



Prevalence and drug resistance of *Escherichia coli* among patients with orthopaedic surgical site infections in China: A systematic review and meta-analysis

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ABSTRACT

To summarize current prevalence and drug resistance rate of *Escherichia coli* (*E. coli*) among orthopaedic surgical site infections (SSIs) in China from English and Chinese language sources. Online databases were searched to collect related researches. A meta-analysis was performed to analyse prevalence and 95 % confidence interval (CI) of *E. coli* among patients with orthopaedic surgical site infections. Meta-regression analysis was used to investigate the difference in the prevalence and antimicrobial resistance of *E. coli* among different subgroups. A total of 52 studies were enrolled into our meta-analysis, with a total of 31,285 strains isolated. The overall *E. coli* prevalence was 13.4 % (95 % CI 11.6–15.5). Study design ($R^2 = 8.98$) and sample size ($R^2 = 10.95$) might be potential sources of heterogeneity and there were no significant differences in risk of bias ($R^2 = 0.28$), study time ($R^2 < 0.01$), region ($R^2 = 2.46$) and hospital level ($R^2 = 1.42$). *E. coli* resistance were reported in 43 of the 52 papers. Antimicrobial resistance of *E. coli* to Ampicillin [87.9 % (95 % CI 83.7–91.1)] before 2015 was higher than that after 2015 [80.3 % (95 % CI 75.0–84.7)] ($R^2 = 30.93$, $P = 0.033$). While, resistance rate to Cefepime and Amikacin was significantly higher before 2015 ($R^2 = 17.25$ and 6.54 , $P = 0.043$ and 0.048), i.e., 46.4 % (36.3–56.9), 19.9 % (13.8–27.7) and 29.1 % (19.4–41.2), 8.6 % (4.4–16.2) in 2015 and after. It is necessary to carry out long-term monitoring to understand the actual prevalence and antimicrobial resistance of *E. coli* to develop appropriate health care mechanisms.

1. Introduction

Surgical site infection (SSI) is the most common complications of surgical patients. It not only brings psychological trauma and prolonged postoperative recovery time, but also increases mortality and medical expenses (Birhanu and Endalamaw, 2020). *Escherichia coli* (*E. coli*) is one of the most prevalent commensal inhabitants of human and the most important conditioned pathogen. Though it rarely causes disease, *E. coli* is responsible for a broad spectrum of diseases including various SSI, such as orthopaedic SSI (Li, 2022; Wang, 2022); laparotomy SSI (Huda et al., 2022), laparoscopic hysterectomy (Salmanov et al., 2022) and so

on (Nejad et al., 2021). In recent years, drug resistant *E. coli*, even multidrug resistant *E. coli* has gradually increased with the wide use of antibiotics in clinical treatment, which has caused great clinical difficulties and attracted more and more researchers' attention (Hou et al., 2020 Jul; Deka et al., 2020 Dec 21).

According to recent researches, the proportions of orthopaedic surgery patients with SSI are quite different, and *E. coli* is always one of the most significant related pathogens (Xu, 2021; Yu and Zheng, 2021; Chen et al., 2022). In 2014, the National Nosocomial Infection Surveillance System in China found that the total prevalence of SSIs *E. coli* was 20.75 % in all the nosocomial infections, and *E. coli* causes 26.92 % SSIs

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infections in all the 33 pathogens (Nan Ren and Anhua, 2016). And Zheng et al found drug resistant *E. coli* existed in orthopedic trauma, and multidrug resistant *E. coli* was also found in China (Zheng, 2021). Therefore, for an orthopedic surgeon, it is necessary to get a better understand of the epidemiological characteristics and antibiotic resistance of *E. coli* in orthopaedic SSI, which will help to develop more effective prevention and treatment measures.

The purpose of this systematic review and meta-analysis was to summarize the current prevalence and drug resistance rate of *E. coli* among orthopaedic SSIs in China from both English and Chinese language sources, and to provide further guidance for the prevention of SSIs.

2. Methods

We ran this meta-analysis in strict accordance with the guidelines of Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) Checklist (<https://www.prisma-statement.org/statement.htm>).

2.1. Search strategy

We used free words combined with subject terms to conducted a careful search in PubMed, Embase, Web of Science, China Biomedical Literature Database (CBM), China National Knowledge Infrastructure (CNKI), Wanfang and Weipu from January 2010 to July 2022. The search terms included surgery, postoperative, surgical wound infection,

site infection, *E. coli*, and China. All the retrieved papers were imported to EndNote (version 20), and duplicates were eliminated. All references included in the study were also reviewed.

2.2. Inclusion and exclusion criteria

Researches met the following criteria were included in the meta-analysis: (1) patients were clinically diagnosed with orthopaedic SSIs; (2) the study subjects were Chinese; and (3) researches were with sufficient data to calculate the prevalence or antimicrobial resistance of *E. coli*, for example the number of *E. coli* strain or resistant *E. coli* strain isolated. The following exclusion criteria were applied: (1) abstracts, reviews, or communication papers; (2) studies with a small number of detected bacterial strains (at least 200 isolated bacterial strains); (3) lack of sufficient information, including incomplete or unavailable research data, for example the number of *E. coli* strain or resistant *E. coli* strain isolated; and (4) research based on data from the National Nosocomial Infection Surveillance System (NNISS). For multiple reports on the same trial, the one with more details and higher quality was selected for analysis.

