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Risk factors for postoperative pneumonia in patients undergoing hip fracture surgery: a systematic review and meta-analysis

Seung-Beom Han¹, Sang-Bum Kim² and Kyun-Ho Shin^{3*}

Abstract

Background: Postoperative pneumonia (POP) is a devastating complication that can frequently occur after hip fracture surgery. This study aimed to quantitatively and comprehensively summarize the risk factors for POP following hip fracture surgery.

Methods: PubMed, Embase, and Cochrane Library were systematically searched for studies assessing risk factors for POP following hip fracture surgery. The pooled odds ratio (OR) and standardized mean difference (SMD) between patients with and without POP were calculated. Evidence was assessed using the Newcastle–Ottawa scale.

Results: Ten studies including 37,130 patients with hip fractures were selected. POP occurred in 1768 cases with an accumulated incidence of 7.8% (95% confidence interval [CI]: 0.061–0.094). Advanced age (SMD: 0.50, 95% CI: 0.10–0.90), male sex (OR: 1.50, 95% CI: 1.12–2.01), American Society of Anesthesiologists physical status scale ≥ 3 (OR: 3.17, 95% CI: 1.25–8.05), chronic obstructive pulmonary disease (OR: 2.05, 95% CI: 1.43–2.94), coronary heart disease (OR: 1.82, 95% CI: 1.27–2.60), arrhythmia (OR: 1.49, 95% CI: 1.04–2.15), congestive heart failure (OR: 1.41, 95% CI: 1.14–1.75), chronic kidney disease (OR: 2.09, 95% CI: 1.28–3.41), and cerebrovascular accident (OR: 2.14, 95% CI: 1.60–2.85) were risk factors for POP. Hemoglobin (SMD: -0.14, 95% CI: -0.25 to -0.03), albumin (SMD: -0.97, 95% CI: -1.54–0.41), blood urea nitrogen (SMD: 0.20, 95% CI: 0.03–0.37), alanine aminotransferase (SMD: 0.27, 95% CI: 0.10–0.44), arterial oxygen pressure (SMD: -0.49, 95% CI: -0.71–0.27), time from injury to surgery (SMD: 0.13, 95% CI: 0.08–0.17), and surgery within 48 h (OR: 3.74, 95% CI: 2.40–5.85) were associated with the development of POP.

Conclusion: Patients with the aforementioned risk factors should be identified preoperatively, and related prophylaxis strategies should be implemented to prevent POP following hip fracture surgery.

Keywords: Pneumonia, Hip fracture, Hip surgery, Postoperative complications, Predictors, Systematic reviews, Meta-analysis

Background

Hip fractures are a major health problem and the number of hip fractures is expected to increase by approximately 2% annually over the next 30 years [1]. Hip fractures are

associated with increased risk of morbidity and mortality [1–3]. Furthermore, the coronavirus disease pandemic, has forced an unprecedented period of challenge for the management of patients with hip fractures [4].

Postoperative pneumonia (POP) is a devastating complication that can occur after hip fracture surgery [5, 6]. However, few studies have been performed to elucidate this complication and investigate patients with hip fracture and POP. The incidence of hip fracture-related pneumonia has been reported to range from 4 to 15% [7–9].

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Evidence has shown that POP is associated with various predisposing factors, including older age, male sex, multiple medical comorbidities, and hypoalbuminemia [7–14].

With the progress in medical technologies and after-care of patients, clinicians are increasingly focusing on the prevention and treatment of POP. To medically optimize patients and provide better perioperative care, identifying various potential risk factors is important for POP. To the best of our knowledge, no formal systematic review and meta-analysis has investigated and summarized the risk factors for POP following hip fracture surgery. Therefore, this meta-analysis aimed to summarize the risk factors for the development of POP in patients undergoing hip fracture surgery. The results of this study are potentially beneficial for clinicians to identify high-risk patients and help prevent postoperative POP following hip fracture surgery.

Methods

This study followed the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines [15]. Patient consent and ethical approval were not required because this study was a meta-analysis of published studies. Two authors (KHS and SBH) independently searched and reviewed the literature, assessed the quality, and extracted data. Disagreements were resolved through discussions or negotiations with a third independent author (SBK). Inter-reviewer reliability was assessed by study screening and selection, quality assessment, data extraction, and result pooling using the kappa statistic (κ). The κ value for the data extraction ranged from 0.88 to 1.00.

Search strategy

MEDLINE/PubMed, Cochrane Central Register of Controlled Trials, and EMBASE were exhaustively searched to identify original studies that included patients with hip fracture with POP published before January 4, 2022. The search terms, Medical Subject Headings terms, and their combinations searched in the title/abstract field of the search engines were as follows: “hip,” “fracture,” “hip fractures,” “pneumonia,” “lower respiratory tract infection,” “pulmonary infection,” “factor,” “risk,” and “predictor.” No other restrictions, including language, were applied. The references of the selected articles were also reviewed to identify relevant articles.

Eligibility criteria and study selection

Two independent authors (KHS and SBH) screened all the titles and abstracts. Initially selected articles were further reviewed for inclusion according to the following inclusion criteria: (1) Cohort and case-control

studies if they reported analyses of the predictors of POP in patients undergoing hip fracture surgery. (2) POP occurred after hip fracture surgery and recurred patients without pneumonia at baseline, (3) comparison between patients with POP as the case group and patients without POP as the control group; (4) accessible full-text articles; and (5) studies reporting sufficient information to extract and calculate relevant standardized mean difference (SMD) or odds ratio (OR) with 95% confidence interval (CI). The specific reasons for the excluded articles are shown in Fig. 1.

