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Contents

Continuous Publication Volume 11 Number 4 April 18, 2023

REVIEW

Artificial intelligence ecosystem for computational psychiatry: Ideas to practice 79 Liu XQ, Ji XY, Weng X, Zhang YF

MINIREVIEWS

- 92 Lipocalin-2 as a biomarker for diabetic nephropathy Dahiya K, Prashant P, Dhankhar R, Dhankhar K, Kumar S, Vashist S
- Dehydroepiandrosterone sulfate supplementation in health and diseases 102 Jethwani P, Rastogi A, Shukla R

SYSTEMATIC REVIEWS

112 Current approach for Boerhaaves syndrome: A systematic review of case reports Yamana I, Fujikawa T, Kawamura Y, Hasegawa S

META-ANALYSIS

125 Role of baricitinib in COVID-19 patients: A systematic review and meta-analysis Thakur M, Babu A, Khatik GL, Datusalia AK, Khatri R, Kumar A



Contents

Continuous Publication Volume 11 Number 4 April 18, 2023

ABOUT COVER

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The primary aim of World Journal of Meta-Analysis (WJMA, World J Meta-Anal) is to provide scholars and readers from various fields of clinical medicine with a platform to publish high-quality meta-analysis and systematic review articles and communicate their research findings online.

WJMA mainly publishes articles reporting research results and findings obtained through meta-analysis and systematic review in a wide range of areas, including medicine, pharmacy, preventive medicine, stomatology, nursing, medical imaging, and laboratory medicine.

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SYSTEMATIC REVIEWS

Current approach for Boerhaaves syndrome: A systematic review of case reports

Ippei Yamana, Takahisa Fujikawa, Yuichiro Kawamura, Suguru Hasegawa

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Abstract

BACKGROUND

There is no consensus on the appropriate therapeutic strategy for Boerhaave syndrome due to its rarity and changing therapeutic approaches. We conducted a systematic review of case reports documenting Boerhaave syndrome.

AIM

To assess the therapeutic methods and clinical outcomes and discuss the current trends in the management of Boerhaave syndrome.

METHODS

We searched PubMed, Google scholar, MEDLINE, and The Cochrane Library for studies concerning Boerhaave syndrome published between 2017 and 2022.

RESULTS

Of the included studies, 49 were case reports, including a total of 56 cases. The mean age was 55.8 ± 16 years old. Initial conservative treatment was performed in 25 cases, while operation was performed in 31 cases. The rate of conservative treatment was significantly higher than that of operation in cases of shock vital on admission (9.7% vs 44.0%; P = 0.005). Seventeen out of 25 conservative cases (68.0%) were initially treated endoscopic esophageal stenting; 2 of those 17 cases subsequently underwent operation due to poor infection control. Twelve cases developed postoperative leakage (38.7%), and 4 of those 12 cases underwent endoscopic esophageal stenting to stop the leakage. The length of the hospital stay was not significantly different between the conservative treatment and operation cases (operation *vs* conservation: 33.52 ± 22.69 *vs* 38.81 ± 35.28 days; *P* = 0.553).

CONCLUSION

In the treatment of Boerhaave syndrome, it is most important to diagnose the



issue immediately. Primary repair with reinforcement is the gold-standard procedure. The indication of endoscopic esophageal stenting or endoluminal vacuum-assisted therapy should always be considered for patients in a poor general condition and who continue to have leakage after repair.

Key Words: Boerhaave syndrome; Esophageal perforation; Self expandable metalic stent; Minimally invasive surgical procedures; Anastomotic leakage; Shock

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Core Tip: Totally 49 published case reports concerning the Boerhaave syndrome were systematically reviewed. In the treatment of Boerhaave syndrome, it is most important to diagnose the issue immediately. Primary repair with reinforcement is the gold-standard procedure. The indication of endoscopic esophageal stenting or endoluminal vacuum-assisted therapy should always be considered for patients in a poor general condition and who continue to have leakage after repair.