2.3. Data extraction and bias risk assessment

Data were independently extracted by 2 researchers (Luqiao Ou and Gaoming Li) using a unified data table. Author, publication year, study area, research design, hospital level, number of SSIs patients, number of bacterial, *E. coli*, resistant *E. coli* strains isolated, Resistance of drug-

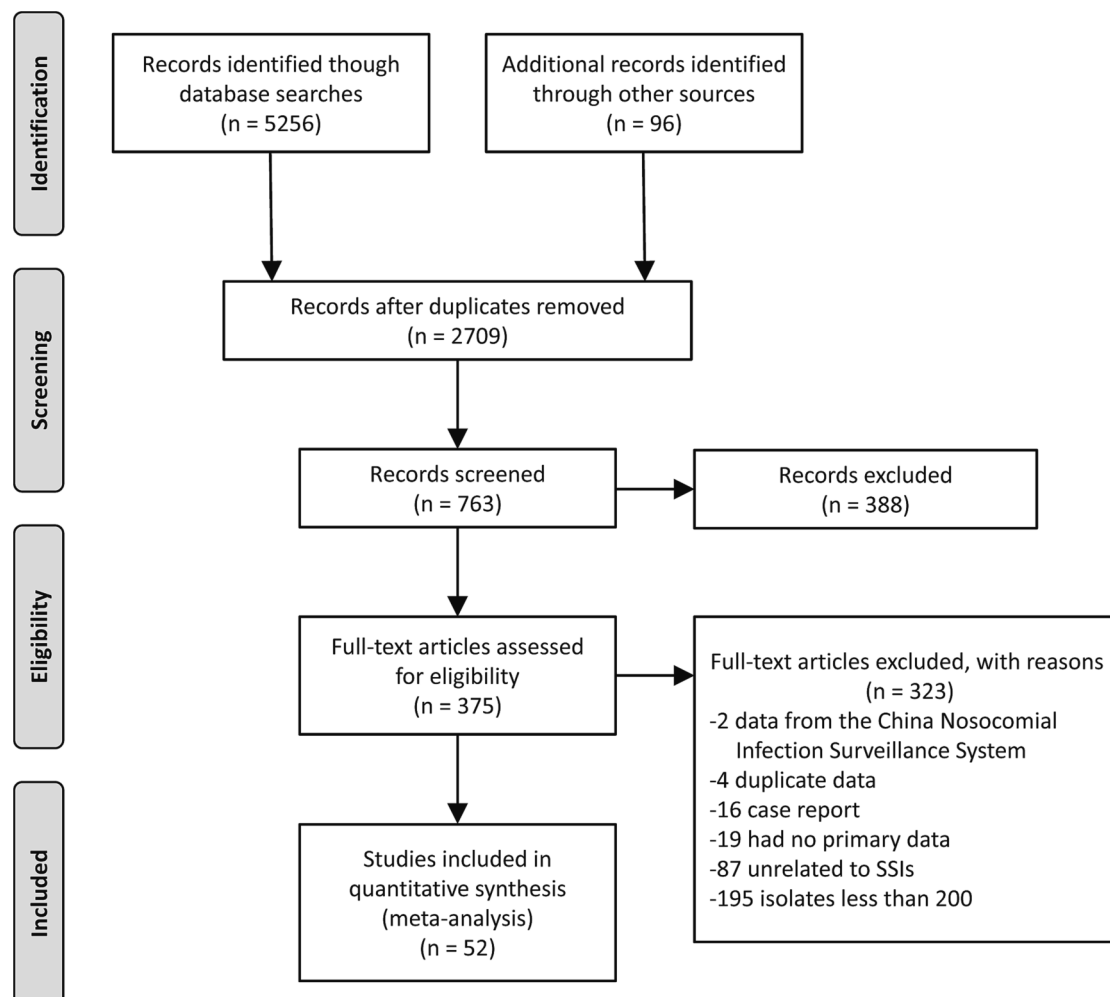


Fig. 1. The flowchart showed the selection of studies for meta-analysis.

Table 1
Characteristics of included studies.

