

Central Lancashire Online Knowledge (CLoK)

| Title | Urinary tract infection is associated with a 2.4-fold increased risk of surgical site infection in hip fracture surgery: a systematic review and meta-analysis |
|----------|---|
| Type | Article |
| URL | https://clok.uclan.ac.uk/id/eprint/47448/ |
| DOI | https://doi.org/10.1016/j.jhin.2023.06.016 |
| Date | 2023 |
| Citation | Suen, Kai Fung Kevin, Low, Joshua Xian Yang and Charalambous, Charalambos Panayiotou (2023) Urinary tract infection is associated with a 2.4-fold increased risk of surgical site infection in hip fracture surgery: a systematic review and meta-analysis. Journal of Hospital Infection, 139. pp. 56-66. ISSN 0195-6701 |
| Creators | Suen, Kai Fung Kevin, Low, Joshua Xian Yang and Charalambous, Charalambos Panayiotou |

It is advisable to refer to the publisher's version if you intend to cite from the work. https://doi.org/10.1016/j.jhin.2023.06.016

For information about Research at UCLan please go to http://www.uclan.ac.uk/research/

All outputs in CLoK are protected by Intellectual Property Rights law, including Copyright law. Copyright, IPR and Moral Rights for the works on this site are retained by the individual authors and/or other copyright owners. Terms and conditions for use of this material are defined in the http://clok.uclan.ac.uk/policies/



Available online at www.sciencedirect.com

Journal of Hospital Infection

journal homepage: www.elsevier.com/locate/jhin



Systematic Review

Urinary tract infection is associated with 2.4-fold increased risk of surgical site infection in hip fracture surgery: systematic review and meta-analysis

K.F.K. Suen a, J.X.Y. Low a, C.P. Charalambous a, b, *

ARTICLE INFO

Article history: Received 24 February 2023 Accepted 15 June 2023 Available online 19 June 2023

Keywords: Hip fracture Surgical site infection Urinary tract infection Urinary catheterization



SUMMARY

Background: There is no consensus regarding whether urinary tract infection (UTI) should be screened for or treated in hip fracture patients.

Aim: To assess the relationship between perioperative UTI and surgical site infection (SSI) in hip fracture patients, and the relationship between urinary catheterization and SSI in these patients.

Methods: PubMed, Embase, CINAHL and Cochrane Library were searched to identify studies that evaluated the relationship between perioperative UTI and SSI and/or between urinary catheterization and SSI. Articles were included if they used the term UTI or specified UTI as symptomatic bacteriuria.

Findings: A total of 4139 records were identified, with eight studies included. Meta-analysis of seven studies which evaluated perioperative UTI and SSI showed an SSI rate of 7.1% (95% confidence interval (CI): 3.8-13.2) among 1217 patients with UTI vs 2.4% (95% CI: 1.0-5.7) in 36,514 patients without UTI (OR: 2.41; 95% CI: 1.67-3.46; P < 0.001). In three studies which specifically defined UTI as symptomatic bacteriuria, the SSI rate among UTI patients was 5.7% (95% CI: 4.0-8.1) vs 1.1% (95% CI: 0.2-5.2) in those without UTI (OR: 3.00; 95% CI: 0.55-16.26; P = 0.20). One study evaluated urinary catheterization and SSI.

Conclusion: Perioperative UTI is associated with a higher risk of SSI among hip fracture patients but the evidence is limited by the heterogeneity in the definition of UTI. We recommend considering the possibility of perioperative UTI in hip fracture patients, with treatment administered as necessary to reduce SSI rates.

© 2023 The Authors. Published by Elsevier Ltd on behalf of The Healthcare Infection Society. This is an open access article under the CC BY license (http://creativecommons.org/licenses/by/4.0/).

E-mail address: cpcharalambous@uclan.ac.uk (C.P. Charalambous).

Introduction

Surgical site infection (SSI) can be a devastating complication in patients with hip fracture, with a reported one-year

^a Department of Orthopaedics, Blackpool Victoria Hospital, Blackpool Teaching Hospitals NHS Foundation Trust, Blackpool, Lancashire, UK

^b School of Medicine, University of Central Lancashire, Preston, Lancashire, UK

^{*} Corresponding author. Address: Department of Orthopaedics, Blackpool Victoria Hospital, Whinney Heys Road, Lancashire, Blackpool, FY3 8NR, UK. Tel.: ± 44 (0)1253 955983.

mortality rate in those with SSI of around 50%, compared with around 30% in those without SSI [1]. The 2021–2022 surveil-lance report of the UK Health Security Agency showed that, among NHS hospitals in England, the SSI rate in total hip replacement surgery for acute trauma and chronic elective indications was 0.5%, whereas the SSI rate of repair neck of femur fracture surgery that included hip hemiarthroplasty and fracture fixation was 0.8% [2]. The SSI of hip fracture patients treated with hemiarthroplasty was reported by research studies as 3–10% compared with about 1% in elective hip arthroplasty patients [3–6]. With about 76,000 hip fractures occurring annually in the UK, SSIs could impact a large group of patients.

According to the 2021–2022 UK Health Security Agency surveillance report, across surgical specialties, Enterobacterales were the most prevalent causative organism for SSIs with an increasing 10-year trend [2]. The most common Enterobacterales species was *Escherichia coli*. Enterobacterales SSIs accounted for 18.3% of superficial and 20.3% of deep infections following hip replacement surgery, and 55% of deep polymicrobial infections following repair neck of femur fracture surgery involved a Gram-negative organism. As Enterobacterales are the main causative organisms in urinary tract infection (UTI), there is concern that UTI may be linked to SSI following hip fracture surgery [7].

