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Robotic Surgery versus Laparoscopy in Colorectal Cancer Resection: A Systematic Review Guilherme Gomes Gil De Menezes Received: 11 September 2021 Accepted: 3 October 2021 Published: 15 October 2021

6 Abstract

14

7 Introduction: Colorectal cancer is a malignant disease, more predominantly observed in men

 $_{\circ}$ and the third most incident tumor among all cancers, with an estimated risk of 26.6 / 100

⁹ thousand. Despite its high incidence and prevalence, it is amenable to treatment, and in most

¹⁰ cases, it is curable -when detected in early stages. Objective: To compare the safety and

¹¹ efficacy of performing robotic surgery with traditional laparoscopic surgery in patients

¹² undergoing colorectal cancer resection regarding the variables: intra and postoperative

¹³ complications, surgical conversion, and mortality.

Objetivo: Comparar a segurança e eficacia da realização da cirurgia robótica com a cirurgia laparoscópica tradicional em pacientes submetidos à ressecção de câncer colorretal, quanto às variáveis: complicações intra e pós-operatórias, conversão cirúrgica e mortalidade.

19 Métodos: Trata-se de uma revisão sistemática caracterizada pela busca de artigos na literatura, com aplicação de metodologia sistematizada, através de bases de dados MEDLINE/PubMed, Scielo, Embase e Cochrane, por 20 21 meio da combinação de descritores, incluindo termos do Medical Subject Headings (MeSH) e dos Descritores em Ciência da Saúde (DECs), incluindo publicações em inglês e português: robotic-assisted conventional laparoscopic 22 surgery colorectal cancer resection, além de busca ativa. Foram incluídos ensaios clínicos randomizados, estudos 23 de coorte e estudos retrospectivos publicados partir de 2010, em português e inglês, que compararam o emprego 24 das técnicas laparoscópicas minimamente invasiva e a ressecção colorretal pela abordagem robótica. Foram 25 excluídos revisões, relatos de casos, série de casos, comentários e correspondências. A análise e aplicação das 26

27 ferramentas CONSORT e STROBE foram feitas por dois avaliadores separadamente.

28 Resultados: Foram encontrados 20 artigos na estratégia de busca, e 07 foram selecionados. As amostras variaram de 56 a 471 participantes (n total = 1589), com variação de idade de 61,2 -69,0. Todos os estudos 29 incluíram ambos os gêneros e, dentre estes, apenas um20 relatou uma proporção maior de mulheres. Dentre os 30 trabalhos selecionados, cinco estudos se caracterizam como coortes retrospectivas e dois estudos como ensaios 31 clínicos randomizados. A variação de duração das intervenções foi de 12 -120 meses. Realizando uma comparação 32 entre as abordagens laparoscópica e robótica acerca da taxa de complicações intraoperatórias, o percentual 33 apresentado pelo grupo da cirurgia robótica (6.0%) foi maior que a taxa de complicações relacionadas à cirurgia 34 Author: e-mail: guilhermemenezes16.2@bahiana.edu.br laparoscópica (5,2%). Sobre as taxas de conversão, 35

a cirurgia robótica apresentou percentual consideravelmente menor: 0% -8,1% contra 0% -37%. Em relação à
morbidade pósoperatória as prevalências foram de 22,6% -60% para a laparoscopia e 8,9% -42,3% para a cirurgia
robótica, sendo observada uma notória variação em ambas as abordagens.

No que tange ás taxas de mortalidade foi identificada prevalência que variou entre 0% -5,6% na cirurgia
laparoscópica, enquanto que na cirurgia robótica as taxas variaram entre 0% e 0,8%.

Conclusão: Frente aos achados descritos, evidências de boa a moderada qualidade, sustentam que a cirurgia robótica para a ressecção de câncer colorretal, apesar de promover melhor ergonomia e conforto para o cirurgião, produz resultados peri e pós-operatórios semelhantes. A cirurgia robótica, no entanto, possui menor taxa de conversão cirúrgica e mortalidade. Contudo, diante de uma literatura ainda carente de evidências mais abrangentes sobre o tema, outros trabalhos se fazem necessários para uma maior constatação das inferências reproduzidas nesse estudo.

¹⁵ Index terms— colorectal neoplasm. colectomy. laparoscopy. robotics.

6 LITERATURE REVIEW A) COLORECTAL CANCER

47 Palavras-Chave: neoplasias colorretais. colectomia. laparoscopia. robótica.

Introduction: Colorectal cancer is a malignant disease, more predominantly observed in men and the third most incident tumor among all cancers, with an estimated risk of 26.6 / 100 thousand. Despite its high incidence and prevalence, it is amenable to treatment, and in most cases, it is curable -when detected in early stages.

⁵¹ Objective: To compare the safety and efficacy of performing robotic surgery with traditional laparoscopic ⁵² surgery in patients undergoing colorectal cancer resection regarding the variables: intra and postoperative

53 complications, surgical conversion, and mortality.