Study ID	Year	Province	Study type	Hospital level	Number of SSIs patients	Isolates	<i>Escherichia coli</i>	Risk of bias
Feng 2012	2009–10	Zhejiang	Retrospective	Tertiary	895	625	140	High
Wang 2012	2009–11	Tianjin	Monitoring	Tertiary	Unclear	1882	99	Low
Wang 2012	2009–11	Jiangxi	Retrospective	Non-tertiary	269	269	46	Low
Gao 2013	2010–12	Hebei	Retrospective	Tertiary	999	1056	86	Low
Han 2013	2010–11	Shanxi	Retrospective	Tertiary	Unclear	619	62	High
Pan 2013	2009–11	Zhejiang	Retrospective	Non-tertiary	Unclear	694	108	High
Shao 2013	2009–11	Zhejiang	Monitoring	Non-tertiary	233	240	51	Low
Cao 2014	2011–13	Shanxi	Retrospective	Non-tertiary	365	210	40	Low
Gao 2014	2008–12	Hebei	Monitoring	Tertiary	Unclear	2456	188	Low
Liu 2014	2008–12	Beijing	Monitoring	Tertiary	587	387	31	Low
Liu 2014	2014	Hunan	Retrospective	Tertiary	243	243	44	High
Xue 2014	2012–13	Henan	Retrospective	Tertiary	946	946	227	Low
Zheng 2014	2012–13	Henan	Retrospective	Tertiary	651	407	86	High
Su 2015	2013–14	Guangdong	Monitoring	Non-tertiary	3317	603	21	Low
Wang 2015	2014	Shandong	Retrospective	Tertiary	360	404	116	High
Dong 2016	2014–15	Hebei	Retrospective	Tertiary	2106	2106	238	Low
Li 2016	2010–14	Xinjiang	Retrospective	Tertiary	576	615	84	Low
Wang 2016	2011–13	Beijing	Retrospective	Tertiary	Unclear	1375	105	Low
Chen 2017	2014–15	Jiangsu	Retrospective	Tertiary	80	200	26	Low
Guo 2017	2014–15	Henan	Retrospective	Non-tertiary	285	285	78	Low
Jin 2017	2014–16	Zhejiang	Retrospective	Non-tertiary	232	296	55	Low
Li 2017	2012–17	Zhejiang	Retrospective	Tertiary	413	281	34	Low
Liu 2017	2012–17	Zhejiang	Retrospective	Non-tertiary	70	327	71	Low
Zeng 2017	2016–17	Guangxi	Retrospective	Non-tertiary	291	341	47	High
Sun 2018	2016–17	Zhejiang	Retrospective	Tertiary	130	Unclear	130	Low
Sun 2018	2014–16	Zhejiang	Retrospective	Non-tertiary	4903	2302	229	Low
Sun 2018	2015–17	Jiangxi	Retrospective	Tertiary	216	270	36	High
Tan 2018	2017	Hubei	Retrospective	Non-tertiary	280	330	45	Low
Xie 2018	2012–16	Yunnan	Retrospective	Tertiary	295	345	62	High
Yang 2018	2015–17	Tianjin	Retrospective	Tertiary	2596	2471	405	Low
Zhang 2018	2015–17	Tianjin	Retrospective	Tertiary	1269	443	38	Low
Zhou 2018	2017–18	Zhejiang	Retrospective	Tertiary	106	251	112	High
Hu 2019	2015–18	Jiangxi	Retrospective	Non-tertiary	36	206	65	Low
Yu 2019	2017–18	Jiangxi	Retrospective	Non-tertiary	72	382	20	Low
Zhao 2019	2017–18	Jilin	Retrospective	Tertiary	Unclear	437	35	High
Chai 2020	2015–18	Henan	Retrospective	Non-tertiary	263	342	25	High
Gong 2020	2015–18	Sichuan	Retrospective	Tertiary	1020	1020	176	Low
Jiang 2020	2017–20	Henan	Retrospective	Non-tertiary	212	236	58	Low
Lin 2020	2016–19	Guangdong	Monitoring	Non-tertiary	409	228	35	Low
Lu 2020	2014–16	Anhui	Retrospective	Non-tertiary	200	213	45	High
Zhang 2020	2017–18	Zhejiang	Retrospective	Tertiary	Unclear	397	21	High
Wang 2021	2020	Liaoning	Monitoring	Tertiary	Unclear	811	56	Low
Wang 2021	2015–19	Henan	Retrospective	Non-tertiary	562	587	92	Low
Xu 2021	2019	Gansu	Retrospective	Tertiary	430	252	42	High
Yang 2021	2017–19	Henan	Retrospective	Tertiary	76	258	62	Low
Ye 2021	2018–19	Fujian	Retrospective	Tertiary	296	296	46	High
Yi 2021	2019–21	Hunan	Retrospective	Tertiary	200	200	10	High
Yu 2021	2015–18	Guangdong	Retrospective	Tertiary	516	516	26	Low
Zheng 2021	2020	Fujian	Retrospective	Tertiary	890	647	37	Low
Chen 2022	2018–19	Henan	Retrospective	Tertiary	265	290	78	Low
Li 2022	2019–21	Henan	Retrospective	Tertiary	95	463	22	Low
Wang 2022	2018–21	Henan	Retrospective	Non-tertiary	370	225	40	Low

resistant *Escherichia coli*, and other essential data were collected as main data. Specified diagnostic criteria of original diseases and SSIs, specified criteria of inclusion, case source, case selection method, specified test for pathogenic bacteria, pathogenic test for all included cases and study type were collected to assess the risk of bias. Any disagreement was solved through discussion or consultation with Bao-chuang Qi.

All the included studies were assessed according to predetermined criteria extracted and modified from a previous case series scale consisting of 9 items. By answering low, high, or unclear to questions, bias can be identified in selected literature. The total score ranges from 0 to 9 points, and the higher the score is, the higher the quality. In this study, R4.1.3 software was used to summarize the risk of bias. A score less than 4 points was defined as high risk, and a score greater than or equal to 4 points was defined as low risk.

2.4. Statistical analysis

All statistical analyses were performed using R4.1.3. Statistical tests were all two-tailed. Unless stated, $P < 0.05$ was considered statistically significant. The prevalence of *E. coli* and antimicrobial-resistant *E. coli* isolates among SSI patients in each study was calculated using the following formula:

$$\text{Prevalence of } E. coli = \frac{\text{Number of } E. coli \text{ isolates}}{\text{Number of all the detected isolates}} \times 100\%$$

$$\begin{aligned} \text{Prevalence of antimicrobial-resistant } E. coli \\ = \frac{\text{Number of detected } E. coli \text{ isolates resistant to a given antibiotic}}{\text{Number of } E. coli \text{ isolates detected}} \\ \times 100\% \end{aligned}$$

The prevalence in each study and its 95 % CI were calculated using logit transformation.

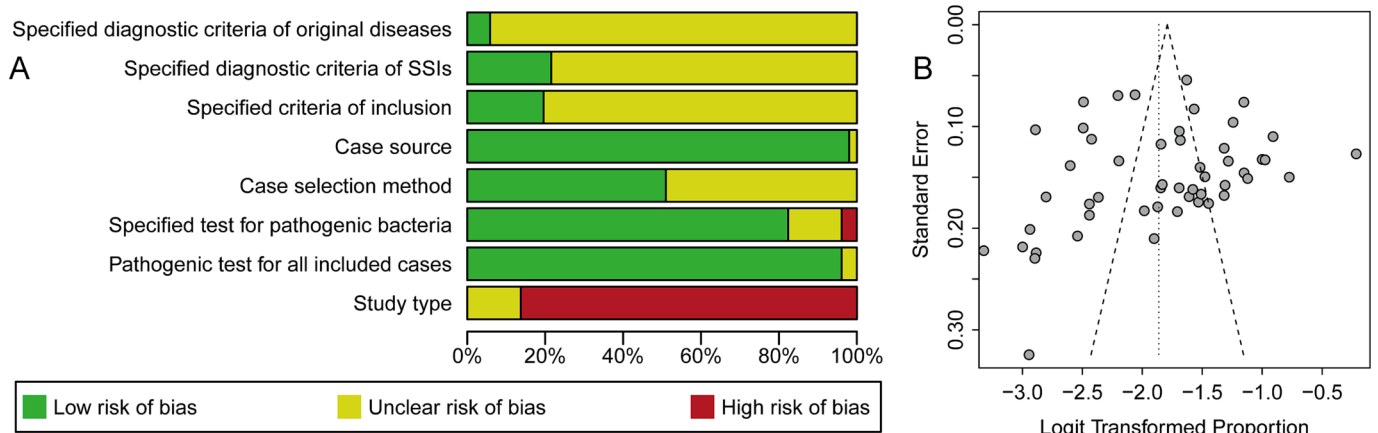


Fig. 2. Risk of bias graph and Funnel plots of included studies. A, Risk of bias graph. B, Funnel plots of included studies.

