



Original Research Article

Assessment of pre-operative factors for predicting a difficult laparoscopic cholecystectomy

Neeraj Kumar Jain¹, Manjari Goel Jain², Sanish Philips¹ and Rinku Yadav^{3,*}

- ¹ Department of Surgery, Chirayu Medical College, Bhopal M.P. India.
- Department of Obstetrics and Gynecology, RKDF Medical College, Bhopal M.P. India.
- ³ Department of Surgery, N.S.C. Govt. Medical College, Khandwa, M.P. India.
- * Correspondence: drrinkugrmc8622@gmail.com

Received: 5 January 2023; Accepted: 10 April 2023; Published: 15 April 2023.

Abstract: Background: Laparoscopic cholecystectomy (LC) is considerably gold standard for symptomatic cholelithiasis. Preoperative prediction of difficult LC and operative grading system may not only improve patient safety but also be beneficial in lessening the overall cost of therapy.

Aim: The present study aimed to predict and analyze risk factors using a scoring system deemed responsible for surgical difficulties in patients undergoing laparoscopic cholecystectomy (LC) for symptomatic cholelithiasis.

Material & Methods: This hospital based prospective study was conducted at Department of Surgery, A tertiary care teaching Hospital, central India. Various factors considered preoperatively were gender, age, previous history of hospitalization, impacted stone, obesity; gall bladder wall thickness, pericholecystic collection; previous abdominal scar and palpable gall bladder were evaluated.

Results: Out of 150 patients, majority of the (34.2%) was 41-50 years age group, predominantly female (62.7%). The scoring system predicted easy LC for 106(70.7%), and difficult for 44(29.3%). No significant association of difficult LC with age group, gender, BMI, h/o hospitalization with acute cholecystitis, abdominal scar, leucocytes count and Hepatic echotexture (P>0.05), whereas palpable gall bladder, GB wall thickness, Distended gallbladder, impacted stone and pericholecystic collection were significantly associated with difficult LC.

Conclusion: Preoperative prediction of the factors leading to difficulty or conversion in cholecystectomy could help plan the surgical strategies and possible outcomes beforehand which can reduce the overall mortality and morbidity.

Keywords: Preoperative scoring system: Gallbladder difficult cholecystectomy; Laparoscopic cholecystectomy.

1. Introduction

holecystectomy is a surgical procedure to remove the gallbladder due to stone or inflammation, and it's the most standard procedure performed in the biliary tract [1]. Laparoscopic cholecystectomy (LC) has become the preferred method and has been accepted as the gold standard for definitive management of symptomatic cholelithiasis or gallstones [2,3]. Laparoscopic cholecystectomy has many advantages over the open cholecystectomy, such as better cosmetic results, short hospital stay, minimal trauma, decreased pain, early recovery and return to physical activity and work, all have contributed to the popularity of this technique [4,5].

Multiple factors that may influence the difficulty of a cholecystectomy have been described, which may be related to the patient, such as male sex, old age, anatomical variations, previous surgeries, obesity, severe inflammation or impacted stones, thickened gall bladder wall, distended gall bladder, pericholecystic collection and external factors such as failure of inappropriate equipment [6,7].

In the early years of the laparoscopic cholecystectomy era, the conversion rate to open procedure was 2-15%. After years of learning and understanding the laparoscopic technique and increasing surgeons'

experience, the conversion rate dropped to approximately 1-6%) is conversion was an attempt to avoid complications due to various difficulties encountered during the procedure [8].

Many surgeons often face difficulties in performing LC due to their inability to precisely identify the anatomy of Calot's triangle as a result of severe inflammation. Therefore, in patients with severe acute cholicystities (AC) the rate of complications, such as bile leakage, common bile duct injury, bleeding, visceral injury and bowel injury, is high after LC [9,10].

Most of the technical disadvantages associated with LC directly or indirectly result from the creation of pneumoperitoneum [11].

2. Aims & Objectives

The present study was undertaken to identify various factors associated with laparoscopic cholecystectomy that may help us in predicting or anticipating a difficult laparoscopic cholecystectomy beforehand in the pre-operative period, hence allowing the surgeon to prepare in advance for the difficulties encountered during the procedure.

