

# **Archives of Orthopaedics**

**Review Article** 

# Surgical Fixation of Severe Rib Fractures: A Systematic Literature Review and Meta-Analysis

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### **Abstract**

**Introduction:** Evidence for treating severe rib fractures without a flail component (non-flail) has not yet been adequately evaluated. This study analyzed contemporary evidence for the surgical versus non-surgical treatment of adults with severe chest rib fractures, with further analysis by the flail component.

**Materials and Methods:** A systematic literature review and meta-analysis included studies evaluating patients with surgical fixation of severe rib fractures. All studies included non-flail patients. Random effects models pooled data for outcomes reported in  $\geq 2$  studies. The primary outcome was the duration of mechanical ventilation (DMV). Secondary outcomes included post-procedural pain, respiratory complications, mortality, tracheostomy, sepsis, intensive care unit (ICU), and hospital length of stay (LOS).

**Results:** Thirty-one studies (n=99,640 patients) evaluating surgical fixation of severe rib fracture patients were included in the meta-analysis. Surgical fixation resulted in statistically significantly shorter DMV (-1.81 days, 95% confidence interval (CI): -3.14 to -0.49 days; p=0.007), lower 2-week pain intensity (SMD -3.29, 95% CI: -5.05 to -1.53; p=0.003), lower risk for atelectasis (RR=0.41, 95% CI: 0.25-0.67; p=0.0003), lower risk for any respiratory complication (RR=0.63, 95% CI: 0.43- 0.92, p=0.02), and lower mortality risk (RR=0.41, 95% CI: 0.23-0.73, p=0.003) compared to non-surgical treatment. Statistically significant differences were not observed for pain 3-day after intervention (SMD -1.28, 95% CI: -3.32 to 0.75; p=0.22); pneumonia (RR=0.66, 95% CI: 0.40- 1.08; p=0.10), acute respiratory distress syndrome (RR 1.19, 95% CI: 0.18-7.96; p=0.85), tracheotomy (RR 0.66, 95% CI: 0.30-1.44, p=0.29), sepsis (RR=0.75, 95% CI: 0.17-3.28, p=0.70), ICU LOS (MD -1.01, 95% CI: -2.42 to 0.939; p=0.16), and hospital LOS (MD-1.52, 95% CI: -3.97 to 0.92; p=0.22).

**Conclusion:** Surgical treatment of patients with severe rib fractures, including a majority of non-flail patients, resulted in statistically significantly shorter DMV, less 2-week pain, lower risk of atelectasis and overall respiratory complications, and reduced mortality compared to non-surgical treatment.

**Keywords:** Rib fractures, Rib fixation, Surgery, Duration of mechanical ventilation

#### Introduction

Rib fractures are a commonly encountered traumatic injury and are associated with significant morbidity and mortality [1,2]. They are a marker of severe injury and can lead to defects in the chest wall and severe pain which may hinder breathing [3]. Patients with rib fracture have an increased risk of developing chest infection, impaired pulmonary function, sepsis, atelectasis, respiratory failure, and other pulmonary pathologies, and they are at risk for prolonged hospitalizations [4]. The contribution of rib fractures to prolonged disability and chronic pain has been found to be greater than traditionally expected [5]. Studies have reported a prevalence of chronic pain of 22% and work disability of 53% among patients with rib fractures [6].

As the number of fractured ribs increases, it is believed that the patient's risk for undesired outcomes is increased not only because of other serious injuries, but also because of the respiratory complications that are a direct consequence of the pain and impaired capacity to ventilate [7-9]. Patients with a 'flail chest' are those who have three or more contiguous ribs fractured in two or more places [10]. Patients with a flail chest, or other forms of severe multiple rib fractures, frequently require mechanical ventilation and are at risk for death [11,12]. Older adults with rib fractures are particularly vulnerable as they have twice the morbidity and mortality of younger adults [13-16].

Surgical stabilization of rib fractures (SSRF) has recently been described as an effective approach to treat flail chest or severe rib fracture injuries [11,17]. Evidence for the effectiveness of SSRF compared to non-surgical treatment has been mounting, particularly over the past decade. The preponderance of evidence has centered around patients with flail chest rib fractures. Multiple published systematic reviews and meta-analyses have demonstrated that SSRF of flail chest may lead to shorter hospital and intensive care unit (ICU) stays, fewer cases of pneumonia, and reduced mortality [18-27]. A recent cost-effectiveness analysis also found that surgical stabilization of rib fractures for patients with flail chest was cost-effective [27].