Test of heterogeneity for each outcome was carried out using Cochran’s Q test and Higgins I-squared statistic. P value (Q test) < 0.05 and/or $I^2 > 50\%$ indicated substantial heterogeneity, and a random effect model (Der Simonian and Laird method) was applied accordingly. In contrast, a fixed effect model (inverse variance method) was employed in the presence of mild or low heterogeneity ($P \geq 0.05$ and/or $I^2 < 50\%$). Publication bias was evaluated by Egger’s linear regression using the funnel plot.

Subgroup analysis and univariate meta-regression were used to evaluate the differences in *E. coli* prevalence and antimicrobial resistance rate. In meta-regression analysis, the dependent variable was *E. coli* prevalence or *E. coli* antibiotic resistance data. The independent variables were region (dummy variable: Eastern region), hospital level (dummy variable: Tertiary), risk of bias (dummy variable: High), study design (dummy variable: Retrospective), sample size (dummy variable: <500 isolates), and study time (dummy variable: Before 2015). In meta-regression analysis, restricted maximum likelihood method was used to estimate the variance between studies, and the proportion of variance explained by any meta-regression model was estimated using R^2 statistic.

3. Results

We retrieved 5256 articles from online database and 96 from other sources. A total of 2643 duplicates were excluded from the initial articles and 375 relevant full texts were obtained according to our criteria. After examining the publication types, titles, as well as abstracts, 52 studies (Li, 2022; Wang, 2022; Xu, 2021; Yu and Zheng, 2021; Chen et al., 2022; Zheng, 2021; Feng and Mao, 2012; Wang et al., 2012; Wang, 2012; Gao et al., 2013; Han and Gao, 2013; Pan et al., 2013; Shao and He, 2013; Cao, 2014; Gao et al., 2014; Liu and Fang, 2014; Liu et al., 2014; Xue et al., 2014; Zheng, 2014; Su et al., 2015; Wang et al., 2015; Dong and Zhang, 2016; Li et al., 2016; Wang et al., 2016; Chen, 2017; Guo, 2017; Jin et al., 2017; Li et al., 2017; Liu et al., 2017; Zeng, 2017; Sun et al., 2018; Sun et al., 2018; Sun et al., 2018; Tan and Ye, 2018; Xie et al., 2018; Yang et al., 2018; Zhang and Yue, 2018; Zhou et al., 2018; Hu, 2019; Yu and Hu, 2019; Zhao et al., 2019; Chai, 2020; Gong et al., 2020; Jiang et al., 2020; Lin et al., 2020; Lu and Yang, 2020; Zhang et al., 2020; Wang et al., 2021; Wang et al., 2021; Yang and Wang, 2021; Ye et al., 2021; Yi and Huang, 2021) were enrolled into our meta-analysis after checking research details. Later, all the references in the 52 studies were checked in case of missing eligible records, but no additional record was found. Fig. 1 showed the flow diagram of this study.

The characteristics of 52 included studies are shown in Table 1. Among them, 51 studies reported the proportion of *E. coli* among SSIs, and bacterial strains isolated ranged from 200 to 2471, with a total of 31,285 strains isolated. Another 1 study reported antimicrobial resistance data for *E. coli*. The risk of bias assessment data is shown in Fig. 2A,

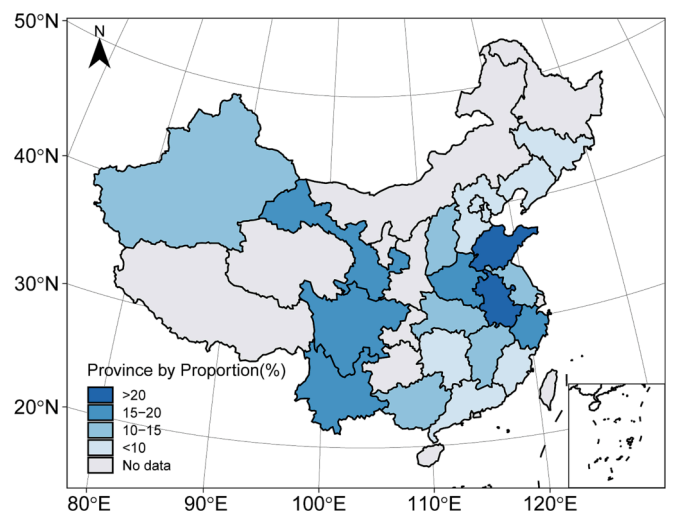


Fig. 3. Geographic distribution of *E. coli* prevalence among patients with orthopaedic surgical site infections in China.

and the details are shown in Table S1. There were 34 high-quality studies (≥ 4), the highest score was 6, and the lowest score was 2 (Table S1). All studies were conducted between 2012 and 2022, and 21 of the 34 provinces in China were represented (Fig. 3). Among them, 25 studies were from the eastern coastal region, 22 from the central region and 5 were from the western region. Most studies were retrospective (45 of 52).