3. Material and Methods

This prospective randomized study was conducted in the department of general surgery, in a tertiary care hospital, central India for the period of two years. The patients having symptomatic cholelithiasis willing to undergo laparoscopic cholecystectomy were enrolled in the study. Informed written consent was obtained from all patients who accepted for participation in the study.

3.1. Inclusion criteria

- Cases of acute or chronic cholecystitis with cholelithiasis who underwent laparoscopic cholecystectomy.
- Age > 18 years.
- Patients willingly giving consent will be included in the study.

3.2. Exclusion criteria

- Age below 18 years.
- Viral marker positive patients (HBs Ag, HCV, HIV).
- Pregnancy.
- Patients with common bile duct (CBD) calculus, dilated CBD.
- Patients who refused laparoscopic cholecystectomy.
- Patients who were not fit for general anesthesia due to various medical illnesses.
- Patient who didn't give informed consent.
- Contraindications to Laparoscopic cholecystectomy like: Cardiovascular and pulmonary disease, coagulopathies and end-stage liver disease (ESLD).

Diagnosis of cholelithiasis was confirmed in patients presenting with abdominal symptoms using an abdominal ultrasonography (USG).

The patients were evaluated prior to undergoing laparoscopic cholecystectomy with a detailed history and examination followed by all the relevant investigations.

The variables assessed included age, sex, body mass index, history of acute cholecystitis attacks, palpable gallbladder mass, scar tissue in the abdominal region, leukocyte count, gallbladder wall thickness, gallbladder distension, Pericholecystic collection, impacted gallstones, and changes in hepatic echotexture. Each variable assessed will be given a weight based on the Risk Scoring System for Laparoscopic Cholecystectomy. The scoring system used in this study was adapted from the scoring system published by Bhondave *et al.*, [12].

The scores for each variable are summarized in Table 1.

The assessment was divided into two categories, namely easy and difficult. The classification system was adapted from the criteria used by Bhondave *et al.*, [12]. A per operative score of 0 - 5 predicted as easy and > 5 considered as difficult.

In this study, we decided that if one of the criteria for laparoscopic cholecystectomy was difficult to fulfill, then the operation was declared difficult. The criteria for each category are summarized in Table 2.

 Table 1. Laparoscopic cholecystectomy surgery risk Preoperative scoring system

Parameter		Score
A co croup	>50 year	1
Age group	= 50 year	0
Gender	Male	1
Gender	Female	0
History of acute cholecystitis attack	Yes	4
	No	0
Body mass index (kg/m2)	>30	1
body mass maex (kg/ mz)	= 30	0
Dalmahla gallhladdar	Yes	1
Palpable gallbladder	No	0
Abdominal scar	Yes	1
Abdomma Scar	No	0
I audio auto accest	>12000	1
Leukocyte count	= 12000	0
Gallbladder wall thickness	>4 mm	2
Gandiauder wan unckriess	= 4 mm	0
Distended or contracted gallbladder	Yes	1
	No	0
Pericholecystic collection	Yes	1
	No	0
Impacted gallstone	Yes	1
	No	0
Changes in honetic ashetouture	Yes	2
Changes in hepatic echotexture	No	0

Table 2. parameters for assessment of prediction of laparoscopic cholecystectomy

Parameter	Classification	
These three criteria must be met:		
Duration of surgery = 60 minutes	East	
No bile acids are spilled into the surrounding tissue	- Easy	
No arterial or ductal damage		
There is one of the following criteria:	- Difficult	
Duration of surgery >60 minutes		
There are bile acids that spill into the surrounding tissue		
There are arterial or ductal damages		
Duration >120 minutes		
Converted to open cholecystectomy		

Patient characteristic		Frequency (N=150)	Percentage (%)	
Age group (in years)	<30	22	14.7%	
	31-40	35	23.3%	
	41-50	52	34.6%	
	51-60	25	16.7%	
	>60	16	10.7%	
Gender	Male	56	37.3%	
	Female	94	62.7%	
H/o hospitalization for acute cholecystitis	Yes	41	27.3%	
	No	109	72.7%	
Body mass index (kg/m2)	Normal <25	7	4.7%	
	Pre obese 25-29.9	60	40%	
	Obese 30-34.5	66	44%	
	Morbid obese >35	17	11.3%	
Palpable gallbladder	Yes	28	18.7%	
	No	122	81.3%	
A1. J	Yes	16	10.7%	
Abdominal scar	No	134	89.3%	

Table 3. Preoperative characteristics of the study patients

Table 4. Outcome of the operation among studied cases

Category of LC	Frequency (N=150)	Percentage (%)
Easy	106	70.7%
Difficult	44	29.3%
Total	150	100%

4. Statistical Analysis

In the present study, statistical analyses of data were carried out using SPSS version 22. Numerical data were expressed as mean \pm standard deviation. P value < 0.05 was considered significant.