54 Methods: This is a systematic review characterized by the

55 1 Introduction

ince the 1980s, when the first robotic surgery was performed 1, much has been said regarding this new technology
and its potential future capabilities. Over the years, robotic surgery has broken the boundaries of innovation in
health technology for better clinical outcomes. Thus, linked to a growing need for more precise and minimally
invasive surgeries, robotics was developed to meet these demands. Nowadays, it performs several functions related

to surgical practice -from assisting in the conduct brain biopsies to performing resection of malignant colorectal
 tumors. Several specialties such as urology, gynecology, cardiology, neurosurgery, and general surgery can use
 robotic surgery 1.

Among the technical advantages offered to surgeons are: the potential for three-dimensional visualization of the structures analyzed, elimination of the physiological tremors produced by the movementsallowing greater accuracy-improved surgical maneuvers permitted by the "robotic wrist" mechanism (positioning of surgical instruments at angulations not previously allowed by the laparoscopic technique), less fatigue of the surgeon, faster surgical recovery and with fewer complications compared to laparoscopy [1][2][3]. However, robotic surgery should be reserved for procedures in which technology can provide maximum benefit, in general when it

69 is necessary to perform precise dissections in confined areas, due to its current high operational cost 3.

This procedure has been becoming more popular since Pigazzi et al. 3 described for the first time the total excision of a malignant rectal tumor performed through robotic surgery in 2006. However, there is still not enough evidence in the literature regarding the safety and effectiveness of robotic surgery compared to traditional

⁷³ laparoscopy in cases of resection of malignant colorectal tumors 2,3.

74 **2** II.

75 3 Objectives a) Primary objective

To compare the safety and effectiveness of robotic surgery with traditional laparoscopic surgery in patients undergoing colorectal cancer resection.

⁷⁸ 4 b) Secondary objective

To compare intraoperative complications rates, surgical conversion, postoperative complications, and mortality
of robotic surgery with laparoscopy in colorectal cancer resection surgeries.

⁸¹ **5 III.**

⁸² 6 Literature Review a) Colorectal cancer

Colorectal cancer is a tumor that affects the large intestine, which is divided into colon and rectum. An essential aspect of this pathology is that the vast majority originates from polyps -small elevations in the colon and/or rectum wall -which grow slowly, starting with an aberrant crypt and developing into a neoplastic precursor lesion and then, finally becoming colorectal cancer. This process can take 10 to 15 years to occur. Thus, these polyps can be palliatively identified and removed before they can even produce malignancy characteristics.

However, some decades ago, colorectal cancer was rarely diagnosed due mainly to a lack of preventive practices
and technological resources. Hence, this pathology used to be diagnosed at extremely advanced stages when no
therapy could reverse the existing problem 4 .

Currently, colorectal cancer is the fourth most lethal cancer globally, causing the death of about 900,000 people each year, accounting for about 10% of the incidence of all cancers diagnosed annually and of cancerrelated deaths worldwide 4,5. It ranks as the second most common cancer among women and the third most common cancer among men. Its major risk factors are lifestyle-related. Intake of red meat, processed meat, fats, sedentariness, obesity, smoking, alcoholism, family predisposition, previous polyps, and age over 50 are conditions that predispose new polyps to appear and consequently increase the likelihood of developing colorectal cancer 4.

The most common signs and symptoms associated with this pathology are hematochezia, anemia with no apparent cause, abdominal discomfort, mild fever, severe weight loss, bowel habit changes, a continued desire to evacuate even after the evacuation, and gas or colic. Nevertheless, colorectal cancer can progress as a silent and asymptomatic disease until it reaches an advanced stage 4.

The diagnosis is based on the association of clinical findings with performing a colonoscopy and other imaging examinations such as computed tomography and laboratory tests such as blood count and concentration of

carcinoembryonic antigen that can be used as complementary tests 4 . Colonoscopy should regularly investigate 103 rectal bleeding in patients over 45 years of age. In younger patients, some additional factors should be considered 104

for increasing diagnostic suspicion: the presence of unfavorable family history, marked and unexplained weight 105 106 loss, and changes in intestinal habit 4.

Through technological advances and the increased possibility of early diagnosis, some cancers are only amenable 107 to local treatment. Incipient polyps can be resected endoscopically, also allowing precise evaluation of risk 108 characteristics, such as the depth of submucosal invasion, lymphatic invasion, presence of the tumor, and its 109 differentiation 4. 110

Surgery is the main therapeutic procedure for treating colorectal cancer, often with radio-and chemotherapy 111 support. The optimal resection of the tumor is fundamental and can be evaluated through safe and objective 112 parameters. Rectal cancer surgery is a complex process because of the difficult access to the surgical site, provided 113 by the limiting pelvic anatomy. Total mesorectal excision is the standard oncologic approach for rectal cancer, 114 and its extent depends mainly on the involvement of the sphincter complex and other surrounding structures. 4 115 Several factors are associated with better prognosis and increased quality of life after surgical treatment. These 116 factors are mostly the same related to colorectal cancer prevention. Thus, patients who adapt to a healthy 117 lifestyle after definitive diagnosis had a 33% lower risk of death during follow-up than those who did not include 118 119 this habit in their daily practices 4.

