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# BMJ Open Systematic review of systematic reviews for effectiveness of internal fixation for flail chest and rib fractures in adults

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## ABSTRACT

**Objectives** Multiple systematic reviews have reported on the impact of rib fracture fixation in the presence of flail chest and multiple rib fractures, however this practice remains controversial. Our aim is to synthesise the effectiveness of surgical rib fracture fixation as evidenced by systematic reviews.

**Design** A systematic search identified systematic reviews comparing effectiveness of rib fracture fixation with non-operative management of adults with flail chest or unifocal non-flail rib fractures. MEDLINE, EMBASE, Cochrane Database of Systematic Reviews and Science Citation Index were last searched 17 March 2017. Risk of bias was assessed using the Risk Of Bias In Systematic reviews (ROBIS) tool. The primary outcome was duration of mechanical ventilation.

**Results** Twelve systematic reviews were included, consisting of 3 unique randomised controlled trials and 19 non-randomised studies. Length of mechanical ventilation was shorter in the fixation group compared with the non-operative group in flail chest; pooled estimates ranged from -4.52 days, 95% CI (-5.54 to -3.5) to -7.5 days, 95% CI (-9.9 to -5.5). Pneumonia, length of hospital and intensive care unit stay all showed a statistically significant improvement in favour of fixation for flail chest; however, all outcomes in favour of fixation had substantial heterogeneity. There was no statistically significant difference between groups in mortality. Two systematic reviews included one non-randomised studies of unifocal non-flail rib fracture population; due to limited evidence the benefits with surgery are uncertain.

**Conclusions** Synthesis of the reviews has shown some potential improvement in patient outcomes with flail chest after fixation. For future review updates, meta-analysis for effectiveness may need to take into account indications and timing of surgery as a subgroup analysis to address clinical heterogeneity between primary studies. Further robust evidence is required before conclusions can be drawn of the effectiveness of surgical fixation for flail chest and in particular, unifocal non-flail rib fractures.

**PROSPERO registration number** CRD42016053494.

## INTRODUCTION

Multiple rib fractures (adjacent unifocal fractures) and flail chest (three or more adjacent bifocal rib fractures with paradoxical chest wall movement) have high mortality (18.7%)

## Strengths and limitations of this study

- Multiple databases were searched for studies and study selection was undertaken by two researchers, reducing the risk of error.
- Risk of bias of studies was assessed using the ROBIS tool by two researchers.
- Primary research within the reviews was mapped to identify if the reviews were based on the same primary evidence.
- Reviews were included regardless of risk of bias.

due to the associated complications of acute respiratory distress syndrome, pneumonia and haemorrhage.<sup>1</sup> Chest trauma accounts for 15% of all trauma admissions<sup>2</sup> most commonly high-energy transfer injury as a result of a road traffic accidents (57.01%) but also from low energy falls (22.96%).<sup>1</sup> Flail chest in particular has a high mortality rate as chest wall disruption causes an increase in the work of breathing for patients who are often in significant pain.

Current treatment options for severe chest injury are mainly supportive, including multimodal analgesia and anaesthesia, as well as non-invasive and invasive ventilation.<sup>3</sup> Surgical fixation is thought to be beneficial to patients with respiratory failure,<sup>4</sup> intractable pain<sup>5</sup> or if failing to wean from invasive ventilation secondary to chest trauma.<sup>6,7</sup> Fixation has potential to restore chest wall biomechanics and reduce the complications associated with poor ventilation and secretion clearance.<sup>7</sup> Due to the rapidly, although heterogeneous, growing evidence base from multiple systematic reviews it is essential to synthesise evidence for this intervention to ascertain safety and efficacy.

This report is part of a wider systematic review to (1) identify and synthesise the evidence of the effectiveness of surgical rib fracture fixation, (2) evaluate the evidence for indications and timing of fixation and (3)

identify the outcomes reported in the literature. Systematic reviews and primary studies were eligible for inclusion. This paper maps and synthesises this evidence from systematic reviews assessing the effectiveness of fixation of rib fractures.

## METHODS

The review was undertaken systematically using the methods described by the Centre for Reviews and Dissemination.<sup>8</sup>

### Eligibility criteria

#### Population

Adults (over 18 years) who have sustained one or more rib fractures following blunt chest trauma, with or without pulmonary contusion were eligible. Single rib fracture, multiple rib fractures and flail chest injuries were included but treated as separate injuries and therefore reported and analysed separately. Patients with penetrating injuries were excluded. Studies of mixed populations with penetrating and non-penetrating injuries were included only if data were presented separately for the two groups. Surgery for chronic non-union was excluded.

#### Intervention

Any method of internal surgical fixation such as plate or strut fixation; metal or synthetic material including intramedullary splints and suture fixation were eligible for inclusion.

#### Comparator

External surgical fixation (traction methods, splints and Hoffman style pin and bar fixation) and non-surgical management (such as supportive ventilation, epidural and regional anaesthesia).

#### Outcomes

All outcomes were eligible (eg, mortality, pain and pneumonia). The primary outcome of interest was duration of mechanical ventilation due to the close relationship with mortality and morbidity of ventilator associated complications.

### Study design

#### Inclusion

Systematic reviews were included if they specified a search strategy in at least one literature database and included primary research. No restrictions were placed on the study design of the primary studies.

#### Exclusion

Literature reviews that did not have a defined research question, search strategy or defined process of selecting articles.

### Search strategy

MEDLINE including PreMEDLINE, EMBASE, Cochrane Database of Systematic Reviews and Science Citation

Index. Clinical guidance, policy documents and relevant databases such as National Institute for Health and Care Excellence (NICE) Evidence, the UK Department of Health policy content, National Clinical Guideline Centre and Scottish Intercollegiate Guidelines Network were searched on 14 December 2016 and updated on 13 March 2017. The Conference Proceedings Citation Index was also searched for unpublished literature.

The start date for the MEDLINE searches was 1976 as that was the year that Advance Trauma Life Support was introduced internationally, incorporating new methods of resuscitation which have significantly improved outcomes.

The search strategy, developed for MEDLINE, is provided in online supplementary file 1 and was adapted to run appropriately on other databases. To identify relevant further reviews reference lists of included studies were assessed for eligibility.

### Selection

Searches were downloaded into Endnote X7 (Clarivate Analytics, V.7.1 release date 2 April 2014) and de-duplicated. Two researchers (HI and EC) independently screened titles and abstracts. Any paper classified as potentially eligible by either reviewer was ordered as a full text and independently screened by both reviewers. It was originally planned to have second screening of only 50% but resources allowed for full duplicate screening. A third researcher reviewed disagreements (CM) where a consensus could not be reached between the researchers.

### Data extraction

Extracted data included study characteristics, patient characteristics, intervention, comparator, outcome measures, duration of follow-up, effect estimates, SE and CIs as available.

One researcher completed data extraction (HI); a second researcher cross-checked 50% (EC). Discrepancies were cross-checked by both researchers at a second review and a consensus reached.

### Risk of bias

Quality assessment with the ROBIS tool<sup>9</sup> was undertaken by one researcher (HI) and checked by a second (CM). Discrepancies were resolved by discussion.

### Data synthesis

All types of internal surgical fixation were synthesised as one group. Flail chest and multiple rib fractures are considered different injuries and were synthesised separately for each outcome extracted. Each outcome was narratively synthesised including number of reviews using the outcome and effect estimates with 95% CIs from the source review. Important numerical data was presented in tables for all outcomes measured. All outcomes that were reported in the reviews were included in the report to avoid reporting bias.<sup>10</sup> Although not fully applicable, reporting was in accordance as much as possible with the

Preferred Reporting Items for Systematic Reviews and Meta-Analyses statement.<sup>11</sup>

### Protocol deviations

The registered protocol encompasses a larger body of work which includes synthesis of primary research for effectiveness, indications for surgery, timing of surgery and mapping of outcome measures. Only the synthesis of systematic reviews is reported here. Although all outcomes were extracted and presented in tables only those that were measured in two or more studies were narratively synthesised.

### Patient involvement

Patients were not involved in the preparation or conduct of this review.