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INTRODUCTION

Since Herman Boerhaave first recognized the disease in 1724, spontaneous esophageal perforation has been described as a medical emergency in the relevant literature[1]. The annual incidence of spontaneous esophageal perforation, also called Boerhaave syndrome, is 3.1 per 1000000; although rare, this condition is associated with high rates of misdiagnosis and mortality[2].

Boerhaave syndrome can be caused by vomiting and is frequently associated with alcohol intoxication[3]. A long period of time between perforation and treatment often results in mediastinitis, followed by septic shock and multiorgan failure[4-10]. Surgery and conservative management are the major treatment options for Boerhaave syndrome. However, few reports have examined whether operation or conservation is the preferred treatment method. Indeed, in the past five years, only one systematic review of Australasian literature on Boerhaave syndrome has been reported[11]. At present, there is no consensus on the optimal therapeutic strategy due to the rarity of Boerhaave syndrome and changing therapeutic approaches.

We therefore reviewed and evaluated 56 cases published in 49 case report articles in PubMed, Google scholar, MEDLINE, and The Cochrane Library in the past 5 years to assess the therapeutic methods and clinical outcomes and discuss the current trends in the management of Boerhaave syndrome.

MATERIALS AND METHODS

Study selection

A case report literature review was conducted using Pubmed, Google scholar, Cochrane Library, and MEDLINE for articles published between October 2017 and October 2022. The search was limited to articles in English. "Boerhaave syndrome" or "spontaneous esophageal perforation" were key words in the search. All titles and abstracts of publications were screened to select articles describing Boerhaave syndrome or spontaneous esophageal perforation. The searches were further broadened by extensively checking all references in the articles retrieved that met the inclusion criteria.

Inclusion and exclusion criteria

The inclusion criterion was patients who underwent operation or conservative therapy for Boerhaave syndrome. The exclusion criteria were meta-analyses, reviews, articles without outcomes reported, articles without the operation method reported, articles involving cases of treatment refusal, articles involving recurrent cases of esophageal perforation, articles involving best supportive care, articles involving pediatric cases, articles focusing on other diseases, and articles in non-English languages.

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Data extraction

The study design, and data on the patients' demographics, interventions, and outcomes were extracted from the included studies. An independent researcher collected the study data using a standard Excel[™] data collection sheet (Microsoft Corporation, Japan). This spreadsheet was used to calculate the descriptive statistics of all parameters that were evaluated in the present study. Continuous and categorical variables were shown as the mean and standard deviation (SD) and range.

Quality appraisal

The overall quality of the cases was classified as good to moderate. The majority of patients adequately described the chief complaint (100%), the patient's medical history (82.1%), the sex (98.2%), the length from symptom onset (98.2%), the length of the hospital stay (76.8%), imaging findings (100%), treatments (100%), and outcomes (100%).

Statistical analyses

All values were presented as the mean \pm SD. Intergroup differences were evaluated by an analysis of variance, while a nonparametric analysis was conducted for data with a skewed distribution. Statistical analyses were performed using EZR (Saitama Medical Center, Jichi Medical University)[12]. EZR for R (The R Foundation for Statistical Computing, version 2.13.0) is a modified version of the R commander (version 1.6–3) that includes statistical functions that are frequently used in biostatistics. *P* values of < 0.05 were considered statistically significant.

RESULTS

The results of the literature search are shown in Figure 1. Through our search, we identified 1310 studies. Of these, 990 studies were excluded by title and abstract. Of the remaining 115 potentially relevant articles, we excluded 48 concerning other diseases, 11 with insufficient data, 3 concerning recurrence cases, 2 involving best supportive care, and 2 pediatric cases. This resulted in the inclusion of 49 case report articles involving 56 cases for this study.

Patients' characteristics

Table 1 shows the details of the included studies. Of the 55 patients whose sex was mentioned, 51 were male, and 4 were female (1 case with no information). The mean age was 55.8 ± 16 years old. Thirty-six of the 55 cases (65.5%) were referred to the hospital within 24 h after symptom onset (1 case with no information). The most common method of the diagnosis was computed tomography (n = 31), followed by esophagography (n = 15), endoscopy (n = 9), and exploratory laparotomy (n = 1). A total of 42 cases (75%) were accurately diagnosed on admission. Fourteen patients (25%) showed shock vitals when they arrived at the hospital. Twelve (21.4%) were intra-mediastinum type, and 44 (78.6%) were extra-mediastinum type. The mean (range) size of the laceration in the 30 cases for which such details were described was 3.8 (1-12) cm (Table 2).