7 b) Robotic versus laparoscopic surgery 120

During the years of development of surgical practice, minimally invasive techniques allowed laparoscopic 121 interventions in the treatment of colorectal cancer patients. Subsequently, several randomized studies have 122 shown that laparoscopic colectomy is associated with lower morbidity rates, less surgical trauma, and better 123 immediate postoperative results, with shorter recovery times and hospital stays compared to surgery performed 124 through laparotomy 6. 125

126 However, a laparoscopic approach in rectal cancer patients is significantly different and more difficult than laparoscopic procedures in patients with colon cancer 6. This is explained by the difficulty of visualization and 127 surgical access at the pelvic anatomic site where the procedure should occur. Deep dissection in the pelvis to 128 perform a total mesorectal excision and obtain a sample with intact margins, making a safe anastomosis are 129 demanding techniques, besides promoting a considerable probability of reoperation 6. 130

Corroborating the hypothesis that the laparoscopic approach for rectal neoplastic procedures is a complex 131 and laborious procedure, the British randomized clinical trial CLASICC 7 in 2006 compared laparotomy and 132 laparoscopy performing 794 colorectal cancer surgeries. This study indicated that rectal excision by laparoscopy 133 resulted in a high conversion rate (38% in the first year, decreasing up to 16% in the last year) and a tendency 134 for greater positivity of the circumferential excision margin. Some other studies also present the same conclusion 135 regarding high conversion rates during colorectal laparoscopic surgery [8][9][10]. 136

The recent introduction of the robotic surgical system has revolutionized the field of minimally invasive 137 surgery. This new technology allows surgeries with a three-dimensional visual field, better ergonomics for the 138 surgeon (by reducing the fatigue), more extensive and better movement amplitudes of the forceps and other 139 surgical instruments, besides eliminating the physiological tremors produced by human arms [1][2][3]11,12. 140 Thus, adopting a robotic surgical system to perform colorectal cancer resection procedures seems attractive from 141 this perspective. Since this new technique can be safer for patients submitted to it -mainly concerning the greater 142 ease of management of an area as confined as the pelvic region -always aiming at a safe surgical procedure, free of 143 complications, with higher overall survival, disease-free survival, and quality of life, which are the most relevant 144 objectives of colorectal cancer treatment. 145

Another advantage related to the robotic surgical procedure is the possibility of using an infrared fluorescent 146 intraoperative imaging system with indocyanine green. This system allows the best identification of noble 147 structures such as vessels, nerves, and lymphatic ducts, thereby facilitating solid organs' partial resection, without 148 damaging their neighboring anatomical structures 13. 149

Nevertheless, robotic surgery still demands a high financial investment to be performed 14, besides counting 150 on some practical obstacles such as the long learning curve, longer surgical time, and size of the robotic system 151 15. Hence, within a publicly funded health system, the replacement of laparoscopic surgeries by I robotic ones 152 in colorectal operation requires a complete and thorough analysis so that their benefits are indeed validated. 153

The current literature evidence is that robotic rectal excision has been verified as feasible and safe, but these 154 conclusions were mostly based on statistically non-significant differences. Therefore, this systematic review has 155 great value to clarify the evidence available in the literature about the advantages of robotic surgery in comparison 156 to traditional laparoscopic surgery in cases of colorectal cancer resections. 157 IV.

158

Material and Methods 8 159

9 a) Study design 160

Systematic literature review. 161

¹⁶² 10 b) Search strategy

The literature review was carried out on September 10, 2019, in the electronic databases MEDLINE/PubMed, Scielo, Embase, and Cochrane, through the combination of descriptors, including terms from Medical Subject Headings (MeSH) and Health Science Descriptors (DECs), using publications in English and Portuguese: roboticassisted conventional laparoscopic surgery colorectal cancer resection. The terms used for the search were related to the population of interest, the parameters to be studied and the outcomes of morbidity and mortality: roboticassisted

AND conventional [All Fields] All **Fields** AND 11 ("la-169 paroscopy"[MeSH Terms] OR "laparoscopy"[All Fields] 170 OR ("laparoscopic" [All Fields] AND "surgery" [All Fields]) 171 OR "laparoscopic surgery" [All Fields]) AND ("colorectal 172 [MeSH Terms] OR ("colorectal" [All Fields] neoplasms" 173 AND "neoplasms" [All Fields]) OR "colorectal neoplasms" 174 [All Fields] OR ("colorectal" [All Fields] AND "cancer" [All 175 Fields]) OR "colorectal cancer"[All 176

Fields]) AND resection[All Fields]. References in the articles identified by the search strategy were also manually
 searched to add to the study and literature review.