Data extraction

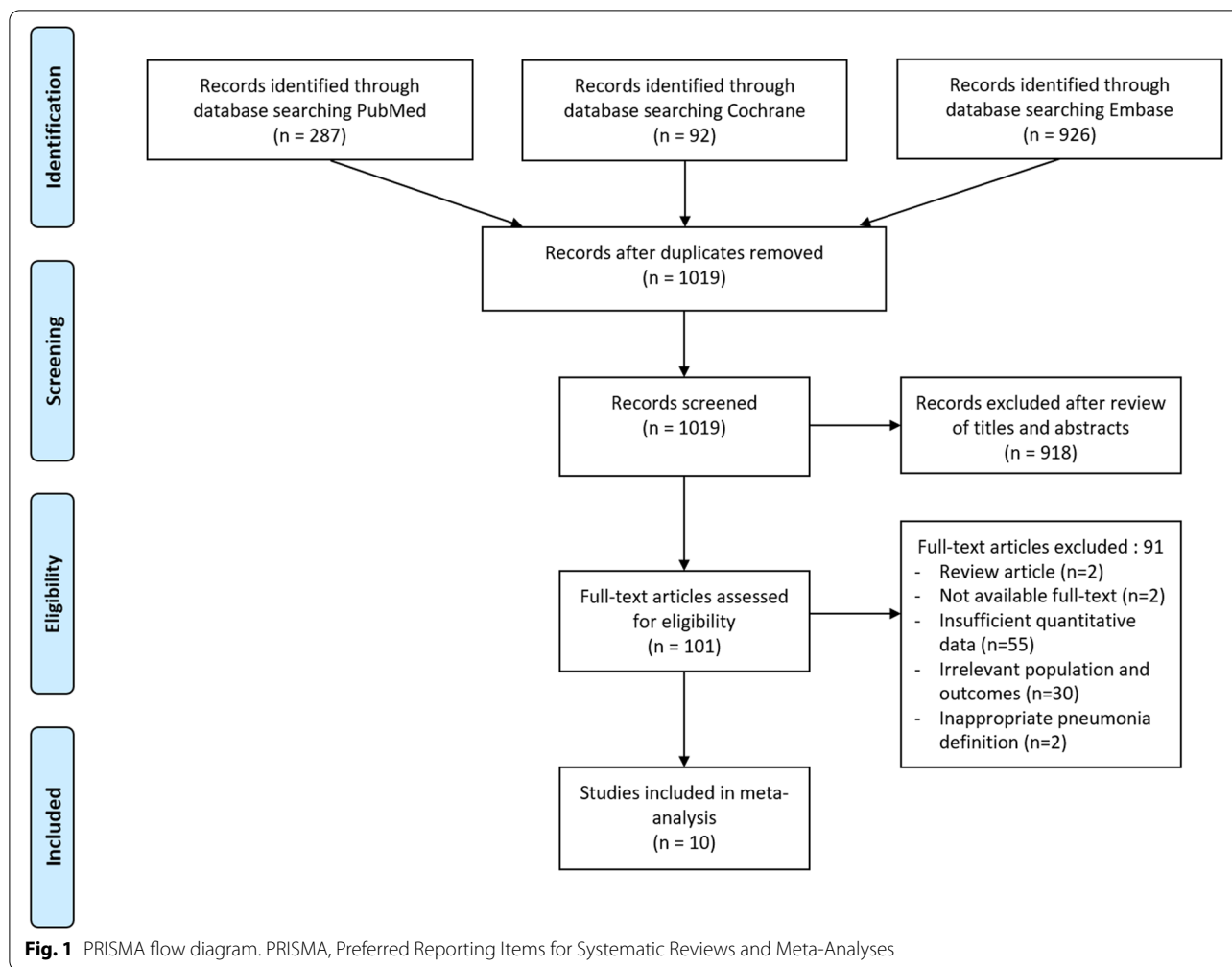
Two independent authors extracted data from the eligible studies (KHS and SBH). Disagreements were resolved through discussion and consensus with the third author (SBK). Data were extracted according to the following descriptive information: (1) study characteristics, including the name of the first author, year of publication, study country, and study design; (2) patient demographics, such as the number of patients with or without POP, age, sex, and the incidence of POP; (3) significant risk factors for POP; and (4) number of citations for each potential risk factor for POP after hip fracture surgery.

Quality assessment

The methodological quality of each included study was evaluated using the Newcastle–Ottawa scale (NOS) [16]. The scale includes selection, comparability, and outcome domains. The selection domain has four categories; comparability domain, two categories; and outcome domain, three categories. A study was awarded a maximum of one star for each category in the selection and outcome domains. A maximum of two stars was assigned to the comparability domain.

Statistical analyses

ORs or SMDs with corresponding 95% CIs were estimated and pooled across studies to assess the association between POP and various potential risk factors. A meta-analysis was performed for each factor ($n \geq 2$), which was presented as an effect size of the 95% CI. The adjusted data were used maximally when available. The inconsistency index (I^2) was determined, and a χ^2 -based test of homogeneity was performed. If I^2 was $< 50\%$, the fixed-effects model (Mantel–Haenszel method) was used due to low heterogeneity. $I^2 \geq 50\%$ was considered a significant heterogeneity. The random-effects model (DerSimonian–Laird method) was used, and a “leave-one-out” sensitivity analysis was performed by sequentially deleting one study to determine the source of heterogeneity [17]. After excluding each study, an analysis was performed to determine the existence of heterogeneity.



When 10 or more studies were included, a small study publication bias was assessed using funnel plot analysis. The significance level was set at $p < 0.05$. All statistical analyses were performed using RStudio v.1.0.143 (RStudio Inc., Boston, MA, USA).

Results

Search results

Figure 1 shows a detailed summary of the study’s identification and selection process. A total of 1305 articles were identified after the initial search. After eliminating 286 duplicates and 918 ineligible articles based on titles and abstracts, the full text of 101 articles were reviewed. After excluding 91 articles without information of inclusion criteria, 10 articles [7–14, 18, 19] were finally selected for the meta-analysis.

Study characteristics

The characteristics of the included studies of POP are shown in Table 1. All studies were published in English

and from 2016 onwards. All the included studies were retrospective cohort studies. A total of 1768 patients with hip fracture had POP and 35,362 patients without POP. The risk factors of POP reported in individual studies are summarized in Table 1.