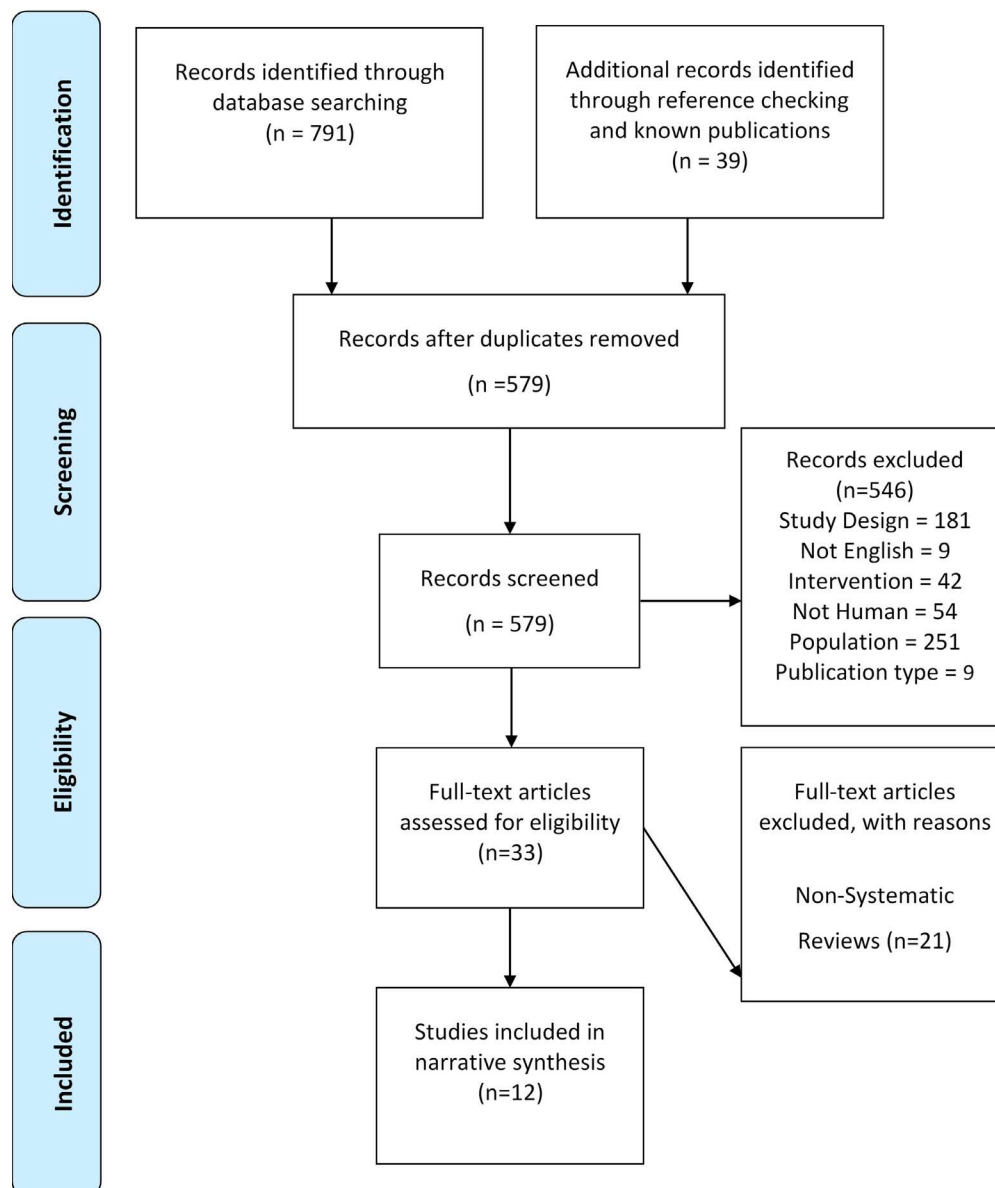
## RESULTS

Electronic searches identified 791 records; an additional 39 records were collected following reference checking. The full text screening identified 12 systematic reviews eligible for inclusion, there were 21 papers excluded because they were not classified as systematic reviews (figure 1). See online supplementary file 2 lists the excluded studies.

### Review characteristics

Eleven systematic reviews and one rapid evidence synthesis,<sup>12</sup> published between 2010 and June 2016, met the inclusion criteria. Table 1 provides a summary of review characteristics. Three of the reviews<sup>13-15</sup> were presented as best evidence topics<sup>16</sup>

Nine reviews<sup>12 13 15 17-22</sup> evaluated the effectiveness of internal surgical fixation in patients with flail chest, two



**Figure 1** Preferred Reporting Items for Systematic Reviews and Meta-Analyses flow diagram.

**Table 1** Review characteristics

Review Year Country	Review aim	Search strategy	Studies and participants	Patient, Intervention, Comparator, Outcome and Study type (PICOS)	Risk of bias	Authors' conclusions
Swart <i>et al</i> <sup>17</sup> 2017 USA	To perform a meta-analysis of high quality literature to evaluate both economic and medical benefits of early fixation of rib fractures in severe chest trauma.	PubMed, Embase, Medline and Scopus. No search start date. Last search date 1 June 2016. Search terms defined, No limitations described. Evidence of hand searching. Eligibility criteria: over 18 years of age and studies comparing operative versus non-operative treatment,	3 RCT n=123 14 Case-control 3 Case series	<b>Population</b> Acute flail chest 18 years or older. <b>Intervention</b> Operative fixation. <b>Comparator</b> Non-operative. <b>Studies type</b> All study designs.	No evidence of quality assessment.	Acute ORIF of rib fractures in patients with flail chest injuries results in reduced mortality and medical complications in conjunction with being cost effective intervention.
Schuermans <i>et al</i> <sup>18</sup> 2017 The Netherlands	Investigate how operative management improves patient care for adults with flail chest.	PubMed, Trip database, Google Scholar. No search start date. Last search date November 2015. Search terms defined, No limitations described. Evidence of reference checking. Eligibility criteria: studies comparing operative versus non-operative treatment, RCT only and English.	3 RCT n=123	<b>Population</b> Acute flail chest. <b>Intervention</b> Operative fixation. <b>Comparator</b> Non-operative. <b>Studies type</b> RCTs.	Quality assessment completed but criteria and explanation unclear.	The operative management group showed a significant lower incidence of pneumonia, whereas mortality rate did not differ between treatment groups.
Schulte <i>et al</i> <sup>13</sup> 2016 UK	In patients with acute flail chest does surgical rib fixation improve outcomes in terms of morbidity and mortality?	OID MEDLINE. Search start date 1946. Last search date January 2016. Search terms defined. Search strategy description minimal, no limitations described. No evidence of reference checking. No specific inclusion or exclusion criteria defined.	1 Meta-analysis by separate author. 1 RCT n=123 (2 further coded as RCT which are non-randomised studies). 3 Retrospective cohort studies.	<b>Population</b> Acute flail chest. <b>Intervention</b> Operative fixation. <b>Comparator</b> Non-operative. <b>Studies type</b> Unclear.	No evidence of quality assessment.	Surgical stabilisation of flail chest in thoracic trauma patients has beneficial effects with respect to reduced ventilatory support, shorter intensive care and hospital stay, reduced incidence of pneumonia and septicaemia, decreased risk of chest deformity and an overall reduced mortality when compared with patients who received non-operative management.
Coughlin <i>et al</i> <sup>19</sup> 2016 UK	Compare the efficacy of flail chest surgical stabilisation to non-operative management.	PubMed MEDLINE, Embase, Cochrane Library, clinical trials.gov. No search start date. Last search date February 2015. Search terms defined, No limitations. Evidence of reference checking. Eligibility criteria: studies comparing operative versus non-operative treatment in flail chest and RCT only.	3 RCT n=123	<b>Population</b> Traumatic flail chest. <b>Intervention</b> Surgical stabilisation of any kind. <b>Comparator</b> Patients treated non-operatively by any other means. <b>Studies type</b> RCTs only.	Clear quality appraisal of the studies.	Surgical stabilisation for a traumatic flail chest is associated with significant clinical benefits including rate of pneumonia, length of hospital an ICU stay and duration of mechanical ventilation in this meta-analysis of three relatively small RCTs.
Unsworth <i>et al</i> <sup>24</sup> 2015 Australia	To review the treatments for blunt chest trauma and their impact on patient and hospital outcomes. Specifically alludes to surgical stabilisation of flail chest.	Cochrane, Medline, EMBASE and CINAHL databases. Search limited to 1990 onwards. Last search date March 2014. Search terms defined. Limited to humans and adults. Evidence of reference checking. Eligibility criteria: original research, blunt chest trauma, intervention for blunt chest trauma including a comparator and contained measured outcomes.	3 RCT n=123 5 Retrospective case-controls n=642. 1 Retrospective cohort n=21.	<b>Population</b> Adult blunt chest trauma. Flail chest. <b>Intervention</b> Multidisciplinary intervention (models of care, management intervention, care practices, care protocols). <b>Comparator</b> Other intervention not specified. <b>Studies type</b> RCTs.	Some quality assessment completed but criteria and explanation unclear.	Across the literature there were consistent improvements in patients with flail chest and surgical fixation with fewer days of mechanical ventilation, ICU-LOS and cost savings compared with non-operative techniques. Three out of nine studies were randomised controlled trials, and the level of evidence in all studies was primarily fair or good.

