whereas only in one case a median minilaparotomy was performed, to allow an extracorporeal anastomosis after right colectomy. In all other cases, the bag was used to temporarily close the abdominal wound, pneumoperitoneum was inducted and an intracorporeal anastomosis was performed.

During hepatic resection, liver ligaments were not sectioned if not strictly necessary, with the utmost respect for anatomy allowed by the laparoscopic caudal approach to the liver. The Pringle maneuver was not routinely used, but a preparation of the hepatic pedicle for clamping was considered to be useful in case of difficult and extensive resections. Intraoperative ultrasound was always performed previous to hepatic resection

for excluding additional metastases and to mark the resection line. The transection was conducted using the device, but we opted for diathermy in 4 cases of small peripheral wedge resection. Minor vessels were divided between clips, major biliovascular structures were closed using Hem-o-lock clips or endoscopic staplers as needed, after accurate identification and isolation from the surrounding parenchyma. The vascular outflow was controlled intraparenchymally using endoscopic staplers, when the transection was almost completed. Hemostasis and biliostasis were accurately achieved using the device and single propylene stitches only when required. In 5 cases (29%) hemostasis was refined using hemostatics (Table 2). The liver specimen was extracted through the incision previously performed and protected by the bag.

### **Results**

All patients underwent a pure laparoscopic approach and the hepatic resection had a curative intent for most of them (13 cases). Two patients underwent a wedge resection in the left segments as first step of a "two stage" hepatectomy. Other 2 patients had a diagnosis of nonresectable bilobar liver metastases, but a wedge resection was requested by the oncologist to obtain a histological examination of the liver nodules for receptor analysis.

There was no 30-day mortality and a global morbidity of 47% (8 patients). Six patients (35%) had minor complications (Clavien–Dindo grade I–II), whereas only 2 patients (12%) had major complications (Clavien–Dindo grade III–IV) (Table 3). One of the patients who underwent subtotal proctectomy with TME and metastasectomy was reoperated for anastomotic leakage and needed recovery in intensive care unit because of sepsis. The other patient was discharged in postoperative day (POD) 8 after subtotal proctectomy with PME and left epatectomy, but he was readmitted 2 weeks later for biliary fistula and treated with endoscopic stent.

The median time of discharge was 8.6 (range 5–36) days. There was no conversion to open surgery. The median operative time was 165 (range 75–320) minutes. The average EBL was 158 (range 10–400) mL, but an intraoperative transfusion was necessary in 2 cases (12%) (Table 4). The rate of postoperative blood transfusion was 12% (2 patients).

We achieved R0 resection margin on the liver specimen in all cases except 1 (94%), a 100% negative circumferential and distal resection margin for the rectum, and a medium number of 20 lymph nodes was retrieved in the colorectal specimen.

Table 3. Postoperative Complications

Clavien–Dindo grade	Complications	Pt	N	
I	Pleural effusion	1	2	
	Bleeding	1		
II	Pneumonia	2	4	
	Ascites	1		
	Atrial fibrillation	1		
III	Biliary fistula	1	1	
IV	Sepsis due to intestinal leakage	1	1	

Pt, patients.

Table 4. Patients' Characteristics, Operating Procedures, Surgical, and Oncologic Outcome

RO	$Y^{a}$	Y	$Y^{a}$	Y	Y	Y	Y	$N/A^b$	X	$N/A^b$	Y	Y	>	-	¥	Z>	<del>-</del> >-
Hospital stay (days)	7	7	7	7	10	9	5	9	8		S		v	)	∞	ر د	36
Morbidity	Nil	Nil	Nil	Nil	Pneumonia	Nil	Pleural effusion	Ascites	Nii	Rectal bleeding	Atrial fibrillation	Nil	N:I	1111	Biliary fistula	Nil	rneumonna Anastomotic leak, sepsis
Postop RBC	Z	Z	Z	Z	X	Z	Z	Z	Z	Z	Z	Z	2	<b>7</b>	Z	ZZ	Z >-
Intraop RBC	Z	Z	Z	Z	Z	Z	Z	Z	Z	Z	Z	Z	2	7	Y	Z	۲Z
EBL (mL)	400	200	20	20	400	200	50	200	150	20	10	150	05	3	200	200	50 50
Hepatic surgery	WR	WR	WR	WR	S	WR	WR	WR	WR	WR	WR	WR	W/D	4	LHep	S	WR WR
Colorectal surgery	TME	ГC	ГС	TC	Miles	TME	ГC	ГС	AR (PME)	RC	AR (PME)	rc or	DG	2	AR (PME)	AR (PME)	AK (FME) TME
cTMN	T4N1	T4N2	T4N2	T4N1	T4N1	T3N0	T4bN1	T4N2	T4N2	T4N2	T2N0	T4N2	CINIT	7714	T4N2	T4N2	T2N0
Previous surgery	Z	Z	Z	Z	Y	Z	Z	Z	Z	Z	Y	Y	>	-	Y	Z	- >
Comorbidity	No	No	No	HHD	COFD Atrial flutter (PM) COPD	Hypothyroidism	CLD HCV+	Diabetes Hypertension PulmM	PM	Hypertension	No	Diabetes	Hypertension	Hypertension Previous PulmS	Atrial fibrillation CLD HCV+	No	HHD Diabetes HHD
BMI	23	21.5	23.5	56	25.6	20	21	23	24	23	19	41	22	3	23	27	32
Sex	M	Σ	Ц	江	M	Ц	Ц	Z	Σ	Σ	Ц	Σ	2	Į.	$\mathbf{Z}$	$\Sigma$	Ĭμ
Age	46	77	55	65	88	61	47	75	89	47	49	62	09	6	92	65	59
Z	_	7	$\alpha$	4	8	9	_	∞	6	10	Ξ	12	5	3	14	15	17