Evidence for the effectiveness of SSRF among patients with non-flail severe rib fractures has been emerging very recently; however, it has not yet been adequately gathered, appraised, and integrated. A meta-analysis from Beks and colleagues [26] evaluated both flail chest and non-flail multiple rib fractures up through June 2017 and found that the evidence for flail chest was strong; however, the authors concluded that the evidence for non-flail multiple rib fractures was insufficient [26]. A systematic review by Ingoe and colleagues [25] evaluated flail chest and multiple rib fractures through March 2017 and found that all outcomes showed a statistically significant improvement in favor of SSRF for flail chest; however, again, the authors concluded that the evidence for multiple, non-flail, rib fractures was limited and the benefits

uncertain. Similarly, the cost-effectiveness analysis by Choi and colleagues [27] evaluated SSRF of rib fracture in patients up through 2014 and found that fixation may be cost-effective in some patients without flail chest; however, the authors concluded that characterizing these patients required further study. Most recently, a 2020 meta-analysis by Long *et al.* evaluating the clinical efficacy of surgical vs non-surgical care for multiple rib fractures, with or without flail chest, concluded that surgical treatment resulted in faster recovery, lower risk of complications and better prognosis than non-surgical care [28].

A better understanding of the current evidence for the clinical and economic value of SSRF for severe rib fractures specifically for non-flail rib fractures would help healthcare providers and payers prioritize resource allocation and develop more effective and targeted interventions. Hence, the objectives of this study were to assess contemporary evidence for the clinical safety, effectiveness and economic outcomes associated with SSRF of severe rib fractures in adults, in studies that include non-flail patients.

#### **Material and Methods**

The systematic literature review and meta-analysis compared SSRF to non-surgical treatment for adult patients with severe chest rib fractures. The systematic literature review was conducted and reported according to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines [29]. The systematic review protocol was registered in the PROSPERO database.

# **Search strategy**

The literature search was conducted on June 22, 2020, by electronic searching of MEDLINE, Embase, Cochrane Central Register of Controlled Trials (CENTRAL), and Cochrane Database of Systematic Reviews databases. The search terms and search strategy utilized were: (multiple AND rib fractur\*) OR ("non flail" OR "not flail" OR "non-flail" OR "nonflail" OR "without flail"). A broad search strategy was used since several relevant articles would have been missed if we narrowed the search using the terms surgery or surgical intervention and displaced or dislocated rib fracture.

#### Identification of studies to be included in the analysis

The search was intended to identify primary clinical studies evaluating outcomes of SSRF for severe rib fractures compared to non-surgical care, in populations that included at least a proportion of patients with a non-flail component. Severe was defined as multiple and/or displaced fractures. Studies included primary surgical centers as well as database analyses.

The search strategies were designed to identify records based on the following PICOTS (patient-intervention-comparator-

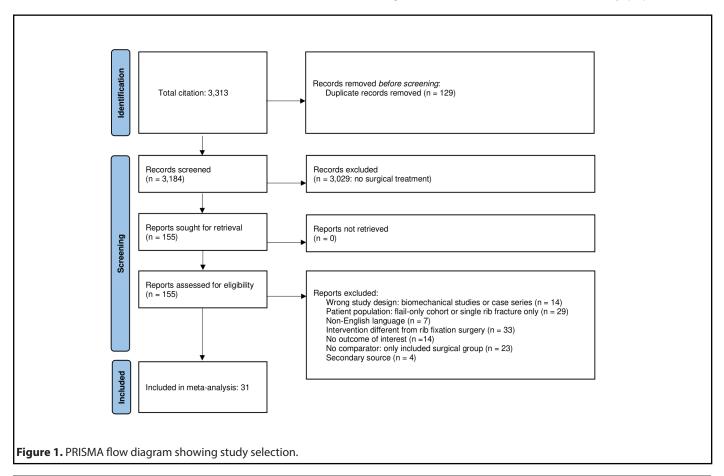
outcome-time-source) framework. Patients: Skeletally-mature patients up to and including 80 years of age, with severe rib fractures (multiple and/or displaced). At least some patients in the study had to be diagnosed/identified as non-flail. This comprised studies evaluating only non-flail patients and studies evaluating mixed cohorts of patients with and without flail chest. Some mixed cohort studies stratified patients by the presence of flail chest, while others did not. Patients diagnosed with a tumor, spinal cord injury, or brain injury of moderate and severe intensity were excluded from the review. Studies evaluating only flail chest patients were excluded. Intervention: SSRF was defined as the intervention of interest. All fixation devices and surgical approaches (open vs minimally invasive) were acceptable. Comparators: Nonsurgical care was defined as the comparator. Outcomes: The primary outcome was the duration of mechanical ventilation (DMV). Secondary outcomes included post-procedural pain, respiratory complications, mortality, tracheostomy, sepsis, and intensive care unit (ICU) and hospital length of stay (LOS). Papers stating at least one of these outcomes were included in the analysis. An effort was made to capture and consider any and all reported major clinical and economic outcomes. Selected outcomes were based on previously conducted metaanalyses on SSRF [18-27,30]. Time: No limit in the postoperative evaluation window was predefined. Outcomes measured from day of surgery onwards were included in the study.

