

# Meta-analysis and systematic review of electronic bronchoscopy in refractory pneumonia

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**Background:** A systematic review and meta-analysis were performed to explore the adoption value of electronic bronchoscopy (EBS) in the diagnosis and treatment of refractory pneumonia and to provide a theoretical basis for the clinical bronchoscopy treatment of patients.

**Methods:** Randomized controlled trials of treatment-resistant pneumonia searched on PubMed, Embase, and other websites before December 31, 2020 were collected. Literature was selected by inclusion and exclusion criteria. Bias risk was assessed using Cochrane intervention system Review manual 5.0.2 and Review Manager 5.3.

**Results:** A total of 6 articles meeting the requirements were included, comprising 796 participants. The results of meta-analysis showed that the clinical efficacy [odds ratio (OR) =3.8; 95% confidence interval (CI): 1.11–12.99; Z=2.13; P =0.03] and white blood cell counts [mean difference (MD) =0.55; 95% CI: -0.57 to 1.67; Z=0.96; P=0.34] of patients both increased. The mortality rate (OR =0.7; 95% CI: 0.3–1.63; Z=0.82; P=0.41), the incidence of infection (OR =0.84; 95% CI: 0.5–1.39; Z=0.69; P=0.49), ICU hospitalization days (OR =0.59; 95% CI: 0.36–0.98; Z=2.04; P=0.04), days of antibiotic use (OR =0.39; 95% CI: 0.18–0.84; Z=2.41; P=0.02), body temperature (MD =–0.2; 95% CI: -0.24 to 0.16; Z=9.5; P<0.0001), and the PaO<sub>2</sub>:FIO<sub>2</sub> ratio (MD =–9.96; 95% CI: -13.31 to –6.61; Z=5.83; P<0.0001) of patients in the experimental group were lower than those of the control group. Differences in white blood cell count, mortality rate, and incidence of infection of patients in the experimental group were not statistically significant compared with those in the control group.

**Discussion:** EBS adopted in the diagnosis and treatment of refractory pneumonia can reduce the occurrence of postoperative adverse reactions, reduce the infection rate, and effectively improve the clinical symptoms. It is therefore suitable for the treatment of this disease.

**Keywords:** Electronic bronchoscopy (EBS); refractory pneumonia; white blood cell count; infection; clinical efficacy

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# Introduction

Refractory pneumonia, as compared to general pneumonia, refers severe pneumonia with no significant improvement or even worsening after anti-infection treatment, along with a poor prognosis (1). Refractory pneumonia is generally treated with macrolide antibiotics. Patients who are resistant to macrolides or stop taking them before they are cured may experience a relapse, even worse than before treatment. Patients may develop lung abscesses, atelectasis, lung necrosis, and pleural effusion. These diseases lead to a very unsatisfactory lung condition, which is prone to deterioration and should be diagnosed as refractory pneumonia. Refractory pneumonia is a disease that is difficult to treat and can have serious consequences if it occurs, and it is difficult to predict the disease in advance because of its unobvious detection symptoms. Therefore, it is very important to find a way to diagnose refractory pneumonia at an early stage (2). With the increase in the rate of bacterial resistance and changes in the structure of the susceptible population in recent years, clinically refractory pneumonia is becoming more common. Causes of the disease can be classified into 3 main factors: bodily factors, microbial factors, and iatrogenic factors. In clinical treatment, the main contradictions should be based on corresponding treatment measures.