<sup>a</sup>First time of "two stage" hepatectomy, macroscopically R2, but microscopically R0.

<sup>b</sup>Hepatic resection for receptor analysis.

AR (PME), anterior resection with partial mesorectal excision; CLD, chronic liver disease; COPD, chronic obstructive pulmonary disease; cTMN, clinical TMN stage; EBL, estimated blood loss; F, female; HHD, hypertensive heart disease; LC, left colectomy; LHep, left hepatectomy; M, male; N, no; N/A, nonapplicable; PM, pacemaker; PulmM, pulmonary metastases; PulmS, pulmonary surgery; RBC, red blood cell; RC, right colectomy; RLS, right lateral sectionectomy; S, segmentectomy; TME, total mesorectal excision; WR, wedge resection; Y, yes.

#### **Discussion**

The management of patients with CRC and resectable SLM is not codified at all. There are several options of treatment for this subset of patients, including "one-stage" surgery, "liver first," and "colon first" approach. The one-stage approach for CRC with SLM has the advantage of solving both colon and liver disease in a single operation, thus avoiding two surgical procedures in a short period of time, which would entail psychological stress for the patient and increased sanitary costs. 4,5,8,10,11,47 The management of patients affected by synchronous colorectal liver metastases depends on a multifactorial evaluation, based on complexity of hepatic and colorectal resection, patient's performance status and underlying comorbidities, besides symptoms, location, and extent of disease.<sup>21</sup> The one-stage approach may not be appropriate if extensive hepatectomy is needed and colorectal resection is expected to be difficult to perform.<sup>2,5</sup> Shubert et al. have demonstrated the reduction of cumulative major morbidity and mortality rates in case of simultaneous approach when a minor hepatic resection is needed, regardless of the complexity of the colorectal resection. 11 Furthermore, a simultaneous major epatectomy can be safe and effective if there are good conditions, such as favorable resection of primary cancer and good performance status of the patient. 4,31 On the contrary, a contraindication to the synchronous resection could be the presence of severe cardiopulmonary comorbidities, especially in elderly patients.<sup>2,48</sup> When the primary cancer is symptomatic and a colorectal resection is recommended to solve or prevent obstruction and/or bleeding, a simultaneous resection could be preferred if a colonic stent is not indicated. In these cases, a one-stage radical operation prevents a delay in the surgical treatment of the secondary tumor, related with an increased risk of metastatic spread.<sup>21</sup>

The aim of laparoscopy is to reduce postoperative pain and wound morbidity, especially when a large incision would be required in open surgery, thus allowing earlier mobilization, a faster recovery of the patient and a prompt start of adjuvant chemotherapy. <sup>13,18,20,21</sup> Laparoscopy for the treatment of CRC has demonstrated better results if compared with the open approach in terms of need for transfusions, shorter recovery, and time of discharge, with similar rate of complications and oncological results. <sup>14–17</sup> Furthermore, in the last years, laparoscopy has gained acceptance also in the treatment of liver metastases, demonstrating better short-term results, without compromising the oncologic outcome. <sup>3,19,20,22</sup> The main obstacle to the development of this challenging surgery is a steep learning curve, in addition to patient's selection. 18 The need for a "change of view" from the open ventral approach to the laparoscopic caudal approach, together with the lack of tactile sensation due to laparoscopy, is the main difficulty encountered by the hepatic surgeon in the era of mini-invasiveness.<sup>49</sup> The Morioka Difficulty Scoring System was elaborated with the intention to assist surgeons in selecting patients eligible for LLR according to the individual learning curve, by providing an objective appreciation of the complexity of a given LLR. 49,50 According to the Southampton Guidelines, laparoscopic left lateral sectionectomy (LLS) and minor resections in anterolateral segments are considered a standard practice.<sup>22</sup> Nonetheless, an adequate experience in both colon and liver laparoscopic surgery is required for simultaneous resections, and this factor is often a limit, because not all centers have surgeons properly skilled in both procedures.<sup>31,51</sup>