5. Results

A total of 150 patients were undergoing laparoscopic cholecystectomy included in this study. Majority of the patients (34.6%) were 41-50 years age group, predominantly female (62.7%). The mean age \pm SD of the patient was 45.62 \pm 15.24 years. Most of the patients 66 (44%) was obese. 41 (27.3%) patients had past history of hospitalization for acute calculus cholecystitis while 16 (10.7%) had prior history of surgery with abdominal scar mostly midline. Gall bladder was palpable clinically in 28 (18.7%) patients. Preoperative characteristics of the study patients are given in Table 3.

Most of the patients had preoperative score of 0-5 and most (70.7%) were easy. However, 44 (29.3%) of patients were labeled difficult laparoscopic cholecystectomy according to pre and intraoperative assessment. Prediction score > 5. Outcome of the operation among studied cases is given in Table 4 and predictive association of risk factors with intraoperative outcome is given in Table 5.

6. Discussion

Prediction of the risk factors for laparoscopic cholecystectomy and safety of the procedure, surgeons could have benefit in deciding the surgical approach, counselling the patients, reducing the risk of complication, reducing the rate of conversion to open cholecystectomy, and reducing overall medical cost. Present study found that the most patients were in the age group of 41-50 years followed by age group of 31-40 years, our results was concordance with the Kulkarni *et al.*, [13] and Abdallah *et al.*, [14].

Majority of patients were in the age group of =50 years and found no significant correlation between age and the difficult laparoscopic cholecystectomy (P>0.05), accordance to the S. Baral *et al.*, [15] and Sandhu *et al.*, [16].

Risk factors		Easy LC (N=106) (70.6%)	Difficult LC (N=44) (29.4%)	χ^2	P value
Age group	>50	31	10	0.665	0.414
(in years)	= 50	75	34	0.003	
Gender	Male	38	18	0.340	0.559
Gender	Female	68	26	0.340 0.339	0.559
Body mass index (Kg/m2)	>30	60	20	1.55 0.2	0.212
body mass maex (Rg/m2)	= 30	46	24	1.55	0.212
History of cholecystitis	Yes	26	15	1.43	0.231
Thistory of cholecystitis	No	80	29	1.43	0.231
Abdominal scar	Yes	10	6	0.576	0.447
Abdominai scai	No	96	38	0.576	0.447
Palpabla gallbladdar	Yes	13	14	8.054	54 0.004
Palpable gallbladder	No	93	30	0.034	
Laukaartaa aaunt	>12000	48	25	1.05	0.198
Leukocytes count	= 12000	58	19	1.65	
Gallbladder wall thickness	>4 mm	46	28	5.096	0.023
Gailbladder wall thickness	= 4 mm	60	16	5.096 0.	0.023
D:-t d- d111-1- d d	Yes	7	9	()(0.012
Distended gallbladder	No	99	35	6.26	
Pericholecystic collection	Yes	9	10	E 607	0.016
	No	97	34	5.697	0.016
Impacted gallstone	Yes	25	27	10 502	0.001
	No	81	17	19.593 0.00	0.001
Hepatic echotexture	Yes	0	2	2.050	0.151
	No	106	42	2.058	0.151

Table 5. Predictive association of risk factors with intraoperative outcome

In this study, gender was not a significant risk factor in predicting difficult laparoscopic cholecystectomy (P>0.05), which tolerates with studies which were conducted by N Gupta *et al.*, [17] and Buono *et al.*, [18], reported that gender did not affect the prediction of difficulty in laparoscopic cholecystectomy.

This current study declares that difficulty in LC was increase with increases in BMI and obesity but statistically not significant (P>0.05), our finding was consistent with the Ahmed M.et al., [19] and Santharaj S,et al., [20].