Since its clinical adoption, electronic bronchoscopy (EBS) has become an indispensable tool for the diagnosis and treatment of adult respiratory diseases, as it is soft and bendable (3). It can visually display the morphology and structure of the bronchus, and it can perform bronchoalveolar lavage and clamp foreign bodies through its biopsy hole to aid in further clinical diagnosis and treatment. Therefore, it is widely used in the diagnosis and pathogenic detection of various intractable respiratory diseases and is used to assist in the treatment of clinical respiratory diseases (4). With the continued development of medical technology and accumulation of clinical experience with this procedure, the clinical adoption of EBS has developed rapidly and gradually become favored by clinicians (5). At present, many methods such as bronchoalveolar lavage, bronchoalveolar brushing, bronchial mucosal biopsy, and transbronchial lung biopsy are used in clinical practice. EBS can be placed deep into the lower respiratory tract to collect specimens, while inflicting little damage and providing accurate sampling, which significantly improves the accuracy of lesion detection. It uniquely contributes to the study of the

occurrence and development of respiratory diseases and the assessment of curative effects and prognosis (6,7).

However, there is a lack of relevant literature on the efficacy and safety of EBS in the treatment of refractory pneumonia. Therefore, with the aim of providing a theoretical basis for the evaluation of the efficacy of EBS in the treatment of refractory pneumonia, clinical randomized controlled trials (RCTs) of EBS in the treatment of refractory pneumonia were screened to further confirm its efficacy, and a meta-analysis was used to conduct a systematic evaluation. The novelty of this study is the examination of the use of a bronchial electron microscope instead of an ordinary fiberoptic bronchoscope. Refractory pneumonia requires early lavage under bronchial electron microscopy to better observe the cleaning effect. Local lavage unblocks the respiratory tract and clears the secretions of the lower respiratory tract and phlegm thrombus. It is helpful for the diagnosis and treatment of refractory pneumonia, especially in children with sputum blockage combined with atelectasis or persistent atelectasis, which can cause refractory pneumonia. Large flake shadows are quickly discerned by imaging. Therefore, the use of bronchial electron microscopy in the early stage can provide better treatment and cleaning of the respiratory tract. We present the following article in accordance with the PRISMA reporting checklist (available at https://dx.doi. org/10.21037/apm-21-2133).

#### **Methods**

#### Literature inclusion and exclusion criteria

The inclusion criteria were as follows: (I) the participants of the study were patients who were clinically diagnosed with refractory pneumonia; (II) the research type was an RCT published in English-language databases; (III) the treatment method of the experimental group was EBS treatment, the surgical treatment of the control group was conventional treatment, and the baseline data of the experimental group and the control group were comparable; and (IV) the evaluation indicators of the study outcomes included postoperative satisfaction of patients and occurrence of adverse reactions.

The exclusion criteria were as follows: (I) research types were non-RCT studies, such as retrospective studies, case reports, and cohort studies; (II) the research objects were animals or cells; (III) the literature types were unpublished documents or non-English documents such as degree

theses; (IV) the treatment method was EBS; (V) the research participants were patients with refractory pneumonia combined with other diseases; and (VI) the literature had incomplete research data from which a corresponding effect index could not be calculated.

#### Literature search

Six English-language databases (PubMed, Embase, MEDLINE, Ovid, Springer, and Web of Science) were searched for literature published up until December 31, 2020. Publicly published RCT studies of EBS in the diagnosis and treatment of refractory pneumonia were retrieved. Literature search terms consisted of subject terms and keywords, including "Refractory pneumonia," "Electronic bronchoscope," and "Conventional drugs". "And" or "or" were used for joint searches among search terms, and the literature search was carried out by 2 researchers using independent search methods.

#### Literature screening

Two researchers independently screened the documents. After the literature search was completed, Note Express 3.2 was used to establish a literature database, and the duplicates of the retrieved literature were then removed. The 2 researchers then manually screened the remaining documents. They were required to read the titles and abstracts of the retrieved documents first and eliminate documents that obviously did not meet the inclusion criteria. The decision whether to include the literature was determined according to the inclusion and exclusion criteria. If the 2 researchers disagreed on the process of document screening, the disagreement would be resolved through discussion. If a consensus was still not reached, a third party would be invited to make a decision after arbitration.