¹⁷⁹ 12 c) Inclusion criteria

There were included studies, with a sample size greater than 30, published from October 2006 to December 2018, comprising patients who underwent robotic or laparoscopic surgery to perform cancer resection in the colorectal region. The clinical outcomes of interest were: surgical time, surgical conversion, other intraoperative and postoperative complications, length of hospital stay, and mortality.

¹⁸⁴ 13 d) Exclusion criteria

185 Reviews, case reports, case series, comments, and correspondence were excluded.

¹⁸⁶ 14 e) Identification and selection of studies

The authors read each pre-selected article's titles and abstracts from the electronic database research to identify only those studies that correctly fulfill the inclusion and exclusion criteria. Subsequently, the full texts were read,

- 189 ensuring the criteria for the systematic review.
- Both authors discussed the divergences trying to respect the inclusion and exclusion criteria previously defined.

¹⁹¹ 15 f) Data extraction

Two authors collected the data using a predefined collection form. The characteristics of interest of the studies 192 included: geographical origin, title, type of study, duration of the study, number of participants, and mean age of 193 the sample. Finally, data were collected related to surgical time, intraoperative and postoperative complications, 194 conversion, length of hospital stay, and mortality. The quality of each study characterized as a randomized clinical 195 trial was evaluated by the Cochrane Tool -Consolidated Standards of Reporting Trials (CONSORT) 16 to assess 196 the risk of bias, which contains the following criteria: adequate randomization; allocation of participants; blinding 197 of participants; blinding of the outcome evaluator; integrity of results; incomplete data; selective outcome reports; 198 and other sources of bias (e.g., the effect of small studies). Strengthening the Reporting of Observational Studies 199 in Epidemiology (STROBE) 17 was used for the methodological evaluation of observational articles. 200 V. 201

202 16 Results

²⁰³ 17 a) Identification and selection of studies

Through the search strategy, 20 records were identified after the exclusion of duplicate studies. Based on the reading of the title and abstract, 8 articles were left for a full reading. Of these, one study was excluded because it did not reach the minimum sample size. Therefore, 7 articles were selected for the systematic review (Figure 1).

²⁰⁸ 18 b) General characteristics of the obtained studies

The samples ranged from 56 to 471 participants (n total = 1589), with an age range of 61.2 -69.0. All studies 209 included both genders, and among these, only one20 reported a higher proportion of women. Among the selected 210 studies, five are characterized as retrospective cohorts and two as randomized clinical The quality assessment of 211 the selected observational studies was performed with the STROBE17 tool, available in the STROBE initiative, 212 verified in Chart 3. 6 conducted an analysis exclusively related to low rectal cancer. The records were collected and 213 prospectively acquired from all patients at Kyungpook University Hospital with rectal cancer located 8 cm from 214 215 the anal margin. After this process, the information was reviewed retrospectively. Patients with tumors causing intestinal obstruction or perforation, local resectable tumor with transanal access, invasion of adjacent organs 216 requiring multiple organ en bloc resection, and distant metastasis were not considered suitable for laparoscopy 217 or robotic surgery. 218

The choice between the two different surgical approaches was based on a joint decision between patients and physicians, and the use of robots did not Levic et al. 19 Ramji et al. 14 Yamaguchi et al. 20 Crolla et al.

221 19 Results

222 20 Participants 13

- 223 Descriptive data 14
- 224 Outcome data 15

225 21 Main results 16

226 Other analysis 17

229

227 22 Discussion

228 23 Key results 18

Limitations 19 Interpretation 20 Generalisability 21 Other Information Funding 22

liver function tests, serum carcinoembryonic antigen (CEA), chest X-ray, and electrocardiogram. Colonoscopy, 230 abdominopelvic computed tomography, and pelvic magnetic resonance imaging were routinely performed to 231 evaluate distant metastases, local infiltration of the disease, and tumor characteristics. This study had limitations 232 due to its retrospective nature and its inherent selection bias. Another established limitation is related to the lack 233 of a detailed economic comparison between the two groups. Some differences in short-term results were considered 234 insufficient to justify the costs of using the new technology. Rodríguez et al. 18, in 2011, besides analyzing 235 rectal cancers, evaluated the occurrence of tumors in the sigmoid. All patients underwent preoperative analysis, 236 including hemogram, liver function, and biochemical tests, chest radiographs, and electrocardiograms. Patients 237 diagnosed with rectal cancer were also submitted to colonoscopy with biopsy for the histological diagnosis of the 238 lesion, accompanied by thoracoabdominal computed tomography, magnetic resonance imaging, and ultrasound 239 examinations. This study analyzed patients' clinical conditions through the American Society of Anesthesiology 240 (ASA) classification and performed histological analyses to define the distance of the distal margin, the total 241 number of resected lymph nodes, and the total length of the sample. This research did not present its eventual 242 limitations. 243