The screening for and management of UTI in patients having elective hip arthroplasty is well studied and it is widely accepted that symptomatic bacteriuria should be treated and elective surgery postponed [8]. Asymptomatic bacteriuria in elective arthroplasty patients is not routinely tested or treated because antibiotic treatment for preoperative asymptomatic bacteriuria has not been shown to reduce any risk for SSI, and because infective agents isolated from prosthetic joint infection tend to differ from those isolated in the urine [9]. However, the relation between UTI and SSI in hip fracture patients is less well understood. Currently, there is no consensus regarding whether UTI should be screened for or treated in patients with hip fractures. Hip fracture patients are often older and with more comorbidities as compared to those having elective hip surgery, which make them more susceptible to the effects of SSI [10,11]. Although elective arthroplasty could be delayed in the presence of symptomatic bacteriuria, hip fractures require urgent surgical intervention, preferably within 36 h from injury, as longer delays are associated with a higher mortality rate [12,13]. Hence, it is not feasible to wait for urine culture results and the clearance of UTI in hip fracture patients. Besides, UTI could arise after the hip fracture

Urinary catheterization is frequently placed in hip fracture patients to prevent or treat urinary retention, accommodate patients' limited mobility, and facilitate the measurement of urine output. Urinary catheterization is a risk factor for UTI as the catheter may act as a nidus for bacteria to settle on, but there is no consensus regarding the association of urinary catheterization (inserted before the fracture or after the fracture) with the risk for SSI among hip fracture patients [14].

In view of the above, this systematic review and metaanalysis aimed to assess, first, the relationship between perioperative UTI and SSI in hip fracture patients and, second, the relationship between urinary catheterization and SSI in such patients. It was hypothesized that UTI and urinary catheterization are associated with a higher SSI risk.

Methods

This systematic review was performed according to the Preferred Reporting Items for Systematic Review and Meta-Analysis (PRISMA) updated guideline in 2020 [15]. The protocol was prospectively registered with PROSPERO (CRD42021273563).

Search strategy

Literature search was performed by two researchers independently in PubMed, Embase, CINAHL, and Cochrane Library databases from their respective inceptions to January 11th, 2022 with no restriction on language. The search commands adopted included: (hip AND fracture AND urinary), (femur AND fracture AND urinary) and (femoral AND fracture AND urinary). References in selected articles found in searches were manually reviewed also to identify any articles missed in the primary search.

Studies on hip fracture patients treated surgically that evaluated the relationship between perioperative UTI and SSI and/or the relationship between urinary catheterization (duration or number of times) and SSI were included. For articles to be included they had to refer to the term UTI or to specify UTI as symptomatic bacteriuria. Exclusion criteria were: (1) studies reporting on elective surgery for non-fracture patients (e.g. osteoarthritis); (2) studies reporting on asymptomatic bacteriuria; (3) systematic reviews/meta-analyses; (4) case series reporting <10 cases; and (5) editorials or commentaries.

Study selection

Two researchers (K.F.K.S. and J.X.Y.L.) independently screened articles by titles and abstracts and then reviewed the full texts to select eligible articles. Disagreement on the selection of articles was settled by discussion between the two researchers and, if necessary, by arbitration by the senior author.

Data extraction

Data were extracted according to a pre-designed proforma, with items including study design, sample size, sex ratio, mean age, patient inclusion and exclusion criteria, surgical intervention, follow-up period, definition and event rate of SSI, definition and event rate of UTI (if any), number of times and time length of urinary catheterization (if any), event rates of SSI in patients with and without UTI (if any), event rates of SSI in patients with and without urinary catheterization (if any), and culture results of SSI and UTI (if any). The risk of bias was assessed with the Newcastle—Ottawa Scale for observational studies.

Statistical analysis

Statistical analysis was performed with R ('meta' and 'metafor' packages), including a meta-analysis to provide an SSI rate and an odds ratio (OR) with confidence interval (CI) calculated, with the examination on the heterogeneity of studies. Meta-

analysis was performed with the random effects model due to the diversity of patient populations among selected articles. Publication bias was assessed by inspection of funnel plots.

Results

The initial literature search produced 4139 records. After removal of duplicated records, screening by title and abstract, and full-text review, seven studies, reporting on a total of 37,731 patients were included which evaluated the relationship between perioperative UTI and SSI (Figure 1). These compared SSI rates between 1217 hip fracture patients with UTI vs 36,514 without UTI [16—22]. However, these studies vary on their definition of UTI and on the inclusion of clinical symptoms for the diagnosis of UTI to be made (Table I). Among these studies, three studies reporting on

1303 patients clearly defined UTI as symptomatic bacteriuria [17—19].

The quality of all included studies was evaluated according to Newcastle—Ottawa Scale (NOS) (Table II).

Among all studies included, the SSI rate among UTI patients was 7.1% (95% CI: 3.8–13.2; heterogeneity: $\chi^2=43.11$ (P<0.0001); $\mathit{l}^2=86.1\%$; $\tau^2=0.4878$) vs 2.4% (95% CI: 1.0–5.7; heterogeneity: $\chi^2=104.40$ (P<0.0001); $\mathit{l}^2=94.3\%$; $\tau^2=1.1757$) in those with no UTI (OR: 2.41; 95% CI: 1.67–3.46 (P<0.01); heterogeneity: $\chi^2=8.39$ (P=0.21); $\mathit{l}^2=25\%$; $\tau^2=0.06$) (Table III, Figure 2).