#### Data extraction

Two hospital researchers were selected to examine patients for disease symptoms, interventions, outcome indicators, and bias assessments based on basic information from the included literature, and data record table was fabricated. Researchers needed to screen literatures from a large number of literatures, extract valid data and information, and screen literatures according to inclusion and exclusion criteria. Researchers can't exchange methods and review criteria. After the two researchers had completed the examination, they can exchange literature. In the literature screening process, if two researchers disagreed, they should discuss and communicate with each other. If there was no result of communication, the third-party researcher shall be asked to appraise again. The data extracted from the documents that met the inclusion criteria mainly included (I) the title of the document, the title of the document, the first author, the year the document was published, and the author's main research contributions; (II) the basic information of patients, such as age, nationality, gender, and intervention measures; (III) different methods in the literature, random methods selected in the literature, and whether the literature was blind; (IV) indicators of disease detection and results of the final incidence and total number of patients.

#### Literature quality assessment

The risk of bias was assessed using the bias analysis provided in 5.0.2 of the Cochrane Intervention Systems Review Manual. Bias risk assessment literature was performed. If the results were good and the risk was low, the literature quality was high. The evaluation indicators were randomization, randomization, blindness, patient awareness, study data integrity, and outcome indicators. If there was no agreement between the two researchers, the two researchers should discuss and communicate with each other. If there was no result, a third researcher was asked to confirm again.

# Statistical analysis

The Cochrane Handbook for Systematic Review of Interventions 5.0.2 was used to assess the risk of literature bias. Stata 11.0 (Texas College Station StataCorp, USA) was employed to consolidate statistics in the included literature. Comprehensive statistical data were analyzed using Review Manager 5.3. Forest maps and funnel plots were used to indicate the accuracy, specificity, and sensitivity of the literature. The relative risk (RR) of binary variables in postoperative adverse event count data should be the same. 95% confidence intervals (95% CI) and continuous variables in the measurements, including heart rate (HR), mean arterial pressure (MAP), and visual analog scale (VAS), were calculated. If the units of measurement indicators were the same, the weighted mean difference (WMD) was used as the effect size. Standardized mean difference (SMD) was

used as the effect size if the measurement measures were in different units. The I<sup>2</sup> test was used to assess heterogeneity in the included literature. The greater the  $I^2$  statistic, the greater the heterogeneity. If  $I^2 > 50\%$  and it failed to explain the source of heterogeneity, a random-effects model (REM) combined with effect size was used for a metaanalysis. If  $I^2 < 50\%$ , which meant good heterogeneity of the literature, and fixed effects model (FIX) was used for analysis combined with effect size. The combined effect size was tested with a 95% confidence interval. P<0.05 indicated a statistically significant difference. Binary variables were tested with 95% confidence intervals. When 95% CI >1 or <1, the data were statistically significant. When the 95% confidence interval contains 1, the data were not statistically significant. Continuous variables were tested with 95% confidence intervals. When 95% CI >0 or <0, the data were statistically significant.

### Sensitivity analysis

Sensitivity analysis was used to evaluate whether the results of meta-analysis were stable and reliable. Specifically, by excluding some controversial studies, low-quality studies, or using different statistical methods/effect models to analyze the same set of data, the changes in the results of metaanalysis were observed. If the sensitivity analysis did not substantially change the results, the results were considered reliable; if the results were substantially changed, this indicated that any interpretation or drawing of conclusions from this data set should be done with caution.

# **Results**

# Literature search results

A preliminary search of literatures in six English databases was conducted, and 455 related literatures were found. After the initial search was completed, 198 literatures remained after the copies were deleted according to inclusion and exclusion criteria. Titles and abstracts were carefully screened, and literatures were screened according to inclusion and exclusion criteria. Literatures with high bias should also be deleted. The 2 researchers then read and cross-examined the text of the literature, and screened and excluded the literature accordingly. Finally, a total of 6 articles were included in this study (*Figure 1*). They were all publicly published RCT studies, and the publication time ranged from 2000 to 2020. The 6 articles contained a total of 796 study participants, and baseline data, such as the age of patients in the experimental group and the control group, were comparable (*Table 1*). The indicators of the 6 included studies were evaluated according to the main intervention: electronic bronchoscopy. The methodological quality of each study in the intervention results was above average and at the same level of literature quality, and there was no methodological heterogeneity. Therefore, sensitivity analysis was not conducted.