²⁴⁴ 24 Medical Research Volume XXI Issue I Version I

Levic et al. 19 conducted a retrospective and multicenter analysis in 2014. The patients considered appropriate 245 for the laparoscopic technique were over 18 years old and had rectal cancer without metastasis. Exclusion criteria 246 were magnetic resonance imaging (MRI) or preoperative computed tomography (CT) showing tumor size >4 cm 247 in diameter or evidence of local invasion (T4 cancer); ASA class IV/V; the anticipated need for intensive care unit 248 (ICU); a history of major anterior abdominal surgery and obese patients with body mass index (BMI) > 32 kg/ 249 m2. Inclusion criteria for robotic surgery were practically the same, except that high BMI was not a reason for 250 exclusion. The tumor staging and preoperative evaluation consisted of a digital rectal examination, proctoscopy, 251 histopathological examination, thoracoabdominal computed tomography, and pelvic magnetic resonance imaging. 252 All patients were discussed at the multidisciplinary team conference before the treatment decision. 253

This study's limitations were the restricted number of patients in each group and the short followup, which made it impossible to reach satisfactory conclusions about the long-term oncologic effects and any possible differences in late complication rates. Moreover, the authors presented the selection bias as a limitation since the study was not randomized, as well as the learning curve of surgeons for both techniques since this can cause distorted results in any direction.

The retrospective study by Ramji et al. 14, in 2015, additionally compared robotic and laparoscopic surgical procedures to laparotomy. The analysis was the only one that compared the economic feasibility between the surgical techniques. This study also analyzed the patients' tumor characteristics according to the ASA classification and comorbidities' existence through the Charlson score. The included cases required a confirmed histological diagnosis of rectal adenocarcinoma and could not be associated with recurrent or synchronous disease.
 Cases with multivisceral involvement and palliative intention were excluded. The study showed limitations related
 to the small number of cases assisted by robotics concluded until its institution.

In their study in 2015, Yamaguchi et al. 20 included all patients who underwent proctectomy for rectal 266 adenocarcinoma at Shizuoka Cancer Center Hospital. Patients undergoing open surgery, high anterior resection, 267 lateral lymph node dissection, or multiple resections were excluded. The preoperative tumor staging was carried 268 out according tocolonoscopy findings, computed tomography, magnetic resonance imaging, and barium enema. 269 The rectal cancers were staged using the tumor-node-metastasis (TNM) classification. The surgical method to be 270 performed was decided through a physician's discussion with the patient. After providing informed consent, the 271 patients selected their preferred approach -however, rectal cancer surgery with lateral lymph node dissemination 272 was performed by the open method if the patient did not desire to undergo robotic surgery -a condition that 273 reproduces a selection bias, somehow restricting the internal validity of the study. The first limitation present in 274 the study was related to the fact that it was a retrospective analysis that potentially included several selection 275 biases. The second limitation was established because of the lack of evaluation of sexual function after surgery. 276

Jayne et al. 21 conducted a randomized multicenter study in 2017, which included 29 different medical 277 centers in 10 countries and 40 surgeons. The patients included were those with an indication for resection and 278 279 were diagnosed with rectal adenocarcinoma. Patients with benign lesions of the rectum, anal canal cancers, 280 locally advanced cancers, or those requiring multivisceral block resection or multiple surgical resections were excluded from the study. The study additionally evaluated bladder function and sexual function through the 281 International Prostate Symptom Score (I-PSS), International Index of Erectile Function (IIEF), and Female 282 Sexual Function Index (FSFI). This research presented limitations related to the low number of patients analyzed 283 -conferring statistically insignificant results among the treatment groups. No blinding was established for this 284 study, consequently affecting the study's primary outcome and mortality measures. 285

In 2018, Crolla et al. 22 carried out their study using a prospectively filled database -with data routinely 286 collected from patients. Multiple organ resections were excluded. Regarding its limitations, this study presented 287 several diagnostic and therapy protocol changes throughout the development period. The randomization process 288 was not performed. This study also did not consider the surgeons' learning curve or adequacy. Besides, the authors 289 showed that confounding factors related to general morbidity might have been neglected. The main intraoperative 290 complications recorded by the studies, besides the surgical conversion, were: significant hemorrhage, need for 291 intraoperative transfusion, injury and/or perforation of the rectum, equipment failure, fecal contamination, and 292 inadvertent perforation of the tumor. Rodríguez et al. 18, in 2011, and Ramji et al. 14, in 2015, did not detail 293 the intraoperative complications analyzed in their studies. 294

The most significant postoperative complications described by the studies included in this review: anastomotic dehiscences, urinary retention, need for reoperation, anemia with the need for transfusion, and infection of the wound or surgical region. Rodríguez et al. 18, in 2011, did not perform an analysis of postoperative morbidities, and Yamaguchi et al. 20, in 2015, did not specify the postoperative morbidities recorded besides anastomotic dehiscence and infection of the surgical site.