The patients of this series were studied preoperatively and selected for laparoscopic synchronous resection according to their performance status and operative risk (ASA score). ASA IV patients were excluded, whereas the age itself was not a contraindication if an ECOG 0-1 performance status was assigned. These criteria had the purpose to select patients who were fit both for laparoscopic surgery and simultaneous resection. They presented with symptoms such as abdominal pain due to stenosis, subocclusion and/or bleeding. The patients with stenosis were not eligible for endoscopic stenting. Hence, we opted for an "upfront" surgery because resection is mandatory before starting systemic chemotherapy among patients with severe intestinal symptoms.<sup>52</sup> None of the patients had a preoperative diagnosis of low rectal cancer, but one patient operated for bleeding and stenosis caused by rectosigmoid junction cancer had an intraoperative diagnosis of synchronous ultralow rectal cancer after exploration in narcosis. Neither the preoperative proctoscopy nor the CT scan had revealed the low rectal cancer, but an intraoperative histological examination confirmed the diagnosis and the patient underwent a Miles operation, because of clinical evidence of sphincters' invasion.

Hepatic or colorectal resection was performed first, according to the more difficult and extensive procedure. Especially when a major hepatectomy eventually requiring the pedicle clamping was performed, the liver was resected before colorectal resection, to avoid the risk of bowel (and anastomotic) congestion. In fact, the surgeon did not use the Pringle maneuver routinely, avoiding the splanchnic blood flow congestion and the consequential edema of the intestinal bowel, which can favor anastomotic leakage.<sup>4</sup>

The most complications observed in the study (35%) were minor (Clavien-Dindo I-II), such as pulmonary infection, cardiac arrhythmia, ascites and rectal bleeding (Table 4). Two patients (12%) experimented major complications (Clavien-Dindo III-IV), one of them having a life threatening complication, but no mortality occurred. A 59-year-old female underwent proctectomy with TME, "ghost ileostomy" and synchronous wedge resection in segment V–VIII. She had a good performance status (ECOG 1), but she also had comorbidities, such as diabetes and hypertensive heart disease (ASA III). In the postoperative course, the patient received multiple transfusions for anemization without evidence of blood loss from the drainages. She was finally reoperated for anastomotic leakage and stayed in intensive care unit because of sepsis. The patient was discharged in POD 36 after complete recovery on the ward. A 76-year-old male who underwent subtotal proctectomy with PME and left hepatectomy was discharged in POD 8, but he was readmitted 2 weeks later with diagnosis of abdominal collection due to a biliary fistula. He was drained by percutaneous approach and then successfully treated with an endoscopic stent.

The overall morbidity rate (47%) observed in the series could be explained by the complexity of this kind of surgery. In fact, the asynchronous risk of morbidity reported in the literature for isolated laparoscopic colorectal and hepatic resection is 19%–45% and 10%–15%, respectively. <sup>13,18,53</sup> Ferretti et al. <sup>54</sup> published the largest series of simultaneous laparoscopic resections in the setting of a multicenter international study. In a retrospective noncomparative analysis, a morbidity rate of 31% was observed. The morbidity rate presented in our series is higher than the most of results found