#### Bias risk assessment of included literature

The Cochrane Handbook of Systematic Evaluation of Interventions 5.0.2 was used to assess the risk of bias for the six articles in this study. The deviation risk maps were output using Review Manager 5.3. The risk assessment of bias included seven items. (I) The six literatures included in this study (8-13) were grouped by "different groups of treatment strategies", indicating low risk. (II) None of the six documents mentioned the existence of "grouping randomization" and did not explain the randomization method, indicating that the risk was not clear. (III) As for whether the subjects were informed, five of the six literatures (8-10,12,13) mentioned that "the patients knew and signed the informed consent form" but did not mention whether the experimenter was blind, so the risk was not clear. (IV) For the blind method of the result assessors, six articles did not mention whether the result assessors were blind, suggesting that the risk was not clear. (V) For the corresponding of methods and results of the literature, the result data of the six articles were complete, indicating a low risk. (VI) For selective reports, there was no selective report in six articles (8-13), indicating a low risk. (VII) For other bias risks, there were three articles (10,11,13) had different numbers in the experimental group and the control group, indicating high risk. In addition, it could not be determined whether there were other deviations in three of the documents (9,12,13), and the risks were considered unclear. The results of the deviation risk assessment were shown in Figures 2 and 3.

# Clinical efficacy

In this study, a total of 3 documents (9,11,13) analyzed the clinical efficacy of EBS in patients. A total of 185 patients with refractory pneumonia were included, including 85 in the experimental group and 100 in the control group. The heterogeneity test results showed

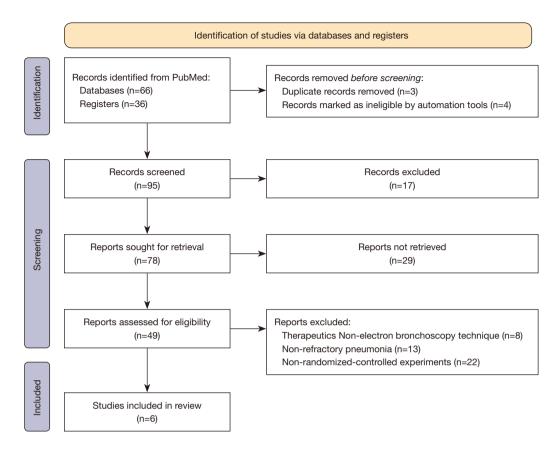
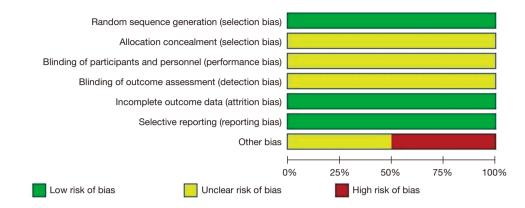
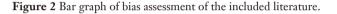


Figure 1 Document retrieval process.

Table 1 Basic characteristics of the included st
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First author	Published year	Group	Sample size	Counter measure	
Mueller (8)	2007	Experimental	52	Electronic bronchoscopy	
		Control	52	Conventional therapy	
Ruiz (9)	2000	Experimental	20	Electronic bronchoscopy	
		Control	20	Conventional therapy	
Fagon (10)	2000	Experimental	204	Electronic bronchoscopy	
		Control	209	Conventional therapy	
Solé Violán (11)	2000	Experimental	43	Electronic bronchoscopy	
		Control	45	Conventional therapy	
Timsit (12)	2001	Experimental	47	Electronic bronchoscopy	
		Control	47	Conventional therapy	
Zhang (13)	2014	Experimental	22	Electronic bronchoscopy	
		Control	35	Conventional therapy	