Study: Randomized controlled trials (RCTs), non-randomized clinical trials or studies, cohort studies, case control studies, registry studies, economic studies (budget impact and costeffective analyses), and case series were included. Excluded studies included: technical articles, animal/cadaver studies, case reports, editorials, commentaries, and letters. Only English language literature was considered for review. The search was not restricted by publication date. Abstracts, case reports and manuscripts were included. Care was taken to ensure no duplication of data: when abstracts were included, subsequent publications were analyzed to ensure that the same patient population was not described twice.

#### Study selection procedure and data extraction

Two reviewers independently applied the PICOTS framework above to screen de-duplicated titles and abstracts obtained from the search strategy. Potentially relevant citations were checked in a full-text screening. Disagreements were resolved through discussion and reasons for exclusion were recorded. **Figure 1** illustrates the study selection process as a PRISMA flow diagram.

Pre-specified data that were extracted from the relevant studies included the journal citation, study objectives, study design and data source, intervention, study population (ie,



baseline demographic and clinical characteristics), sample size, duration of follow-up, primary and secondary outcome measures, and author's conclusions.

#### **Quality assessment of studies**

Studies were appraised for their level of evidence based on the study design and the rigor of methodology used, as well as the ability to prevent and/or control for biases to analyze cause and effect. All included studies were critically appraised and ranked as either low, good, or high-quality evidence using the Evidence level and Quality Guide from John Hopkins Nursing Evidence-Based Practice [31,32].

### **Evidence synthesis and statistical analysis**

The studies were initially summarized descriptively using frequency counts and proportions to describe the different types of studies and the numbers of patients. Qualitative synthesis, including narrative summarization, was conducted for outcomes where the meta-analysis of effect estimates was not appropriate or possible (outcomes not reported in at least two included studies), or because characteristics of studies (such as study designs, intervention types, or outcomes) were too diverse to yield a meaningful summary estimate of effect.

A meta-analysis was performed for outcomes that were reported in at least two distinct studies. For continuous outcome measures (ie, DMV, ICU LOS, and hospital LOS), the inverse variance random effects model (REM) with Dersimonian Lair was used to estimate the pooled mean difference (MD) between treatment groups, across all studies. The pooled standardized mean difference (SMD) was used for pain scores since the studies used different pain scales. The mean and standard deviation (SD) were extracted from individual studies or were derived from medians with interquartile ranges or means with p-values. For dichotomous outcomes, the Mantel-Haenszel REM was used to estimate the pooled risk ratios (RR). For the pooled summary statistics for each outcome in the surgical and non-surgical intervention groups, inverse variance REMs were used. All effect sizes were reported with 95% confidence intervals (CI). The  $\chi^2$  test was used to identify statistical heterogeneity ( $\alpha$ =0.05) and heterogeneity was quantified using the I<sup>2</sup> statistic. Subgroup analyses evaluating the impact of: (a) presence or absence of flail chest patients; (b) study type; and (c) quality of evidence were also conducted. The statistical significance was set at p-value ≤0.05. RevMan was used for all analyses.

### **Results**

### Study identification and selection

The PRISMA flow diagram is shown in **Figure 1**. The literature search yielded 3,313 citations, of which 129 were duplicates leaving 3,184 studies that were screened for inclusion. Of

these, 3,029 articles were excluded because they did not have a surgical treatment group or involved single fractures. Full texts of 155 of the studies were retrieved for further screening, of which 14 were excluded based on study design, 29 based on the patient population, 7 on non-English language, 33 on intervention, 14 due to lack of relevant outcome, 23 due to lack of or different comparator, and 4 that were prior meta-analyses (secondary sources).