Continued

Table 1 Continued

Review Year Country	Review aim	Search strategy	Studies and participants	Patient, Intervention, Comparator, Outcome and Study type (PICOS)	Risk of bias	Authors' conclusions
de Lesquen <i>et al</i> <sup>15</sup> 2015 France	In flail chest is open reduction and internal fixation needed?	Medline and Science Direct. Search start date limited to 1994 onwards. Last search date January 2014. Search Terms defined. No evidence of hand searching or reference checking. Eligibility criteria: Exclusions of both child and vascular injuries.	2 Meta-analysis. 3 RCT n=123. 1 Prospective cohort n=60. 5 Retrospective cohort n=238.	<b>Population</b> Blunt chest trauma. Flail chest. <b>Intervention</b> Open reduction and internal fixation. <b>Comparator</b> Unclear. <b>Studies type</b> Unclear.	No evidence of quality assessment.	For flail chest, early surgical stabilisation can be considered in patients who would require mechanical ventilation for >48 hours.
Cataneo <i>et al</i> <sup>20</sup> 2015 Brazil	To evaluate the effectiveness and safety of surgical stabilisation compared with clinical management for people with flail chest.	Cochrane Injuries Group Specialised Register, CENTRAL, Medline, Embase, CINAHL, SCI, CPCI-S, Clinical trials.gov, ICTR. No search start date. Last search Date 12 May 2014. Search terms defined, no limitations. Evidence of reference checking. Eligibility criteria: RCTs.	3 RCTs n=123.	<b>Population</b> Adults or children with flail chest. <b>Intervention</b> Surgical stabilisation of any kind. <b>Comparator</b> Clinical management included any type of chest wall stabilisation without surgical intervention such as straps or bags and any type of ventilatory assistance. <b>Studies type</b> RCTs only.	Clear quality appraisal of the studies.	There was no evidence that surgical intervention reduced mortality in people with FC compared with non-surgical management. There was some evidence that surgical intervention could reduce the risk of developing pneumonia and thoracic deformity; need for tracheostomy; duration of mechanical ventilation, length of ICU stay and hospital stay; and chronic pain, but the trials to date have been small. There is an urgent need for larger high-quality randomised con-trolled trials.
de Jong <i>et al</i> <sup>23</sup> 2014 The Netherlands	To specify indications for rib fracture fixation of non-flail chests.	Medline, Cochrane, Embase. Search start date limited to 2010. Last search date December 2013. Search terms defined, limited to year 2000 onwards. Evidence of reference checking. Eligibility criteria: Studies included at least 10 participants who were surgically treated for non-flail chest rib fractures. Reported in English, Dutch or German. Excluded were case reports, biomechanical studies, animal studies and expert opinions.	1 Case-control n=60. 2 Cohort studies n=47.	<b>Population</b> Traumatic non-flail chest. <b>Intervention</b> Surgical treatment of non-flail chest. <b>Comparator</b> Unclear. <b>Studies type</b> All studies with at least 10 surgically treated.	No evidence of quality assessment.	The evidence for surgical treatment of non-flail chest rib fractures is limited.
Slobogean <i>et al</i> <sup>21</sup> 2013 Canada	Compare the critical care outcomes of surgical fixation to non-operative management in patients with flail chest injuries.	Medline, Embase, Cochrane Database of Systematic Reviews and the Cochrane Central, Register of Controlled Trials (CENTRAL). No search start date. Last search date May 2011. No limitations. No evidence of reference checking or hand searching. Eligibility criteria: Comparator studies with more than 10 cases.	2 RCTs. 1 Case-control n=60. 8 Cohort n=676.	<b>Population</b> Acute flail chest. <b>Intervention</b> Operative fixation. <b>Comparator</b> Conservative management. <b>Studies Type</b> RCTs.	No evidence of quality assessment.	Improved outcomes of multiple critical care outcomes with narrow CIs but based on small retrospective studies. Suggests prospective RCT to overcome potential biases.

Continued

Table 1 Continued

Review Year Country	Review aim	Search strategy	Studies and participants	Patient, Intervention, Comparator, Outcome and Study type (PICOS)	Risk of bias	Authors' conclusions
Leinicke <i>et al</i> <sup>22</sup> 2013 USA	Comparing operative to non-operative therapy in adult flail chest patients.	MEDLINE (1966–2012), Embase (1947–2012), Scopus (all years), Cochrane Databases and ClinicalTrials.gov Last search date February 2012. Search terms defined, limited to English and human studies. Evidence of reference checking. Eligibility criteria: Studies comparing operative versus non-operative treatment in patients with flail chest. Excluded case reports and case series.	2 RCTs. 3 Case-control n=158. 4 Cohort n=303.	<b>Population</b> Flail chest. <b>Intervention</b> Operative fixation. <b>Comparator</b> Non-operative. <b>Studies type</b> RCTs, cohort and case-control trials.	Clear quality appraisal of the studies.	As compared with non-operative therapy, operative fixation of FC is associated with reductions in DMV, LOS, mortality and complications associated with prolonged MV. These findings support the need for an adequately powered clinical study to further define the role of this intervention.
Girsowicz <i>et al</i> <sup>14</sup> 2012 France	In patients over 45 years old with isolated, movable and painful rib fractures without true flail chest is surgical stabilisation superior to non-operative management in improving outcomes?	OID Medline 1948–2011. Last search date June 2011. Search terms defined, limited to Human and English language. Evidence of reference checking. Eligibility criteria: Excluded flail chest but inclusions not well described.	4 Retrospective cohort n=107. 1 Non-systematic review. 1 Case-control=30. 2 Case report n=2.	<b>Population</b> Over 45 years old with isolated, movable and painful. Rib fractures without true flail chest. <b>Intervention</b> surgical stabilisation. <b>Comparator</b> Non-operative management. <b>Studies type</b> Unclear.	Some comments on strengths and weaknesses but no quality or risk of bias assessment.	Surgical stabilisation in the management of isolated multiple non-flail and painful rib fractures improved outcomes (pain, respiratory function, quality of life and reduced socio-professional disability). Studies provided a low level of evidence (small studies with few numbers of patients and short-term follow-up or case reports). Large prospective controlled trials are thus necessary to confirm these encouraging results.
NICE Evidence <sup>12</sup> 2010 UK	To make recommendations about the safety and efficacy of surgical rib fracture fixation in flail chest.	MEDLINE, PREMEDLINE, EMBASE, Cochrane Library. No search start date. Last search date May 2010. Search terms defined. No limitations. No evidence of reference checking but other searches performed. Eligibility criteria: clinical studies of patients with flail chest operated with metal rib reinforcements and published in English. Excluded conference abstracts and reviews.	1 RCT. 2 Non-randomised studies. Four case series. Total 225 patients.	<b>Population</b> Flail chest. <b>Intervention</b> Insertion of metal rib reinforcements. <b>Comparator</b> Unclear. <b>Studies type</b> Clinical studies were included. Abstracts were excluded where no clinical outcomes were reported or where the paper was a review, editorial or a laboratory or animal study. Conference abstracts were also excluded.	No evidence of quality assessments.	Surgical rib fracture fixation should be consider in patients with flail chest.