In the three studies which specifically defined UTI as symptomatic bacteriuria, the SSI rate among UTI patients was 5.7% (95% CI: 4.0–8.1; heterogeneity: $\chi^2 = 2.42$ (P = 0.30); $I^2 = 17.3\%$; $\tau^2 < 0.0001$) vs 1.1% (95% CI: 0.2–5.2; heterogeneity: $\chi^2 = 13.16$ (P = 0.0014); $I^2 = 84.8\%$; $\tau^2 = 1.4930$) in those with

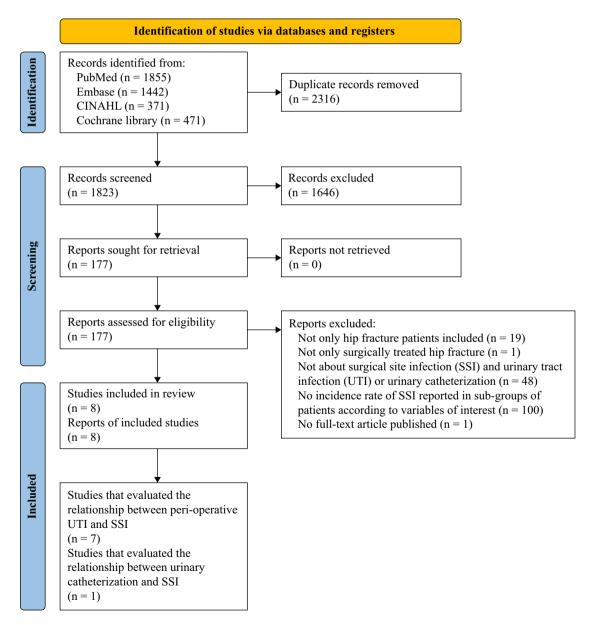


Figure 1. PRISMA flow diagram 2020.

Table ICharacteristics of included studies reporting on the relationship between UTI and SSI rates

| Study | Design, centre, duration, country | Sample size | Age (years), mean (SD), unless otherwise specified | Sex | Inclusion criteria, in addition to traumatic hip fracture treated surgically | Exclusion criteria, in addition to not meeting inclusion criteria | • | Definition of 'UTI' or 'symptomatic bacteriuria' (time of test/diagnosis) | Definition of SSI (timeframe of diagnosis) | Antibiotics prophylaxis for surgery and antibiotics for UTI | Incidence of SSI among UTI patients | Incidence of SSI among non-UTI patients |
|-------------------------|---|----------------|---|--|---|--|------------------|---|--|---|--|---|
| Craxford et al. [16] | Retrospective cohort study, single centre, 11 years (Aug 2007 —Aug 2018), UK | 3966 | Among SSI: 82.20 (10.06) Among no SSI: 82.45 (8.89) | Among SSI: M: 20 (29.0%), F: 69 (71.0%) Among no SSI: M: 1118 (28.7%), F: 2779 (71.3%) | Hemiarthroplasty | Revision surgery for periprosethetic fracture or dislocation | НА | Unspecified definition of 'UTI' (pre- and postoperative) | CDC criteria (2008) (12 months) | Unspecified. For most patients, gentamicincontaining cement was used | 17/444 (3.82%) | 52/3522 (1.48%) |
| Crouser et al. [17] | Retrospective cohort study of database (multi- centre), 2 years (2015–2016), USA | 31,621 | 65–79 years: 10,388 80–89 years: 13,948 ≥90 years: 7285 | M: 9171 (29.0%) F: 22,450 (71.0%) | - | / | HA, THR, ORIF | Characteristic symptoms of UTI present, but not essentially with bacteriuria shown microbiologically ^a (at the time of surgery) | CDC criteria (30 days) | Unspecified | 24/410 (5.85%) | 1159/31,211 (3.71%) |
| group 2 of | Retrospective cohort study, single centre, 31 months (Sep 2014—Mar 2017) for Group 2, Germany | | 83.93 (7.70) for Group 2 | Group 2: M: 120 (27.2%), F: 321 (72.8%) | Age ≥65 years | Deep surgical site infection after revision surgery | HA, THR, ORIF | Bacteriuria shown microbiologically, with data about the subgroup of patients with symptomatic bacteriuria (preoperative at admission) | , , | Cefuroxime at time of surgery + preoperative ciprofloxacin for few days if bacteriuria | 0/141 (0.00%) Among those with symptomatic bacteriuria: 0/77 (0.00%) | 2/300 (0.67%) Among those without symptomatic bacteriuria: 2/364 (0.55%) |
| Bliemel et al. [19] | Prospective cohort, single centre, 2.5 years (Apr 2009 —Sep 2011), Germany | 402 | 81 (8) | M: 109 (27.1%) F: 293 (72.9%) | Age ≥60 years | Polytrauma, malignancy- related fracture | HA, THR, ORIF | Characteristic symptoms of UTI present, but not essentially with | Clinical, laboratory and sonographic examinations (unspecified time) | Yes, unspecified antibiotics. UTI was treated according to local guidelines | 6/97 (6.19%) | 1/305 (0.33%) |