The prevalence of *E. coli* included ranged from 3.5 % (95 % CI 2.3–5.3) to 44.6 % (95 % CI 38.6–50.8) (Fig. 4), indicating significant heterogeneity among these studies. The Higgin’s I^2 value was 95.8 (Q test $p < 0.001$), and the overall *E. coli* prevalence was 13.4 % (95 % CI 11.6–15.5). The prevalence was 11.6 % (95 % CI 9.3–14.5) in the eastern region, 15.4 % (95 % CI 12.5–18.9) in the central region and 15.9 % (95 % CI 14.2–17.8) in the western region. The prevalence of *E. coli* exceeded 20 % in the 2 provinces: 28.7 % (95 % CI 24.5–33.3) in Shandong and 21.1 % (95 % CI 16.2–27.1) in Anhui. Univariate random effects meta-regression analysis indicated that the study design ($R^2 = 8.98$) and sample size ($R^2 = 10.95$) might be potential sources of heterogeneity and there were no significant differences in risk of bias ($R^2 = 0.28$), study time ($R^2 < 0.01$), region ($R^2 = 2.46$) and hospital level ($R^2 = 1.42$) (Table 2). Egger’s test indicated no evidence of publication bias regarding the total prevalence of *E. coli* ($t = -0.727, p = 0.471$; Fig. 2B).

We further analyzed *E. coli* resistance to different antibiotics, and 43 of the 52 papers reported *E. coli* resistance to 56 antibiotics. Among

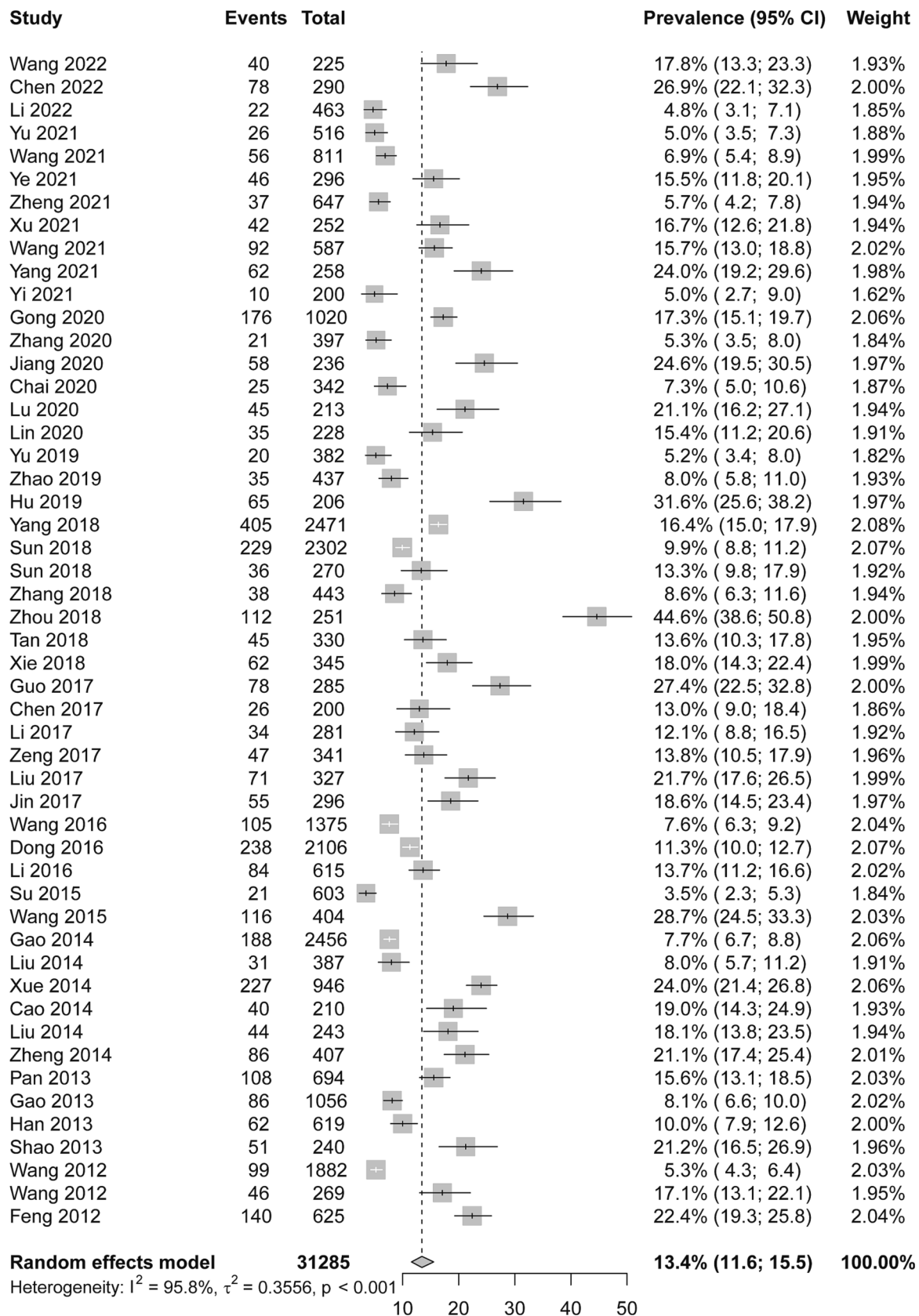


Fig. 4. The prevalence of *E. coli* among patients with orthopaedic surgical site infections.

Table 2
Prevalence of *E. coli* in different subgroups of patients with orthopedic SSI.