DMV, duration of mechanical ventilation; FC, flail chest; ICU, Intensive care unit; LOS, length of stay; MV, mechanical ventilation; NICE, National Institute for Health and Care Excellence; ORIF, open reduction internal fixation; RCT, randomised controlled trial.

included patients with multiple rib fractures<sup>14 23</sup> and one included all rib fractures but only reported outcomes for flail chest.<sup>24</sup>

Three reviews<sup>18–20</sup> included only RCTs and eight included other study designs<sup>12–15 17 21–23</sup> (two systematic reviews, 19 non-randomised studies, 11 case series and two case reports) (table 2). As would be expected, there was overlap across the reviews in the included primary studies. The total number of patients who had internal fixation in primary studies (excluding duplicate studies) was 1036, and there were 1187 controls.

The rapid evidence synthesis by NICE<sup>12</sup> was the first review published in 2010, consisting of seven primary studies including one RCT published in 2001.<sup>25</sup> Cataneo *et al.*<sup>20</sup> was the first meta-analysis published (in 2015) and included three RCTs.<sup>25–27</sup> Two further systematic reviews published since then<sup>18 19</sup> identified the same three RCTs and repeated the same meta-analyses for the same review question.

### Risk of bias

Seven reviews rated as low risk of bias,<sup>12 15 18–22</sup> three as unclear,<sup>14 17 23</sup> and two as high.<sup>13 24</sup> (table 3) The high risk of bias rating was due to lack of detail in the search strategy, no attempts to minimise errors of data extraction and no quality assessment of included studies.

The only review for which a protocol was identified was the Cochrane review undertaken by Cataneo *et al.*<sup>20</sup>

### Outcome evaluation

All reviews undertook a narrative synthesis with six also including a meta-analysis.<sup>17–22</sup> Table 4 summarises the meta-analyses for flail chest, table 5 summarises the narrative syntheses for flail chest and table 6 the narrative syntheses for multiple rib fractures. Across all the reviews, 18 outcomes were reported and 11 outcomes were reported by more than one review.

### Primary outcome: length of mechanical ventilation (days)

#### Flail chest

Ten systematic reviews reported length of mechanical ventilation; six undertook a meta-analysis<sup>17–22</sup> four<sup>17–20</sup> of which included the same three RCTs.<sup>25–27</sup>

There was substantial variation across the reviews in the pooled estimates for this outcome, related to pooling different sets of studies. The largest reduction in duration of mechanical ventilation when surgery is compared with no surgery was reported by Slobogean *et al.*<sup>21</sup> pooling two RCTs<sup>25 26</sup> and six non-randomised studies<sup>28–33</sup> (mean difference [MD] –7.5 days, 95% CI –9.9 to –5.5) (table 4). The MD was 3 days greater than the pooled estimates from Leinicke *et al.*<sup>22</sup> (–4.52 days, 95% CI –5.54 to –3.50) and Swart *et al.*<sup>17</sup> (–4.57 days, SD 0.59).

There were differences in the data reported across the four meta-analyses<sup>17–20</sup> that included the same three RCTs. Schuurmans *et al.*<sup>18</sup> extracted median duration from the Marasco *et al.* RCT,<sup>27</sup> producing an estimate MD –6.53 days, 95% CI –11.88 to –1.18. In contrast, Coughlin

*et al.*,<sup>19</sup> Cataneo *et al.*<sup>20</sup> and Swart *et al.*<sup>17</sup> report the total mean time on mechanical ventilation which they state was obtained directly from the authors producing an estimate MD –6.30 days, 95% CI –12.16 to –0.43.

Variations also arose in relation to the extraction of data from the RCT by Granetzny *et al.*<sup>26</sup> who did not report SDs for mechanical ventilation. Slightly different SD values are found in all six meta-analyses<sup>17–22</sup> which may have arisen from different methods of imputation and all give slightly different estimates.

Substantial heterogeneity was seen in all meta-analyses reporting this outcome<sup>18 19 21 22</sup> ( $I^2=48\%$  to 95%). Only one study<sup>20</sup> did not pool due to heterogeneity.

Narrative synthesis from two reviews concluded that surgery reduces the length of mechanical ventilation compared with no surgery<sup>15 24</sup> (table 5).

### Multiple rib fractures

Two reviews<sup>14 23</sup> included one primary study<sup>33</sup> that had matched non-operative controls<sup>33</sup> (non-operative treatment not described) and reported a statistically significant reduction in postoperative ventilator days ( $p=0.02$ ) in favour of the fixation group (table 6) but no statistically significant difference in total ventilator days ( $p=0.12$ ).

### Mortality

#### Flail chest

Seven reviews reported mortality; six undertook a meta-analysis.<sup>15 17–22</sup> Three reviews<sup>18–20</sup> which pooled the same three RCTs<sup>25–27</sup> showed a non-statistically significant reduction in mortality with surgery compared with no surgery (risk ratio [RR] 0.56, 95% CI [0.13 to 2.42]<sup>18 20</sup> and RR 0.57, 95% CI [0.13 to 2.52]<sup>19</sup> [table 4]).

Three reviews pooled randomised and non-randomised studies.<sup>17 21 22</sup> Estimates were RR 0.19, 95% CI 0.13 to 0.26<sup>21</sup>; (RR 0.43, 95% CI 0.28 to 0.69<sup>22</sup> and RR 0.44, SD 0.09.<sup>17</sup> Overall, statistical heterogeneity was low ( $I^2=0\%$ ) for this outcome in all studies that presented this data.<sup>17–22</sup>

#### Multiple rib fractures

Mortality was not assessed by de Jong *et al.*<sup>23</sup> or Girsowicz *et al.*<sup>14</sup>

### Length of intensive care unit stay (days)

#### Flail chest

Eight reviews<sup>12 15 17–22</sup> assessed length of intensive care unit (ICU) stay; six undertook a meta-analysis.<sup>17–22</sup> Pooled estimates ranged from –3.25 days (SD 1.29)<sup>17</sup> to –6.46 days, 95% CI –9.73 to –3.19<sup>19</sup> and were all in favour of surgical fixation compared with a variety of comparators (table 4). The range in pooled estimates may be partly explained by the pooling of different sets of studies. However, differences occurred as some pooled median length of ICU stay and others pooled the mean. Furthermore, some used postoperative time spent in ICU and others the total time spent in ICU.<sup>17–19</sup>

Variation also arose across reviews in the data extracted from a trial that did not report SDs in the primary publication.<sup>26</sup> Values were imputed or the raw data obtained