(continued on next page)

| Study | Design, centre, duration, country | Sample size | Age (years), mean (SD), unless otherwise specified | | Inclusion criteria, in addition to traumatic hip fracture treated surgically | in addition to not | • | Definition of 'UTI' or 'symptomatic bacteriuria' (time of test/diagnosis) | Definition of SSI (timeframe of diagnosis) | Antibiotics prophylaxis for surgery and antibiotics for UTI | Incidence of SSI among UTI patients | Incidence of SSI among non-UTI patients |
|--------------------------|--|----------------|---|---|---|---|------------------|--|--|--|---|---|
| Yassa et al. [20] | Retrospective cohort study, single centre, 1 year (unspecified time), UK | 460 | 80.9 | M: 124 (27%) F: 336 (73%) | / | / | HA, THR, ORIF | 'UTI' was defined as bacteriuria shown by positive dipstick, but their database did not specify whether 'UTI' cases were symptomatic or not ^c (routine dipstick at admission) | Unspecified (unspecified time) | - | 23/99 (23.2%) | 34/361 (9.42%) |
| Capdevila et al. [21] | Retrospective cohort study, single centre, 16 months (Jun 2012–Sep 2013), Spain | 657 | 83.1 (10.4) | M: 188 (28.6%) F: 469 (71.4%) | / | / | HA, THR, ORIF | Unspecified definition of 'UTI' | Superficial and deep SSI, CDC criteria (1999) (1 year) | Cefuroxime and teicoplanin at time of surgery. For cemented prosthesis, cement without antibiotics was used. | 4/78 (5.13%) | 9/579 (1.55%) |
| Westberg et al. [22] | Retrospective cohort study, single centre, 2 years (2018 –2019), Norway | | With PJI: 79 (9) Without PJI: 81 (10) | With PJI: M: 6 (35%), F: 11 (65%) Without PJI: M: 52 (31%), F: 115 (69%) | Arthroplasty | Patients from outside the hospital catchment area, or pathological fractures | HA, THR | Unspecified definition of 'UTI' | PJI defined as deep SSI of CDC criteria (1999) and Tsukayama's criteria for timing for early infection (4 weeks) | Cephalotin at time of surgery and 3 more doses over postoperative 24 h. For most | 2/12 (16.7%) | 15/172 (8.02%) |

UTI, urinary tract infection; SSI, surgical site infection; THR, total hip replacement; HA, hemiarthroplasty; ORIF, open reduction internal fixation; PJI, prosthetic joint infection; APJI, acute prosthetic joint infection.

^a Diagnosis of 'UTI' in these studies required the presence of characteristic UTI symptom(s) and required either positive culture growth OR positive urine analysis nitrite and/or leucocytes.

b Diagnosis of 'bacteriuria' required positive urine sediment and microscopy analysis results; diagnosis of 'UTI' required 'bacteriuria' with characteristic UTI symptoms.

^c Diagnosis of 'UTI' required positive urine dipstick and did not require microbiological evidence or characteristic UTI symptoms.

Table IIThe Newcastle—Ottawa scale score of included studies

| Cohort study | Representativeness of the exposed cohort | Selection of the non-exposed cohort | Ascertainment of exposure | Demonstration that outcome of interest was not present at start of study | Comparabilit cohorts on basis of the de analysis | the sign or | Assessment of outcome | Follow-up long enough for outcomes to occur | Adequacy of follow-up of cohorts | Total |
|--------------------------|---|---|---------------------------|--|--|----------------|------------------------------|--|----------------------------------|-------|
| Craxford et al. [16] | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 9/9 |
| Crouser et al. [17] | 1 | 1 | 1 | 1 | 0 | 0 | 1 | 1 | 1 | 7/9 |
| Langenhan et al. [18] | 1 | 1 | 1 | 1 | 0 | 0 | 1 | 1 | 1 | 7/9 |
| Bliemel et al. [19] | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 9/9 |
| Yassa et al. [20] | 1 | 1 | 1 | 1 | 0 | 0 | 1 | 1 | 1 | 7/9 |
| Capdevila et al. [21] | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 9/9 |
| Westberg et al. [22] | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 9/9 |
| Case—control study | Adequate case definition | Representativeness of the cases | Selection of controls | Definition of controls | Comparability of cases and controls on the basis of the design or analysis | | Ascertainment of exposure | Same method of ascertainment for cases and controls | Non-response rate | Total |
| Cumming and Parker [23] | 1 | 1 | 0 | 1 | 1 | 0 | 1 | 1 | 0 | 6/9 |

Table III
SSI rates among UTI patients and among non-UTI patients

| Definition of UTI | No. of | | UT | Ί | No UTI | | | |
|--|---------|-----------------|----------------------|--|-----------------|----------------------|--|--|
| | studies | No. of patients | SSI rate (95% CI) | Heterogeneity: χ^2 (<i>P</i> -value), I^2 , τ^2 | No. of patients | SSI rate (95% CI) | Heterogeneity: χ^2 (<i>P</i> -value), I^2 , τ^2 | |
| Studies referring to the term 'UTI' or to the term 'symptomatic bacteriuria' | 7 | 1217 | 7.1% (3.8–13.2) | 43.11 (<i>P</i> < 0.0001), 86.1%, 0.4878 | 36,514 | 2.4% (1.0–5.7) | 104.40 (<i>P</i> < 0.0001), 94.3%, 1.1757 | |
| Studies referring to 'symptomatic bacteriuria' | 3 | 584 | 5.7% (4.0–8.1) | 2.42 (<i>P</i> = 0.30), 17.3%, <0.0001 | 31,880 | 1.1% (0.2—5.2) | 13.16 (<i>P</i> = 0.0014), 84.8%, 1.4930 | |

SSI, surgical site infection; UTI, urinary tract infection; CI, confidence interval.

0.55–16.26 (P=0.20); heterogeneity: $\chi^2=5.35$ (P=0.07); $\ell^2=63\%$; $\tau^2=1.39$) (Table IV, Figure 3).