Group	Prevalence of <i>Escherichia coli</i>					Univariate meta-regression			
	Studies	Isolates	<i>Escherichia coli</i>	Estimate	I^2	β (SE)	OR (95 % CI)	p value	R^2
Region									
Eastern region	25	21,294	2374	11.6(9.3–14.5)	96.9	Ref	Ref	Ref	2.46
Central region	21	7418	1216	15.4(12.5–18.9)	93.0	0.322(0.199)	1.380(0.933–2.040)	0.106	
Western region	5	2573	411	15.9(14.2–17.8)	33.5	0.356(0.328)	1.428(0.751–2.713)	0.277	
Hospital Level									
Tertiary	32	22,969	2830	12.4(10.1–15.0)	96.6	Ref	Ref	Ref	1.42
Non-tertiary	19	8316	1171	15.5(12.5–19.0)	93.1	0.256(0.196)	1.292(0.880–1.896)	0.192	
Risk of bias									
High	17	6336	1037	15.1(11.7–19.4)	94.6	Ref	Ref	Ref	0.28
Low	34	24,949	2964	12.7(10.6–15.1)	96.0	−0.203(0.202)	0.816(0.549–1.214)	0.316	
Study Design									
Retrospective	44	24,678	3520	14.5(12.5–16.7)	95.0	Ref	Ref	Ref	8.98
Non-retrospective ^b	7	6607	481	8.4(5.7–12.2)	94.2	−0.614(0.265)	0.541(0.322–0.910)	0.021	
Sample size ^c									
<500	33	9954	1622	15.6(13.0–18.6)	93.3	Ref	Ref	Ref	10.95
≥500	18	21,331	2379	10.3(8.2–12.9)	96.9	−0.477(0.188)	0.621(0.430–0.897)	0.011	
Year									
Before 2015	26	19,386	2372	14.1(11.6–17.2)	96.2	Ref	Ref	Ref	<0.01
2015 or after	25	11,899	1629	12.7(10.1–15.9)	95.4	−0.123(0.192)	0.884(0.607–1.288)	0.521	

them, 20 antibiotics were reported in 10 or more studies (Table 3, Fig. 5). We performed a meta-analysis of these antibiotics and the antimicrobial resistance rates for *E. coli* were also compared before and after 2015. The results indicated that compared with data before 2015 [80.3 % (95 % CI 75.0–84.7)], antimicrobial resistance of *E. coli* to Ampicillin [87.9 % (95 % CI 83.7–91.1)] was significantly higher in studies after 2015 ($R^2 = 30.93$, $P = 0.033$). While, the rate of resistance to Cefepime and Amikacin was significantly higher before 2015 ($R^2 = 17.25$ and 6.54 , $P = 0.043$ and 0.048), i.e., 46.4 % (36.3–56.9), 19.9 % (13.8–27.7) before 2015 and 29.1 % (19.4–41.2), 8.6 % (4.4–16.2) in 2015 and after. In addition, *E. coli* resistance to Cefotaxime, Ceftazidime and Tobramycin showed decreasing trends, and *E. coli* resistance to Ceftriaxone showed increasing trends, but the differences were not statistically significant. Notably, *E. coli* resistance to ampicillin exceeded 80.0 %.

4. Discussion

In this study, we conducted a meta-analysis of *E. coli* epidemiological characteristics and antimicrobial resistance among orthopaedic SSIs in China. The total prevalence of *E. coli* among SSIs (13.4 %) was significantly lower than the NNISS data (20.75 %) (Nan Ren and Anhua, 2016), and we believe the discrepancy may be caused by the following factors. First, the NNISS data include all nosocomial infections, and the number of bacteria isolated from SSIs was 1638, far less than the 31,285 strains in our study. In addition, the 52 studies included in our study involved more than 62 hospitals distributed in 21 different provinces, including tertiary hospitals and nontertiary hospitals. Second, the studies included in this study were all conducted after 2010, and some (39) were even completed after 2015. However, the NNISS data were collected in 2014. Over time, *E. coli* prevalence among SSI samples may fluctuate, leading to the difference between our results and the NNISS data.

E. coli is resistant to a large number of antimicrobial drugs (Kresken et al., 2023), and empirical treatment is dependent on microbiological test results. However, the high resistance of pathogen to antibiotics may increase the possibility of inappropriate empirical treatment, resulting in poor clinical outcomes and increased financial burden. Therefore, in addition to taking appropriate measures in a timely manner and following the antibiotics using guidelines, it is also necessary to design an empirical treatment plan based on antimicrobial susceptibility tests. In addition to intrinsic antimicrobial resistance, *E. coli* can also acquire antimicrobial resistance through other mechanisms and develop into multidrug-resistant or pandrug-resistant bacteria, leading to life-threatening serious infections (Kresken et al., 2023; Nkansa-Gyamfi et al., 2019). Our results showed that the resistance of *E. coli* to

Ampicillin was significantly increased, and the resistance to Cefepime and Amikacin was decreased, which provided a reference for clinical prevention and control of *E. coli* infection among SSIs.

In this study, the *E. coli* resistance rate to the third generation cephalosporins, ceftriaxone (65.6 %, 2015 or after) and cefotaxime (46.8 %, 2015 or after), was slightly different from that reported by the Chinese Antimicrobial Resistance Surveillance System (CARSS) in 2020 (51.6 %) (<https://www.carss.cn/Report/Details?aId=808>). The *E. coli* resistance rate to carbapenems, including imipenem (2.5 %, 2015 or after) and meropenem (2.5 %, 2015 or after) is comparable to the CARSS data in 2020 (1.6 %). And when comes to quinolone drugs, including levofloxacin (48.4 %, 2015 or after) and ciprofloxacin (54.0 %, 2015 or after), the situation is also similar (the CARSS data 50.7 %).

This study has the following limitations. First, the heterogeneity among the included studies was considerable, subgroup analysis and meta-regression analysis could not fully explain the heterogeneity. Second, studies with small sample size are prone to producing accidental results, and neglect of these studies may result in a lack of some potential important data. Third, the majority of regions included were the central and eastern coastal areas of China, and SSI monitor was more popularized. Therefore, the estimated *E. coli* prevalence may not reflect the overall situation in China. Fourth, SSIs sample collection, strains identification including *E. coli* detection, antimicrobial resistance evaluation methods may be not strictly consistent in collected researches, for example the strains identification and antimicrobial resistance evaluation were completed by different equipment or even by artificial approach, which would lead to a different frequency. Finally, we only described the antimicrobial resistance status of *E. coli* in SSIs without covering related mechanism, further research might as well pay attention to the expression of drug resistance genes, which would also be reference for medication strategies and prognostic indicator selection (Halaji et al., 2022; Halaji et al., 2020; Halaji et al., 2020).