**Table 2** Primary studies included in each review and the number of included patients

Review	Leiniche et al <sup>16</sup>	Slobogean et al <sup>17</sup>	Thiela et al <sup>18</sup>	Ginnatray et al <sup>19</sup>	Miracco et al <sup>20</sup>	Paris et al <sup>21</sup>	Kim et al <sup>22</sup>	Karee <sup>23</sup>	Ahmed and Mohyuddin <sup>24</sup>	Voggenreiter et al <sup>25</sup>	Bell et al <sup>26</sup>	Ting et al <sup>27</sup>	Nurula et al <sup>28</sup>	Alhazem et al <sup>29</sup>	De Moys et al <sup>30</sup>	Gravité and Pabouat <sup>31</sup>	Dobler et al <sup>32</sup>	Jajay et al <sup>33</sup>	Pierucci et al <sup>34</sup>	Zhang et al <sup>35</sup>	Wada et al <sup>36</sup>	Xu et al <sup>37</sup>	Majumdar et al <sup>38</sup>	Delfosse et al <sup>39</sup>	Chvapart et al <sup>40</sup>	Hilberg et al <sup>41</sup>	Mensel et al <sup>42</sup>	Morton et al <sup>43</sup>	Cacchione et al <sup>44</sup>	Landouk et al <sup>45</sup>	Khanjani et al <sup>46</sup>	Gaspert et al <sup>47</sup>	Bonelli and Azam <sup>48</sup>	Comballi et al <sup>49</sup>	Mahery et al <sup>50</sup>	Richardson et al <sup>51</sup>	Morino Chi Lu et al <sup>52</sup>			
Intervention patients		18	20	23	18	18	40	26	20	27	32	30	22	20	16	60	10	10	35	24	84	17	38	41	14	10	18	23	1	66	40	1	127	32	46	7	22			
Control patients		19	20	23	11	45	93	38	22	37	28	30	28	32	153	11	10	35	15	420	15	57	45	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-		
Swart et al <sup>17</sup>		•	•	•		•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•	•																
Schuurmans et al <sup>17</sup>		•	•	•																																				
Schulte et al <sup>17</sup>		•												•	•	•	•	•	•	•	•	•																		
Coughlin et al <sup>17</sup>			•	•	•																																			
Unsworth et al <sup>17</sup>			•	•	•			•	•				•	•	•		•																							
de Lesquen et al <sup>17</sup>	•	•	•	•	•		•	•	•	•			•	•																										
Cattaneo et al <sup>17</sup>																																								
de Jong et al <sup>17</sup>													•																							•	•			
Slobogean et al <sup>17</sup>		•	•			•	•	•	•	•	•	•	•	•											•															
Leiniche et al <sup>17</sup>		•	•				•		•	•			•	•	•																									
Gronowicz et al <sup>17</sup>													•																											
NICE <sup>17</sup>		•			•				•																	•	•	•												
	Systematic review	Randomised control trial	Non-randomised study																			Case series or report																		

NICE, National Institute for Health and Care Excellence.

**Table 3** Risk of bias using ROBIS tool

Studies	Study eligibility criteria	Identification and selection of studies	Data collection and study appraisal	Synthesis and findings	Risk of bias in the review
Swart <i>et al</i> <sup>17</sup>	Low	Unclear	High	High	Unclear
Schuurmans <i>et al</i> <sup>18</sup>	Low	Unclear	High	Low	Low
Schulte <i>et al</i> <sup>13</sup>	High	High	High	High	High
Coughlin <i>et al</i> <sup>19</sup>	Low	Low	Low	Low	Low
Unsworth <i>et al</i> <sup>24</sup>	Low	Low	Unclear	Unclear	High
de Lesquen <i>et al</i> <sup>15</sup>	Unclear	High	Unclear	Unclear	Low
Cataneo <i>et al</i> <sup>20</sup>	Low	Low	Low	Low	Low
de Jong <i>et al</i> <sup>23</sup>	High	Unclear	High	High	Unclear
Slobogean <i>et al</i> <sup>21</sup>	Low	Low	High	Low	Low
Leinicke <i>et al</i> <sup>22</sup>	Low	Low	Low	Low	Low
Girsowicz <i>et al</i> <sup>14</sup>	High	High	High	High	Unclear
NICE <sup>12</sup>	Low	Unclear	Unclear	Low	Low

NICE, National Institute for Health and Care Excellence.

from the authors resulting in SD values ranging from 0.7 to 4.4 and 2.2 to 7.3 in the operative and non-operative groups respectively. There was also a substantial difference in the effect estimate for this trial.<sup>26</sup> In one review,<sup>22</sup> the effect estimate of -10 days, 95% CI -15.41 to -4.59, was 5 days greater than the estimate used from the same trial in other reviews. It is the same as the as length of mechanical ventilation effect estimate reported in the same study<sup>26</sup> so is possibly a transcription error.

Statistical heterogeneity ranged from substantial to none<sup>34</sup> ( $I^2=74.9\%$ ,<sup>22</sup>  $40\%$ ,<sup>18</sup>  $35\%$ <sup>19</sup> and  $0.1\%$ .<sup>21</sup> The narrative syntheses concluded that in patients with flail chest undergoing surgical fixation length of ICU stay was reduced compared with non-operative management.<sup>15 24</sup>

### Multiple rib fractures

A single review<sup>14</sup> included one non-randomised study reporting no statistically significant difference in ICU days ( $p=0.51$ ), the MD and 95% CI was not reported.<sup>33</sup>

### Length of hospital stay (days)

#### Flail chest

Nine reviews<sup>12 15 17-22 24</sup> reported length of hospital stay, six undertook a meta-analysis.<sup>17-22</sup> Two reviews<sup>17 20</sup> pooled the same two trials<sup>25 26</sup> and found a significantly shorter hospital length of stay in favour of surgery compared with non-operative management (MD -11.39 days 95% CI -12.39 to -10.38). When non-randomised studies were included in the meta-analysis the pooled effects were smaller -3.83 days, 95% CI -7.12 to -0.54<sup>22</sup>; -4 days, 95% CI -7.4 to -0.7<sup>21</sup> and -4.48 days, SD 1.9<sup>17</sup> in favour of fixation (table 4).

Heterogeneity ranged from low ( $I^2=0$ ,<sup>18 19</sup> meta-analyses of RCTs only) to moderate or substantial ( $I^2=89\%$ ,<sup>17</sup>  $I^2=68.9\%$ <sup>22</sup> and  $I^2=33\%$ ).<sup>21</sup>

### Multiple rib fractures

Two systematic reviews<sup>14 23</sup> (table 6) included a single non-randomised study<sup>33</sup> reporting no statistically significant difference in hospital stay with surgery (mean 18.8 days [SD 1.8]) compared with the non-operative management (21.1 days [SD 3.9]),  $p=0.59$ ).

### Pneumonia

#### Flail chest

Ten reviews,<sup>12 15 17-22 35</sup> reported the risk of developing pneumonia, six undertook a meta-analysis.<sup>13 17-22 24</sup> Three RCTs<sup>25-27</sup> were pooled in two of the reviews<sup>15 17</sup> and they found a RR of 0.36, 95% CI 0.15 to 0.85, in favour of fixation compared with non-operative management. When non-randomised studies were combined the RR ranged from 0.31, 95% CI 0.21 to 0.41<sup>21</sup> to 0.45, 95% CI 0.29 to 0.70<sup>36</sup> in favour of fixation (table 4). Substantial heterogeneity was seen in meta-analyses for this outcome<sup>18-20</sup> that included the three RCTs<sup>25-27</sup> ( $I^2=66\%$  to  $74\%$ ). In the reviews that pooled the RCTs alongside the non-randomised studies<sup>21 22</sup> there were lower levels of heterogeneity ( $I^2=4\%$  and  $I^2=31\%$ , respectively).

Two narrative syntheses report that among patients with flail chest, risk of pneumonia was reduced in the surgery group compared with the no surgery group (table 5).<sup>15 24</sup>

### Tracheostomy

#### Flail chest

Five reviews reported a meta-analysis for tracheostomy.<sup>17 18 20-22</sup> Pooled RRs ranged from 0.25, 95% CI 0.13 to 0.47 to 0.40, 95% CI 0.2 to 0.7 (table 4). Moderate and substantial heterogeneity was seen in two reviews ( $I^2=42\%$ ,<sup>17</sup>  $I^2=64\%$ ),<sup>20</sup> low in two reviews<sup>21 22</sup> ( $I^2=0\%$ ) and one did not report heterogeneity.<sup>18</sup>

**Table 4** Results of individual reviews that report a meta-analysis for flail chest**Total length of invasive mechanical ventilation (days)**