Table 5. Authors' Results $^{26-43,51,54}$ 

Hospital stay (days) <sup>c</sup>	8.6 (5-36) 6.0 (5-9) 8.5 (5-22) 11.6 ± 5.2 6.4 ± 0.8 9.0 (8-12) 8.0 (8-12) 9.0 (4-17) 8.0 (4-30) 8.0 (4-30) 8.0 (4-30) 8.0 (5-23) 13.5 (10-18) 13.5 (10-45) 16.2 ± 6.1 6.0 (5-8) 17.5 (30-5) 8.5 ± 1.9 7.0 (3-54) 10.5 (7-15) 9.0 (3-37)
Mortality	$\begin{array}{cccccccccccccccccccccccccccccccccccc$
Morbidity Mortality	8/17 14/61 2/22 4/16 1/14 3/20 5/10 6/25 4/142 5/35 4/24 2/10 1/7 1/4 1/2 1/13 3/13 1/10 No
$EBL (mL)^{b}$	158 200 100 369 347 175 105 200 200 200 325 245 400 152 475 50 500 1500 1500
Operative time (minutes) <sup>b</sup>	165 206 135 320 321 246 261 420 360 240 290 452 550 472 495 313 313 310 401
Major Hep	1 / 2 8 2 4 / 9 / / 4 / 2 / 1 / 1
Minor Hep	16 61 20 20 13 12 16 10 10 10 7 7
Miles	7
Rectal resection	8
Subtotal CR	~~~~~~~~
_	941 / X / × × × × × × × × × × × × × × × × ×
Right-sided Left-sided CR CR	22.8 4 N
$N^a$	17/17 61/122 22/123 16/38 14/43 20/20 10/10 25/25 127/142 35/35 24/24 8/10 10/10 10/10 1/14 4/5 2/5 13/28 13/28 10/10 10/10
Study	CSS
Year	2019 2019 2018 2018 2018 2017 2018 2018 2019 2019 2010 2010 2010 2000 2000
Author	Bizzoca et al Chen et al. 21 Shim et al. 26 Chen et al. 27 Gorgun et al. 28 Xu et al. 29 Ivanecz et al. 30 Ratti et al. 31 Ferretti et al. 32 Iung et al. 33 Inoue et al. 34 Inoue et al. 36 Ida et al. 35 Hoekstra et al. 36 Polignano et al. 37 Polignano et al. 37 Polignano et al. 38 Hu et al. 39 Patriti et al. 40 Patriti et al. 42 Patriti et al. 43

<sup>a</sup>No. of pure laparoscopic approach/No. of cases in the series.

<sup>b</sup>Median values.

<sup>c</sup>Values are presented as mean ± standard deviation, or median (range).

CM, case matched; CS, case series; CR, colonic resection; EBL, estimated blood loss; Hep, hepatectomy; MIS, multicenter international study; N/A, not available.

in literature (Table 5), probably because all cases considered technically feasible in laparoscopy were included in the study. Even though only 12% of major complications occurred, this brought us to evaluate if a better selection of the patients could increase the postoperative results. The limited number of patients of the series and the retrospective nature of the study did not allow us a further analysis.

Despite the complications, no mortality occurred and the median hospital stay was 8.6 days, similar to other authors' results<sup>26,29–33,39,42,43,54</sup> (Table 5). In fact, minor complications (grade I–II) usually do not considerably affect either the postoperative course or the time of discharge. The simultaneous resections were carried out safely, with no need for conversion to open surgery. The median operative time was 165 (range 75–320) minutes, in contrast with the evidence of long operative time needed in case of laparoscopic performances. <sup>28,31,38,54</sup> An average EBL of 158 (range 10–400) mL was observed, and this result is comparable to other experiences found in literature, or even more favorable 26,28,31,36,54 (Table 5). These results are probably a consequence of the surgical skills as well as the prevalence of wedge resections in the series. Nonetheless, in 2 cases an intraoperative blood transfusion was needed, but both patients had low preoperative hemoglobin and cardiovascular comorbidities. These data confirm the evidence that laparoscopy allows optimal hemostasis, thanks to a better-magnified view and the effect of pneumoperitoneum.49

In the last few years, oncologic outcome has been a major concern about LLR, as well as the completeness of TME in case of rectal resection. In both cases, there is evidence in the literature that laparoscopy can offer the same results as the open approach. <sup>22,50,55</sup> In our series, a R0 resection on the liver specimen was achieved in 94% of cases, as well as a 100% negative distal and circumferential resection margin in case of rectal surgery. Moreover, a medium number of 20 lymphnodes were retrieved in the colorectal specimen, according to the guidelines for CRC resection. <sup>56,57</sup>

This retrospective analysis of simultaneous laparoscopic liver and colorectal resection is supported by several international experiences, even if all the published studies are retrospective and the most of the series are small. Lupinacci et al.<sup>47</sup> analyzed 14 articles, which included 39 laparoscopic simultaneous resections, and concluded that LLS associated with CRC resection is safe and feasible and should be routinely proposed.

More recently, Moris et al.<sup>58</sup> reviewed the literature and selected 12 studies, 8 of them retrospectively comparing laparoscopic versus open simultaneous resection. The short-term and oncologic results were analyzed, and similar outcomes for open and mini-invasive resections were observed, with a trend favoring the laparoscopic approach in terms of length of stay and EBL.