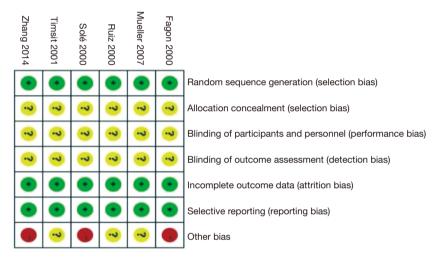


Figure 3 Assessment chart of the risk of bias in the included literature.

that  $I^2=65\%$  and P=0.06, which suggested that there was heterogeneity in the articles. Therefore, the REM was used for analysis (results in *Figure 4*). The results showed that the combined effect of meta-analysis was as follows: OR =3.8, 95% CI: 1.11–12.99, Z=2.13, and P=0.03. The diamond in the forest plot was on the right side of the vertical line, which indicated that the clinical efficacy of EBS in patients with refractory pneumonia was higher than that in the control group.

#### Mortality rate

A total of 3 articles (8-10) in this study analyzed the mortality rate of patients, and 557 patients with refractory pneumonia were included, including 276 in the experimental group and 281 in the control group. The heterogeneity test results showed that  $I^2=0\%$  and P=0.48, which suggested that there was heterogeneity in the articles. Therefore, FEM was used for analysis (results in *Figure 5*). The results showed that the combined effect of metaanalysis was the following: OR =0.7, 95% CI: 0.3–1.63, Z=0.82, and P=0.41. The diamond in the forest plot was on the left of the vertical line, which indicated that the mortality rate of patients with refractory pneumonia treated by EBS was lower than that of the control group.

### Infection

In this study, a total of 3 documents (8,11,12) analyzed patients' infection status. A total of 286 patients with

	Experim	ental	Contr	ol		Odds Ratio	Odds Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Ruiz 2000	13	20	7	20	32.6%	3.45 [0.94, 12.65]	
Solé 2000	32	43	29	45	40.0%	1.61 [0.64, 4.02]	
Zhang 2014	20	22	14	35	27.4%	15.00 [3.02, 74.53]	
Total (95% CI)		85		100	100.0%	3.80 [1.11, 12.99]	
Total events	65		50				
Heterogeneity: Tau <sup>2</sup> :							
Test for overall effect: Z = 2.13 (P = 0.03)							[experimental] [control]

Figure 4 Forest plot of clinical efficacy of patients with refractory pneumonia treated by EBS. EBS, electronic bronchoscopy.

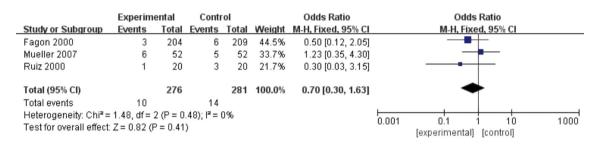


Figure 5 Forest plot of the mortality rate of patients with refractory pneumonia treated by EBS. EBS, electronic bronchoscopy.

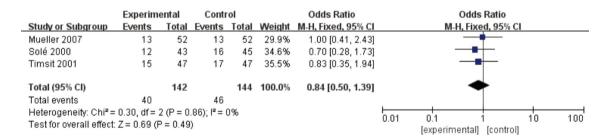


Figure 6 Forest plot of the infection rate of patients with refractory pneumonia treated by EBS. EBS, electronic bronchoscopy.

refractory pneumonia were enrolled, including 142 in the experimental group and 144 in the control group. The heterogeneity test results were  $I^2=0\%$  and P=0.86, which suggested that there was heterogeneity in the articles. Therefore, the FEM was used for analysis (results in Figure 6). The combined effect of meta-analysis was as follows: OR =0.84, 95% CI: 0.5-1.39; Z=0.69, and P=0.49. The diamond in the forest plot was on the left side of the vertical line, which indicated that the infection rate of patients with refractory pneumonia treated by EBS was lower than that of the control group.