Only one study was identified which evaluated the relationship between urinary catheterization and SSI in hip fracture patients (Table V), so meta-analysis was not performed with regard to this evaluation. In that study, Cumming *et al.* showed that deep wound infection in hip fracture patients was associated with more than two catheterizations and long-term catheters (>21 days of catheterization or being discharged from hospital with a catheter *in situ*) [23].

Discussion

The current meta-analysis showed a high rate of SSI among hip fracture patients who have perioperative UTI, estimated at 7.1%, while the risk for SSI among UTI patients was estimated at 2.4-fold higher compared with non-UTI patients.

Several mechanisms have been postulated to explain the relationship between UTI and SSI. Understanding such mechanisms is vital as it may shed further light on the pathogenesis of SSI and identify a possible target for intervention to reduce SSI rates and thus mortality in hip fracture patients [1]. The relation between UTI and SSI may be due to direct contamination of the wound by infected urine. External surgical wound contamination by urine micro-organisms could also be a source of SSI in hip fracture patients. Reduced mobility and cognitive status are also risk factors for UTI [24]. Dementia, cognitive impairment, and delirium are common among hip fracture patients, which may lead to contamination of the surgical wound by unintentional physical contact with urine [25,26].

Contamination may also occur via a haematogenous route from the urinary tract to the surgical wound. It was previously shown that haematoma at the site of a hip fracture commonly contained bacteria even before surgery and the presence of Gram-negative bacilli in the haematoma was a risk factor for early postoperative prosthetic joint infection [27]. The researchers postulated that this could be due to UTI contaminating the wound via bacterial translocation. Bacterial translocation is a well-recognized phenomenon which refers to the passage of gut microflora from the gastrointestinal tract to other tissues and organs, due to increased intestinal permeability at times of physiological stress and could be the mechanism behind postoperative sepsis after major gastrointestinal surgery [28-30]. A similar process of translocation of uropathogens through the uroepithelium, into the renal interstitium and subsequently into the bloodstream is well studied in UTI, and this process involves the formation of intracellular bacterial communities within uroepithelial cells [31]. There is also limited evidence about bacterial translocation in asymptomatic bacteriuria: in a small animal study, uropathogenic Escherichia coli cultured from five asymptomatic bacteriuric humans were seen to form intracellular bacterial communities to cause cystitis in a mouse model [32].

Alternatively, UTI may cause immunosuppression that indirectly leads to a higher risk of SSI [20]. This proposed mechanism of immunosuppression due to UTI may be particularly important in hip fracture patients due to their greater burden of comorbidities. Such a mechanism could explain the mismatch between UTI and SSI culture isolates [16,18,20].

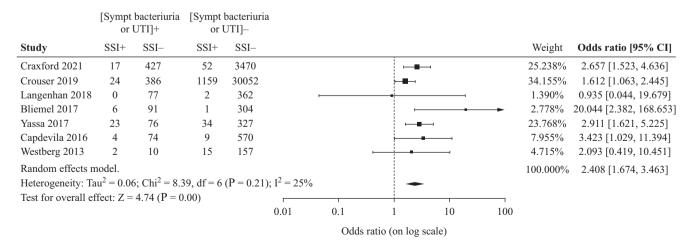


Figure 2. Forest plot of odds ratio for surgical site infection among urinary tract infection patients.

Table IVOdds ratio for SSI in UTI patients vs non-UTI patients

| Definition of UTI | No. of studies | No. of patients | Odds ratio (95% CI), <i>P</i> -value | Heterogeneity: χ^2 (<i>P</i> -value), I^2 , τ^2 , |
|---|----------------|-----------------|---|--|
| Studies referring to 'UTI' or 'symptomatic bacteriuria' | 7 | 37,731 | 2.41 (1.67–3.46) P < 0.01 | 8.39 (<i>P</i> = 0.21) 25%, 0.06 |
| Studies referring to 'symptomatic bacteriuria' | 3 | 32,444 | 3.00 (0.55–16.26) P = 0.20 | 5.35 (<i>P</i> = 0.07) 63%, 1.39 |

SSI, surgical site infection; UTI, urinary tract infection; CI, confidence interval.

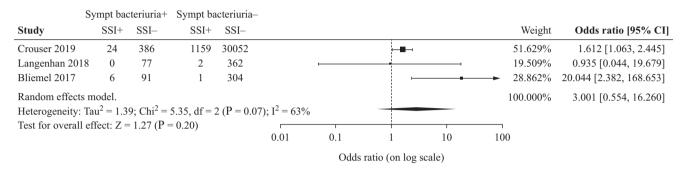


Figure 3. Forest plot of odds ratio for surgical site infection among patients with urinary tract infection specifically defined as symptomatic bacteriuria.

Diagnosis of UTI is challenging, as clinical symptoms and urine culture samples may not be in line. Some patients may experience urinary symptoms but have a negative urine culture, some may have no urinary symptoms but generalized symptoms of infection with a positive or negative urine culture, whereas some may have asymptomatic bacteriuria. Asymptomatic bacteriuria is generally considered a separate entity as compared to UTI as it may simply reflect colonization of the urinary tract, hence articles reporting on asymptomatic bacteriuria were excluded from our analysis.

Although we established a statistical association between UTI and SSI, as the included studies were observational, a causal relationship could not be established. Nevertheless, there is some evidence suggesting that treatment of UTI reduces the SSI rate. Langenhan $et\ al.$ retrospectively collected data in a group of patients who received a single dose of 1.5 g of cefuroxime as preoperative prophylaxis and then prospectively collected data in another group which, in addition to the preoperative cefuroxime, received ciprofloxacin if bacteriuria was found by urine sediment analysis at admission [18]. At three-month follow-up, they showed a deep SSI rate of about 2.1% in the former group but only 0.45% in the latter group 2 (P=0.02). However, it was pointed out that ciprofloxacin might only delay the presentation of SSI to beyond the study's surveillance period, instead of the SSI rate [33,34].

