POSTGRADUATE PROGRAM:

'MINIMALLY INVASIVE SURGERY, ROBOTIC SURGERY & TELESURGERY'

NATIONAL AND KAPODESTRIAN UNIVERSITY OF ATHENS

MEDICAL SCHOOL

MASTER THESIS

COMPARISON OF POSTOPERATIVE AND ONCOLOGIC OUTCOMES IN LAPAROSCOPIC AND OPEN RIGHT COLECTOMY FOR COLON CANCER. A 5-YEAR EXPERIENCE

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ATHENS

FEBRUARY 2022

ΜΕΤΑΠΤΥΧΙΑΚΟ ΠΡΟΓΡΑΜΜΑ ΣΠΟΥΔΩΝ

' ΕΛΑΧΙΣΤΑ ΕΠΕΜΒΑΤΙΚΗ ΧΕΙΡΟΥΡΓΙΚΗ, ΡΟΜΠΟΤΙΚΗ ΧΕΙΡΟΥΡΓΙΚΗ & ΤΗΛΕΧΕΙΡΟΥΡΓΙΚΗ'

ΕΘΝΙΚΟ ΚΑΙ ΚΑΠΟΔΙΣΤΡΙΑΚΟ ΠΑΝΕΠΙΣΤΗΜΙΟ ΑΘΗΝΩΝ ΙΑΤΡΙΚΗ ΣΧΟΛΗ

ΔΙΠΛΩΜΑΤΙΚΗ ΕΡΓΑΣΙΑ

ΘΕΜΑ: ΣΥΓΚΡΙΣΗ ΜΕΤΕΓΧΕΙΡΗΤΙΚΩΝ ΚΑΙ ΟΓΚΟΛΟΓΙΚΩΝ ΑΠΟΤΕΛΕΣΜΑΤΩΝ ΣΤΗ ΛΑΠΑΡΟΣΚΟΠΙΚΗ ΚΑΙ ΑΝΟΙΚΤΗ ΔΕΞΙΑ ΚΟΛΕΚΤΟΜΗ ΓΙΑ ΚΑΡΚΙΝΟ ΤΟΥ ΠΑΧΕΟΣ ΕΝΤΕΡΟΥ. ΜΙΑ ΕΜΠΕΙΡΙΑ 5 ΕΤΩΝ

ΜΕΤΑΠΤΥΧΙΑΚΗ ΦΟΙΤΗΤΡΙΑ: ΣΤΑΥΡΟΥ ΕΥΦΡΟΣΥΝΗ

ΕΠΙΒΛΕΠΩΝ:

ΚΑΘΗΓΗΤΗΣ ΔΗΜΗΤΡΙΟΣ ΔΗΜΗΤΡΟΥΛΗΣ

ΑΘΗΝΑ ΦΕΒΡΟΥΑΡΙΟΣ 2022

<u>Περίληψη</u>

Εισαγωγή/ Σκοπός: Η. λαπαροσκοπική κολεκτομή είναι μια επέμβαση η οποία πραγματοποιείται τις τελευταίες τρεις δεκαετίες και συνεχώς κερδίζει έδαφος συγκριτικά με την παραδοσιακή ανοικτή κολεκτομή. Η μελέτη αυτή διεξήχθη με σκοπό τη σύγκριση μετεγχειρητικών και ογκολογικών αποτελεσμάτων στη λαπαροσκοπική και ανοικτή δεξιά κολεκτομή για καρκίνο του παχέος εντέρου.

Ασθενείς και Μέθοδοι: Η εν λόγω εργασία είναι μια αναδρομική μελέτη από ένα κέντρο γενικής χειρουργικής και αφορά το διάστημα από το 2015 έως το 2020. Οι παράμετροι που μελετήθηκαν μετεγχειρητικά είναι η CRP (C-Αντιδρώσα πρωτεΐνη), CPK(κινάση κρεατινίνης), LDH(γαλακτική αφυδρογονάση), ο αριθμός των εξαιρεθέντων λεμφαδένων και η διάρκεια νοσηλείας.