#### ICU hospitalization days

A total of 4 articles (8-10,13) in this study analyzed the length of stay of patients in the ICU. A total of 574 patients with refractory pneumonia were enrolled, including 278 in the experimental group and 296 in the control group. The heterogeneity test results were  $I^2=17\%$  and P=0.3, which suggested that there was heterogeneity in the articles. Therefore, the FEM was used for analysis (results in Figure 7). The combined effect of meta-analysis was as follows: OR =0.59; 95% CI: 0.36-0.98, Z=2.04, and P=0.04. The diamond in the forest plot was on the left side of the

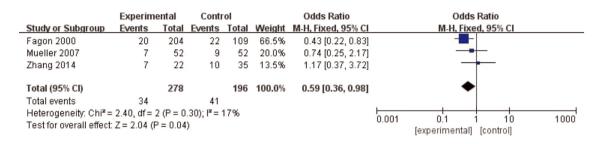


Figure 7 Forest plot of ICU hospitalization days for patients with refractory pneumonia treated by EBS. EBS, electronic bronchoscopy.

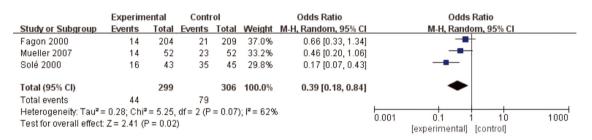


Figure 8 Forest plot of the number of days of antibiotic use in patients with refractory pneumonia treated by EBS. EBS, electronic bronchoscopy.

vertical line, which indicated that the number of days of ICU hospitalization for patients with refractory pneumonia treated by EBS was lower than that of the control group.

# Days of antibiotic use

In this study, a total of 3 documents (8,10,11) analyzed the number of days patients took antibiotics. A total of 605 patients with refractory pneumonia were enrolled, including 299 in the experimental group and 306 in the control group. The heterogeneity test results were  $I^2=62\%$ and P=0.07, suggesting that there was a certain degree of heterogeneity among the studies. Therefore, the REM was used for analysis, (results in *Figure 8*). The combined effect of meta-analysis was as follows: OR =0.39, 95% CI: 0.18–0.84, Z=2.41, and P=0.02. The diamond in the forest plot was on the left side of the vertical line, which indicated that the number of days of antibiotic use in patients with refractory pneumonia treated by EBS was lower than that in the control group.

#### Body temperature

In this study, a total of 3 documents (8,11,12) analyzed the

body temperature of patients during treatment, and 286 patients with refractory pneumonia were enrolled, including 142 in the experimental group and 144 in the control group. The heterogeneity test results showed that  $I^2=0\%$  and P=0.48, which suggested that there was heterogeneity in the articles. Therefore, the FEM was used for analysis (results in *Figure 9*). The combined effect of meta-analysis was as follows: MD =-0.20, 95% CI: -0.24 to -0.16, Z=9.5, and P<0.00001. The diamond in the forest plot was on the left side of the vertical line, which indicated that the body temperature of patients with refractory pneumonia treated with EBS was lower than that of the control group.

# White blood cell count

In this study, a total of 3 documents (8,11,12) analyzed the patients' white blood cell counts, and 286 patients with refractory pneumonia were enrolled, including 142 in the experimental group and 144 in the control group. The heterogeneity test results showed that  $I^2=0\%$  and P=0.93, which suggested that there was heterogeneity in the articles. Therefore, the FEM was used for analysis (results in *Figure 10*). The combined effect of meta-analysis was as follows: MD =0.55, 95% CI: -0.57 to 1.67, Z=0.96, and

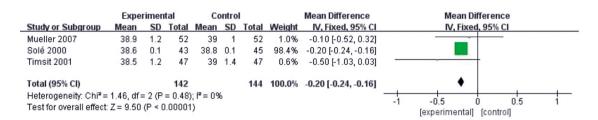


Figure 9 Forest plot of the body temperature of patients with refractory pneumonia treated by EBS. EBS, electronic bronchoscopy.