Risk of bias analysis

The risk of bias assessment of the included studies is summarized in Table 2. The NOS scores of the selected studies ranged from 8 to 9. Methods of the cohort selection and outcome assessment were clearly stated in all studies. Most studies excluded persons with pneumonia preoperatively. Most studies accounted for confounding factors using standard statistical regression techniques.

Meta-analysis results

Crude accumulated incidence of POP was 4.8% (1768/35,362) with an accumulated incidence of 7.8% (95% CI: 0.061–0.094; $I^2 = 94\%$). Heterogeneity could

Table 1 Characteristics of the included studies

First author (year)	Country	Study design	Sample size (n)		Age (years)		Male sex (%)		Significant factors		
			Total	Pneumonia	No pneumonia	Pneumonia	No pneumonia	Pneumonia		No pneumonia	Incidence (%)
Lv et al. 2016 [8]	China	Retrospective cohort study	1429	70	1359	Median 82	Median 74	22 (31.4)	575 (42.3)	70/1429 (4.9)	Age, male sex, fracture type, number of comorbidities, ASA ≥ 3, surgical type, preoperative hypoalbuminemia, high Cr, high RDW, preoperative mechanical ventilation
Bohl et al. 2018 [7]	USA	Retrospective cohort study	29,377	1191	28,186	NA	NA	NA	NA	1911/28,186 (4.1)	Age, male sex, COPD, low BMI, CHF, dyspnea on exertion, functional status, anemia
Chang et al. 2018 [10]	China	Retrospective cohort study	240	25	215	NA	NA	9 (36.0)	68 (31.6)	15/240 (6.3)	Age, CVA, cancer, low platelet, high blood glucose
Wang et al. 2019 [12]	China	Retrospective cohort study	720	54	666	82.3	77.5	20 (37.0)	27 (41.0)	54/720 (7.5)	COPD, CVA, preoperative hypoalbuminemia, time from injury to surgery
Salarbaks et al. 2020 [9]	Netherlands	Retrospective cohort study	407	62	345	Median 84	Median 83	29 (46.8)	98 (28.4)	62/407 (15.2)	Male sex, COPD
Shin et al. 2020 [11]	South Korea	Retrospective cohort study	1155	59	1096	83.1	77.9	21 (35.6)	295 (26.9)	59/1155 (5.1)	Age, cardiovascular disease, early postoperative hypoalbuminemia
Wang et al. 2020 [19]	China	Retrospective cohort study	293	33	260	84.5	85.1	20 (60.6)	76 (29.2)	33/293 (11.3)	Male sex, smoking, preoperative hypoalbuminemia, low arterial oxygen saturation
Xiang et al. 2020 [13]	China	Retrospective cohort study	1113	166	947	86.4	78.8	53 (31.9)	331 (35.0)	166/1113 (14.9)	Low BMI, preoperative hypoalbuminemia, high CRP, functional status, time from injury to surgery

Table 1 (continued)

First author (year)	Country	Study design	Sample size (n)		Age (years)		Male sex (%)		Incidence (%)	Significant factors
			Total	No pneumonia	Pneumonia	No pneumonia	Pneumonia	No pneumonia		
Zhao et al. 2020 [14]	China	Retrospective cohort study	1495	1442	NA	NA	28 (52.8)	483 (33.5)	53/1495 (3.5)	Age, male sex, chronic respiratory disease, liver disease, urinary tract infection, high CK-MB, high BNP, high D-dimer
Ji et al. 2021 [18]	China	Retrospective cohort study	901	846	81.6	78.5	23 (41.8)	280 (33.1)	55/901 (6.1)	Age, COPD, CVA, hypoxemia, time from injury to surgery

ASA American Society of Anesthesiologists physical status, Cr Creatinine, RDW Red blood cell distribution width, COPD Chronic obstructive pulmonary disease, BMI Body mass index, CHF Congestive heart failure, CVA Cerebrovascular accident, CRP C-reactive protein, CK-MB creatine kinase MB, BNP B-type natriuretic peptide

Table 2 Quality assessment of included studies

First author (year)	Selection				Comparability		Outcomes		
	Representativeness of the exposed cohort	Selection of the nonexposed cohort	Ascertainment of exposure	Demonstration that the outcome of interest was not present at the start of the study	Controlled for age and comorbidities	Controlled for any additional factors	Assessment of outcomes	Sufficient follow-up	Adequacy of follow-up
Lv et al. 2016 [8]	★	★	★	★	★	★	★	★	★
Bohl et al. 2018 [7]	★	★	★	★	★	★	★	★	★
Chang et al. 2018 [10]	★	★	★		★	★	★	★	★
Wang et al. 2019 [12]	★	★	★	★	★	★	★	★	★
Sal- arbaks et al. 2020 [9]	★	★	★	★	★	★	★		★
Shin et al. 2020 [11]	★	★	★	★	★	★	★		★
Wang et al. 2020 [19]	★	★	★	★	★	★	★		★
Xiang et al. 2020 [13]	★	★	★	★	★	★	★		★
Zhao et al. 2020 [14]	★	★	★	★	★	★	★		★
Ji et al. 2021 [18]	★	★	★	★	★	★	★	★	★

not be resolved using sensitivity analyses. Potential risk factors were classified into four categories: basic demographic predictors, medical comorbidity predictors, surgical characteristic predictors, and baseline laboratory predictors. Detailed results for each factor are presented in Tables 3 and 4.

Basic demographic predictors

Advanced age (SMD: 0.50; 95% CI: 0.108–0.90; $p=0.01$; $I^2=90\%$), male sex (OR: 1.50; 95% CI: 1.12–2.01; $p<0.01$; $I^2=72\%$), and the American Society of Anesthesiologists physical status (ASA) scale ≥ 3 (OR: 3.17; 95% CI: 1.25–8.05; $p=0.02$; $I^2=90\%$) were significantly associated with a high risk of POP (Table 3). Significant heterogeneity was found for the pooled results of advanced age, male sex,

body mass index, and ASA scale. After sensitivity analyses, heterogeneity was resolved, and the significance did not change (Additional file 1). A funnel plot of sex was symmetrical and suggested a low risk of publication bias (Fig. 2).