Αποτελέσματα: Συλλέξαμε στοιχεία από 21 ανοικτές και 17 λαπαροσκοπικές δεξιές κολεκτομές σε διάρκεια 5 ετών. Οι μετρήσεις τη δεύτερη μετεγχειρητική ημέρα των τιμών της CRP και CPK ήταν σημαντικά χαμηλότερες στους ασθενείς μετά από λαπαροσκοπική κολεκτομή, ενώ οι τιμές της LDH δεν έδειξαν ιδιαίτερη διαφορά μεταξύ των δυο μεθόδων. Ο αριθμός των εξαιρεθέντων λεμφαδένων ήταν μεγαλύτερος σε όσους υπεβλήθησαν σε ανοικτή κολεκτομή. Η αναλγησία στους ασθενείς μετά από λαπαροσκοπική κολεκτομή. Η αναλγησία στους ασθενείς μετά από ανοικτή κολεκτομή επετεύχθη με τη χρήση πεθιδίνης και τραμαδόλης τις πρώτες τρεις μετεγχειρητικές ημέρες, ενώ μετά από λαπαροσκοπική κολεκτομή χορηγήθηκε συστηματικά παρακεταμόλη και τραμαδόλη κατ΄ επίκληση. Η διάρκεια νοσηλείας ήταν σημαντικά βραχύτερη στους ασθενείς που χειρουργήθηκαν λαπαροσκοπικά.

Συμπέρασμα: Η λαπαροσκοπική δεξιά κολεκτομή υπερέχει της ανοικτής όσον αφορά τη μετεγχειρητική αναλγησία και τη διάρκεια νοσηλείας, καθώς επίσης και στις μετρήσεις συγκεκριμένων παραμέτρων. Όσον αφορά την ογκολογική επάρκεια, η λαπαρσκοπική κολεκτομή υστερούσε έναντι της ανοικτής.

Λέξεις κλειδιά: Λαπαροσκοπική δεξιά κολεκτομή, Καρκινος δεξιού κόλου, μετεγχειρητικές παράμετροι, εξαίρεση λεμφαδένων

<u>Abstract</u>

Background/Aim: Laparoscopic colectomy is a procedure which is being performed for three decades and is gaining popularity continuously over the traditional open colectomy. This study was conducted in order to compare postoperative and oncologic results based on several factors in laparoscopic and open right colectomy for right colon cancer.

Patients and Methods: This is a retrospective study of right colectomy at a single institution from 2015 until 2020. The factors that were studied included postoperative values of C-reactive protein (CRP), lactate dehydrogenase (LDH), creatine phosphokinase (CPK), the number of excised lymph nodes, the use of postoperative analgesics and the length of hospital stay.

Results: We collected data from 21 open and 17 laparoscopic right colectomies through a 5-year period. Measurements on the second postoperative day revealed mean CRP and CPK values significantly lower in the laparoscopic group compared to the open group, while LDH levels did not affirm major differences between the two groups. The mean number of lymph nodes excised during the open procedure was superior to those harvested in the laparoscopic group. The use of analgesics throughout the entire hospital stay was a combination of pethidine and tramadol for the first three postoperative days in open procedures, while paracetamol and, occasionally, tramadol were administered upon patient request following laparoscopic procedures. The mean hospital stay was substantially shorter in the laparoscopic group compared to the open surgery group.

Conclusion: Laparoscopic right colectomy is superior compared to open right colectomy with regards to postoperative analgesia and length of hospital stay, but also in certain postoperative laboratory values. Despite these there was no supremacy considering oncologic clearance.

Key Words: Laparoscopic right colectomy, right colon cancer, postoperative values, lymph node excision.

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Introduction

Colorectal cancer is the most common malignancy in the Western World and the second cause of death (1). Right-sided colon cancer represents one third of all cases and is traditionally treated by right hemicolectomy (2). Laparoscopic colorectal surgery was first described by Jacobs *et al.* (3) in 1991 and is widely accepted for the treatment of both benign and malignant left colon and rectal pathologies (4). Laparoscopic approach in the right colon still remains controversial, probably due to the complexity of right colon laparoscopic anatomy and the vulnerability of the vascular peduncles that require a greater attention and laparoscopic experience compared to left colon and rectal surgery (5).

The purpose of this study was to compare the postoperative and oncological results after laparoscopic and open right colectomy for right-sided adenocarcinoma, and decipher the advantages of the laparoscopic technique over the conventional open one.

Patients and Methods

Patients. We carried out a retrospective analysis of patients with right sided colonic adenocarcinoma who underwent either open, or laparoscopic right colectomy in a single institution during a 5-year period, from January 2015 until June 2020.

Several patients were excluded from the study, such as those who were i) immunocompromised, ii) with distant metastases, iii) with lesions other than colonic adenocarcinoma, iv) with acute complications, such as obstruction, or rupture, requiring an emergency operation, and v) cases in which laparoscopy was converted into laparotomy.