#### **Descriptive characterization of studies**

A total of 31 studies were included in the analysis, describing outcomes of 99,640 patients, of which 4,675 underwent SSRF and 94,965 were treated non-surgically. The mean age of patients varied from 38 to 70 years. All of the included studies focused on surgical treatment and compared SSRF with nonsurgical approaches. Whereas all studies described outcomes of non-flail patients, 8 studies included only non-flail patients, all other reported on a mix of flail and non-flail patients [12,33-39]. When available, the percentage of patients with non-flail component in the surgical and control groups are reported in **Table 1.** Overall, of the 4,675 SSRF patients, at least 82% (3,829/4,675) were identified as non-flail. In the control group, the proportion of identified non-flail patients was 97% (92,383/94,965). Studies included one randomized controlled trial (RCT) [40] and one hybrid RCT/prospective controlled cohort study [36]. In addition to these 2 studies, 3 cohort studies were prospectively designed [12,41,42]. All other studies were retrospective case-control or controlled cohort studies. The outcomes of 4 studies were presented in abstracts only [33,35,43,44], all other studies were reported in manuscripts. Eighteen of 31 studies were published between 2018 and 2020.

Three studies were used for narrative synthesis only [33,35,45], all other studies contributed outcomes data to the meta-analyses. The primary study outcome, the DMV, was reported in 16 studies, pain was described in 4 studies, respiratory complications in 17 studies, mortality in 19 studies, tracheotomy and sepsis in 8 and 7 studies, respectively, and hospital LOS and ICU LOS in 18 and 16 studies, respectively.

The studies were analyzed and ranked according to the John Hopkins Nursing Evidence-based Practice Level and Quality guidelines [31,32]. A summary of the quality assessment of each of the included clinical studies is available in **Table 1**. Visual inspection of funnel plots was also used to determine possible publication bias. Funnel plots were generated for outcomes with at least 10 included studies. The funnel plots for the secondary outcome measures are available as **Figures S1-S5** of the **Supplemental Files**.

#### **Meta-analyses**

**Duration of Mechanical Ventilation:** 16 studies contributed to the DVM analysis, as shown in **Figure 2**. The aggregate

			Jo CN	Mean	Fol-	Surgica	Surgical Cohort	Non-Su	Non-Surgical Cohort		Level and
Study	Year	Study design	patients	mean age <sup>†</sup>	low-up (months)	z	Of which: non-flail	z	Of which: non-flail	Outcome <sup>‡</sup>	Quality of Evidence
Nirula <i>et al. [63]</i>	2006	Cohort Study	09	51	NR	30	50% (15/30)	30	70% (21/30)	AGH	III - Low
de Moya <i>et al. [52]</i>	2011	Case Control Study	48	46	NR	16	43% (7/16)	32	66% (21/32)	ACDFGH	III - Low
Balci <i>et al.* [44]</i>	2013	Cohort Study	178	NR	25.9	92	Unknown	98	Unknown	D	III - Low
Majercik <i>et al. [64]</i>	2015	Case Control Study	411	56	NR	137	26% (36/137)	274	68% (187/274)	ACEGH	III - Good
Okoye <i>et al.* [35]</i>	2015	Cohort Study	13,853	48	NR	1339	100%	12514	100%		III - Low
Wu et al. [40]	2015	Single-center RCT	164	51	2	75	59% (44/75)	89	61% (54/89)	ABCDEGH	I-Low
Fagevik Olsen <i>et al. [45]</i>	2016	Cohort Study	61	89	12	31	Unknown	30	Unknown		III - Low
Metin <i>et al. [65]</i>	2016	Case Series	44	57	36	17	Unknown <sup>II</sup>	27	Unknown <sup>II</sup>	I	III - Low
Qiu <i>et al. [37]</i>	2016	Cohort Study	124	38	9	9	100%	65	100%	СДН	III - Good
Pieracci <i>et al. [42]</i>	2016	Cohort Study	70	51	NR	35	20% (7/35)	35	69% (24/35)	ABCDEGH	II- Good
Tarng <i>et al. [66]</i>	2016	Case Series	65	99	24	12	25% (3/12)	53	Unknown	ADEFGH	III - Low
Velasquez <i>et al. [67]</i>	2016	Cohort Study	40	49	NR	20	Unknown	20	Unknown	ACDFGH	III - Low
Uchida <i>et al. [48]</i>	2017	Cohort Study	20	09	NR	10	40% (4/10)	10	Unknown	ACDEFG	III - Low
Ali-Osman <i>et al. [68]</i>	2018	Cohort Study	199	70	NR	64	Unknown	135	Unknown	АСДВН	III - Good
Kane <i>et al. [69]</i>	2018	Cohort Study	1,116	48	NR	116	35% (41/116)	1000	Unknown	CDEGH	III - Good
Achary <i>et al.</i> * [43]	2019	Cohort Study	691	NR	NR	31	Unknown	099	Unknown	AFGH	III - Low
Azim <i>et al.*</i> [33]	2019	Cohort Study	474	NR	N R	237	100%	237	100%		III - Good