In conclusion, this retrospective study on a series of 17 patients operated by a single surgeon suggests that simultaneous mini-invasive colorectal and liver resection is a challenging but feasible procedure. In our experience, the advantages of mini-invasiveness and of treating primary cancer and metastases in the same recovery justify the high morbidity rate, considering that most of the complications are minor and no cases of mortality occurred. Further studies with a larger number of patients are required to better understand if a more accurate selection of patients could further improve the surgical outcome and reduce the morbidity rate.

# Acknowledgment

The authors thank Nicole Ferreri for the English Language support.

#### **Authors' Contributions**

I confirm that all authors qualify for authorship according to the Uniform requirements for the articles published by ICMJE.

#### **Disclosure Statement**

No competing financial interests exist.

#### References

- Bray F, Ferlay J, Soerjomataram I, Siegel RL, Torre LA, Jemal A. Global cancer statistics 2018: GLOBOCAN estimates of incidence and mortality worldwide for 36 cancers in 185 countries. CA Cancer J Clin 2018;68:394–424.
- Adam R, de Gramont A, Figueras J, Kokudo N, Kunstlinger F, Loyer E, Poston G, Rougier P, Rubbia-Brandt L, Sobrero A, Teh C, Tejpar S, Van Cutsem E, Vauthey JN, Påhlman L; of the EGOSLIM (Expert Group on OncoSurgery management of LIver Metastases) group. Managing synchronous liver metastases from colorectal cancer: A multidisciplinary international consensus. Cancer Treat Rev 2015;41:729-741.
- 3. Abu Hilal M, Underwood T, Zuccaro M, Primrose J, Pearce N. Short- and medium-term results of totally laparoscopic resection for colorectal liver metastases. Br J Surg 2010;97: 927–933.
- Garritano S, Selvaggi F, Spampinato MG. Simultaneous minimally invasive treatment of colorectal neoplasm with synchronous liver metastasis. Biomed Res Int 2016;2016: 9328250.
- Gonzalez HD, Figueras J. Practical questions in liver metastases of colorectal cancer: General principles of treatment. HPB 2007;9:251–258.
- Martin R, Paty P, Fong Y, Grace A, Cohen A, DeMatteo R, Jarnagin W, Blumgart L. Simultaneous liver and colorectal resections are safe for synchronous colorectal liver metastasis. J Am Coll Surg 2003;197:233–241; discussion 241– 242.
- Weber JC, Bachellier P, Oussoultzoglou E, Jaeck D. Simultaneous resection of colorectal primary tumour and synchronous liver metastases. Br J Surg 2003;90:956–962.
- Jaeck D, Bachellier P, Weber JC, Boudjema K, Mustun A, Pâris F, Schaal JC, Wolf P. [Surgical strategy in the treatment of synchronous hepatic metastases of colorectal cancers. Analysis of a series of 59 operated on patients]. Chirurgie 1999;124:258–263.
- Martin RC, Augenstein V, Reuter NP, Scoggins CR, McMasters KM. Simultaneous versus staged resection for synchronous colorectal cancer liver metastases. J Am Coll Surg 2009;208:842–850; discussion 850–852.
- Reddy SK, Pawlik TM, Zorzi D, Gleisner AL, Ribero D, Assumpcao L, Barbas AS, Abdalla EK, Choti MA, Vauthey JN, Ludwig KA, Mantyh CR, Morse MA, Clary BM. Simultaneous resections of colorectal cancer and synchronous liver metastases: A multi-institutional analysis. Ann Surg Oncol 2007;14:3481–3491.
- Shubert CR, Habermann EB, Bergquist JR, Thiels CA, Thomsen KM, Kremers WK, Kendrick ML, Cima RR, Nagorney DM. A NSQIP review of major morbidity and