Initial treatment for Boerhaave syndrome

Conservative treatment was performed in 25 cases, while operation was performed in 31 cases. Conservative treatment included endoscopic esophageal stents in 17 cases, endoscopic clipping in 5, thoracic drainage in 21, and endoluminal vacuum-assisted (EVAC) therapy in 1. The operation approach was trans-thoracic and trans-abdominal approaches in 18 and 10 cases, respectively; a combined trans-abdominal and trans-thoracic approach was performed in 3 cases. In the trans-thoracic approach, minimally invasive surgery was performed in 5 cases (23.8%). In the trans-abdominal approach, minimally invasive surgery was performed in 8 cases (61.5%). The operation methods were primary repair only in eight cases, primary repair with omentoplasty in six cases, primary repair with fundus pouch in six cases, primary repair with intercostal muscle pouch in five cases, and esophagostomy in one case. Twelve out of 31 cases (38.7%) developed postoperative leakage. Two of those cases underwent EVAC therapy, and four of the cases underwent endoscopic esophageal stenting. Seven out of the 56 total cases (12.5%) died following treatment for Boerhaave syndrome; notably, 4 of those 7 cases (57.1%) had already had shock vitals on arrival at the hospital (Table 3).

Endoscopic esophageal stenting

Seventeen cases underwent endoscopic esophageal stenting initially, and 14 of them (82.4%) had severe comorbidities. Ten of the 17 cases (58.8%) who underwent endoscopic esophageal stenting had had shock vitals on arrival at the hospital. One case (14.3%) was the intra-mediastinum type, while the other 16 (85.7%) were the extra-mediastinum type. Two of the 17 cases who underwent endoscopic esophageal stenting had initially undergone operation due to poor infection control.