Medical comorbidity predictors

Patients with anemia (OR: 1.55; 95% CI: 1.16–2.08; $p<0.01$; $I^2=85\%$), chronic obstructive pulmonary disease (COPD) (OR: 2.05; 95% CI: 1.43–2.94; $p<0.01$; $I^2=52\%$), coronary heart disease (OR: 1.82; 95% CI: 1.27–2.60; $p<0.01$; $I^2=56\%$), arrhythmia (OR: 1.49; 95% CI: 1.04–2.15; $p=0.03$; $I^2=0\%$), congestive heart failure (OR: 1.41; 95% CI: 1.14–1.75; $p<0.01$, $I^2=5\%$), chronic kidney disease (OR: 2.09; 95% CI: 1.28–3.41; $p<0.01$; $I^2=0\%$), and cerebrovascular accident (OR: 2.14; 95% CI:

Table 3 Pooled risk of demographic characteristics and comorbidities for postoperative pneumonia following hip fracture surgery

	No. of studies	OR or SMD ^a	LL 95% CI	UL 95% CI	p value	Heterogeneity (%)	Analysis model
Age	5	0.50 ^a	0.10	0.90	0.01	90	Random
Male	10	1.50	1.12	2.01	< 0.01	72	Random
BMI	5	-0.32 ^a	-0.90	0.25	0.27	97	Random
Dependent functional status	2	1.87	0.84	4.13	0.12	88	Random
ASA scale ≥ 3	4	3.17	1.25	8.05	0.02	90	Random
Smoking	6	1.15	0.82	1.60	0.43	0	Fixed
Anemia	3	1.55	1.16	2.08	< 0.01	85	Random
Hypertension	7	1.08	0.89	1.30	0.45	0	Fixed
Diabetes mellitus	9	1.11	0.91	1.37	0.30	7	Fixed
COPD	8	2.05	1.43	2.94	< 0.01	52	Random
Coronary heart disease	6	1.82	1.27	2.60	< 0.01	56	Random
Arrhythmia	4	1.49	1.04	2.15	0.03	0	Fixed
Congestive heart failure	3	1.41	1.14	1.75	< 0.01	5	Fixed
Chronic kidney disease	4	2.09	1.28	3.41	< 0.01	0	Fixed
Cerebrovascular accident	6	2.14	1.60	2.85	< 0.01	22	Fixed
Dementia	3	2.03	0.87	4.71	0.10	59	Random
Cancer	3	1.56	0.93	2.63	0.09	23	Fixed

OR Odds ratio, SMD Standardized mean difference, LL Lower limit, CI Confidence interval, UL Upper limit, ASA American Society of Anesthesiologists physical status, COPD Chronic obstructive pulmonary disease

^a Results of pooled standardized mean difference

Table 4 Pooled risk of baseline laboratory data and surgical characteristics of pneumonia following hip fracture surgery

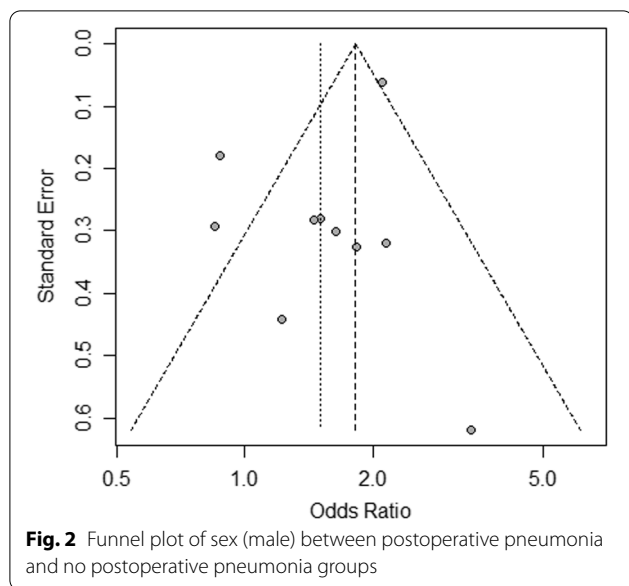
	No. of studies	OR or SMD ^a	LL 95% CI	UL 95% CI	p value	Heterogeneity (%)	Analysis model
Hemoglobin	5	-0.14 ^a	-0.25	-0.03	0.01	46	Fixed
Albumin	4	-0.97 ^a	-1.54	-0.41	< 0.01	95	Random
Blood urea nitrogen	3	0.20 ^a	0.03	0.37	0.02	35	Fixed
Creatinine	5	0.22 ^a	-0.01	0.46	0.06	76	Random
Aspartate aminotransferase	3	0.00 ^a	-0.17	0.17	0.99	27	Fixed
Alanine aminotransferase	3	0.27 ^a	0.10	0.44	< 0.01	0	Fixed
Total bilirubin	2	-0.05 ^a	-0.24	0.14	0.62	0	Fixed
Arterial O ₂ pressure	2	-0.49 ^a	-0.71	-0.27	< 0.01	0	Fixed
Arterial CO ₂ pressure	2	0.04 ^a	-0.18	0.26	0.70	9	Fixed
Intertrochanteric fracture	3	0.90	0.64	1.27	0.55	0	Fixed
Arthroplasty	7	1.24	0.78	1.97	0.35	64	Random
Delayed surgery of over 48 h	3	3.74	2.40	5.85	< 0.01	0	Fixed
Time from injury to surgery	4	0.39 ^a	-0.02	0.79	0.06	94	Random
General anesthesia	5	0.99	0.76	1.30	0.95	44	Fixed
Surgical duration	5	-0.01 ^a	-0.11	0.10	0.91	0	Fixed
Intraoperative blood loss volume	3	-0.01 ^a	-0.17	0.15	0.90	0	Fixed
Perioperative transfusion rate	2	0.98	0.68	1.40	0.91	0	Fixed

OR Odds ratio, SMD Standardized mean difference, LL Lower limit, CI Confidence interval, UL Upper limit

^a Results of pooled standardized mean difference

1.60–2.85; $p < 0.01$; $I^2 = 22\%$) were more likely to develop POP after hip fracture surgery (Table 3). Significant heterogeneity was found for anemia, COPD, coronary heart

disease, and dementia. After sensitivity analyses, heterogeneity was resolved, and the significance did not change (Additional file 1).