Studies reporting outcome	No of studies (no of participants in analysis)	Study types		Details of meta-analysis	Results	I <sup>2</sup>
		RCT	NR			
Cataneo <i>et al</i> <sup>20</sup>	3 (123)	3	0	MD (IV, fixed, 95% CI)	Results not pooled	–
Coughlin <i>et al</i> <sup>19</sup>	3 (123)	3	0	MD (IV, random, 95% CI)	–6.30 (–12.16 to –0.43)	95
Leinicke <i>et al</i> <sup>22</sup>	8 (474)	2	6	MD (IV, random 95% CI)	–4.52 (–5.54 to –3.50)	48.6
Schuurmans <i>et al</i> <sup>18</sup>	3 (123)	3	0	MD (IV, random, 95% CI)	–6.53 (–11.88 to –1.18)	93
Slobogean <i>et al</i> <sup>21</sup>	8 Studies (563)	2	6	MD (IV, fixed, 95% CI)	–7.5 (–9.9 to –5.0)	48
Swart <i>et al</i> <sup>17</sup>	18 Studies (1150)	3	15	MD (IV, random, SD)	–4.57 (0.59)	83
<b>Mortality (frequency)</b>						
Cataneo <i>et al</i> <sup>20</sup>	3 (123)	3	0	RR (M-H, fixed, 95% CI)	0.56 (0.13 to 2.42)	0
Coughlin <i>et al</i> <sup>19</sup>	2 (86)	2	0	RR (M-H, random, 95% CI)	0.57 (0.13 to 2.52)	0
Leinicke <i>et al</i> <sup>22</sup>	5 (343)	1	0	RR (95% CI)	0.43 (0.28 to 0.69)	0
Schuurmans <i>et al</i> <sup>18</sup>	2 (86)	2	0	RR (M-H, fixed, 95% CI)	0.56 (0.13 to 2.42)	0
Slobogean <i>et al</i> <sup>21</sup>	7 (582)	2	5	OR (M-H, fixed, 95% CI)	0.31 (0.20 to 0.48)	–
Slobogean <i>et al</i> <sup>21</sup>	7 (582)	2	5	RR (M-H, fixed, 95% CI)	0.19 (0.13 to 0.26)	0
Swart <i>et al</i> <sup>17</sup>	13 (1263)	3	10	RR (M-H, random, SD)	0.44 (0.09)	0
<b>Total length of stay in intensive care unit (days)</b>						
Cataneo <i>et al</i> <sup>20</sup>	2 (77)	2	0	MD (IV, fixed, 95% CI)	Results not pooled	–
Coughlin <i>et al</i> <sup>19</sup>	3 (123)	3	0	MD (IV, random, 95% CI)	–6.46 (–9.73 to –3.19)	35
Leinicke <i>et al</i> <sup>22</sup>	5 (235)	2	3	MD (IV, random, 95% CI)	–3.4 (–6.01 to –0.80)	74.9
Schuurmans <i>et al</i> <sup>18</sup>	3 (123)	3	0	MD (IV, fixed, 95% CI)	–5.18 (–6.17 to –4.19)	40
Slobogean <i>et al</i> <sup>21</sup>	4 (261)	2	2	MD (IV, fixed, 95% CI)	–4.8 (–7.9 to –1.6)	0.1
Swart <i>et al</i> <sup>17</sup>	14 (840)	3	11	MD (IV, random, SD)	–3.25 (1.29)	91
<b>Total length of stay in hospital (days)</b>						
Coughlin <i>et al</i> <sup>19</sup>	2 (86)	2	0	MD (IV, random, 95% CI)	–11.39 (–12.39 to –10.38)	0
Leinicke <i>et al</i> <sup>22</sup>	5 (262)	1	4	MD (IV, random 95% CI)	–3.83 (–7.12 to –0.54)	68.9
Schuurmans <i>et al</i> <sup>18</sup>	2 (86)	2	0	MD (IV, fixed, 95% CI)	–11.39 (–12.39 to –10.38)	0
Slobogean <i>et al</i> <sup>21</sup>	4 (404)	1	3	MD (IV, fixed, 95% CI)	–4.0 (–7.4 to –0.7)	33
Swart <i>et al</i> <sup>17</sup>	11(438)	1	10	MD (IV, random, SD)	–4.48 (1.98)	89
<b>Pneumonia (frequency)</b>						
Cataneo <i>et al</i> <sup>20</sup>	3 (123)	3	0	RR (M-H, random, 95% CI)	0.36 (0.15 to 0.85)	66
Coughlin <i>et al</i> <sup>19</sup>	3 (123)	3	0	RR (M-H, random, 95% CI)	0.36 (0.15 to 0.85)	66
Leinicke <i>et al</i> <sup>22</sup>	4 (260)	1	3	RR (95% CI)	0.43 (0.28 to 0.69)	31
Schuurmans <i>et al</i> <sup>18</sup>	2 (83)	2	0	RR (M-H, fixed, 95% CI)	0.45 (0.29 to 0.7)	74

Continued

**Table 4** Continued

**Total length of invasive mechanical ventilation (days)**

Studies reporting outcome	No of studies (no of participants in analysis)	Study types		Details of meta-analysis	Results	I <sup>2</sup>
		RCT	NR			
Slobogean <i>et al</i> <sup>21</sup>	8 (816)	2	6	OR (M-H, fixed, 95% CI)	0.18 (0.11 to 0.32)	4
Slobogean <i>et al</i> <sup>21</sup>	8 (816)	2	6	RR (M-H, fixed, 95% CI)	0.31 (0.21 to 0.41)	4
Swart <i>et al</i> <sup>17</sup>	15 (1005)	3	12	RR (M-H, random, SD)	0.59 (0.10)	55
Tracheostomy (frequency)						
Cataneo <i>et al</i> <sup>20</sup>	2 (83)	2	0	RR (M-H, random, 95% CI)	0.38 (0.14 to 1.02)	64
Leinicke <i>et al</i> <sup>22</sup>	4 (215)	1	3	RR (95% CI)	0.25 (0.13 to 0.47)	0
Schuurmans <i>et al</i> <sup>18</sup>	2 (83)	2	0	RR (M-H, fixed, 95% CI)	0.4 (0.2 to 0.7)	Not reported
Slobogean <i>et al</i> <sup>21</sup>	3 (165)	1	2	OR (M-H, fixed, 95% CI)	0.12 (0.04 to 0.32)	0
Slobogean <i>et al</i> <sup>21</sup>	3 (165)	1	2	RR (M-H, fixed, 95% CI)	0.34 (0.10 to 0.57)	0
Swart <i>et al</i> <sup>17</sup>	11 (975)	2	9	RR (M-H, random, SD)	0.52 (0.07)	42
Sepsis (frequency)						
Slobogean <i>et al</i> <sup>21</sup>	4 (345)	0	4	OR (M-H, fixed, 95% CI)	0.36 (0.19 to 0.71)	0
Slobogean <i>et al</i> <sup>21</sup>	4 (345)	0	4	RR (M-H, fixed, 95% CI)	0.14 (0.56 to 0.23)	0
Spirometry (percentage of predicated)						
Coughlin <i>et al</i> <sup>19</sup>	–	–	–	–	–	–
FVC	2 (74)	2	0	MD (IV, random, 95% CI) p value	1.53 (–13.49 to 16.55) p=0.84	Not reported
FEV1	2 (74)	2	0	MD (IV, random, 95% CI) p value	–0.42 (–4.83 to 3.98) p=0.85	Not reported
TLC	2 (74)	2	0	MD (IV, random, 95% CI) p value	3.69 (–3.08 to 10.46) p=0.29	Not reported
PEFR	2 (74)	2	0	MD (IV, random, 95% CI) p value	0.38 (–0.76 to 1.53) p=0.51	Not reported
Chest deformity (frequency)						
Cataneo <i>et al</i> <sup>20</sup>	2 (86)	2	0	RR (M-H, fixed, 95% CI)	0.13 (0.03 to 0.67)	0
Slobogean <i>et al</i> <sup>21</sup>	4 (228)	1	3	OR (M-H, fixed, 95% CI)	0.11 (0.02 to 0.60)	2.1
Slobogean <i>et al</i> <sup>21</sup>	4 (228)	1	3	RR (M-H, fixed, 95% CI)	0.30 (0.00 to 0.60)	2.1
Dyspnoea (frequency)						
Slobogean <i>et al</i> <sup>21</sup>	3 (135)	1	2	OR (M-H, fixed, 95% CI)	0.40 (0.16 to 1.01)	0
Slobogean <i>et al</i> <sup>21</sup>	3 (135)	1	2	RR (M-H, fixed, 95% CI)	0.15 (0.09 to 0.39)	0
Chest pain (frequency)						
Slobogean <i>et al</i> <sup>21</sup>	2 (71)	1	1	OR (M-H, fixed, 95% CI)	0.40 (0.01 to 12.60)	0
Slobogean <i>et al</i> <sup>21</sup>	2 (71)	1	1	RR (M-H, fixed, 95% CI)	0.18 (–0.46 to 0.83)	0

FEV1, forced expiratory volume; FVC, force vital capacity; IV, inverse variance; M-H, Mantel-Haenszel; MD, mean difference; NR, non-randomised study; PEFR, peak expiratory flow rate; RCT, randomised controlled trial; RR, risk ratio; TLC, total lung capacity.