8 BIZZOCA ET AL.

- mortality of synchronous liver resection for colorectal metastasis stratified by extent of liver resection and type of colorectal resection. J Gastrointest Surg 2015;19:1982–1994.
- Tohme S, Goswami J, Han K, Chidi AP, Geller DA, Reddy S, Gleisner A, Tsung A. Minimally invasive resection of colorectal cancer liver metastases leads to an earlier initiation of chemotherapy compared to open surgery. J Gastrointest Surg 2015;19:2199–2206.
- 13. Fretland ÅA, Kazaryan AM, Bjørnbeth BA, Flatmark K, Andersen MH, Tønnessen TI, Bjørnelv GM, Fagerland MW, Kristiansen R, Øyri K, Edwin B. Open versus laparoscopic liver resection for colorectal liver metastases (the Oslo-CoMet Study): Study protocol for a randomized controlled trial. Trials 2015;16:73.
- Tanis PJ, Buskens CJ, Bemelman WA. Laparoscopy for colorectal cancer. Best Pract Res Clin Gastroenterol 2014; 28:29–39.
- Rondelli F, Trastulli S, Avenia N, Schillaci G, Cirocchi R, Gullà N, Mariani E, Bistoni G, Noya G. Is laparoscopic right colectomy more effective than open resection? A meta-analysis of randomized and nonrandomized studies. Colorectal Dis 2012;14:e447–e469.
- Chen K, Cao G, Chen B, Wang M, Xu X, Cai W, Xu Y, Xiong M. Laparoscopic versus open surgery for rectal cancer: A meta-analysis of classic randomized controlled trials and high-quality Nonrandomized Studies in the last 5 years. Int J Surg 2017;39:1–10.
- Tjandra JJ, Chan MK. Systematic review on the short-term outcome of laparoscopic resection for colon and rectosigmoid cancer. Colorectal Dis 2006;8:375–388.
- Tranchart H, Dagher I. Laparoscopic liver resection: A review. J Visc Surg 2014;151:107–115.
- 19. Wakabayashi G, Cherqui D, Geller DA, Buell JF, Kaneko H, Han HS, Asbun H, O'Rourke N, Tanabe M, Koffron AJ, Tsung A, Soubrane O, Machado MA, Gayet B, Troisi RI, Pessaux P, Van Dam RM, Scatton O, Abu Hilal M, Belli G, Kwon CH, Edwin B, Choi GH, Aldrighetti LA, Cai X, Cleary S, Chen KH, Schön MR, Sugioka A, Tang CN, Herman P, Pekolj J, Chen XP, Dagher I, Jarnagin W, Yamamoto M, Strong R, Jagannath P, Lo CM, Clavien PA, Kokudo N, Barkun J, Strasberg SM. Recommendations for laparoscopic liver resection: A report from the second international consensus conference held in Morioka. Ann Surg 2015;261:619–629.
- Ciria R, Cherqui D, Geller DA, Briceno J, Wakabayashi G. Comparative short-term benefits of laparoscopic liver resection: 9000 cases and climbing. Ann Surg 2016;263:761–777.
- Cipriani F, Rawashdeh M, Stanton L, Armstrong T, Takhar A, Pearce NW, Primrose J, Abu Hilal M. Propensity scorebased analysis of outcomes of laparoscopic versus open liver resection for colorectal cancer metastases. Br J Surg 2016;103:1504–1512.
- 22. Abu Hilal M, Aldrighetti L, Dagher I, Edwin B, Troisi RI, Alikhanov R, Aroori S, Belli G, Besselink M, Briceno J, Gayet B, D'Hondt M, Lesurtel M, Menon K, Lodge P, Rotellar F, Santoyo J, Scatton O, Soubrane O, Sutcliffe R, Van Dam R, White S, Halls MC, Cipriani F, Van der Poel M, Ciria R, Barkhatov L, Gomez-Luque Y, Ocana-Garcia S, Cook A, Buell J, Clavien PA, Dervenis C, Fusai G, Geller D, Lang H, Primrose J, Taylor M, Van Gulik T, Wakabayashi G, Asbun H, Cherqui D. The Southampton Consensus Guidelines for laparoscopic liver surgery: From indication to implementation. Ann Surg 2018;268:11–18.

 Allaix ME, Degiuli M, Giraudo G, Marano A, Morino M. Laparoscopic versus open colorectal resections in patients with symptomatic stage IV colorectal cancer. Surg Endosc 2012;26:2609–2616.