From the studies added to the systematic review, Park et al. 6, in 2010, Ramji et al. 14, in 2015, and 300 Yamaguchi et al. 20, in 2015, classified patients through Clavien-Dindo postoperative complications severity 301 classification system, verified in Table 2. Park et al. 6, in 2010, divided patients into two groups: the first 302 integrating classifications I and II, while the second joined classifications III and IV. The other researches did 303 not make any reference to this classification tool. The most used type of surgery among the studies was the 304 low anterior resection, followed by the abdominoperineal resection, shown in Table 4. Most studies included 305 only rectal cancer in their analysis. Park Robotic Surgery versus Laparoscopy in Colorectal Cancer Resection: 306 A Systematic Review et al. 6, in 2010, were even more specific and analyzed only low rectal cancers. Only 307 Rodríguez et al. 18, in 2011, additionally analyzed colon cancers in their studytotaling 44 patients. 308

309 25 Rodríguez

et al. 18 Levic et al. 19 Ramji et al. 14 Yamaguchi et al. 20 Jayne et al. 21 Crolla et al. 22 Low anterior resection

312 26 Discussion

The present study aimed to select four main variables related to the efficacy and safety of different surgical approaches: prevalence of intraoperative complications, surgical conversions, postoperative morbidities, and mortality. This systematic review obtained a total sample of 1,589 patients submitted to colorectal cancer surgery, either by laparoscopic or robotic technique.

The prevalence of intraoperative complications from laparoscopy ranged from 0% to 14.8%, and the most prevalent among the complications mentioned in the studies were: significant hemorrhage, damage to some organ or structure, low rate of anal sphincter preservation and surgical equipment failure. Yamaguchi et al. 20, in 2015, and Crolla et al. 22, in 2018, showed no results for this variable. Park et al. 6, in 2010, Ramji et al. 14, in 2015, and Levic et al. 19, in 2014 presented prevalence below the average of studies included in the review, while in the studies by Rodríguez et al. 18, in 2011, and Jayne et al. 21, in 2017, showed above average results. The

reason for Rodríguez et al. 18, in 2011 and Jayne et al. 21, in 2017 being the only studies with an above-average 323 prevalence of intraoperative complications is because most studies did not present an adequate sample size in 324 order to obtain statistically significant results and avoid type II error -this being the main limitation mentioned 325 in the studies. Thus, Jayne et al. 21, in 2017, probably because of a more significant sample number (230), 326 was the study that most closely resembled the data available in the literature, which present an approximate 327 average prevalence of intraoperative complications of 16.5% [23][24][25][26][27]. The prevalence of intraoperative 328 complications related to robotic surgery ranged from 0% to 15.3%. Yamaguchi et al. 20, in 2015, and Crolla 329 et al. 22, in 2018, also showed no results for this variable. Park et al.6, in 2010 and Ramji et al. 14, in 2015 330 reported no intraoperative complications related to robotic surgery, while Rodríguez et al. 18, in 2011, Levic 331 et al. 19, in 2014, and Jayne et al. 21, in 2017 presented similar results with those found in the literature, 332 which has an average prevalence of approximately 14% [28][29][30][31][32]. The rationale used by these studies is 333 related to the lack of tactical sensitivity that the robotic system transmits to the surgeon, especially to those who 334 are at the beginning of their learning curve, consequently causing damage to the patient's organs and structures. 335 To prove this rationale, Rodríguez et al. 18, in 2011, went further, and performed a brief review on the possible 336 causes of intraoperative complications in robotic surgery, finding a result that corroborates with the rationale 337 mentioned above. 338

339 27 Medical

Comparing the laparoscopic and robotic approaches concerning the rate of intraoperative complications, the rate related to the robotic surgery group (6.0%) was higher than the rate of complications related to laparoscopic surgery (5.2%), being registered 7 more cases.

As mentioned by Crolla et al. 22, in 2018, "a low conversion rate is important because, in general, the 343 conversion is associated with more complications, longer hospital stay and worse long-term outcome". Thus. 344 regarding surgical conversion rates during laparoscopic surgeries, a prevalence ranging from 0% to 37% was 345 found. Park et al. 6 , in 2010, and Levic et al. 19 , in 2014, registered no surgical conversion. Rodríguez et 346 al. 18, in 2011, had a conversion rate of 7.14% and reported no statistical differences about robotic surgery. 347 Other studies that found significant differences about laparoscopy varied their prevalence between 3.3% -37% 348 and reported that the main reasons for the occurrence of surgical conversions in this type of technique were: 349 difficulty of visualization, visible anastomotic leaks, adhesions, stapler complications, tumor invasion of adjacent 350 structures and difficulty in manipulating the target organ. 351