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Table 1 Descriptive comparative characteristics of all included 49 studies								
Ref.	Age	Sex	Accurate diagnosis	Rupture type	Shock vital	Laceration size (cm)	Treatment	Prognosis
Jahangir <i>et al</i> [4], 2021	64	М	Yes	Intrapleural type	Yes	1	Stent, thoracic drainage	Death
Issa <i>et al</i> [<mark>23</mark>], 2019	32	М	Yes	Intrapleural type	No	2	Stent, thoracic drainage	Alive
Tan <i>et al</i> [5], 2022	84	М	No	Intrapleural type	No	Unknown	Thoracotomy, primary repair only	Death
Chang <i>et al</i> [13], 2021	67	М	Yes	Intrapleural type	No	3	Thoracopy, primary repair only, feeding jejunostomy	Alive
Chang <i>et al</i> [<mark>13</mark>], 2021	62	М	Yes	Intrapleural type	No	2	Thoracopy, primary repair only, feeding jejunostomy	Alive
Sheshala <i>et al</i> [<mark>24</mark>], 2021	39	М	No	Intrapleural type	Yes	Unknown	Stent, thoracic drainage	Alive
Matsumoto <i>et al</i> [25], 2019	60	М	No	Intrapleural type	Yes	Unknown	Stent, thoracic drainage	Alive
Ayazi <i>et al</i> [<mark>6</mark>], 2021	22	М	Yes	Intrapleural type	Yes	Unknown	Thoracotomy, esophagectomy, gastrostomy	Death
Maki <i>et al</i> [<mark>44</mark>], 2022	76	М	Yes	Intramediastinal type	No	7	Transhiatal approach, primary repair plus omentoplasty, feeding jejunostomy	Alive
Ioannidis <i>et al</i> [<mark>39</mark>], 2021	83	F	Yes	Intrapleural type	No	Unknown	Thoracic drainage	Alive
Y K et al[<mark>26</mark>], 2018	86	М	Yes	Intrapleural type	Yes	5	Stent, thoracic drainage, feeding jejunostomy	Alive
Czopnik <i>et al</i> [3], 2017	47	М	Yes	Intrapleural type	No	5	Transhiatal approach, primary repair, gastrostomy	Alive
Awadelkarim <i>et</i> al[<mark>27]</mark> , 2021	36	М	Yes	Intrapleural type	Yes	2	Stent, thoracic drainage	Alive
Chalikonda <i>et al</i> [<mark>28</mark>], 2019	74	М	No	Intrapleural type	Yes	Unknown	Stent, thoracic drainage	Alive
Śnieżyński <i>et al</i> [<mark>29</mark>], 2021	53	М	Yes	Intrapleural type	No	3	Stent, thoracic drainage	Alive
Matsuura <i>et al</i> [<mark>21</mark>], 2022	69	М	Yes	Intramediastinal type	No	Unknown	Endoscopic clipping	Alive
Chen <i>et al</i> [<mark>19</mark>], 2021	57	М	No	Intramediastinal type	No	Unknown	Transhiatal approach, primary repair only, feeding jejunostomy	Alive
Truyens <i>et al</i> [<mark>30]</mark> , 2020	66	М	Yes	Intramediastinal type	Yes	Unknown	Antibiotic administration	Alive
Truyens <i>et al</i> [<mark>30]</mark> , 2020	77	М	Yes	Intramediastinal type	No	Unknown	Stent	Alive
Swol <i>et al</i> [7], 2019	70	М	Yes	Intramediastinal type	No	2	Transhiatal approach, primary repair plus fundus pauch	Death
Park <i>et al</i> [<mark>45</mark>], 2021	Unknown	Unknown	Yes	Intramediastinal type	No	5	Laparoscopic transhiatal approch, primary repair plus omentoplasty \rightarrow endoscopic clipping, stent	Alive
Rahman <i>et al</i> [49], 2021	53	М	Yes	Intrapleural type	No	Unknown	Thoracotomy, primary repair plus intercostal muscle pauch, gastrojujuno tube \rightarrow stent	Alive
Nachiappan <i>et</i> al[46], 2022	59	М	No	Intrapleural type	No	1,5	Endscopic clipping, stent \rightarrow laparo- scopic transhiatal approach, primary repair plus omentoplasty	Alive
Pasternak <i>et al</i> [<mark>14</mark>], 2019	37	М	Yes	Intrapleural type	No	Unknown	Thoracotomy, primary repair only, gastrostomy	Alive
Kita <i>et al</i> [<mark>55</mark>], 2022	46	М	Yes	Intramediastinal type	No	4	Laparoscopic transhiatal approch, primary repair plus fundus pauch	Alive