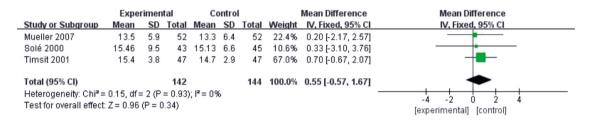


Figure 10 Forest plot of white blood cell counts in patients with refractory pneumonia treated by EBS. EBS, electronic bronchoscopy.

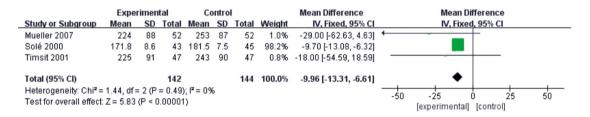


Figure 11 Forest plot of the PaO2:FIO2 ratio in patients with refractory pneumonia treated by EBS. EBS, electronic bronchoscopy.

P=0.34. The diamond in the forest plot was on the right side of the vertical line, which indicated that the white blood cell count of patients with refractory pneumonia treated with EBS was higher than that of the control group.

### PaO2:FIO2 ratio

In this study, a total of 3 documents (8,11,12) analyzed the PaO<sub>2</sub>:FIO<sub>2</sub> ratio of patients, and 286 patients with refractory pneumonia were enrolled, including 142 in the experimental group and 144 in the control group. The heterogeneity test results showed that  $I^2=0\%$  and P=0.49, which suggested that there was heterogeneity in the articles. Therefore, the FEM was used for analysis (results in *Figure 11*). The combined effect of meta-analysis was as follows: MD =-9.96, 95% CI: -13.31 to -6.61, Z=5.83, and P<0.00001. The diamond in the forest plot was on the left side of the vertical line, which indicated that the PaO<sub>2</sub>:FIO<sub>2</sub> ratio of patients with refractory pneumonia treated by EBS was lower than that of the control group.

#### Publication bias analysis

The results of postoperative adverse reaction indexes were analyzed in terms of publication bias using Review Manager 5.3 (results in *Figure 12*). The clinical efficacy, mortality, infection, ICU hospitalization days, antibiotic use days, body temperature, white blood cell count, and PaO<sub>2</sub>:FIO<sub>2</sub> ratio were basically distributed within the credible interval, and the literature bias was low.

#### Discussion

Refractory pneumonia is not a single disease, and the reasons for intractable pneumonia vary from person to

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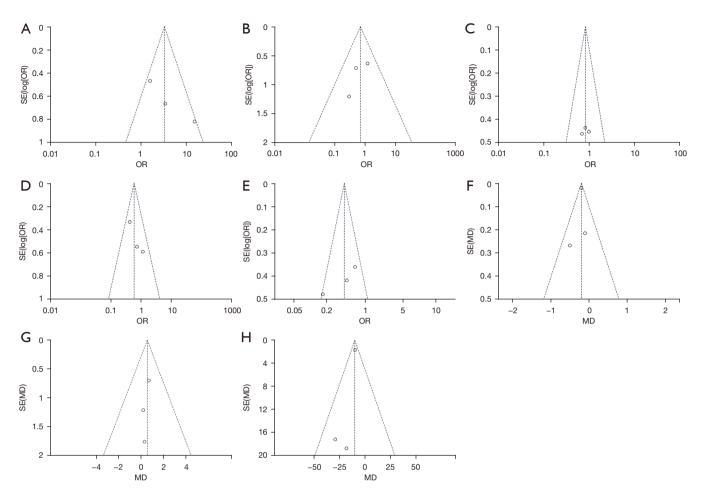


Figure 12 Funnel plots of various evaluation indicators. (A) Clinical efficacy. (B) Mortality. (C) Infection. (D) ICU hospitalization days. (E) Days of antibiotic use. (F) Body temperature. (G) White blood cell count. (H) PaO<sub>2</sub>:FIO<sub>2</sub> ratio.

person. Basic diseases or immune damage often make the clinical manifestations of pneumonia imperceptible and atypical, which can delay diagnosis and treatment or lead to more complications and increased complexity of treatment (14-16). In clinical treatment, the pathogens must first be identified to avoid bacterial infection in the upper respiratory tract. Bacterial sampling must be carried out in the lower respiratory tract, so precision instruments are required (17).