The robotic technique's prevalence of surgical conversion rates was found to vary between 0% -8.1%. Park 352 353 et al. 6, in 2010, and Yamaguchi et al. 20, in 2015, did not report surgical conversion. The study by Levic et al. 19, in 2014, was the only one that presented more conversions (3 versus 0) during robotic surgery. The 354 355 other studies always showed a lower conversion rate compared to laparoscopic surgeries. The leading causes for 356 surgical conversion during the robotic approach were: the presence of severe fibrosis in the pelvis as a sequel to 357 radiotherapy with a rectal lesion, tumor fixation, and perforation of the rectum due to a narrow pelvis. The studies justified a better performance of robotic surgery in this field by the improved visualization with the 3D 358 359 camera and a better capacity to maneuver the surgical instruments. Jayne et al. 21, in 2017, still mention that the benefits of robotic surgery for surgical conversion rates are enhanced when surgeons already have some 360 experience in the practice of robotic surgery itself, i.e., when they are at a high level in their learning curve. 361

By making a parallel between the two approaches analyzed, robotic surgery compared to conventional laparoscopic surgery in colorectal cancer improved the conversion rate, presenting a considerably lower percentage. However, the authors showed that although data related to robotic surgery have achieved better blood loss rates and fewer conversions compared to laparoscopy, this may be less a reflection of the surgical tools used and more a result of the surgeon's improved skill and experience in minimally invasive surgery, which can be considered a confounding bias.

The prevalence of morbidity after surgery related to laparoscopy ranged from 22.6% to 60%. Rodríguez et al. 368 18, in 2011, presented no results for this variable. he other studies presented similar prevalence, ranging from 369 22.6% to 31.7% -except Crolla et al. 22, in 2018, who reported a rate of about 60% -well above the average 370 of approximately 29% found in the literature 21, [31][32][33]. This discrepant result was established due to the 371 introduction of an additional variable combined with the postoperative complications mentioned, called by the 372 study of "any other complications" without, however, describing what these possible complications would be. 373 The most mentioned postoperative complications in the analyzed studies were: anastomotic dehiscence, urinary 374 retention and other urinary complications, the need for reoperation, infection of the surgical site, bleeding with 375 376 the need for transfusion, and cardiorespiratory complications.

377 Regarding postoperative morbidity related to robotics, the prevalence varied between 8.9% -42.3%. Rodríguez 378 et al. 18, in 2011, presented no data for this variable. Yamaguchi et al. 20, in 2015, showed the lowest 379 prevalence of morbidity. The study identified fewer occurrences of urinary retention, wound infection, small bowel obstruction, anastomotic dehiscence, intraabdominal or intraluminal bleeding, and enteritis. Among these 380 complications, the least recurrence of urinary retention was emphasized, and the rationale found for such an event 381 was that "[...] This is probably due to the superior free-moving multi-joint forceps, highquality three-dimensional 382 imaging, and steady "traction and counter-traction" allowing easier recognition and preservation of the pelvic 383 splanchnic nerves and inferior hypogastric plexus". Crolla et al. 22, in 2018, presented a 42.3% prevalence 384

-a result above the average found in the literature of approximately 27% 28,34,35 Furthermore, it is worth
mentioning that colorectal cancer surgery is a high-risk intervention, which depends significantly on the patient's
tumor characteristics and good general condition. Therefore, it is expected that about 1/3 of the patients present
postoperative complications in less than 30 days 21. This data agrees with the selected studies' variation and is
valid for both the laparoscopic and robotic techniques, with no significant difference being observed concerning
general postoperative morbidity.

The mortality variable in the laparoscopic surgery subgroup was identified as a prevalence ranging from 0%391 to 5.6%, similar to data found in the literature, which defines average mortality of 2% 33,36. Park et al. 6, in 392 2010, Rodríguez et al. 18, in 2011, Ramji et al 14, in 2015, and Yamaguchi et al. 20, in 2015, reported no 393 deaths, while Levic et al. 19, in 2014, Jayne et al. 21, in 2017, and Crolla et al. 22, in 2018, showed a mortality 394 rate of 5.6%, 0.9% and 4.9%, respectively. Levic et al. 19, in 2014, despite recording the highest percentage 395 of mortality, presented only two deaths in a total of 36 patients, not representing statistical significance. In all 396 studies reported in this review, there were a total of 13 deaths related to laparoscopy. Most of the deaths were 397 associated with organ and structure perforation-causing extensive hemorrhage during surgery and postoperative 398 sepsis. 399

Regarding robotic surgery, mortality prevalence variation was between 0% -0.8%, a result compatible with the average found in the literature of about 1%28, 37. The only studies reported deaths were conducted by Jayne et al. 21, in 2017, and Crolla et al. 22, in 2018, recording 3 negative outcomes. The rationales for the deaths involving robotic surgery were the same as for laparoscopy.