Kita et al <mark>[55</mark>], 2022	48	М	Yes	Intramediastinal type	No	3	Laparoscopic transhiatal approch, primary repair plus fundus pauch	Alive
Kita <i>et al</i> [<mark>55</mark>], 2022	65	М	Yes	Intramediastinal type	No	5	Laparoscopic transhiatal approch, primary repair plus fundus pauch	Alive
Saffo <i>et al</i> [<mark>8</mark>], 2021	76	М	No	Intrapleural type	Yes	Unknown	Stent, thoracic drainage	Death
Kochar <i>et al</i> [<mark>50</mark>], 2019	40	М	Yes	Intrapleural type	Yes	Unknown	Thoracotomy, primary repair plus intercostal muscle pauch, intraop- erative stent, thoracic drainage	Alive
Bury et al[<mark>51</mark>], 2022	50	М	No	Intrapleural type	No	4	Thoracotomy, primary repair plus intercostal muscle pauch, thoracic drainage	Alive
Aref <i>et al</i> [<mark>47</mark>], 2019	32	М	Yes	Intramediastinal type	No	2	Laparoscopic transhiatal approach, primary repair plus omentoplasty	Alive
Bani Fawwaz et al[15], 2022	63	М	Yes	Intrapleural type	Yes	3	Stent, thoracic drainage	Alive
Bani Fawwaz et al[15], 2022	56	F	Yes	Intrapleural type	No	Unknown	Thoracotomy, primary repair plus T tube, Belsey fundoplication, intraop- erative stent, thoracic drainage, gastrostpomy	Alive
Xu et al <mark>[22]</mark> , 2021	63	М	Yes	Intrapleural type	No	Unknown	Endoscopic clipping	Alive
Tuñon <i>et al</i> [<mark>57</mark>], 2021	24	М	Yes	Intrapleural type	No	4	Endoluminal vacuum therapy \rightarrow endoscopic clipping	Alive
Lee <i>et al</i> [<mark>58</mark>], 2018	52	М	Yes	Intrapleural type	No	Unknown	Thoracoscopic approach, primary repair only \rightarrow endoluminal vacuum therapy, thoracic drainage	Alive
He <i>et al</i> [<mark>54</mark>], 2018	57	М	Yes	Intramediastinal type	No	6	Endoscopic clipping	Death
Kim et al <mark>[59]</mark> , 2019	56	М	Yes	Intrapleural type	No	Unknown	Thoracotomy, primary repair only \rightarrow endoluminal vacuum therapy, thoracic drainage	Alive
Shennib <i>et al</i> [<mark>52</mark>], 2021	47	М	No	Intrapleural type	Yes	5	Thoracotomy, primary repair plus pericardial pauch, gastrostomy, feeding jejunostomy	Alive
Agrawal <i>et al</i> [40], 2019	26	М	No	Intrapleural type	No	Unknown	thoracic drainage	Alive
Sato <i>et al</i> [<mark>31</mark>], 2018	52	М	Yes	Intrapleural type	No	Unknown	Tho racotomy, primary repair only \rightarrow stent, tho racic drainage	Alive
Sato <i>et al</i> [<mark>31</mark>], 2018	53	М	No	Intrapleural type	Yes	Unknown	Stent, thoracic drainage	Alive
Ali et al <mark>[16]</mark> , 2020	30	F	No	Intrapleural type	No	4	Thoracotomy, primary repair only	Alive
Anand <i>et al</i> [<mark>48</mark>], 2022	64	М	Yes	Intrapleural type	No	2	Thoracotomy, primary repair plus intercostal muscle pauch, thoracic drainage	Alive
Barakat <i>et al</i> [<mark>32</mark>], 2017	62	М	Yes	Intrapleural type	No	1	Stent, endoscopic clipping	Alive
Alakkari <i>et al</i> [<mark>17</mark>], 2019	69	F	Yes	Intrapleural type	No	Unknown	Thoracotomy, primary repair plus T tube	Alive
Zhu et al <mark>[18]</mark> , 2021	33	М	No	Intrapleural type	No	Unknown	Stent, PEG \rightarrow thoracotomy, drainage	Alive
Sekiya et al <mark>[56]</mark> , 2019	61	М	Yes	Intrapleural type	No	3	Thoracoscopic and laparoscopic approach, primary repair plus pericardial pauch, gastrostomy	Alive
Sekiya <i>et a</i> l[<mark>56</mark>], 2019	64	М	Yes	Intrapleural type	No	4	Thoracoscopic and laparoscopic approach, primary repair plus pericardial pauch, feeding jejunostomy	Alive
Olivero et al	67	М	No	Intrapleural type	No	2	Thoracotomy, primary repair plus	Alive



[<mark>53</mark>], 2019							pericardial pauch, thoracic drainage	
Felipe <i>et al</i> [<mark>38</mark>], 2021	47	М	Yes	Intrapleural type	No	12	Thoracotomy and laparotomy approach, esophagostomy, gastrostomy \rightarrow stent	Alive
Ahmad <i>et al</i> [<mark>33</mark>], 2018	63	М	Yes	Intrapleural type	No	2.5	Stent, thoracic drainage	Alive
Hashmi <i>et al</i> [<mark>10]</mark> , 2021	83	М	Yes	Intrapleural type	Yes	Unknown	Antibiotic administration	Death
Teblick <i>et al</i> [34], 2018	74	М	Yes	Intrapleural type	No	Unknown	Stent, thoracic drainage	Alive
He et al[<mark>9</mark>], 2018	27	М	Yes	Intrapleural type	No	6	Thoracotomy, primary repair plus pleural flap, feeding jejunostomy	Alive

PEG: Percutaneous endoscopic gastrostomy; M: Male; F: Female.