The emergence of bronchoscopy has provided convenience in clinical diagnosis and treatment, but early bronchoscopy has disadvantages including large diameter and poor lighting. With the rise in improvement of glass optical fiber technology, the progress of related technologies, and the demand of clinical needs, the resolution of bronchoscopes has continued to increase, the field of view has widened, and the diameter of the insertion point has grown smaller. At present, the smallest diameter of the bronchoscope can be 1.2 mm (18-20). As clinical experience in China has grown and medical technology has advanced, EBS has been able to provide increasing benefit in the diagnosis and treatment of clinical symptoms, with greater higher safety (21,22). Patients with refractory mycoplasma pneumoniae pneumonia often have respiratory mucus obstruction, atelectasis, and other conditions. Early bronchoscopy and bronchoalveolar lavage treatment to clear respiratory secretions and phlegm thrombus has an important role in reducing symptoms such as high fever, promoting lung recruitment, and reducing the occurrence of sequelae. After bronchoscopy lavage treatment is applied in patients with refractory pneumonia, the symptoms of fever and cough are significantly reduced, and the absorption time of lung shadows and the length of hospitalization are significantly shortened.

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To systematically evaluate the application value of EBS in the diagnosis and treatment of refractory pneumonia, a total of 6 reports were included in this study. A meta-analysis was performed to systematically evaluate the efficacy of EBS in the treatment of refractory pneumonia. The results showed that the clinical efficacy of treatment in the experimental group was significantly higher than that of the control group, the adverse outcomes, such as infection and death, were lower than those of the control group. Studies have revealed that EBS directly inspect the bronchial lumen of grade III to IV pneumonia (23). Direct suction and removal of sputum and secretions at the target location can be realized by EBS, and the lavage of sputum stasis with normal saline can dilute the sputum to stimulate cough and facilitate the expectoration and aspiration of sputum (24), which greatly reduces the survival rate and reproduction of bacteria. EBS thus facilitates the rapid recovery of patients (25,26). It was also found that the body temperatures of patients in the experimental group were lower during treatment, and the results of biochemical examinations were better than those of the control group, indicating that EBS had a good therapeutic effect and resulted in less damage to the body and stress responses, which was conducive to the improvement of the disease. Bronchial electron microscopy can also be used in patients with COVID-19. Patients with COVID-19 are weak in coughing and sputum production, which leads to a further decline in lung oxygenation. In critically ill patients, phlegm and mucus plugs cause treatment difficulties and aggravate their condition. Sputum suction and lavage with the assistance of bronchoscopy are important treatments for symptomatic relief and recovery of critically ill patients. Most critically ill patients with COVID-19 require tracheal intubation or tracheotomy for ventilator-assisted ventilation, and bronchoscopy can be used for guidance and monitoring.

### Conclusions

In this study, a total of 6 articles on EBS diagnosis and treatment of refractory pneumonia were included in the meta-analysis, involving 796 patients. The results showed that EBS can significantly reduce the incidence of adverse postoperative outcomes and treatment time for patients compared with conventional treatment, while improving the clinical efficacy; this demonstrates the significant advantages of EBS treatment. However, this study also has certain limitations, which mainly includes the considerable publication bias of some of the included literature. In addition, due to differences in the research directions of the authors of the literature, some analysis indicators contained a small number of samples, and the meta-analysis results are thus not sufficiently accurate. Therefore, in follow-up studies, more large samples and high-quality articles will be included to verify the clinical effect of EBS in the treatment of refractory pneumonia.

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#### Footnote

*Reporting Checklist:* The authors have completed the PRISMA reporting checklist. Available at https://dx.doi. org/10.21037/apm-21-2133

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*Ethical Statement:* The authors are 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.

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