In summary, *E. coli* prevalence and antimicrobial resistance vary with region and time, which need to be monitored at all times. Compared with the NNISS data, *E. coli* prevalence among SSIs in this study was lower. Therefore, it is necessary to carry out long-term monitoring to understand the actual prevalence and antimicrobial resistance of *E. coli* to develop appropriate health care mechanisms. According to findings of the current research, the antibiotic therapy concerning orthopaedic SSIs may require minor adjustments in China. We therefore recommend initiating appropriate infection prevention measures, strengthening existed antimicrobial stewardship programmes, and conducting regular antimicrobial surveillance to prevent antimicrobial-resistant *E. coli* infection in hospitals.

Table 3
Combined prevalence of drug-resistant *E. coli* at different times.

	Year	Prevalence of resistant <i>Escherichia coli</i>				Univariate meta-regression				
		Studies	<i>Escherichia coli</i>	Resistant <i>Escherichia coli</i>	Estimate	<i>I</i> ²	β (SE)	OR (95 % CI)	p value	<i>R</i> ²
Ampicillin	Before 2015	9	538	439	80.3 (75.0–84.7)	37.2	Ref	Ref	Ref	30.93
	2015 or after	16	1035	911	87.9 (83.7–91.1)	61.1	0.506 (0.238)	1.659 (1.041–2.644)	0.033	
Ampicillin-sulbactam	Before 2015	4	309	165	52.5 (26.1–77.7)	95.0	Ref	Ref	Ref	<0.01
	2015 or after	6	333	161	38.5 (25.3–53.7)	81.7	−0.594 (0.572)	0.552 (0.180–1.694)	0.299	
Aztreonam	Before 2015	11	767	316	40.8 (34.1–47.9)	69.7	Ref	Ref	Ref	<0.01
	2015 or after	11	644	299	46.3 (35.9–57.1)	79.6	0.216 (0.293)	1.242 (0.699–2.205)	0.460	
Cefazolin	Before 2015	9	915	607	63.2 (46.7–77.1)	94.2	Ref	Ref	Ref	<0.01
	2015 or after	11	723	508	71.1 (59.9–80.3)	87.3	0.380 (0.431)	1.462 (0.629–3.400)	0.378	
Cefepime	Before 2015	13	1216	581	46.4 (36.3–56.9)	91.0	Ref	Ref	Ref	17.25
	2015 or after	15	771	236	29.1 (19.4–41.2)	86.4	−0.748 (0.371)	0.473 (0.229–0.978)	0.043	
Cefoperazone-sulbactam	Before 2015	6	634	103	17.6 (9.1–31.4)	90.1	Ref	Ref	Ref	
	2015 or after	10	624	64	7.5(3.7–14.7)	79.4	−0.944 (0.588)	0.389 (0.123–1.232)	0.108	
Cefotaxime	Before 2015	7	669	373	56.6 (48.5–64.3)	73.3	Ref	Ref	Ref	11.82
	2015 or after	7	280	121	46.8 (34.9–59.1)	72.9	−0.414 (0.285)	0.661 (0.378–1.155)	0.146	
Ceftazidime	Before 2015	16	1322	541	43.6 (34.6–53.0)	89.7	Ref	Ref	Ref	2.85
	2015 or after	17	959	339	32.8 (24.4–42.5)	86.5	−0.457 (0.295)	0.633 (0.355–1.128)	0.121	
Ceftriaxone	Before 2015	8	412	216	51.8 (39.5–63.9)	82.8	Ref	Ref	Ref	7.79
	2015 or after	15	973	642	65.6 (55.5–74.4)	87.9	0.574 (0.353)	1.776 (0.889–3.544)	0.104	
Cefuroxime	Before 2015	5	670	433	66.7 (52.2–78.7)	91.8	Ref	Ref	Ref	<0.01
	2015 or after	7	420	242	60.5 (48.7–71.2)	80.5	−0.265 (0.358)	0.767 (0.381–1.548)	0.460	
Imipenem	Before 2015	18	1442	40	3.3(2.1–5.3)	43.9	Ref	Ref	Ref	<0.01
	2015 or after	21	1178	89	2.5(0.7–8.0)	90.4	0.253 (0.561)	1.288 (0.428–3.869)	0.653	
Meropenem	Before 2015	12	1125	43	2.2(0.8–6.1)	85.3	Ref	Ref	Ref	<0.01
	2015 or after	11	579	18	2.5(0.5–11.2)	84.7	0.285 (0.824)	1.330 (0.265–6.692)	0.729	
Piperacillin	Before 2015	14	1394	886	64.3 (51.2–75.5)	94.1	Ref	Ref	Ref	<0.01
	2015 or after	10	440	247	62.0 (51.7–71.3)	69.1	−0.134 (0.423)	0.875 (0.382–2.006)	0.752	
Piperacillin-tazobactam	Before 2015	9	939	97	8.3(4.6–14.4)	83.8	Ref	Ref	Ref	<0.01
	2015 or after	10	481	18	5.6(2.9–10.6)	41.1	−0.504 (0.493)	0.604 (0.230–1.589)	0.307	
Amikacin	Before 2015	15	1319	278	19.9 (13.8–27.7)	87.9	Ref	Ref	Ref	6.54
	2015 or after	15	853	85	8.6(4.4–16.2)	85.2	−0.869 (0.439)	0.419 (0.177–0.992)	0.048	
Gentamycin	Before 2015	17	1467	734	48.9 (42.8–55.0)	79.3	Ref	Ref	Ref	<0.01
	2015 or after	15	814	376	45.1 (39.6–50.8)	54.9	−0.165 (0.187)	0.848 (0.588–1.223)	0.378	
Tobramycin	Before 2015	5	396	164	44.6 (32.4–57.6)	83.0	Ref	Ref	Ref	15.35
	2015 or after	10	747	203	26.2 (17.4–37.4)	87.3	−0.824 (0.429)	0.439 (0.189–1.017)	0.055	
Ciprofloxacin	Before 2015	12	1210	711	58.7 (49.8–67.0)	87.8	Ref	Ref	Ref	<0.01
	2015 or after	15	866	458	54.0 (46.6–61.2)	74.9	−0.192 (0.254)	0.825 (0.501–1.359)	0.450	