Table 2 Characteristics of the patients with Boerhaave syndrome included in the review, n (%) Sex ^aMale 51, female 4 55.8 ± 16 Age ^b36 (65.5) The length from symptom within 24 h The method of diagnosis CT 31 Esophagography 15 Endoscopy 9 Exploratory laparotomy 1 Accurate diagnosis on admission 42 (75) Shock vital on admission 14 (25) Intramediastinal Rupture type 12 (21.4) Extramediatinal 44 (78.6) °3.8 (1-12) Lacelation size (cm) (range)

^aOne case with no information.

^bOne case with no information

^c26 cases with no information.

CT: Computed tomography.

Four of the cases who initially underwent operation consequently underwent endoscopic esophageal stenting to stop leakage.

Minimally invasive surgery

Eleven out of 31 cases (35.5%) underwent minimally invasive surgery. Seven of the 13 cases (53.8%) who underwent the trans-abdominal approach received the trans-hiatal approach specifically with laparoscopic surgery. Five of the 21 cases (23.8%) who underwent the trans-thoracic approach received thoracoscopic surgery. The length of the hospital stay after surgery tended to be shorter with minimally invasive surgery than with non-minimally invasive surgery [minimally invasive surgery (n = 10) vs nonminimally invasive surgery (*n* = 15): 25.5 ± 17.1 *vs* 38.86 ± 24.85 d; *P* = 0.153] (Figure 2A).

Conservative treatment vs surgery

Table 4 shows the differences in details between patients who underwent an operation and those who received conservative treatment. The sex, age, rate of patients admitting within 24h after symptom onset, rupture type, and rate of survival did not significantly differ between patients who underwent an operation and those who received conservative treatment. The rate of patients with shock vitals on admission did differ significantly between patients who underwent an operation and those who received conservative treatment (9.7% vs 44.0%; P = 0.005). The length of hospital stay was not significantly different among the 43 cases (operation vs conservative treatment: $33.52 \pm 22.09 vs$ $38.81 \pm 2000 vs$ 35.28 d; *P* = 0.55) (Figure 2B).



Table 3 Initial treatment for Boerhaave syndrome
--

			The number do not add up because of duplication case
Conservation ($n = 25$) ^a	Esophageal stent		17
	Clipping		5
	Thoracic drainage		21
	EVAC ^b		1
Operation $(n = 31)$	Approach	Trans-thoracic approach	18
		Trans-abdominal approach	10
		Trans-thoracic and abdominal approach	3
	Method	Primary repair only	8
		Primary repair with omentoplasty	6
		Primary repair with fundus pauch	6
		Primary repair with intercostal muscle pauch	5
		Primary repair with pericardial fat pauch	5
		T tube	2
		Esophagectomy	1
		Esophagostomy and gastrostomy	1

^aDuplication exist.

^bEndoluminal vacuum-assisted therapy.

EVAC: Endoluminal vacuum-assisted.

Table 4 The length of hospital stay was not significantly different among the 43 cases

Factor	Group	Operation (<i>n</i> = 31)	Conservation (<i>n</i> = 25)	P value
Sex (%)	М	27 (90.0)	24 (96.0)	0.617
	F	3 (10.0)	1 (4.0)	
mean ± SD		53.17 (14.68)	59.00 (18.14)	0.193
The length from symptom within 24 h (%)	Yes	10 (32.3)	9 (36.0)	1
	No	20 (64.5)	16 (64.0)	
Shock vital on admission (%)	Yes	3 (9.7)	11 (44.0)	0.005
	No	28 (90.3)	14 (56.0)	
rupture type	Intramediastinal (%)	8 (25.8)	4 (16.0)	0.516
	Extramediastinal (%)	23 (74.2)	21 (84.0)	
Alive (%)	Yes	28 (90.3)	21 (84.0)	0.688
	No	3 (9.7)	4 (16.0)	

F: Female; M: Male.