- Nguyen KT, Marsh JW, Tsung A, Steel JJ, Gamblin TC, Geller DA. Comparative benefits of laparoscopic vs open hepatic resection: A critical appraisal. Arch Surg 2011;146: 348–356.
- Nomi T, Fuks D, Kawaguchi Y, Mal F, Nakajima Y, Gayet B. Laparoscopic major hepatectomy for colorectal liver metastases in elderly patients: A single-center, case-matched study. Surg Endosc 2015;29:1368–1375.
- 26. Shim JR, Lee SD, Park HM, Lee EC, Park B, Han SS, Kim SH, Park SJ. Outcomes of liver resection in patients with colorectal liver metastases by laparoscopic or open surgery. Ann Hepatobiliary Pancreat Surg 2018;22:223–230.
- Chen YW, Huang MT, Chang TC. Long term outcomes of simultaneous laparoscopic versus open resection for colorectal cancer with synchronous liver metastases. Asian J Surg 2019;42:217–223.
- Gorgun E, Yazici P, Onder A, Benlice C, Yigitbas H, Kahramangil B, Tasci Y, Aksoy E, Aucejo F, Quintini C, Miller C, Berber E. Laparoscopic versus open 1 stage resection of synchronous liver metastases and primary colorectal cancer. Gland Surg 2017;6:324–329.
- 29. Xu X, Guo Y, Chen G, Li C Wang H, Dong G. Laparoscopic resections of colorectal cancer and synchronous liver metastases: A case controlled study. Minim Invasive Ther Allied Technol 2018;27:209–216.
- Ivanecz A, Krebs B, Stozer A, Jagric T, Plahuta I, Potrc S. Simultaneous pure laparoscopic resection of primary colorectal cancer and synchronous liver metastases: A single institution experience with propensity score matching analysis. Radiol Oncol 2017;52:42–53.
- Ratti F, Catena M, Di Palo S, Staudacher C, Aldrighetti L. Impact of totally laparoscopic combined management of colorectal cancer with synchronous hepatic metastases on severity of complications: A propensity-score-based analysis. Surg Endosc 2016;30:4934–4945.
- 32. Berti S, Francone E, Minuto M, Bonfante P, Sagnelli C, Bianchi C, Tognoni A, Falco E. Synchronous totally laparoscopic management of colorectal cancer and resectable liver metastases: A single center experience. Langenbecks Arch Surg 2015;400:495–503.
- 33. Jung KU, Kim HC, Cho YB, Kwon CH, Yun SH, Heo JS, Lee WY, Chun HK. Outcomes of simultaneous laparoscopic colorectal and hepatic resection for patients with colorectal cancers: A comparative study. J Laparoendosc Adv Surg Tech A 2014;24:229–235.
- 34. Inoue A, Uemura M, Yamamoto H, Hiraki M, Naito A, Ogino T, Nonaka R, Nishimura J, Wada H, Hata T, Takemasa I, Eguchi H, Mizushima T, Nagano H, Doki Y, Mori M. Short-term outcomes of simultaneous laparoscopic colectomy and hepatectomy for primary colorectal cancer with synchronous liver metastases. Int Surg 2014;99:338–343
- 35. Ida S, Oki E, Ando K, Kimura Y, Yamashita Y, Saeki H, Ikegami T, Yoshizumi T, Watanabe M, Morita M, Shirabe K, Kusumoto T, Ikeda T, Baba H, Maehara Y. Pure laparoscopic right-sided hepatectomy in the semi-prone position for synchronous colorectal cancer with liver metastases. Asian J Endosc Surg 2014;7:133–137.
- 36. Takasu C, Shimada M, Sato H, Miyatani T, Imura S, Morine Y, Ikemoto T, Kanamoto M, Kurita N, Eto S,

- Utsunomiya T. Benefits of simultaneous laparoscopic resection of primary colorectal cancer and liver metastases. Asian J Endosc Surg 2014;7:31–37.
- Spampinato MG, Mandalá L, Quarta G, Del Medico P, Baldazzi G. One-stage, totally laparoscopic major hepatectomy and colectomy for colorectal neoplasm with synchronous liver metastases: Safety, feasibility and short-term outcome. Surgery 2013;153:861–865.
- Hoekstra LT, Busch OR, Bemelman WA, van Gulik TM, Tanis PJ. Initial experiences of simultaneous laparoscopic resection of colorectal cancer and liver metastases. HPB Surg 2012;2012:893956.
- Hu MG, Ou-yang CG, Zhao GD, Xu DB, Liu R. Outcomes of open versus laparoscopic procedures for synchronous radical resection of liver metastatic colorectal cancer: A comparative study. Surg Laparosc Endosc Percutan Tech 2012;22:364–369.
- Polignano FM, Quyn AJ, Sanjay P, Henderson NA, Tait IS. Totally laparoscopic strategies for the management of colorectal cancer with synchronous liver metastasis. Surg Endosc 2012;26:2571–2578.
- 41. Lee JS, Hong HT, Kim JH, Lee IK, Lee KH, Park IY, Oh ST, Kim JG, Lee YS. Simultaneous laparoscopic resection of primary colorectal cancer and metastatic liver tumor: Initial experience of single institute. J Laparoendosc Adv Surg Tech A 2010;20:683–687.
- Patriti A, Ceccarelli G, Bartoli A, Spaziani A, Lapalorcia LM, Casciola L. Laparoscopic and robot-assisted one stage resection of colorectal cancer with synchronous liver metastases: A pilot study. J Hepatobiliary Pancreat Surg 2009; 16:450–457.
- Sasaki A, Nitta H, Otsuka K, Takahara T, Nishizuka S, Wakabayashi G. Ten-year experience of totally laparoscopic liver resection in a single institution. Br J Surg 2009; 96:274–279.
- 44. Zeng Y, Tian M. Laparoscopic versus open hepatectomy for elderly patients with liver metastases from colorectal cancer. J BUON 2016;21:1146–1152.
- Dindo D, Demartines N, Clavien PA. Classification of surgical complications: A new proposal with evaluation in a cohort of 6336 patients and results of a survey. Ann Surg 2004;240:205–213.
- 46. Hamady ZZ, Cameron IC, Wyatt J, Prasad RK, Toogood GJ, Lodge JP. Resection margin in patients undergoing hepatectomy for colorectal liver metastasis: A critical appraisal of the 1 cm rule. Eur J Surg Oncol 2006;32:557– 563.
- Lupinacci RM, Andraus W, De Paiva Haddad LB, Carneiro D' Albuquerque LA, Herman P. Simultaneous laparoscopic resection of primary colorectal cancer and associated liver metastases: A systematic review. Tech Coloproctol 2014; 18:129–135.
- 48. Vassiliou I, Arkadopoulos N, Theodosopoulos T, Fragulidis G, Marinis A, Kondi-Paphiti A, Samanides L, Polydorou A, Gennatas C, Voros D, Smyrniotis V. Surgical approaches of resectable synchronous colorectal liver metastases: Timing considerations. World J Gastroenterol 2007;13:1431–1434.