(continued on next page)

Table 3 (continued)

	Year	Prevalence of resistant <i>Escherichia coli</i>				Univariate meta-regression				
		Studies	<i>Escherichia coli</i>	Resistant <i>Escherichia coli</i>	Estimate	<i>I</i> ²	β (SE)	OR (95 % CI)	p value	<i>R</i> ²
Levofloxacin	Before 2015	11	868	445	48.3 (40.8–55.8)	77.2	Ref	Ref	Ref	<0.01
	2015 or after	16	860	411	48.4 (37.7–59.1)	86.8	0.027 (0.344)	1.028 (0.524–2.017)	0.937	
Bactrim	Before 2015	7	507	322	63.9 (54.1–72.7)	73.8	Ref	Ref	Ref	<0.01
	2015 or after	8	435	272	63.3 (56.2–69.8)	45.3	–0.008 (0.270)	0.992 (0.585–1.684)	0.977	

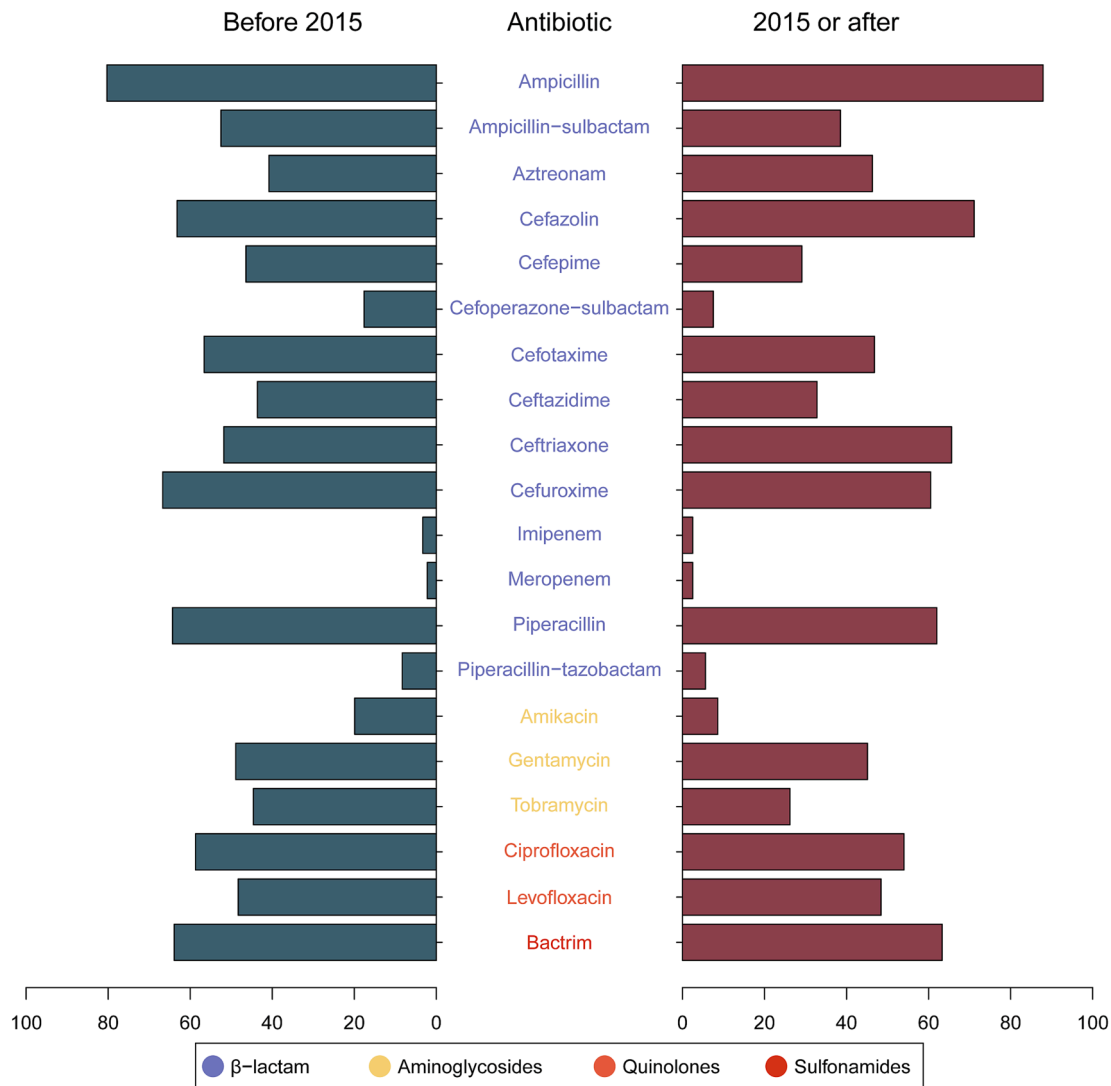


Fig. 5. Antibiotic resistance rates for *E. coli* for different subgroups of years.

CRedit authorship contribution statement

Luqiao Pu: Methodology, Validation, Formal analysis, Investigation, Resources, Data curation, Writing – original draft, Visualization. **Gaoming Li:** Methodology, Validation, Formal analysis, Investigation, Resources, Data curation, Visualization. **Baochuang Qi:** Methodology, Validation, Formal analysis, Investigation, Resources, Data curation, Visualization. **Chuan Li:** Writing – original draft. **Pengfei Bu:** Writing – original draft. **Yapin Li:** Writing – original draft. **Ze Xu:** Writing – original draft. **Yan Bai:** Writing – review & editing. **Dehong Yin:**

Writing – review & editing. **Jian Wang:** Conceptualization, Supervision, Project administration. **Yongqing Xu:** Conceptualization, Funding acquisition, Supervision, Project administration.

Declaration of Competing Interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

Data availability

Data will be made available on request.

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Appendix A. Supplementary data

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