DISCUSSION

Primary surgical repair has been the gold-standard treatment for esophageal perforation for a long time [13-19]. Primary repair of the esophagus conducted with mediastinal and thoracic drainage is reported to have a 90% success rate. Cases in which esophageal rupture is diagnosed at an early stage (within 24 h) without associated esophageal disease are reported to show a particularly high success rate[20].





Figure 1 PRISMA flow diagram demonstrating articles selection process.



Figure 2 Comparison of length of hospital stay. A: The length of hospital stay in the non-minimally and minimally invasive surgery groups; B: The length of hospital stay in the operation and conservative treatment groups.

There has been a recent trend toward more non-operative management^[21,22], such as esophageal stent replacement via an endoscopic approach 23-32]. The indications for esophageal stenting include multiple comorbidities, advanced mediastinal sepsis, hemodynamic compromise, and clinical intolerance of extensive surgical repair[33]. In our review, the rate of conservation was significantly higher than that of operation in instances of shock vital on admission (44.0% vs 9.7%; P = 0.005).

Esophageal stenting was able to be attempted for patients who were in a bad general condition or intolerant to surgery [34]. Endoscopic esophageal stenting was also performed for cases of postoperative leakage. Kauer et al[35] in 2008 first described the usefulness of stent placement in the management of thoracic anastomotic leakage after esophagectomy. An interval approach utilizing covered metallic stent was then introduced for the management of anastomotic leakage after esophagectomy[36]. However, no prospective clinical study comparing the outcomes of esophageal stenting to that of conservative/ surgical treatment has yet been performed. Bi *et al*[37] reported that the efficacy of the three-tube method, (tube drainage of the abscess, placement of a jejunal feeding tube, and placement of a

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gastrointestinal decompression tube, with implantation of a covered metallic stent) for the management of anastomotic leakage following esophagectomy. This means that it is important not only to place esophageal stents but also to provide adequate drainage, a concept that can also be applied for treating Boerhaave syndrome.

Surgical approaches differed among facilities in our review. The operation approach in our evaluated studies was the trans-thoracic approach in 18 cases, trans-abdominal approach in 10 cases, and combined trans-thoracic and trans-abdominal approach in 3 cases. The approach seemed to differ depending on laceration site, the patient's general condition, and whether the operator was a thoracic surgeon or a gastrointestinal surgeon. The reported operative methods for Boerhaave syndrome include primary repair (with/without reinforcement), an exclusion diversion operation[38], esophageal resection, and simple thoracic drainage[39-40]. Previous reports mentioned that reinforcement with vascularized tissue was associated with reduced fistula formation and mortality rates in comparison to repair without reinforcement[41-43]. In the case of friability of the tissue, primary repair with reinforcement, such as omental flaps[44-47], intercostal muscle flaps[48-51], and pericardial flaps[52-54], should be performed. A comprehensive evaluation of the degree of laceration, extent of laceration, and general condition required for deciding the repair method should be conducted.

There have been a few recent reports concerning minimally invasive surgery for Boerhaave syndrome. Kita *et al*[55] suggested that a good clinical course can be obtained by laparoscopic transhiatal esophageal repair for Boerhaave's syndrome with localized mediastinal collections to avoid surgical invasion due to thoracotomy. Sekiya *et al*[56] reported the convenience and usefulness of minimally invasive surgery *via* an abdominal and left thoracic approach, which provides excellent visualization of the abdominal and thoracic cavities and facilitates quick switching between views. The authors further suggested that, in cases with an interval to the diagnosis < 24 h, no severe comorbidities, and a perforation site in the left lower esophagus, a trans-hiatal approach for minimally invasive surgery is feasible to repair the laceration and ameliorate the infection[56]. In our systematic review, the length of hospital stay after minimally invasive surgery tended to be shorter than after non-minimally invasive surgery (25.5 ± 17.1 *vs* 38.86 ± 24.85 d; *P* = 0.153). Minimally invasive surgery is useful for its cosmetic aspect, camera magnification effect, and ease of suturing, especially a laparoscopic trans-hiatal approach.