In our study, the previous attacks of acute cholecystitis is found to be directly proportional to difficult LC but statistically not significant (P>0.05), similar finding reported by Maehira H, *et al.*, [21] and Madan *et al.*, [22]. The repeated episodes of acute cholecystitis are associated with excessive fibrosis in GB bed that fix GB to the surroundings, making Calot's triangle frozen with difficult dissection.

No significant correlation (P>0.05] between past abdominal surgeries (indicated by abdominal scar) and difficulties encountered during LC., concordance with the Sharma M *et al.*, [23] and Roy *et al.*, [24].

In our study we have find statistically significant association [p = 0.004] between palpable gall bladder and difficult LC, similar finding also reported by A Iqbal *et al.*, [25] and Agrawal *et al.*, [26]. If gall bladder is palpable then it would be difficult, as there may be residual inflammation adhesion. The total leukocyte count did not show a statistically significant association (p>0.05) with LC in the present study, accordance to the Wibowo *et al.*, [27] and Nidoni *et al.*, [28].

In this study, we had a statistically significant association of difficult LC with the thickness of the GB wall (P= 0.02), our results are comparable with the study conducted by Giraldo *et al.*, [29] and Lal P *et al.*, [30]. The thickness of GB walls (>4 mm) may leads to difficult grasping of the GB with dense adhesions, and this added to the difficulty of procedure.

We found that impacted stone at the neck of GB is significantly associated with difficult LC (p<0.05), same observation also reported by many other studies like Arone *et al.*, [31] and Bhardwaj R,*et al.*, [32]. Stone impacted in the neck of GB hinders grasping the GB neck during dissection; moreover, the GB is usually mucocele with difficult grasping.

We also recorded that distended GB to be significantly associated with difficult LC, which agreed with what was mentioned by Cho *et al.*, [33]. In our study significant association was seen with difficulty in laparoscopy with Pericholecystic collection (p<0.05) which was consistent with that Hasan *et al.*, [34] and Godre *et al.*, [35].

According to a preoperative scoring method formulated by Bhondave *et al.*, [12] for predicting difficult LC, we have found prediction that 70.7% was easy and 29.3 % of difficult LC, our finding was consistent with the Paul S, *et al.*, [36] and Ahmed M *et al.*, [19].

7. Conclusion

The difficult laparoscopic cholecystectomy can be predicted preoperatively based on number of previous attacks of cholecystitis, gall bladder wall thickness, and presence or absence of pericholecystic collection and palpable gall bladder /or impacted stones This may be helpful to determine the risk of conversion of laparoscopic cholecystectomy to open cholecystectomy beforehand and allow patients to be better prepared for surgery. Also, such prediction may allow a surgeon to be better prepared, to take extra precautions and reduce intra-operative complications and to convert to open cholecystectomy at an earlier stage.

Author Contributions: All authors contributed equally to the writing of this paper. All authors read and approved the final manuscript.

Conflicts of Interest: "Authors declare that they do not have any conflict of interests."