Fokin <i>et al.</i> [71] 2019	Conort Study	332	00	W W	92	43% (28/65)	267	79% (212/267)	ACGH	- Good
	Cohort Study	174	56	N.R.	87	49% (44/87)	87	(28/82) %06	AD	- Good
Haddadin <i>et al. [34]</i> 2019	Cohort Study	65,337	53	NR	1240	100%	64097	100%	O	III - Low
Jiang et al. [72]	Cohort Study	167	55	8	75	Unknown	92	Unknown	СДН	- Good
Marasco <i>et al. [73]</i> 2019	Cohort Study	1,482	54	24	29	48% (32/67)	1415	90% (1277/1415)	ADG	m - Good
Mullens <i>et al. [74]</i> 2019	Cohort Study	12,910	57	< 1	57	82% (47/57)	12853	100%	AEGH	III - Low
Turner <i>et al. [75]</i> 2019	Cohort Study	64	NR	NR	32	Unknown	32	Unknown	D	III - Low
Xiong <i>et al.</i> [76] 2019	Cohort Study	123	47	N. R.	89	Unknown	55	Unknown	СН	poo5 - III
Zhang <i>et al.</i> [39]	Cohort Study	78	50	9	39	100%	39	100%	BDEF	poo5 - III
Dorman <i>et al. [77]</i> 2020	Cohort Study	96	65	N. R.	54	28% (15/54)	42	57% (24/42)	O	III - Low
Dubrov <i>et al. [41]</i> 2020	Cohort Study	41	40	NR	17	53% (9/17)	24	63% (15/24)	ACDFGH	- Good
Li et al. [12]	Cohort Study	86	55	12	99	100%	32	100%	Q	- Good
Pieracci <i>et al. [36]</i> 2020	Cohort Study⁰	110	55	2	51	100%	59	100%	BCD	- Good
Xiao <i>et al. [38]</i> 2020	Cohort Study	1,010	50	-	430	100%	580	100%	ACDGH	poo5 - III

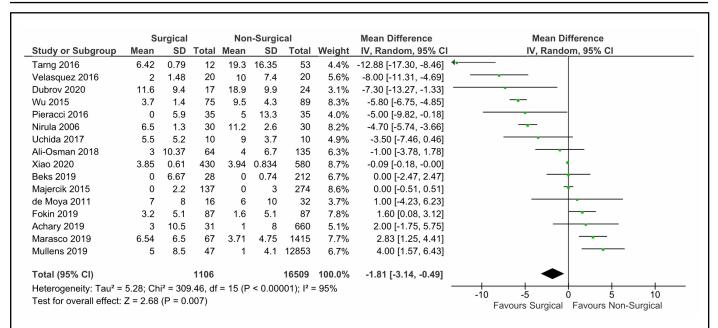
\*Abstract only - no publication

<sup>†</sup> As reported (mean or median) or calculated from reported ages of surgical and non-surgical cohorts

<sup>||</sup> non-flail percent provided for entire cohort: 68% (30/44)

<sup>♦</sup> Mixed design: prospective cohort and RCT

assessments (Table S1 in Supplemental Files) have no associated letters. Abbreviations: A, duration of mechanical ventilation; B, pain scores; C, respiratory complications; D, mortality, E, Tracheotomy; F, Sepsis; G, length of hospital stay; H, length of ICU stay. # The contribution of each study to the specific meta-analyses is shown with letters (A-H) indicative of each outcome type. Studies that only contributed to the narrative