Our study encompassed 38 patients, 17 of whom belonged to the laparoscopic group and 21 to the open group (Table I).

Preoperative procedures. All patients had preoperative colonoscopy with removal of biopsies to confirm the presence, exact location and type of the tumour, while the marking of the tumour with methylene blue was only performed in patients who would undergo a laparoscopic procedure. All patients underwent abdominal ultrasonography and CT examination of the abdomen and thorax as well as standard laboratory tests, such as complete blood count, tumour markers (carcinoembryonic antigen, CA 19-9), thyroid hormones, and basic biochemical values (urea, creatinine, Na+, K+, blood glucose). Bowel preparation was achieved using disodium phosphate dodecahydrate and preoperative broad-spectrum antibiotics were initiated 24 h before the operation.

Open and laparoscopic surgical procedures. In both groups, a Foley catheter and a nasogastric tube were routinely used, which were inserted following the induction of general anaesthesia at the operating theatre.

Patients undergoing right laparoscopic colectomy were placed in lithotomy position. Pneumoperitoneum was achieved using a Verres needle inserted to the left hypochondrium and the abdomen was insufflated to 12 mm Hg. The 12 mm trocar was placed supra-umbilically and a 10mm one was placed at the left hypochondrium, at the site of the Verres needle insertion. Another two 5 mm trocars were placed in the lower abdomen, one 2-3 cm anteriorly and medially to the anterior superior iliac spine on the left side and one in the right lower abdomen on the right side. After the port placement, the patients were tilted 20 degrees to the left and in slight Trendelenburg position. In all operations the terminal ileum, cecum, and ascending colon were completely mobilized up to the level of hepatic flexure, with particular attention to safeguard the ureter and the duodenum in a medial to lateral approach. Ileocolic, right colic and the right branch of the middle colic vessels were ligated with endoscopic clips and with the use of bipolar cautery. The anastomosis was in all cases extracorporeal laterolateral ileotransverse as well as isoperistaltic, by extending the supraumbilical incision to about 6-7 cm.

The open group underwent typical right hemicolectomy with midline incision and medial to lateral colon mobilization. The anastomosis was laterolateral ileotransverse, same as in the laparoscopic group, but it was made either iso- or antiperistaltic, depending on the surgeon's preference. The midline incision was closed in layers using a separate popropylene suture for each layer.

A penrose drain was placed in all patients, in the rectouterine space in females and in the rectovesical space in males, regardless of the type of operation.

Postoperative procedures. The following parameters (Table II) were measured prospectively: i) C-reactive protein (CRP), ii) creatine phosphokinase (CPK) and iii) lactate dehydrogenase (LDH), all on the second postoperative day, iv) the number of lymph nodes harvested, v) analgesic requirements and vi) the duration of hospital stay (Table II)

Statistics. All our data are expressed as mean \pm SD and Unpaired t-test through GraphPad software was used to analyze several quantitative variables. We considered *p*<0.05 as statistically significant.

Results

The patients that were included in our study were all diagnosed with a right-sided colon adenocarcinoma with similar demographic data and comorbidities (Table I).

On the second postoperative day all patients were tested for LDH, CRP and CPK. There was no significant difference between the open and the laparoscopic group regarding the postoperative levels of LDH; our measurements revealed a mean LDH level of 222.80 in open and 184.82 in laparoscopic group (p=0.0593). The values of postoperative CRP and CPK were considerably lower in the laparoscopic group compared to those of the open surgery group, which, as is widely accepted, are significant predictors of postoperative survival, complications and mortality (6, 7). We measured a mean CRP value of 132.14 in open and 102.23 in laparoscopic surgery (p=0.0049), as well as a mean value of CPK levels of 470.42 in open and 162.76 in laparoscopic surgery, respectively (p=0.0208).

The number of excised lymph nodes showed that patients who underwent open right colectomy had superior oncologic clearance, with a mean number of lymph nodes excised 27.95 compared to those who underwent laparoscopic surgery who had a mean number of lymph nodes equal to 19.52

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(p=0.0035), which shows great significance. In one patient from the laparoscopic group only 10 lymph nodes were harvested, even though the accepted number of lymph nodes during oncologic operations is 12, according to international literature (8).

The length of postoperative hospital stay was undoubtedly shorter in the laparoscopic group (mean \pm SD=5.76 \pm 0.93 days), when compared to the open group (mean \pm SD=8.80 \pm 341 days) (*p*=0.0011).