References

- [1] Veerank, N., & Togale, M. D. (2018). Validation of a scoring system to predict difficult laparoscopic cholecystectomy: A one-year cross-sectional study. *Journal of the West African College of Surgeons*, 8(1), 23-39.
- [2] De, U. (2004). Evolution of cholecystectomy: A tribute to Carl August Langenbuch. *Indian Journal of Surgery*, 66(2), 97-100.
- [3] Corr, P., Tate, J. J., Lau, W. Y., Dawson, J. W., & Li, A. K. (1994). Preoperative ultrasound to predict technical difficulties and complications of laparoscopic cholecystectomy. *The American Journal of Surgery*, 168(1), 54-56.
- [4] Radunovic, M., Lazovic, R., Popovic, N., Magdelinic, M., Bulajic, M., Radunovic, L., ... & Radunovic, M. (2016). Complications of laparoscopic cholecystectomy: our experience from a retrospective analysis. *Open Access Macedonian Journal of Medical Sciences*, 4(4), 641-646.
- [5] Bhondave, S., Dash, N., Thipse, M. V., & Gadekar, J. (2017). Proposed diagnostic scoring system to predict difficult laparoscopic cholecystectomy. *Journal of Medical Science And clinical Research*, 5(12), 31682-31688.
- [6] Bourgouin, S., Mancini, J., Monchal, T., Calvary, R., Bordes, J., & Balandraud, P. (2016). How to predict difficult laparoscopic cholecystectomy? Proposal for a simple preoperative scoring system. *The American journal of surgery*, 212(5), 873-881.
- [7] Rose, J. B., & Hawkins, W. G. (2017). Diagnosis and management of biliary injuries. *Current Problems in Surgery*, 54(8), 406-435.
- [8] Al-Bahlooli, S. H., Al-Malahi, A., Ghallab, N. H., Al-Dain, A. S. A., & Al Sabahi, A. A. (2009). Conversion rate of laparoscopic to open cholecystectomy. *Yemeni Journal for Medical Sciences*, 3(1), 1-8.
- [9] Inoue, K., Ueno, T., Douchi, D., Shima, K., Goto, S., Takahashi, M., ... & Naito, H. (2017). Risk factors for difficulty of laparoscopic cholecystectomy in grade II acute cholecystitis according to the Tokyo guidelines 2013. *BMC surgery*, 17(1), 1-8.
- [10] Persson, G., Strömberg, J., Svennblad, B., & Sandblom, G. (2012). Risk of bleeding associated with use of systemic thromboembolic prophylaxis during laparoscopic cholecystectomy. *Journal of British Surgery*, 99(7), 979-986.
- [11] Joris, J. L., Noirot, D. P., Legrand, M. J., Jacquet, N. J., & Lamy, M. L. (1993). Hemodynamic changes during laparoscopic cholecystectomy. *Anesthesia & Analgesia*, 76(5), 1067-1071.
- [12] Bhondave, S., Dash, N., Thipse, M. V., & Gadekar, J. (2017). Proposed diagnostic scoring system to predict difficult laparoscopic cholecystectomy. *Journal of Medical Science and Clinical Research*, 5(12), 31682-31688.
- [13] Abd-El-Aal, A. S., & Abdallah, H. A. (2018). Evaluation of preoperative predictive factors for difficult laparoscopic cholecystectomy in comparison with intraoperative parameters. *The Egyptian Journal of Surgery*, *37*(4), 504-511.
- [14] Kulkarni, S. V., & Kumar, S. S. (2018). Preoperative predictors of a difficult laparoscopic cholecystectomy. *International Surgery Journal*, *5*(2), 608-613.
- [15] Baral, S., Thapa, N., & Chhetri, R. K. (2020). Validation of a preoperative scoring system to predict difficult laparoscopic cholecystectomy: a nepalese perspective. *Journal of Lumbini Medical College*, 8(1), 3-9.
- [16] Sandhu. G., Rana, M. L., & Singh, K. (2016). Preoperative prediction of difficult laparoscopic cholecystectomy: A scoring method. *Indian Journal of Applied Research*, 6(6), 1-6.
- [17] Gupta, N., Ranjan, G., Arora, M. P., Goswami, B., Chaudhary, P., Kapur, A., ... & Chand, T. (2013). Validation of a scoring system to predict difficult laparoscopic cholecystectomy. *International Journal of Surgery*, 11(9), 1002-1006.