Regarding urinary catheterization, it is a well-established risk factor for UTI, as catheters provide a direct conduit for rectal and peri-urethral microbes to reach the bladder [35]. The National Institute of Clinical Excellence in England defined catheter-associated UTI as the presence of symptoms or signs compatible with UTI in people with a catheter with no other identified source of infection, plus significant levels of bacteria in a catheter or a midstream urine specimen when the catheter has been removed within the previous 48 h [36]. Currently,

there is no standardization or consensus among hospitals about the protocol of urinary catheterization regarding antibiotic prophylaxis, duration of catheterization, and procedures of peri-urethral cleansing [37]. In the context of prosthetic joint infection, it has been assumed that catheter insertion could cause local trauma and hence haematogenous spreading of infection to prosthetic joint, but a systematic review on elective knee and hip arthroplasty concluded that there was no evidence for any association between urinary catheterization and an increased risk for prosthetic joint infection [38]. We found only one study investigating the relationship between urinary catheterization and SSI in hip fracture patients, which showed that deep wound infection was associated with more than two catheterizations and with prolonged catheter use (i.e. having >21 days of catheterization or being discharged home with a catheter in situ) [23].

Based on our findings and the benefit of treating UTI shown by Langenhan *et al.*, we recommend considering the possibility of perioperative UTI in hip fracture patients with treatment administered as necessary. As the surgical treatment for hip fracture could not be deferred until the definite diagnosis and clearance of UTI, the possibility of UTI in hip fracture patients has to be evaluated early and carefully, without causing any delay to the hip fracture's surgical management.

Evaluating hip fracture patients for UTI may be undertaken with targeted clinical history-taking, clinical examination, and routine urine evaluation before sending off the urine specimen for conventional urine culture in the laboratory. The evaluation on urine could be done by point-of-care tests of UTI, such as semi-automated urine analysers based on the level of colony-forming units/mL and culture-based devices, to speed up the diagnosis [39,40]. In the circumstances where these point-of-care tests are not available, alternatives include inspection of the urine for cloudiness, urine dipstick, and microscopy.

Table V Characteristics of included studies reporting on the relation between urinary catheterization and SSI rates

| Study | Design, centre, duration, country | Sample size | Age (mean (range) | Sex | Inclusion criteria, in addition to traumatic hip fracture treated surgically | Exclusion criteria, in addition to not meeting inclusion criteria | | Definition of SSI (timeframe of diagnosis) | Antibiotic prophylaxis for surgery and for catheterization | Having urinary catheter perioperatively or within 5 days of surgery | Having more than 2 episodes of urinary catheterization | Long-term cathet- erization ^a |
|----------------------------------|---|--|---|--|--|--|------------------|--|--|---|---|--|
| Cumming and Parker [23] | Retrospective case—control study, single centre, about 10 years (Jun 1994—Sep 2004), UK | patients with hip fracture. 18 patients | Among SSI: 79.8 (60–94) Among control: 81.9 (47–95) | SSI: M: 3 (16.7%), F: 15 (83.3%) Among control: M: 7 | group: for each patient | N/A | HA, THR, ORIF | Deep wound infection (used interchangeably with 'deep sepsis'), defined as presence of clinical evidence of infection below the deep fascia, with or without microbiological confirmation (1 year from injury) | Three doses of cefuroxime at the time of surgery and postoperatively. No prophylactic antibiotics as standard on insertion or removal of the catheter. | Deep sepsis: 12/18 (66.7%) No sepsis: 18/ 36 (50.0%) | Deep sepsis: 6/ 18 (33.3%) No sepsis: 3/36 (8.33%) | Deep sepsis: 7/18 (38.9%) No sepsis: 3/36 (8.33%) |

THR, total hip replacement; HA, hemiarthroplasty; ORIF, open reduction internal fixation.

a Long-term catheterization was defined as either catheterization of >21 days, or having a catheter *in situ* when discharged.

The urine specimen could be collected via a urinary catheter, if necessary due to patient's immobility, but the catheter should be removed as early as possible to reduce the risk of SSI associated with catheterization [23]. If point-of-care tests for UTI show positive results, or if the urine is cloudy, or if the dipstick and microscopy are positive, then antibiotics for UTI may be considered and continued until the results of the conventional urine culture in the laboratory become available. In deciding to initiate antibiotic treatment, it must also be considered that patients with reduced cognition may have difficulty expressing themselves when they suffer from UTI symptoms, so UTI could be under-diagnosed [24,41]. However, the consideration of treatment must be balanced against potential side-effects of antibiotics such as causing diarrhoea which could lead to wound contamination.

Regarding urinary catheterization and its relationship to SSI in hip fracture patients, we identified only one relevant article addressing this issue, so we recommend avoiding the risk factors reported in that study. These risk factors were more than two catheterizations and prolonged catheter use (>21 days of catheterization or being discharged from hospital with a catheter *in situ*) [23].