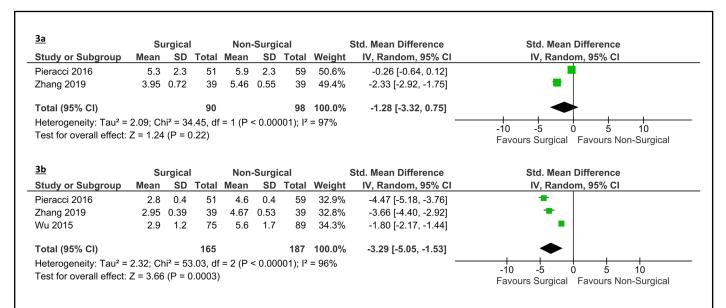
**Figure 2**. Forest plot of the meta-analysis of surgical vs non-surgical care in patients with severe rib fracture on duration of mechanical intervention (in days).

mean DVM in the surgical group was 3.9 (SD 0.6) and in the non-surgical group, 5.5 (SD 0.6), for a mean difference of -1.8 (95% confidence intervals (Cl): -3.1 to -0.5). This difference was significant (p<0.00001). Significant heterogeneity was identified across all studies ( $I^2 = 95\%$ ).

**Post-Treatment Pain:** Two different measures of pain were available: pain after 3 days (**Figure 3a**) vs. after 2 weeks post-intervention (**Figure 3b**). Only 2 papers described pain within 3 days of admission. In those papers, pain in the surgical cohort was lower (aggregate VAS score: 4.6 (SD: 0.7)) compared to that

of patients in the non-surgical group (aggregate VAS score: 5.6 (SD: 0.2)). The mean difference of -1.28 (95%CI: -3.3 to + 0.75) was not significant [39,42]. For the 2 weeks post-intervention pain analysis, 3 studies contributed data. The difference in pain at that time was significant at -3.3 (95%CI: -5.0 to -1.5, p < 0.00001), suggesting lower pain for patients surgically treated [36,39,40]. For both analyses, significant heterogeneity was observed (I<sup>2</sup>>95%).

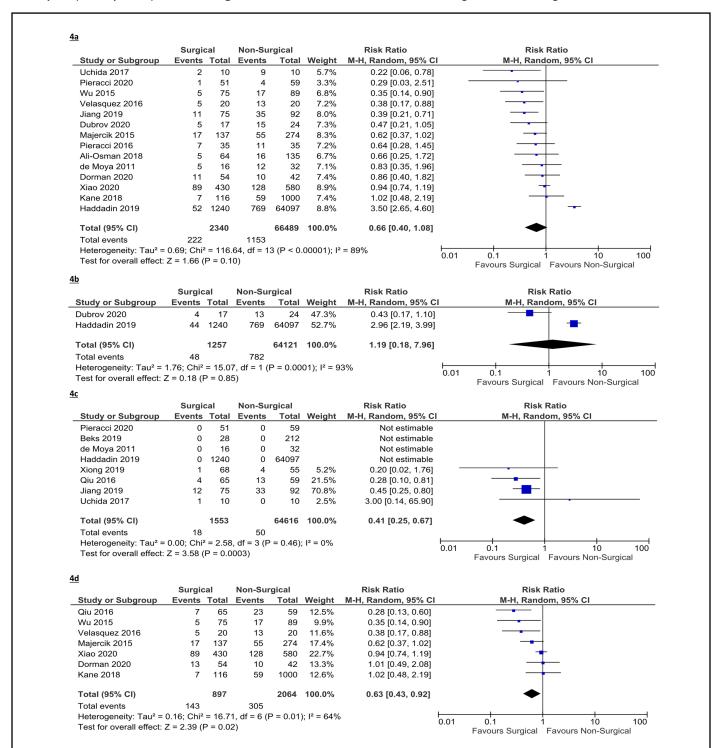
**Respiratory Complications:** 17 studies reported on respiratory complications, but not all studies reported on



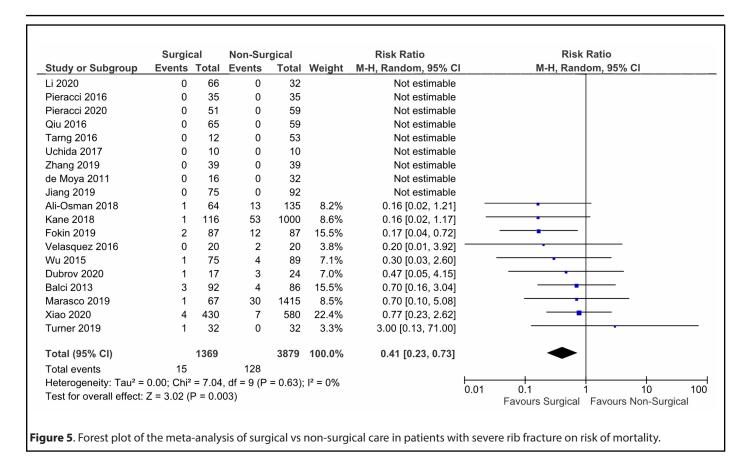
**Figure 3.** Forest plot of the meta-analysis of surgical vs non-surgical care in patients with severe rib fracture on pain at 3-days and 2 weeks post-intervention. **3a:** Effect on pain scores at 3 days post-intervention. **3b:** Effect on pain scores at 2 weeks post-intervention.