- [18] Di Buono, G., Romano, G., Galia, M., Amato, G., Maienza, E., Vernuccio, F., ... & Agrusa, A. (2021). Difficult laparoscopic cholecystectomy and preoperative predictive factors. *Scientific Reports*, 11(1), 1-6.
- [19] Hassan, A. M. (2021). Preoperative predictive risk factors of difficult laparoscopic cholecystectomy. *The Egyptian Journal of Surgery*, 40(2), 536-543.
- [20] Santharaj, S., & Marahanumaiah, S. (2022). Pre-operative predictors of difficult laparoscopic cholecystectomy: a comparative study between two scoring systems. *International Surgery Journal*, *9*(5), 960-966.
- [21] Maehira, H., Kawasaki, M., Itoh, A., Ogawa, M., Mizumura, N., Toyoda, S., ... & Kameyama, M. (2017). Prediction of difficult laparoscopic cholecystectomy for acute cholecystitis. *Journal of Surgical Research*, 216, 143-148.
- [22] Madan, A. K., Aliabadi-Wahle, S., Tesi, D., Flint, L. M., & Steinberg, S. M. (2002). How early is early laparoscopic treatment of acute cholecystitis?. *The American Journal of Surgery*, 183(3), 232-236.
- [23] Sharma, M., Muthuraman, S., Anand, S., & Minhas, S. S. (2020). Preoperative ultrasonography as a predictor of difficult laparoscopic cholecystectomy: A prospective study. *Annals of International Medical and Dental Research*, 6(3), RD01-RD04
- [24] Roy, A., Mukhopadhyay, M., Mukherjee, S., & Rahman, Q. M. (2020). Prediction of difficult laparoscopic cholecystectomy by adoption of a pre-operative scoring system. *Journal of Evidence-Based Medicine and Healthcare*, 7(41), 2319-2324.
- [25] Iqbal, A., Perveen, S., Khan, I., Ahmed, T., Khan, M. I., & Tunio, M. (2020). Preoperative assessment of scoring system designed for prediction of difficult cholecystectomy in patients with symptomatic gall stones. *Pakistan Armed Forces Medical Journal*, 70(2), 610-16.
- [26] Agrawal, N., Singh, S., & Khichy, S. (2015). Preoperative prediction of difficult laparoscopic cholecystectomy: a scoring method. *Nigerian Journal of Surgery*, 21(2), 130-133.
- [27] Ary Wibowo, A., Tri Joko Putra, O., Noor Helmi, Z., Poerwosusanta, H., Kelono Utomo, T., & Marwan Sikumbang, K. (2022). A Scoring System to Predict Difficult Laparoscopic Cholecystectomy: A Five-Year Cross-Sectional Study. *Minimally Invasive Surgery*, 2022, Article ID 3530568.
- [28] Nidoni, R., Udachan, T. V., Sasnur, P., Baloorkar, R., Sindgikar, V., & Narasangi, B. (2015). Predicting difficult laparoscopic cholecystectomy based on clinicoradiological assessment. *Journal of Clinical and Diagnostic Research*, 9(12), PC09-PC12.
- [29] Ramirez-Giraldo, C., Alvarado-Valenzuela, K., Isaza-Restrepo, A., & Navarro-Alean, J. (2022). Predicting the difficult laparoscopic cholecystectomy based on a preoperative scale. *Updates in Surgery*, 74(3), 969-977.
- [30] Lal, P., Agarwal, P. N., Malik, V. K., & Chakravarti, A. L. (2002). A difficult laparoscopic cholecystectomy that requires conversion to open procedure can be predicted by preoperative ultrasonography. *JSLS: Journal of the Society of Laparoendoscopic Surgeons*, 6(1), 59-63.
- [31] Arone, S., Raghunath, K. J., Venkatasubramanian, R., Muralidharan, M., & Chordia, D. (2022). Role of Preoperative Ultrasonography Findings in Predicting Difficult Laparoscopic Cholecystectomy. *World*, 15(3), 229-234.
- [32] Bhardwaj, R., Bali, R. S., & Zahoor, Y. (2018). Pre-operative factors for predicting a difficult laparoscopic cholecystectomy. *International Surgery Journal*, 5(9), 2991-2994.
- [33] Cho, K. S., Baek, S. Y., Kang, B. C., Choi, H. Y., & Han, H. S. (2004). Evaluation of preoperative sonography in acute cholecystitis to predict technical difficulties during laparoscopic cholecystectomy. *Journal of Clinical Ultrasound*, 32(3), 115-122.
- [34] Hasan, M. M., Muhsin, S. M., Alam, M. E., Anwar, S. M., Mostafa, F., Momin, M. N., & Khan, M. R. (2021). Predicting Difficult Laparoscopic Cholecystectomy Based on Clinicoradiological Assessment. *IAHS Medical Journal*, 4(2), 70-73.
- [35] Godre, P., & Krishnanand, D. S. (2022). An observational study of pre-operative predictors of difficult laparoscopic cholecystectomy using clinical, haematological, radiological parameters. *International Journal of Contemporary Medicine, Surgery and Radiology*, 7(2), B8-B14.
- [36] Paul, S., Khataniar, H., Ck, A., & Rao, H. K. (2022). Preoperative scoring system validation and analysis of associated risk factors in predicting difficult laparoscopic cholecystectomy in patients with acute calculous cholecystitis: A prospective observational study. *Turkish Journal of Surgery*, 38(4), 375-381.



© 2023 by the authors; licensee PSRP, Lahore, Pakistan. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC-BY) license (http://creativecommons.org/licenses/by/4.0/).