**Table 5** Results of individual reviews that report a narrative synthesis for flail chest

Study details	Included studies	Outcomes assessed	Narrative synthesis
<b>Author</b> Unsworth <i>et al</i> <sup>24</sup> <b>Year</b> 2015 <b>Country</b> Australia	RCT=2 Granetzny (40) Tanaka (37) Marasco (46) Non-randomised=6 Ahmed (64) Althausen (50) Doben (21) De Moya (48) Nirula (60) Voggenreiter (42) Total number of patients=408	<ul style="list-style-type: none"> <li>▶ Mortality</li> <li>▶ Pneumonia</li> <li>▶ Pneumothorax and haemothorax</li> <li>▶ Hospital length of stay</li> <li>▶ ICU stay</li> <li>▶ Costings</li> <li>▶ Treatment outcome</li> </ul>	<ul style="list-style-type: none"> <li>▶ Significant decrease in mechanical ventilation requirements after surgical fixation. decreasing in ventilator-acquired pneumonia after surgical fixation.</li> <li>▶ Decrease in ICU-LOS, fewer days of mechanical ventilation and cost savings compared with non-operative management.</li> <li>▶ Decreased days of ventilator dependence and shorter ICU-LOS.</li> <li>▶ Lower incidence of pneumonia, a higher return to full time work at 6 months.</li> <li>▶ Less persistent pain at six and 12 months in those receiving surgery.</li> <li>▶ Significantly fewer days of mechanical ventilation and a shorter hospital and ICU-LOS.</li> <li>▶ The estimated cost savings ranged from US\$10 000 to \$A14 443 per patient with surgical rib fixation as a result of the decrease in ICU-LOS.</li> <li>▶ None of the studies were large enough to draw conclusions on the effect of this intervention on thromboembolism and death.</li> </ul>
<b>Author</b> de Lesquen <i>et al</i> <sup>15</sup> <b>Year</b> 2015 <b>Country</b> France	Meta-analysis=2 Leinicke nine studies (538 patients) Slobogean 11 studies (732 patients) RCT=3 Marasco (46) Granetzny (40) Tanaka (37) Non-randomised=6 Ahmed (64) Karev (40) Voggenreiter (20) Balci (64) Nirula (60) Althausen (50) Total number of patients=421	<ul style="list-style-type: none"> <li>▶ Duration of IMV</li> <li>▶ LOS-ICU</li> <li>▶ Pneumonia</li> <li>▶ Mortality</li> </ul>	For flail chest, early surgical stabilisation can be considered in patients who would require mechanical ventilation for >48 hours. (Grade B, extrapolated recommendations from level I evidences.)

Continued

**Table 5** Continued

Study details	Included studies	Outcomes assessed	Narrative synthesis
<b>Author</b> NICE <sup>12</sup> <b>Year</b> 2010 <b>Country</b> UK	RCT=1 Tanaka (37) Non-randomised=2 Voggenreiter (42) Paris (29) Case Series=4 Lardinois (66) Mouton (23) Menard (18) Hellberg (10) Total number of patients=225 Intervention group=173 Control group=52	<ul style="list-style-type: none"> <li>▶ Duration of IMV</li> <li>▶ Mortality</li> <li>▶ LOS ICU</li> <li>▶ Pneumonia</li> <li>▶ Lung function</li> <li>▶ Return to Employment</li> <li>▶ Sepsis</li> <li>▶ Pain or discomfort requiring removal of plates</li> </ul>	Surgical stabilisation with metal rib reinforcements aims to allow earlier weaning from mechanical ventilation, reduce acute complications and avoid chronic pain sometimes associated with permanent malformation of the chest wall. Kirschner wire may be used on its own, but this method of rib stabilisation is not covered by this guidance.
<b>Author</b> Schulte <i>et al</i> <sup>13</sup> <b>Year</b> 2016 <b>Country</b> UK	Systematic review=1 Slobogean (753) RCT=1 Marasco (23,23) Non-randomised studies=9 Jayle (10,10) Pieracci (35,35) Zhang (24,15) Wada (84,336) Granhed (60,153) Doben (10,11) Xu (17,15) Althausen (22,28) De Moya (16,32) Total number of patients=1712 Intervention group=301 Control group=658	<ul style="list-style-type: none"> <li>▶ Duration of IMV</li> <li>▶ Mortality</li> <li>▶ LOS hospital</li> <li>▶ LOS-ICU</li> <li>▶ Pneumonia</li> </ul>	Surgical stabilisation of flail chest in thoracic trauma patients has beneficial effects with respect to reduced ventilatory support, shorter intensive care and hospital stay, reduced incidence of pneumonia and septicaemia, decreased risk of chest deformity and an overall reduced mortality when compared with patients who received non-operative management.

ICU, intensive care unit; IMV, invasive mechanical ventilation; LOS, length of stay; NICE, National Institute for Health and Care Excellence; RCT, randomised controlled trial.

**Table 6** Results of individual reviews that report a narrative synthesis for multiple rib fractures

Study details	Included studies	Outcomes assessed	Narrative synthesis
<b>Author</b> de Jong <i>et al</i> <sup>23</sup> <b>Year</b> 2014 <b>Country</b> The Netherlands	RCT=0 Non-randomised=1 Nirula (60) Case series=2 Campbell (32) Mayberry (46, 15 non-flail) Total number of patients=138 Intervention group=108 Control group=30	<ul style="list-style-type: none"> <li>▶ LOS hospital</li> <li>▶ Duration of IMV</li> <li>▶ Time of operation</li> <li>▶ Chronic pain</li> </ul>	Only Nirula <i>et al</i> <sup>33</sup> concluded that rib fracture fixation showed a trend toward fewer total ventilator days. Mayberry <i>et al</i> <sup>41</sup> investigated the quality of life after rib fixation, and they concluded that there was low long-term morbidity and pain. Campbell <i>et al</i> <sup>40</sup> demonstrated low levels of pain and satisfactory rehabilitation.
<b>Author</b> Girsowicz <i>et al</i> <sup>14</sup> <b>Year</b> 2012 <b>Country</b> France	Non-systematic review=1 Nirula and Mayberry Case Comparator=1 Nirula (30,30) Case Series=4 Mayberry (46) Richardson (7) Barajas (22) Campbell (32) Case report=3 Gasparri (1) Cacchione (1) Kerr-Valentic (1) Total number of patients=169 Intervention group=139 Control group=30	<ul style="list-style-type: none"> <li>▶ Pain</li> <li>▶ Disability</li> <li>▶ Respiratory function</li> <li>▶ Number of days lost from work</li> </ul>	In general, of the nine studies presented, all indicated that surgical stabilisation in the management of isolated multiple non-flail and painful rib fractures improved outcomes. Indeed, the interest and benefit was shown not only in terms of pain and respiratory function but also in improved quality of life and reduced socio-professional disability. Hence, the current evidence shows surgical stabilisation to be safe and effective in alleviating post-operative pain and improving patient recovery, thus enhancing the outcome of the procedure. However, retrieved studies provided a low level of evidence (small studies with few numbers of patients and short-term follow-up or case reports). Large prospective controlled trials are thus necessary to confirm these encouraging results.

IMV, invasive mechanical ventilation; LOS, length of stay; RCT, randomised controlled trial.