By comparing the mortality rates of the two surgical techniques, robotics presented 3 deaths out of a total of 789 patients included in the study, which represents a percentage of 0.38%, while laparoscopy showed a total of 13 deaths out of a universe of 800 patients, representing a percentage of 1.6%. Thus, robotic surgery proved promising since the researches revealed a lower mortality rate compared to laparoscopic surgery.

The different methodologies presented by the studies, besides the low sample value of some of them during the analysis of the variables, indicate the need for additional research on the comparison between laparoscopy and robotic surgery in colorectal cancer resection. Larger samples and clearly defined methodological criteria are needed to establish the safety and efficacy of each approach. Also, this present study has not been extended to a meta-analysis to obtain a better statistical result is defined as a limitation.

413 28 VII. Conclusion

Given the described findings, evidence of good to moderate quality supports that robotic surgery for colorectal cancer resection produces similar perioperative and postoperative results, even though it promotes better ergonomics and comfort for the surgeon. However, robotic surgery reflects lower surgical conversion and mortality rates. In the face of the literature still lacking more extensive evidence on the topic, other studies are necessary for more significant verification of the inferences reproduced in this study.

419 29 Medical



Figure 1: Figure 1 :



Figure 2:

420

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2	Figure 3: Chart 2 :
3	Figure 4: Chart 3 :
	Figure 5:
	Figure 6:
	Figure 7:
	Figure 8:

Park et al. 6 Retrospective cohort			South	41	82	(73/50)	63.0	61.2	120
			Korea						
Rodríguez et al. 18	Randomized clinical tr	ial 2011	Spain	28	28	(29/27)	61.5	68.0	19
Levic et al. 19	Retrospective cohort	Denmark	36	56	(51/41)	69.0	65.0	24	
Ramji et al. 14	Retrospective cohort 2015		USA	27	26	(38/15)	63.7	62.1	24
Yamaguchi et al. 20	Retrospective cohort	2015	Japan	239	203	(294/148)	65.9	64.8	45
Jayne et al. 21	Randomized clinical trial 2017		United	234	237	(234/237)	65.5	64.4	12
			King-						
			dom						
Crolla et al. 22	Retrospective cohort	2018 N	etherlands	184	168	(216/136)	68.1	67.0	60

Figure 9: Mean age (years) Study time (months) Laparoscopy Robotics Laparoscopy Robotics

Introduction	
Background/Rationale	2
Objectives	3
Methods	
Study design	4
Setting	5
Participants	6
Variables	7
Data	8
source/Measurement	
Bias	9
Study size	10
Quantitative variables	11
Statistical methods	12

Figure 10: 22 Title and abstract 1

1

Source: The author (2020) Legend: NR: Not referred

Figure 11: Table 1 :

Study	Surgical time		Surgical conversion		Ν
	(in minutes $)$		(n)		р
	Laparoscopy	Robotics Lapar	roscopy Robotics Laparoscopy Robotics Lapare	oscopy Robotics	
Park et	168.6	231.9	0	0	1
al. 6					
Rodríguez	135.1	159.4	2	2	Ν
et al. 18					
Levic et	295	247	0	3	1
al. 19					
Ramji et	240	407	10	3	$\overline{7}$
al. 14					
Yamaguch	i 227.6	232.9	8	0	5
et al. 20					
Jayne et	$261\ 172$	$298.5\ 219$	28	19 3	$\overline{7}$
al. 21 al.			23		
22 Crolla					
et					

Study Park et Length of hospital stay (days) Laparoscopy Robotics 9.4 9.9 Postoperative Cla al. 6

Rodríguez et	$9.2\ 7\ 11.3$	9.3		N
al. 18 Levic et al. 19 Ramji		$\frac{8.0}{7}$	I:6 II: 0 III: 0	0 I
et al. 14				3
			IV: 4	Γ
Yamaguchi et	9.3	7.3	0: 1	0
al. 20			I: 107	I
			II: 41	Ι
			III: 73	Ι
			IV: 17	Γ
Jayne et al. 21	8.2	8	NR	Ν
Crolla et al.	7	6	NR	Ν
22				

Figure 12:

 $\mathbf{2}$

[Note: Source: The author (2020) Legend: NR: Not referred]

Figure 13: Table 2 :

 $\mathbf{4}$

Types

Figure 14: Table 4 :

Figure 15: of surgery Study Park et al. 6

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Figure 16:

Surgery time (min) Laparos copy Robo tics		oRobo	Postoperat Laparos copy	tive morbidities (Extra which ones?) Table? Describe Robo tics
1 168. 6 231. 9	0	0	19	12
$\begin{array}{cccccccccccccccccccccccccccccccccccc$	2	2	x	x
3 295 247	0	3	10	12
4 240 407	10	3	1	0
5 227. 232. 6 9	8	0	54	18
6 261 298. 5	28	19	73	78
7 172 219	23	3	223	178

Figure 17:

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430 .2 Appendixes

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