Baseline laboratory predictors

Lower preoperative hemoglobin (SMD: -0.14; 95% CI: -0.25 to -0.03; $p=0.01$; $I^2=46\%$), lower preoperative serum albumin (ALB) (SMD: -0.97; 95% CI: -1.54 to -0.41; $p<0.01$; $I^2=95\%$), higher preoperative blood urea nitrogen (BUN) (SMD: 0.20; 95% CI: 0.03–0.37; $p=0.02$; $I^2=35\%$), higher preoperative alanine aminotransferase (SMD: 0.27; 95% CI: 0.10–0.44; $p<0.01$; $I^2=0\%$), and lower partial pressure of oxygen in arterial blood (SMD: -0.49; 95% CI: -0.71–-0.27; $p<0.01$; $I^2=0\%$) indicated an increased risk of POP (Table 4). Significant heterogeneity was observed in the meta-analysis of ALB and creatinine levels. After sensitivity analyses, heterogeneity was resolved for the results of serum creatinine levels without changing the significance (Additional file 1). However, sensitivity analyses could not determine an influential study with high heterogeneity in the ALB level.

Surgical characteristic predictors

Patients who underwent hip fracture surgery that was delayed for >48 h from admission or injury had a significantly higher risk of developing POP (OR: 3.74; 95% CI: 2.40–5.85; $p<0.01$; $I^2=0\%$) (Table 4). Significant heterogeneity was found for surgery type (arthroplasty vs. osteosynthesis) and the time from injury to surgery. After sensitivity analyses, the heterogeneity was resolved, and the intergroup difference in time from injury to surgery was significant (SMD: 0.13; 95% CI: 0.08–0.17; $p<0.01$; $I^2=0\%$) (Additional file 1).

Discussion

The present study extensively reviewed and summarized the predictors of POP in patients undergoing hip fracture surgery. A total of 34 predictors were available

for meta-analysis, of which 15 predictors, namely, male sex, advanced age, ASA scale ≥ 3 , anemia, COPD, coronary heart disease, arrhythmia, congestive heart failure, chronic kidney disease, cerebrovascular accident, time from injury to surgery, delayed surgery >48 h after admission or injury, lower preoperative hemoglobin and ALB levels, lower partial pressure of oxygen in arterial blood, and higher BUN and alanine aminotransferase levels, were statistically significant.

POP occurs frequently in patients undergoing hip fracture surgery, particularly in older patients. Results of this meta-analysis revealed that the overall prevalence of POP was 4.8%, which was comparable to the previously reported range of 4.1–15.3% in patients with hip fracture [7–9, 11]. POP is closely associated with prolonged hospital stay and significantly increased mortality [5–8]. It is directly associated with patient prognosis. Therefore, identification and medical optimization of high-risk patients associated with these risk factors are increasingly important.

Advanced age and male sex have long been associated with adverse postoperative morbidities, including POP, in non-cardiac and orthopedic surgeries [20–23]. Airway inflammation and pneumonia increase with age because of swallowing and immune dysfunctions [24–26]. In addition, impaired spirometric lung age, which is correlated with advanced chronological age, is a well-known risk factor for POP [27]. Furthermore, male patients might have more extensive smoking histories, which can modify lung cell biology and impair mucociliary clearance by the increased number of abnormal cilia. In the same context as impaired lung function, the present study found that patients with lower partial pressure of oxygen in arterial blood were more susceptible to POP development.

In terms of basic demographic data predictors, this meta-analysis also found that ASA scale ≥ 3 was a significant risk factor for POP following hip fracture surgery, consistent with results of previous studies [28, 29]. Therefore, it is needed to give more attention to monitor elderly male patients, particularly those with current status of smoking, dependent functional status, and higher ASA scale, so that early detection could be achieved and prevention strategies could be implemented to reduce POP incidence.

The presence of medical comorbidities has a significant impact in the development of POP after hip fracture surgery. The present study found that anemia, COPD, coronary heart disease, arrhythmia, congestive heart failure, chronic kidney disease, and cerebrovascular accidents were significant risk factors for POP. In particular, comorbid COPD dramatically increases the risk of POP development in patients undergoing hip fracture surgery.

COPD is a common condition in elderly patients with hip fractures, and is associated with increased risk of death and postoperative complications [30, 31]. Patients with COPD are in a state of chronic systemic/vascular inflammation and immune system derangements with upregulated C-reactive protein and increased production of inflammatory cytokines and tissue factors [32–34]. Additionally, limited gas exchange and impaired mucociliary clearance of pathogens can predispose patients with COPD to postoperative pulmonary complications [35, 36]. Targeted interventions to reduce the risk of pneumonia are essential in patients with COPD. Potential interventions for COPD include the use of incentive spirometry, elevation of the head of the bed, early ambulation with pain control, and institution of oral hygiene with chlorhexidine [37].

Previous evidence has suggested that anemia is a significant risk factor for postoperative complications, including POP and increased mortality [38, 39]. Consistent with previous studies, the present study showed that patients with comorbid anemia had an increased risk of POP. In the same context, the pooled results showed an increased risk of POP in patients with lower baseline hemoglobin levels. Thus, medical care in the perioperative period, including patient blood management, should be optimized in patients with comorbid anemia to decrease complications, including POP following hip fracture surgery [40].

Evidence suggests that pneumonia is associated with various medical comorbidities, including coronary heart disease, arrhythmia, congestive heart failure, and chronic kidney disease [41–46]. Cerebrovascular accidents are well-known risk factors for dysphagia and pneumonia [8, 47, 48]. Consistent with previous evidence, the pooled results of the present study showed that coronary heart disease, arrhythmia, congestive heart failure, chronic kidney disease, and cerebrovascular accident were significant risk factors for POP following hip fracture surgery. Generally, co-existing medical morbidities are unmodifiable. However, clinicians should have detailed information on coexisting diseases to assess the risk of POP and identify high-risk patients to apply preventive strategies.