all complication types. Fourteen studies described risk of pneumonia (**Figure 4a**), 2 for risk of acute respiratory distress syndrome (ARDS) (**Figure 4b**), 8 studies described risk for atelectasis (**Figure 4c**), and 7 provided an aggregate risk for any respiratory complications (**Figure 4d**). The risk for

pneumonia was not different between the surgical and nonsurgical cohorts (risk ratio: 0.66 (95%CI: 0.40 to 1.08, p=0.1). The risk for ARDS was also not significantly different between groups. However, the risk for atelectasis was significantly lower in the surgical vs non-surgical cohort (risk ratio: 0.41



**Figure 4.** Forest plot of the meta-analysis of surgical vs non-surgical care in patients with severe rib fracture on risk of respiratory complications. **4a:** Effect on risk of pneumonia. **4b:** Effect on risk of acute respiratory distress syndrome (ARDS). **4c:** Effect on risk of atelectasis. 4d: Effect on overall risk of respiratory complications.



(95%Cl: 0.25 to 0.67, p = 0.0003), and the aggregate risk for any respiratory complications was significantly lower in the surgical vs the non-surgical cohort (risk ratio: 0.63 (95%Cl: 0.43 to 0.92), p = 0.02). High heterogeneity was observed for pneumonia, ARDS and the aggregate respiratory complication analyses ( $l^2 > 63\%$ ) but low heterogeneity was observed for risk of atelectasis ( $l^2 = 0\%$ ).

**Mortality:** Nineteen studies evaluated the impact of SSRF on mortality, of which 10 studies reported actual numerical outcomes (**Figure 5**). The duration of follow-up ranged from the duration of the hospital stay to three months. The meta-analysis found that surgically treated patients had a statistically significantly lower risk of mortality (RR 0.41 (95% CI 0.23 to 0.73, p=0.003).

**Tracheotomy and Sepsis:** As shown in **Figures S6 and S7** in the **Supplemental Files**, no difference in rate of tracheotomy or sepsis were observed between SSRF and non-surgically treated patients (for tracheotomy: RR 0.66 (95%CI: 0.30-1.44, p = 0.29) – for sepsis: RR 0.75 (95%CI: 0.17 to 3.28, p = 0.70).

**Length of Stay (LOS)** – **Intensive Care Unit (ICU) and Hospital:** Differences in LOS, both in ICU and in hospital overall, were not significantly different between SSRF and nonsurgically treated patients, showing a trend towards shorter LOS in SSRF patients, as shown in **Figures S8(a) and S8(b).** (Difference between SSRF and non-surgically treated LOS, in

days: ICU: - 1.01 (95%Cl: -2.42 to +0.39, p = 0.16 - Hospital: - 1.52 (95%Cl: -3.97 to +0.92, p = 0.22))

#### Discussion

Patients with non-flail, severe rib fractures, particularly older adults [13-16], frequently require mechanical ventilation and are at increased risk for death [11,12]. Many of these patients could benefit from SSRF [36,46-48], however, a lack of consensus regarding the appropriate indications for which to initiate surgical treatment typically leads to nonsurgical management [49-52]. As surgical techniques and the technology of internal fixation devices have advanced, SSRF has become safer, easier to perform, and more efficient, and the debate about whether non-flail chest patients would benefit from SSRF continues to evolve as additional evidence emerges [30].

The current study showed that SSRF is beneficial to patients with severe rib fractures, including non-flail cases, as it results in a statistically significant shorter DMV, lower 2-week pain intensity, lower risk of atelectasis and overall respiratory complications, and reduced mortality, compared to non-surgical treatment. Although trending in favor of SSRF, differences between SSRF and non-surgical treatment were not statistically significant for pain three days after the intervention, pneumonia, sepsis, ICU LOS, and hospital LOS.