Analgesics were used as a combination of tramadol and pethidine in patients after open surgery for the first three postoperative days, subsequently reducing the use of pethidine and replacing it with paracetamol. Patients in the laparoscopic group were provided with paracetamol for the first three postoperative days and tramadol only upon patient request.

Discussion

The first successful laparoscopic colectomy was reported by Jacob *et al.* in 1991 (3) and since then there has been a tremendous development of laparoscopic colonic surgery, especially during the past decade (9). The laparoscope itself is undoubtedly a tool to i) explore thoroughly the abdominal cavity, ii) exclude liver and peritoneal metastases, and iii) prove useful in mobilization and resection of select lesions.

Laparoscopic right colectomy has many technical difficulties and a steep learning curve for the surgeon in order to achieve advanced laparoscopic skills. It also requires specialized equipment and a high number of similar cases designated to the surgeons so as for them to attain familiarity with the specific instruments and for the entire therapeutic team to learn how to cooperate efficiently during this procedure (10). The risk of dissemination and port site metastases from laparoscopic colon surgery have markedly decreased since they were first reported in 1993 (11).

Although the mechanism of port-site recurrence is not completely clear yet, and was first noted in thoracoscopic surgery without CO_2 insufflation, it is probably caused by a combination of the following factors: i) tumor manipulation, ii) failure to isolate the tumor, and iii) forceful extraction of the surgical specimen (12).

There are only a few publications which specifically compare the outcomes of patients who underwent laparoscopic right hemicolectomy for colon cancer with those who were treated with the open approach (13). Our study confirms that the laparoscopic approach is superior to open surgery regarding the length of hospital stay and, thus, the overall costs of hospitalization. The use of narcotics was minimal in the laparoscopic group, also leading to faster recovery, bowel movement and subsequent defecation. Although not included in this study, minimization of postoperative pain due to the length of the incision (<7 cm) leads to faster patient mobilization, reduction of postoperative respiratory complications, and surgical site infections (SSIs) (14).

Moreover, the levels of CPK and CRP as major indicators of tissue and muscle damage, postoperative tissue inflammation and destruction (6), as well as the predictive value of CRP concerning anastomotic complications (7), provided important parameters for assuring the safe discharge of our patients from the hospital. Importantly, in our study both these factors were found at significantly lower levels following laparoscopic right colectomy compared to the open surgery group. The levels of LDH, which is also an indicator of tissue injury (15) was not significantly lower in the laparoscopic group compared to the open surgery group.

A more extensive nodal resection has been associated with lower rates of cancer recurrence, as it allows for accurate cancer staging and a more appropriate use of adjuvant chemotherapy for nodepositive patients. Importantly, it has also been associated with improved survival following resection for colon cancer (16). A resection of minimum 12 nodes has been endorsed as a consensus standard following safe colectomy for colon cancer (17). Lymph node excision in this study and, thus, possibly the oncologic outcomes may still be more advantageous following open surgery compared to laparoscopy, since the number of overall lymph nodes harvested was higher during the former than the latter.

In conclusion, our study concludes that laparoscopic right colectomy is superior to open surgery concerning postoperative analgesia and length of hospital stay, and also with respect to certain postoperative laboratory values, such as CRP and CPK. Unfortunately, laparoscopy remains less efficient when it comes to oncologic clearance.

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<u>Appendix</u>

	No of	Type II	AFib	DLP	HTN	Age (mean in	BMI
	patients	diabetes				years)	(mean in
							kg/m ²)
Males	23	16	3	17	11	60.1±15.2	24.6± 6.3
Females	15	10	4	12	8	67.5±13.4	27.9±7.8

Table I. Demographic data and comorbidities of patients.

AFib: Atrial fibrillation; DLP: dyslipidemia; HTN: hypertension; BMI: body mass index.

	Laparoscopic group	Open group	<i>p</i> -Value
CRP	102.23±36.55	132.14±24.80	0.0049**
СРК	162.76±10.17	470.42±51.52	0.0208*
LDH	184.82±47.20	222.80±68.18	0.0593
No of lymph nodes	19.52±6.49	27.95±9.47	0.0035**
Days of hospital stay	5.76±0.93	8.80±3.41	0.0011**

Table II. All the parameters studied during the operation and postoperatively. Values are expressed as mean \pm standard deviation. **p*>0.05, ***p*>0.005.

CRP: C-reactive protein; CPK: creatine phosphokinase; LDH: lactate dehydrogenase.