In our systematic review, 12 out of 31 cases (38.7%) developed postoperative leakage. Two of those 13 Leakage cases underwent EVAC therapy. Recently, the efficacy of EVAC therapy for esophago-pleural fistula after an operation for Boerhaave syndrome was reported[57-59]. EVAC therapy can be applied in postoperative management according to the principle applied for external wounds that provide wound drainage and tissue granulation. EVAC therapy can be applied to conservatively treat cases where primary surgical repair of esophageal perforation is unsuccessful. Moreover, with the use of an S-B tube, the patient can simultaneously receive intraluminal EVAC therapy with enteral nutrition in a non-invasive manner[58]. This may accelerate the healing of the injured esophagus and reduce the duration of hospitalization.

We suggest an algorithm that might be useful in the treatment of Boerhaave syndrome in Figure 3, with reference to our systematic review. If Boerhaave syndrome is suspected on computed tomography, esophagography or upper gastrointestinal endoscopy should be performed immediately. The treatment of Boerhaave syndrome is basically primary repair with reinforcement. If postoperative leakage occurs, endoscopic esophageal stenting or EVAC therapy should be considered. If the patient is inoperable (severe shock vitals, super-elderly patients, severe comorbidities, *etc.*), endoscopic esophageal stenting and thoracic drainage should be considered.

Several limitations associated with the present study warrant mention. Importantly, due to its rarity, there are few large case series on Boerhaave syndrome. Furthermore, the therapeutic strategies for Boerhaave syndrome have changed over time, with new approaches being developed recently. We reviewed and analyzed 49 articles; however, the review process may have included various publication biases.

CONCLUSION

In the treatment of Boerhaave syndrome, it is most important to diagnose the issue immediately. Primary repair with reinforcement is the gold-standard procedure. The optimal treatment should be determined according to the etiology, general physical condition of the patient, and site of perforation, as well as the extent of contamination, as determined by radiology. The indication of endoscopic esophageal stenting or endoluminal vacuum-assisted therapy should always be considered for patients in a poor general condition and who continue to have leakage after repair.

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Figure 3 Algorithm for the treatment of Boerhaave syndrome with reference to the systematic review findings. CT: Computed tomography; EVAC: Endoluminal vacuum-assisted

ARTICLE HIGHLIGHTS

Research perspectives

As far, it has long been reported that Boerhaave syndrome has a poor prognosis when diagnosed late. However, no consensus has been reached concerning the appropriate therapeutic strategy for Boerhaave syndrome because of the rarity of the disease and the changing therapeutic trends.

Research conclusions

We assess the therapeutic methods [operation vs drainage vs stent vs endoluminal vacuum-assisted (EVAC), etc.] and clinical outcomes and discuss the current trends in the management of Boerhaave syndrome.

Research results

We believe that this systematic review will be useful in future treatment of Boerhaave syndrome when there is doubt as to whether conservative treatment or surgery should be done, as well as the method of surgery.

Research methods

We searched PubMed, Google scholar, MEDLINE, and The Cochrane Library for studies concerning Boerhaave syndrome published between 2017 and 2022.

Research objectives

In results, the key to treatment of Boerhaave syndrome was early diagnosis. In addition, although surgery was the basic treatment, esophageal stents and drainage may be useful for patients with intolerance. Furthermore, for postoperative leakage, esophageal stents, drainage, and EVAC were useful.

Research motivation

In the treatment of Boerhaave syndrome, it is most important to diagnose the issue immediately. Primary repair with reinforcement is the gold-standard procedure. The indication of endoscopic esophageal stenting or endoluminal vacuum-assisted therapy should always be considered for patients in a poor general condition and who continue to have leakage after repair.

Research background

Because Boerhaave syndrome is a rare disease, observational studies should be conducted in collaboration with other centers. We hope that this will result in a high-quality strategy.

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FOOTNOTES

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