This study has several limitations, so the results should be interpreted with caution. There are differences among included studies, such as population demographics, surgical modalities and definitions or diagnostic criteria of SSI and UTI. Thus, although the odds ratio was statistically significant when comparing SSI rates in UTI and that in non-UTI groups, this did not hold when analysing the three studies that used a specific definition of symptomatic bacteriuria for UTI. Nevertheless, in the latter comparison, there was a clear trend of a higher risk of SSI with UTI (OR: 3.0; 95% CI: 0.55-16.26) and the nonsignificance may be related to the smaller number of studies available for analysis. Regarding SSI, studies have adopted different definitions and timeframes for diagnosis. In addition, all included studies are observational, with no high-quality randomized controlled trial available. Furthermore, the use of indwelling urinary catheters was not well documented in many studies, so we were unable to assess the relationship between urinary catheterization and SSI.

Future work in the form of high-quality randomized controlled trials comparing the SSI rates between treated and untreated preoperative UTI may provide evidence as to a causal link between the two. Further work on the relationship between urinary catheterization and SSI rates is also warranted. The relation between asymptomatic bacteriuria and SSI rates, which was not examined in this study, also remains to be determined.

In conclusion, our review has revealed an association of perioperative UTI with a higher risk of SSI among hip fracture patients treated surgically, but there is substantial heterogeneity among the available studies examined. Further work is needed to refine the diagnostic criteria of UTI, especially in patients with poor cognitive function, and to determine whether treatment of UTI reduces SSI rates. However, until such high-quality evidence is gathered, given the devastating effects of SSI on hip fracture patients, we recommend considering the possibility of perioperative UTI in this highly vulnerable patient group, with treatment administered as necessary to reduce SSI rates. UTI should also be considered when assessing and comparing SSI rates, either at a local hospital or wider healthcare level.

Author contributions

K.F.K.S.: Article screening and selection, data collection, statistical analysis, writing the original draft, reviewing and editing the manuscript. J.X.Y.L.: Article screening and selection, writing the original draft. C.P.C.: Conceptualization, statistical analysis, reviewing and editing the manuscript, supervision and mentorship. All authors made the final approval of the version to be published and agreed to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

Conflict of interest statement None declared.

Funding sources None.

References

- [1] Edwards C, Counsell A, Boulton C, Moran CG. Early infection after hip fracture surgery: risk factors, costs and outcome. J Bone Joint Surg Br 2008;90:770–7.
- [2] UK Health Security Agency. Surveillance of surgical site infections in NHS hospitals in England — April 2020 to March 2021. 2022. Available at: https://www.gov.uk/government/publications/ surgical-site-infections-ssi-surveillance-nhs-hospitals-in-england [last accessed June 2023].
- [3] Masters J, Metcalfe D, Ha JS, Judge A, Costa ML. Surgical site infection after hip fracture surgery: a systematic review and meta-analysis of studies published in the UK. Bone Joint Res 2020:9:554–62.
- [4] de Jong L, Klem T, Kuijper TM, Roukema GR. Factors affecting the rate of surgical site infection in patients after hemiarthroplasty of the hip following a fracture of the neck of the femur. Bone Joint J 2017;99-B:1088–94.
- [5] Dyck M, Embil JM, Trepman E, Bohm E. Surgical site infection surveillance for elective primary total hip and knee arthroplasty in Winnipeg, Manitoba, Canada. Am J Infect Control 2019;47:157–63.
- [6] Kurtz SM, Lau E, Schmier J, Ong KL, Zhao K, Parvizi J. Infection burden for hip and knee arthroplasty in the United States. J Arthroplasty 2008;23:984—91.
- [7] Gajdacs M, Abrok M, Lazar A, Burian K. Comparative epidemiology and resistance trends of common urinary pathogens in a tertiarycare hospital: a 10-year surveillance study. Medicina (Kaunas) 2019;55(7).
- [8] David TS, Vrahas MS. Perioperative lower urinary tract infections and deep sepsis in patients undergoing total joint arthroplasty. J Am Acad Orthop Surg 2000;8:66-74.
- [9] Sousa RJG, Abreu MA, Wouthuyzen-Bakker M, Soriano AV. Is routine urinary screening indicated prior to elective total joint arthroplasty? A systematic review and meta-analysis. J Arthroplasty 2019;34:1523—30.
- [10] Yoon RS, Mahure SA, Hutzler LH, Iorio R, Bosco JA. Hip arthroplasty for fracture vs elective care: one bundle does not fit all. J Arthroplasty 2017;32:2353—8.
- [11] Le Manach Y, Collins G, Bhandari M, Bessissow A, Boddaert J, Khiami F, et al. Outcomes after hip fracture surgery compared with elective total hip replacement. JAMA 2015;314:1159–66.
- [12] NHS. National. Tariff payment system 2022/23 Annex C: guidance on best practice tariffs. 2022. Available at: https://www.england.nhs.uk/wp-content/uploads/2020/11/22-23NT_Annex-C-Best-practice-tariffs.pdf [last accessed October 2022].
- [13] Department of Health. Payment by results guidance for 2010—11. Leeds: payment by results team. Department of Health; 2010.