Measurement of ALB level can provide an index of severity of protein-energy malnutrition in patients with hip fractures [49]. Preoperative hypoalbuminemia is a well-described risk factor for perioperative morbidity and mortality in patients undergoing orthopedic surgery [50]. In addition, BUN level is frequently elevated in patients with pneumonia because of hydration and increased reabsorption of urea by the kidneys [51, 52]. An elevated BUN/ALB level has also been reported as an independent predictor of mortality and pneumonia

severity [51, 53]. Abnormal liver function test results are common in patients with pneumonia. Patients with low ALB or elevated alanine aminotransferase levels show increased mortality and length of stay [54]. Several lines of evidence suggest that the lung liver axis is characterized by a shared and prominent feature of pneumonia with a hepatic acute-phase response [55, 56]. In the same context, the present study found that lower ALB, higher BUN, and higher alanine aminotransferase levels as baseline laboratory predictors were associated with POP development.

Importantly, the present study also found that the time from injury to surgery and delayed surgery for over 48 h after admission or injury were significantly associated with the development of POP. The impact of delays in hip fracture surgery on postoperative complications and mortality has been the object of scientific discussion. Most studies have shown that delays in surgery can lead to worse outcomes, such as mortality, pain, complications, and length of stay [57–62]. Therefore, the international clinical practice guidelines recommend early hip surgery within 48 h of admission, if possible [63].

This study has several strengths. This systematic review and meta-analysis is the first to investigate risk factors for POP in patients undergoing hip fracture surgery. In addition, this meta-analysis was based on the most recent studies published within the last 5 years. Nevertheless, this study has several limitations. First, only retrospective studies with low levels of evidence were included. A general limitation of meta-analyses of observational studies is that the result may be a precise, but biased estimate due to inherent biases and confounding in the original studies. We assessed carefully the quality of the component studies and performed sensitivity analyses excluding studies with a high risk of bias. Second, some of our findings showed a significant heterogeneity and require careful interpretation. However, after sensitivity analyses, the heterogeneity was resolved ($I^2 < 50\%$) for most results, except for some variables, such as alcohol consumption, ALB level, and the time from injury to surgery. Third, the small sample size might limit the generalizability of the results. Well-designed studies with a large sample and high quality are required in the future.

Conclusions

This study summarizes numerous predictors of POP in patients undergoing hip fracture surgery. The results can be used to predict the risk of POP development after hip fracture surgery and also provide foundation for future studies. Advanced age, male sex, anemia, diabetes, COPD, coronary heart disease, arrhythmia, congestive heart failure, chronic kidney disease, cerebrovascular

accident, surgery over 48 h after injury or admission, lower preoperative serum hemoglobin or ALB levels, lower partial pressure of oxygen in arterial blood, and higher BUN or alanine aminotransferase levels might contribute to the development of POP after hip fracture surgery.

Abbreviations

POP: Postoperative pneumonia; PRISMA: Preferred Reporting Items for Systematic Reviews and Meta-Analyses; SMD: Standardized mean difference; OR: Odds ratio; CI: Confidence interval; NOS: Newcastle–Ottawa assessment scale; ASA: American Society of Anesthesiologists physical status; COPD: Chronic obstructive pulmonary disease.

Supplementary Information

The online version contains supplementary material available at <https://doi.org/10.1186/s12891-022-05497-1>.

Additional file 1. Results of sensitive analysis for variables.

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None.

Authors' contributions

KHS was the project leader and participated in all aspects of the study, including planning, design, literature search, data screening and extraction, quality appraisal, and management of all aspects of manuscript preparation and submission. SBH and SBK contributed to the study design, literature search, data screening and extraction, quality appraisal, and manuscript editing. All authors have read and approved the final manuscript.

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Availability of data and materials

All data generated or analyzed during this study are included in this published article and its additional information files.

Declarations

Ethics approval and consent to participate

Not applicable.

Consent for publication

Not applicable.

Competing interests

The authors declare that they have no conflicts of interest.