## Sepsis

### Flail chest

One review,<sup>21</sup> pooling four non-randomised studies<sup>28 30 37 38</sup> estimated a RR of 0.14, 95% CI 0.56 to 0.23 with  $I^2=0\%$  in favour of fixation compared with non-operative management for sepsis. The estimate RR reported is not possible given the CI does not include the estimated value, 0.14. The lower interval of 0.56 could possibly be  $-0.56$  creating a wider CI and would suggest that the author's conclusion was correct and there was a statistically significant difference in favour of fixation.

## Spirometry

### Flail chest

One review<sup>19</sup> reported a meta-analysis of spirometry data which included two RCTs<sup>26 27</sup> with spirometry measured at two different time points (three and 2 months respectively). No statistically significant differences in any spirometry data were seen between surgery and no surgery (table 4).

## Chest deformity

### Flail chest

Two reviews reported a meta-analysis of chest deformity,<sup>20 21</sup> both reported a statistically significant difference in favour of surgery compared with no surgery (RR

0.30, 95% CI 0.00 to 0.60,  $I^2=2.1\%$  and RR 0.13, 95% CI 0.03 to 0.67,  $I^2=0\%$ ).

## Dyspnoea

### Flail chest

One review<sup>21</sup> pooled an RCT<sup>25</sup> and two non-randomised studies.<sup>37 39</sup> for dyspnoea (RR 0.15, 95% CI 0.09 to 0.39 in favour of fixation). Duration of follow-up was 1 year for two of the primary studies<sup>25 39</sup> and unclear in the third.<sup>37</sup> It was unclear how dyspnoea was measured or defined in the three primary studies.

## Chest pain

### Flail chest

Chest pain was reported in one review<sup>21</sup> which pooled one RCT<sup>25</sup> and one non-randomised study<sup>39</sup> suggesting a benefit in favour of fixation (RR 0.18, CI 95% -0.46 to 0.83).

## Other reported outcomes

Several other outcomes were reported within the systematic reviews however no others have been pooled in a meta-analysis. A narrative synthesis was not completed on the outcomes: wound infection, pain-requiring removal of metalwork, return to work, socio-professional disability cost, pulmonary embolism, pneumothorax and haemothorax. In

the reviews, data on these additional outcomes was minimal and presented as a narrative synthesis without presenting numerical data (tables 5 and 6).

## DISCUSSION

Twelve systematic reviews on the effectiveness of surgery for flail chest and multiple rib fractures published between 2010 and 2016. This is the first systematic review of reviews and highlighted that there are a large number of reviews with same aims and including the same primary studies.

### Flail chest

Six<sup>17–22</sup> of the 12 systematic reviews presented meta-analyses for flail chest based on overlapping primary studies. They reported reductions in length of mechanical ventilation, length of stay, pneumonia and tracheostomy rates with surgery compared with non-surgical management and inconsistent results for mortality. Across many of the meta-analyses there was moderate to high levels of heterogeneity and variation in the effect estimates.

A single systematic review found reductions in sepsis, dyspnoea, chest deformity and chest pain with surgery compared with no surgery management. Nevertheless, as the outcome measures were not defined it is difficult to know whether the reductions are clinically significant. Reporting of adverse outcomes was infrequent across the reviews, which could reflect lack of measurement and/or reporting of adverse events in the primary studies or the systematic reviews. Therefore, the benefits of surgery could be overestimated in light of the potential risks not being considered. Synthesising multiple meta-analyses data that include overlapping primary studies has the potential to overestimate the strength of the findings therefore it is important to be mindful of the limited evidence on which our conclusions are based. In addition, significant heterogeneity for several of the outcomes that were pooled makes drawing firm conclusions difficult.

### Multiple rib fractures

Evidence in support of multiple rib fracture fixation in the absence of flail chest is limited. Two systematic reviews<sup>14 23</sup> reported on one non-randomised study<sup>33</sup> that recruited between 1996 and 2000, four case series<sup>40–43</sup> and two case reports.<sup>44 45</sup> Hence, due to limited evidence no conclusive statements on effectiveness can be drawn.

### Review quality

A significant amount of effort and time is required to conduct a high quality systematic review and should only be undertaken when there is sufficient cause<sup>46 47</sup> (eg, to incorporate the findings of a new RCT or to address an evidence gap). Eight of the systematic reviews were published within 18 months although none were registered on PROSPERO<sup>48</sup> so it is possible the authors were unaware of each other's research. Registering reviews allows transparency of methods and also reduces research

waste.<sup>49</sup> As similar search strategies and search dates were used in each systematic review, inevitably many of the included studies were the same across reviews.

Only two of the 12 systematic reviews formally appraised the quality of the included studies, therefore 10 of the reviews were not in a position to fully consider the impact of risk of bias on their conclusions. High or unclear risk of bias within reviews have affected the conclusions drawn from this evidence synthesis. In a systematic review of 106 emergency surgery systematic reviews, a low risk of bias was found in 53.8%, identifying a common problem of poor quality reviews conducted in emergency surgery.<sup>50</sup>

### Heterogeneity and meta-analysis errors

The  $I^2$  value describes the percentage of total variation across studies that is due to heterogeneity rather than chance.<sup>51</sup> Examining the meta-analyses including RCTs highlights moderate to high levels of statistical heterogeneity.

There was also clinical variation in the primary studies in terms of indications and timing of surgery and it is possible that these between study differences could be a source of the substantial heterogeneity. For example, in one RCT<sup>25</sup> patients were randomised after 5 days of invasive ventilation, whereas another RCT<sup>26</sup> randomised and fixed within 24 to 72 hours regardless of initial intubation state. Also, many reviews define the comparator as usual care or non-operative care but do not elaborate on what encompasses this care. Differences in how outcomes were measured may also have contributed to between study heterogeneity. It was unknown due to lack of reporting whether the outcomes were equivalent in the pooled primary studies or overall between systematic reviews.

In all systematic reviews with meta-analyses, they reported that two reviewers were involved in the data extraction to minimise errors.<sup>17 19–22</sup> Despite attempts to minimise errors and therefore an apparent low risk of bias, some errors (up to an MD of 10 days in the measurement of length of intensive care stay) were identified across reviews. It is worth noting that there were no significant changes in the conclusions drawn from these analyses. Although there was substantial statistical and clinical heterogeneity and lack of consideration of risk of bias in many of the reviews, conclusions tended to be similar and in the direction of benefit with fixation suggesting that further high quality RCTs investigating the effectiveness (including adverse effects) of internal surgical fixation over non-operative management are warranted.

### Strengths

Multiple databases were searched for studies and study selection was undertaken by two researchers, reducing the risk of error and bias. Although only English language studies were included, some sources of unpublished studies were searched. A mapping of the studies included in the reviews was undertaken to take into account individual studies being included in multiple reviews and hence double counting studies.



## Limitations

All systematic reviews were included irrespective of their risk of bias scoring. It could be argued that several reviews were stretching the traditional definition of a systematic review however they did hold to the protocol definition with an electronic database search strategy and included primary evidence. Due to best evidence topics and rapid evidence synthesis being included it was then difficult to apply the ROBIS tool consistently. The ROBIS tool is not designed for rapid evidence synthesis and therefore this type of review showed high risk of bias as they were being assessed against a tool designed for full systematic reviews. Rapid evidence syntheses, by their nature address a trade-off between time and methodological rigour and comprehensiveness.<sup>52</sup>

## CONCLUSION

The considerable duplication of work across reviews could be mitigated through protocol registration and greater attention to establishing whether a review is necessary by scoping the literature before commencing a new review. Despite this review identifying 12 systematic reviews they only included 37 unique primary studies, only three of which were RCTs. Synthesis of the reviews has shown some potential improvement in patient outcomes with flail chest after surgical intervention. However, there were differences in indications and timing of interventions in the primary studies and moderate to high levels of heterogeneity across reviews. For future review updates, meta-analysis for effectiveness may need to take into account indications and timing of surgery as a subgroup analysis to address clinical heterogeneity between primary studies. Further robust evidence is required before conclusions can be drawn of the effectiveness of surgical fixation for flail chest and in particular, multiple rib fractures.

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