- [14] Jacobsen SM, Stickler DJ, Mobley HL, Shirtliff ME. Complicated catheter-associated urinary tract infections due to *Escherichia coli* and *Proteus mirabilis*. Clin Microbiol Rev 2008;21:26—59.
- [15] Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. BMJ 2021;372:n71.
- [16] Craxford S, Marson BA, Nightingale J, Ikram A, Agrawal Y, Deakin D, et al. Deep infection after hip hemiarthroplasty: risk factors for infection and outcome of treatments. Bone Jt Open 2021;2:958-65.
- [17] Crouser N, Malik AT, Phieffer LS, Ly TV, Khan SN, Quatman CE. Urinary tract infection (UTI) at time of geriatric hip fracture surgery increases the risk of experiencing adverse 30-day outcomes. J Clin Orthop Trauma 2019;10:774—8.
- [18] Langenhan R, Bushuven S, Reimers N, Probst A. Peri-operative antibiotic treatment of bacteriuria reduces early deep surgical site infections in geriatric patients with proximal femur fracture. Int Orthop 2018;42:741—6.
- [19] Bliemel C, Buecking B, Hack J, Aigner R, Eschbach DA, Ruchholtz S, et al. Urinary tract infection in patients with hip fracture: an underestimated event? Geriatr Gerontol Int 2017;17:2369–75.
- [20] Yassa RR, Khalfaoui MY, Veravalli K, Evans DA. Pre-operative urinary tract infection: is it a risk factor for early surgical site infection with hip fracture surgery? A retrospective analysis. JRSM Open 2017;8:2054270416675083.
- [21] Capdevila A, Navarro M, Bori G, Tornero E, Camacho P, Bosch J, et al. Incidence and risk factors for infection when teicoplanin is included for prophylaxis in patients with hip fracture. Surg Infect (Larchmt) 2016;17:381—4.
- [22] Westberg M, Snorrason F, Frihagen F. Preoperative waiting time increased the risk of periprosthetic infection in patients with femoral neck fracture. Acta Orthop 2013;84:124—9.
- [23] Cumming D, Parker MJ. Urinary catheterisation and deep wound infection after hip fracture surgery. Int Orthop 2007;31:483—5.
- [24] Kostakopoulos NA, Karakousis ND, Moschotzopoulos D. Frailty associated urinary tract infections (FaUTIs). J Frailty Sarcopenia Falls 2021;6:9—13.
- [25] Seitz DP, Adunuri N, Gill SS, Rochon PA. Prevalence of dementia and cognitive impairment among older adults with hip fractures. J Am Med Dir Assoc 2011;12:556—64.
- [26] Mosk CA, Mus M, Vroemen JP, van der Ploeg T, Vos DI, Elmans LH, et al. Dementia and delirium, the outcomes in elderly hip fracture patients. Clin Interv Aging 2017;12:421–30.
- [27] Font-Vizcarra L, Zumbado A, Garcia S, Bosch J, Mensa J, Soriano A. Relationship between haematoma in femoral neck fractures contamination and early postoperative prosthetic joint infection. Injury 2011;42:200—3.
- [28] MacFie J. Current status of bacterial translocation as a cause of surgical sepsis. Br Med Bull 2004;71:1—11.

- [29] Doudakmanis C, Bouliaris K, Kolla C, Efthimiou M, Koukoulis GD. Bacterial translocation in patients undergoing major gastrointestinal surgery and its role in postoperative sepsis. World J Gastrointest Pathophysiol 2021;12:106—14.
- [30] Owrangi B, Masters N, Kuballa A, O'Dea C, Vollmerhausen TL, Katouli M. Invasion and translocation of uropathogenic Escherichia coli isolated from urosepsis and patients with communityacquired urinary tract infection. Eur J Clin Microbiol Infect Dis 2018;37:833—9.
- [31] Lewis AJ, Richards AC, Mulvey MA. Invasion of host cells and tissues by uropathogenic bacteria. Microbiol Spectr 2016;4(6).
- [32] Garofalo CK, Hooton TM, Martin SM, Stamm WE, Palermo JJ, Gordon JI, et al. *Escherichia coli* from urine of female patients with urinary tract infections is competent for intracellular bacterial community formation. Infect Immun 2007;75:52—60.
- [33] Probst A, Langenhan R. Letter to the Editor on 'Is routine urinary screening indicated prior to elective total joint arthroplasty? A systematic review and meta-analysis'. J Arthroplasty 2019;34:2193—4.
- [34] Sousa RJG, Abreu MA. Reply to the Letter to the Editor on 'Is routine urinary screening indicated prior to elective total joint arthroplasty? A systematic review and meta-analysis'. J Arthroplasty 2019;34:2194—5.
- [35] Werneburg GT. Catheter-associated urinary tract infections: current challenges and future prospects. Res Rep Urol 2022;14:109—33.
- [36] National Institute for Health and Care Excellence (NICE). NICE guideline [NG113] Urinary tract infection (catheter-associated): antimicrobial prescribing. 2018. Available at: https://www.nice. org.uk/guidance/ng113 [last accessed June 2023].
- [37] Gad MH, AbdelAziz HH. Catheter-associated urinary tract infections in the adult patient group: a qualitative systematic review on the adopted preventative and interventional protocols from the literature. Cureus 2021;13:e16284.
- [38] Roberts T, Smith TO, Simon H, Goodmaker C, Hing CB. Antibiotic prophylaxis for urinary catheter manipulation following arthroplasty: a systematic review. ANZ J Surg 2021;91:1405—12.
- [39] Thomas ST, Heneghan C, Price CP, Van den Bruel A, Plüddemann A. Horizon Scan Report 0045: point-of-care testing for urinary tract infections. 2016. Available at: https://www. community.healthcare.mic.nihr.ac.uk/reports-and-resources/ horizon-scanning-reports/point-of-care-testing-for-urinary-tractinfections [last accessed June 2023].
- [40] Arienzo A, Cellitti V, Ferrante V, Losito F, Stalio O, Murgia L, et al. A new point-of-care test for the rapid detection of urinary tract infections. Eur J Clin Microbiol Infect Dis 2020:39:325—32.
- [41] Woodford HJ, George J. Diagnosis and management of urinary tract infection in hospitalized older people. J Am Geriatr Soc 2009;57:107—14.