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References

- Brauer CA, Coca-Perrillon M, Cutler DM, Rosen AB. Incidence and mortality of hip fractures in the United States. *JAMA*. 2009;302:1573–9.
- Lunde A, Tell GS, Pedersen AB, Scheike TH, Apalset EM, Ehrenstein V, et al. The role of comorbidity in mortality after hip fracture: a nationwide Norwegian study of 38,126 women with hip fracture matched to a general-population comparison cohort. *Am J Epidemiol*. 2019;188:398–407.
- Stewart NA, Chantrey J, Blankley SJ, Boulton C, Moran CG. Predictors of 5 year survival following hip fracture. *Injury*. 2011;42:1253–6.
- Ciatti C, Maniscalco P, Quattrini F, Gattoni S, Magro A, Capelli P, et al. The epidemiology of proximal femur fractures during COVID-19 emergency in Italy: a multicentric study. *Acta Biomed*. 2021;92:e2021398.
- Lo IL, Siu CW, Tse HF, Lau TW, Leung F, Wong M. Pre-operative pulmonary assessment for patients with hip fracture. *Osteoporos Int*. 2010;21:5579–86.
- Vestergaard P, Rejnmark L, Mosekilde L. Increased mortality in patients with a hip fracture—effect of pre-morbid conditions and post-fracture complications. *Osteoporos Int*. 2007;18:1583–93.
- Bohl DD, Sershon RA, Saltzman BM, Darrith B, Della Valle CJ. Incidence, risk factors, and clinical implications of pneumonia after surgery for geriatric hip fracture. *J Arthroplast*. 2018;33:1552–6.
- Lv H, Yin P, Long A, Gao Y, Zhao Z, Li J, et al. Clinical characteristics and risk factors of postoperative pneumonia after hip fracture surgery: a prospective cohort study. *Osteoporos Int*. 2016;27:3001–9.
- Salarbaks AM, Lindeboom R, Nijmeijer W. Pneumonia in hospitalized elderly hip fracture patients: the effects on length of hospital-stay, in-hospital and thirty-day mortality and a search for potential predictors. *Injury*. 2020;51:1846–50.
- Chang SC, Lai JI, Lu MC, Lin KH, Wang WS, Lo SS, et al. Reduction in the incidence of pneumonia in elderly patients after hip fracture surgery: an inpatient pulmonary rehabilitation program. *Medicine (Baltimore)*. 2018;97:e11845.
- Shin KH, Kim JJ, Son SW, Hwang KS, Han SB. Early postoperative hypoalbuminaemia as a risk factor for postoperative pneumonia following hip fracture surgery. *Clin Interv Aging*. 2020;15:1907–15.
- Wang Y, Li X, Ji Y, Tian H, Liang X, Li N, et al. Preoperative serum albumin level as a predictor of postoperative pneumonia after femoral neck fracture surgery in a geriatric population. *Clin Interv Aging*. 2019;14:2007–16.
- Xiang G, Dong X, Xu T, Feng Y, He Z, Ke C, et al. A nomogram for prediction of postoperative pneumonia risk in elderly hip fracture patients. *Risk Manag Healthc Policy*. 2020;13:1603–11.
- Zhao K, Zhang J, Li J, Guo J, Meng H, Zhu Y, et al. In-hospital postoperative pneumonia following geriatric intertrochanteric fracture surgery: incidence and risk factors. *Clin Interv Aging*. 2020;15:1599–609.
- Moher D, Liberati A, Tetzlaff J, Altman DG, PRISMA Group. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *BMJ*. 2009;339:b2535.
- Stang A. Critical evaluation of the Newcastle–Ottawa scale for the assessment of the quality of nonrandomized studies in meta-analyses. *Eur J Epidemiol*. 2010;25:603–5.
- Cumpston M, Li T, Page MJ, Chandler J, Welch VA, Higgins JP, et al. Updated guidance for trusted systematic reviews: a new edition of the Cochrane handbook for systematic reviews of interventions. *Cochrane Database Syst Rev*. 2019;10:ED000142.
- Ji Y, Li X, Wang Y, Cheng L, Tian H, Li N, et al. Partial pressure of oxygen level at admission as a predictor of postoperative pneumonia after hip fracture surgery in a geriatric population: a retrospective cohort study. *BMJ Open*. 2021;11:e048272.
- Wang X, Dai L, Zhang Y, Lv Y. Gender and low albumin and oxygen levels are risk factors for perioperative pneumonia in geriatric hip fracture patients. *Clin Interv Aging*. 2020;15:419–24.
- Gupta H, Gupta PK, Schuller D, Fang X, Miller WJ, Modrykamien A, et al. Development and validation of a risk calculator for predicting postoperative pneumonia. *Mayo Clin Proc*. 2013;88:1241–9.
- Bohl DD, Ahn J, Rossi VJ, Tabaraee E, Grauer JN, Singh K. Incidence and risk factors for pneumonia following anterior cervical decompression and fusion procedures: an ACS-NSQIP study. *Spine J*. 2016;16:335–42.
- Nagle RT, Leiby BE, Lavu H, Rosato EL, Yeo CJ, Winter JM. Pneumonia is associated with a high risk of mortality after pancreaticoduodenectomy. *Surgery*. 2017;161:959–67.
- Ally SA, Foy M, Sood A, Gonzalez M. Preoperative risk factors for postoperative pneumonia following primary total hip and knee arthroplasty. *J Orthop*. 2021;27:17–22.
- Carpagnano GE, Turchiarelli V, Spanevello A, Palladino GP, Barbaro MP. Aging and airway inflammation. *Aging Clin Exp Res*. 2013;25:239–45.