Differentiating pain levels at three days is challenging as it is expected to be very severe for all patients with severe rib fractures and, hence, it may be too early to detect differences between treatments utilizing validated instruments. For pneumonia, the reduced risk with SSRF approached statistical significance. Also, a subgroup analysis only including the studies with a quality rating of "good" (n=8) showed significantly lower risk of pneumonia (RR = 0.67, 95% CI= 0.51 to 0.87, p = 0.002,  $I^2=28\%$ ) for SSRF vs non-surgically treated patients. The occurrence of sepsis was very infrequent, hence showing a statistically significant difference would require large sample sizes. It is also challenging to show statistically significant differences in resource utilization such as the mean hospital and ICU LOS given the variability and unpredictability among patients and the presence of outliers. Studies also included populations from different geographical locations (outside the United States) with potentially different protocols for duration of hospital care and discharge.

There were not adequate data to conduct meta-analyses for dose and duration of pain medicines, rates of emphysema, pneumothorax, hemothorax, pulmonary effusion, and readmission, cost of treatment, functional outcomes, and QoL. Evidence for the effectiveness of SSRF for particular outcomes has been mounting over the past decade, and it would be important to collect data for these outcomes as they have been identified as important considerations in the published medical literature [30].

The majority of studies evaluating SSRF included non-flail, severe patients, with some flail chest cases; however, the population with flail chest, in the SSRF arm, made up less than 5% of the overall sample. Our findings are therefore generalizable to most patients with severe, non-flail rib fractures.

Limitations of the current study include the heterogeneity of the patient populations evaluated, the surgical techniques and technologies employed, and the definitions of outcomes used in the analyses (eg, DMV may be defined as continuous positive airway pressure [CPAP] or tracheostomy and a mechanical ventilator). Heterogeneity is likely to arise when there are differences in patient populations, treatments, study design, outcomes, and data quality and is expected when pooling observational (real-world) data [53]. In the field of orthopedics, a growing opinion suggests that inclusion of observational studies in meta-analyses might lead to more robust conclusions without compromising the quality of the results [54,55]. The current study was conducted in line with recommendations available in the literature for the use of real-world evidence in meta-analyses [56]. Statistical heterogeneity was evaluated using Cochran's Q test ( $\chi^2$  test) and the I<sup>2</sup> statistic. Since Q was significant and I<sup>2</sup> was >50%, it was appropriate to use the random-effects model (REM) to calculate pooled summary estimates. The range of I<sup>2</sup> values observed in the current study (0% to 97%) is consistent with the range of those observed in other published meta-analyses

of SSRF (0% to 95%) [18-27].

Aside from SSRF, other factors that may influence the outcomes of patients with multiple severe rib fractures are the presence of lung contusion, the presence of pleural effusion including hemothorax and pneumothorax, a change of respiratory mechanics (eg, flail chest), compromised respiratory function due to pain, polytrauma, and the patient's baseline health conditions [16,49,57-60]. The role of lung contusion in influencing the decision to proceed with surgical treatment is unclear [61]. Lung contusion is considered as a key risk factor for the need for ventilation and for near-term and long-term respiratory function and relevant complications [62]. The current study did not attempt to ascertain the influence of lung contusion on outcome given that the published data were not stratified by the presence of contusion and the studies were not randomized; hence, this variable or other unknown variables may have confounded results. However, it is also plausible that patients who received SSRF were more severe, in which case the bias would be against those treated surgically. A better understanding of the indications and circumstances under which surgical management of rib fractures is the optimal treatment option is needed.

Additional prospective and randomized controlled trials should be conducted among patients with multiple non-flail displaced rib fractures to validate the findings of this meta-analysis. Further studies should focus on the effect of injury severity, surgical technique, timing of surgery, follow-up duration, and the effectiveness of SSRF. Evaluation of economic benefits would also be beneficial to better understand the cost-effectiveness of SSRF in patients with non-flail rib fractures.

## **Conclusion**

In conclusion, SSRF for patients with multiple displaced non-flail rib fractures resulted in shorter DMV, lower 2-week pain intensity, lower risk of atelectasis, lower risk of overall respiratory complications, and reduced mortality compared to non-surgical treatment; SSRF might be indicated in a broader range of cases than is currently performed. Additional studies would be beneficial in more clearly elucidating the specific patient populations that would benefit the most from SSRF.

#### **Conflict of Interest**

SW, TG, AW, RES, CEH and MV were employees of Johnson & Johnson (JnJ) at the time of the study. RS was working with ClinChoice as a contractor to JnJ at the time of the study. AMS is a consultant for DePuy Synthes, a J&J company, and a consultant for Globus Medical.

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