- 49. Wakabayashi G. What has changed after the Morioka consensus conference 2014 on laparoscopic liver resection? Hepatobiliary Surg Nutr 2016;5:281–289.
- 50. Hallet J, Beyfuss K, Memeo R, Karanicolas PJ, Marescaux J, Pessaux P. Short and long-term outcomes of laparoscopic compared to open liver resection for colorectal liver metastases. Hepatobiliary Surg Nutr 2016;5:300–310.
- 51. Van der Poel MJ, Tanis PJ, Marsman HA, Rijken AM, Gertsen EC, Ovaere S, Gerhards MF, Besselink MG, D'Hondt M, Gobardhan PD. Laparoscopic combined resection of liver metastases and colorectal cancer: A multicenter, case-matched study using propensity scores. Surg Endosc 2018. DOI: 10.1007/s00464-018-6371-1.
- Yoshida Y, Hoshino S, Shiwaku H, Beppu R, Tanimura S, Tanaka S, Yamashita Y. Early start of chemotherapy after resection of primary colon cancer with synchronous multiple liver metastases: A case report. Case Rep Oncol 2011; 4:250–254.
- 53. Morneau M, Boulanger J, Charlebois P, Latulippe JF, Lougnarath R, Thibault C, Gervais N; Comité de l'Évolution des Pratiques en Oncologie. Laparoscopic versus open surgery for the treatment of colorectal cancer: A literature review and recommendations from the Comité de l'évolution des pratiques en oncologie. Can J Surg 2013;56:297– 310.
- 54. Ferretti S, Tranchart H, Buell JF, Eretta C, Patriti A, Spampinato MG, Huh JW, Vigano L, Han HS, Ettorre GM, Jovine E, Gamblin TC, Belli G, Wakabayashi G, Gayet B, Dagher I. Laparoscopic simultaneous resection of colorectal primary tumor and liver metastases: Results of a Multicenter International Study. World J Surg 2015;39: 2052–2060.
- Vennix S, Pelzers L, Bouvy N, Beets GL, Pierie JP, Wiggers T, Breukink S. Laparoscopic versus open total mesorectal excision for rectal cancer. Cochrane Database Syst Rev 2014;CD005200. DOI: 10.1002/14651858.CD005200.pub3.
- NCCN (2018) Clinical Practice Guidelines in Oncology (NCCN Guidelines) Colon Cancer Version 4.2018. www .nccn.org (accessed October 19, 2018).
- 57. NCCN (2018) Clinical Practice Guidelines in Oncology (NCCN Guidelines) Rectal Cancer Version 3.2018. www .nccn.org (accessed August 7, 2018).
- Moris D, Tsilimigras DI, Machairas N, Merath K, Cerullo M, Hasemaki N, Prodromidou A, Cloyd JM, Pawlik TM. Laparoscopic synchronous resection of colorectal cancer and liver metastases: A systematic review. J Surg Oncol 2019;119:30–39.

Address correspondence to: Cinzia Bizzoca, MD General Surgery "Balestrazzi" Polyclinics of Bari Piazza G. Cesare, 11 Bari 70124

E-mail: cinziabiz84@gmail.com