25. Ebihara S, Ebihara T, Kohzuki M. Effect of aging on cough and swallowing reflexes: implications for preventing aspiration pneumonia. *Lung*. 2012;190:29–33.
26. Janssens JP, Krause KH. Pneumonia in the very old. *Lancet Infect Dis*. 2004;4:112–24.
27. Okamura A, Watanabe M, Mine S, Nishida K, Kuroguchi T, Imamura Y. Spirometric lung age predicts postoperative pneumonia after esophagectomy. *World J Surg*. 2016;40:2412–8.
28. Hackett NJ, De Oliveira GS, Jain UK, Kim JY. ASA class is a reliable independent predictor of medical complications and mortality following surgery. *Int J Surg*. 2015;18:184–90.
29. Yang CK, Teng A, Lee DY, Rose K. Pulmonary complications after major abdominal surgery: National Surgical Quality Improvement Program analysis. *J Surg Res*. 2015;198:441–9.
30. de Luise C, Brimacombe M, Pedersen L, Sorensen HT. Chronic obstructive pulmonary disease and mortality following hip fracture: a population-based cohort study. *Eur J Epidemiol*. 2008;23:115–22.
31. Regan EA, Radcliff TA, Henderson WG, Cowper Ripley DC, Maciejewski ML, Vogel WB, et al. Improving hip fractures outcomes for COPD patients. *COPD*. 2013;10:11–9.
32. Bhat TA, Panzica L, Kalathil SG, Thanavala Y. Immune dysfunction in patients with chronic obstructive pulmonary disease. *Ann Am Thorac Soc*. 2015;12(Suppl 2):S169–75.
33. Gan WQ, Man SF, Senthilvelan A, Sin DD. Association between chronic obstructive pulmonary disease and systemic inflammation: a systematic review and a meta-analysis. *Thorax*. 2004;59:574–80.
34. Thomsen M, Dahl M, Lange P, Vestbo J, Nordestgaard BG. Inflammatory biomarkers and comorbidities in chronic obstructive pulmonary disease. *Am J Respir Crit Care Med*. 2012;186:982–8.
35. Smetana GW, Lawrence VA, Cornell JE. American College of Physicians. Preoperative pulmonary risk stratification for noncardiothoracic surgery: systematic review for the American College of Physicians. *Ann Intern Med*. 2006;144:581–95.
36. Vogelmeier CF, Criner GJ, Martinez FJ, Anzueto A, Barnes PJ, Bourbeau J, et al. Global strategy for the diagnosis, management and prevention of chronic obstructive lung disease 2017 report: GOLD executive summary. *Respirology*. 2017;22:575–601.
37. Kazaure HS, Martin M, Yoon JK, Wren SM. Long-term results of a postoperative pneumonia prevention program for the inpatient surgical ward. *JAMA Surg*. 2014;149:914–8.
38. Foss NB, Kristensen MT, Kehlet H. Anaemia impedes functional mobility after hip fracture surgery. *Age Ageing*. 2008;37:173–8.
39. Yombi JC, Putineanu DC, Cornu O, Lavand'homme P, Cornette P, Castanares-Zapatero D. Low haemoglobin at admission is associated with mortality after hip fractures in elderly patients. *Bone Joint J*. 2019;101-B:1122–8.
40. Suh YS, Nho JH, Seo J, Jang BW, Park JS. Hip fracture surgery without transfusion in patients with hemoglobin less than 10 g/dL. *Clin Orthop Surg*. 2021;13:30–6.
41. Aliberti S, Ramirez JA. Cardiac diseases complicating community-acquired pneumonia. *Curr Opin Infect Dis*. 2014;27:295–301.
42. Chou CY, Wang SM, Liang CC, Chang CT, Liu JH, Wang IK, et al. Risk of pneumonia among patients with chronic kidney disease in outpatient and inpatient settings: a nationwide population-based study. *Medicine (Baltimore)*. 2014;93:e174.
43. James MT, Quan H, Tonelli M, Manns BJ, Faris P, Laupland KB, et al. CKD and risk of hospitalization and death with pneumonia. *Am J Kidney Dis*. 2009;54:24–32.
44. Kim J, Park SJ, Choi S, Seo WW, Lee YJ. Hospitalization for acute coronary syndrome increases the long-term risk of pneumonia: a population-based cohort study. *Sci Rep*. 2021;11:9696.
45. Mor A, Thomsen RW, Ulrichsen SP, Sorensen HT. Chronic heart failure and risk of hospitalization with pneumonia: a population-based study. *Eur J Intern Med*. 2013;24:349–53.
46. Zhu J, Zhang X, Shi G, Yi K, Tan X. Atrial fibrillation is an independent risk factor for hospital-acquired pneumonia. *Plos One*. 2015;10:e0131782.
47. Hibberd J, Fraser J, Chapman C, McQueen H, Wilson A. Can we use influencing factors to predict aspiration pneumonia in the United Kingdom? *Multidiscip Respir Med*. 2013;8:39.
48. Ding R, Logemann JA. Pneumonia in stroke patients: a retrospective study. *Dysphagia*. 2000;15:51–7.
49. O'Daly BJ, Walsh JC, Quinlan JF, Falk GA, Stapleton R, Quinlan WR, et al. Serum albumin and total lymphocyte count as predictors of outcome in hip fractures. *Clin Nutr*. 2010;29:89–93.
50. Bohl DD, Shen MR, Hannon CP, Fillingham YA, Darrih B, Della Valle CJ. Serum albumin predicts survival and postoperative course following surgery for geriatric hip fracture. *J Bone Joint Surg Am*. 2017;99:2110–8.
51. Ugajin M, Yamaki K, Iwamura N, Yagi T, Asano T. Blood urea nitrogen to serum albumin ratio independently predicts mortality and severity of community-acquired pneumonia. *Int J Gen Med*. 2012;5:583–9.
52. Woodford-Williams E. Respiratory tract disease. Diagnosis and management of pneumonia in the aged. *Br Med J*. 1966;1:467–70.
53. Feng DY, Zhou YQ, Zou XL, Zhou M, Yang HL, Chen XX, et al. Elevated blood urea nitrogen-to-serum albumin ratio as a factor that negatively affects the mortality of patients with hospital-acquired pneumonia. *Can J Infect Dis Med Microbiol*. 2019;2019:1547405.
54. Jinks MF, Kelly CA. The pattern and significance of abnormal liver function tests in community-acquired pneumonia. *Eur J Intern Med*. 2004;15:436–40.
55. Quinton LJ, Jones MR, Robson BE, Mizgerd JP. Mechanisms of the hepatic acute-phase response during bacterial pneumonia. *Infect Immun*. 2009;77:2417–26.
56. Weber M, Lambeck S, Ding N, Henken S, Kohl M, Deigner HP, et al. Hepatic induction of cholesterol biosynthesis reflects a remote adaptive response to pneumococcal pneumonia. *FASEB J*. 2012;26:2424–36.
57. Novack V, Jotkowitz A, Etzion O, Porath A. Does delay in surgery after hip fracture lead to worse outcomes? A multicenter survey. *Int J Qual Health Care*. 2007;19:170–6.
58. Orosz GM, Magaziner J, Hannan EL, Morrison RS, Koval K, Gilbert M, et al. Association of timing of surgery for hip fracture and patient outcomes. *JAMA*. 2004;291:1738–43.
59. Radcliff TA, Henderson WG, Stoner TJ, Khuri SF, Dohm M, Hutt E. Patient risk factors, operative care, and outcomes among older community-dwelling male veterans with hip fracture. *J Bone Joint Surg Am*. 2008;90:34–42.
60. Siegmeth AW, Gurusamy K, Parker MJ. Delay to surgery prolongs hospital stay in patients with fractures of the proximal femur. *J Bone Joint Surg Br*. 2005;87:1123–6.
61. Simunovic N, Devereaux PJ, Sprague S, Guyatt GH, Schemitsch E, Debeer J, et al. Effect of early surgery after hip fracture on mortality and complications: systematic review and meta-analysis. *CMAJ*. 2010;182:1609–16.
62. Smektala R, Endres HG, Dasch B, Maier C, Trampisch HJ, Bonnaire F, et al. The effect of time-to-surgery on outcome in elderly patients with proximal femoral fractures. *BMC Musculoskelet Disord*. 2008;9:171.
63. Roberts KC, Brox WT, Jevsevar DS, Sevarino K. Management of hip fractures in the elderly. *J Am Acad Orthop Surg*. 2015;23:131–7.

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