



# Incidence rate of pneumonia after post-stroke dysphagia: A meta-analysis

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## Abstract

**Objective:** To figure out the latest incidence rate of pneumonia after post-stroke-dysphagia (PSD), related literature was included, then a meta-analysis was completed.

**Methods:** Articles on the incidence rate of pneumonia after PSD was searched by manual and electrical databases. Data were selective by screening the abstract, browsing the full text according to diagnostic criteria, and extracting the data on incidence rate of pneumonia. Subsequently, a meta-analysis was performed using Stata 16.

**Results:** 1900 studies were initially retrieved and 14 studies met the inclusion criteria for meta-analysis after extraction. The meta-analysis showed that the pooled incidence rate of pneumonia was 39% and the overall effect was 11.046 with high heterogeneity ( $I^2 > 75%$ ) using random effects. Furthermore, the incidence rate of pneumonia in the dysphagia group was 4.39 times higher than in the non-dysphagia group, with medium heterogeneity  $I^2 = 60.5%$ . To find the heterogeneity, a subgroup analysis was carried out and the results indicated that the incidence rate of pneumonia was 37% in the prospective group and 26% in the retrospective group, but all with high heterogeneity ( $I^2 > 75%$ ). Finally, the combined incidence rate of hospital-acquired pneumonia was 0.26 without heterogeneity, which could be associated with sources and types of literature.

**Conclusion:** Our results have analyzed the incidence rate of pneumonia after PSD in 14 related literature, and the results showed that there was a 39% incidence rate of pneumonia in PSD patients, which might have certain guiding significance for early treatment and prognosis of pneumonia.

**Keywords:** Pneumonia; Dysphagia; Stroke; Incidence Rate; Meta analysis

## Introduction

Stroke is one of the most common neurogenic diseases characterized by loss of neurological function caused by hemorrhage or ischemia in brain tissue. Among them, swallowing dysfunction was the most common complication after stroke, defined as post-stroke-dysphagia (PSD)<sup>1</sup>. According to statistics, up to 80% of stroke patients encounter dysphagia<sup>2</sup>, the incidence rate is associated with early dysphagia screening, therapies, and

postural control management, etc.<sup>3</sup>. For the other complications after stroke, malnutrition and dehydration are also common, especially pneumonia attributed to aspiration<sup>4</sup>. Research showed that there were 4% to 10% of patients who had pneumonia after stroke, which was related to dysphagia with a high mortality rate.

Pneumonia is one of the most common complication after stroke, ranging from 5% to 22%<sup>5,6</sup>, which may affect the mortality in stroke patients, reaching 3.06 times independently associated. Pneumonia is closely associated with aspiration, and the presence of tracheal intubation after stroke is also an important factor for pneumonia, although it avoids a certain amount of food aspiration<sup>7</sup>. Regarding the cause of pneumonia, the decline in the immune system after stroke could increase the incidence rate of infection. Furthermore, the response of tracheal epithelial cells caused by the immunomodulatory response after stroke will altered, resulting in a reduction in lung clearance and an increase in the appearance of aspiration<sup>5</sup>. In fact, the severity of pneumonia was closely related to the severity of stroke, age, and consciousness<sup>8</sup>. Pneumonia after PSD is the main cause of death in stroke, associated with early assessment, age, complications, or more severe neurological impairment.

Pneumonia after stroke can influence the severe rate and

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mortality during hospital stay, which also increases the medical and economic burden to the country. Among those, dysphagia was a risk factor for pneumonia after stroke. As previous studies reported, the nasogastric tubes were associated with respiratory infections, and the diagnostic criteria was also closely related to the occurrence of pneumonia in clinical stroke<sup>9</sup>. Thus, the reported incidence rate of pneumonia after stroke in the literature was highly variable due to a variety of factors, of which dysphagia was a contributing factor. However, there was no existing statistical data showing the incidence rate of pneumonia after PSD recently. In our study, literature related to pneumonia after PSD was searched from the electrical databases, then a meta-analysis was performed to view the incidence rate of pneumonia after PSD. It would provide theoretical and data guidance for clinicians on reducing the incidence rate of pneumonia after PSD.

## Methods

The method for this meta-analysis was guided by the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) statement for systematic reviews<sup>10</sup>. The study protocol was registered on PROSPERO (an international prospective registry of systematic reviews) with the registration number CRD42023390385.

The PICOS framework was used to formulate the study protocol. Population: All patients diagnosed with dysphagia after stroke. Outcome: The incidence rate of pneumonia after PSD was the primary outcome measure. The type of study was limited to cohort study. This review did not have an intervention. Partial research has been grouped into the control group without dysphagia.

### Study eligibility

**Inclusion criteria** The studies included met the following criteria: Cohort study; the subjects were human; the retrieval time was conducted from the beginning of the database to September 24, 2023; Age > 18; definite diagnosis of stroke; patients diagnosed with swallowing disorder through Videofluoroscopic Swallowing Study (VFSS), Flexible Endoscopic Evaluation Swallowing (FEES), or other evaluation methods, not limited to Standard Swallowing Assessment (SSA); patients diagnosed with pneumonia (clinical symptoms and signs, chest X-ray or CT inspection); contained complete data of incidence rate.

**Exclusion criteria** Patients without dysphagia after diagnosis of VFSS or FEES; patients with leakage or aspiration and without inflammatory infiltration with chest X-ray; review article; meta-analysis article.

### Data sources

The academic databases used for the systematic review included Web of Science, Embase, China National Knowledge Infrastructure (CNKI), China Biomedical Literature Database (CBM), Wanfang database, China Science and Technology Journal Database (cqvip) and PubMed. Keywords related to

the population and outcome used synonyms for each disease according to the respective medical subject headings (MeSH). The search strategy was as follows. Web of Science: TS=(pneumonia\* OR pulmonary infection) AND TS=(cerebral apoplexy\* OR cerebrovascular accident OR stroke) AND TS=(dysphagia) AND TS=(incidence); Embase: ('stroke'/exp OR 'cerebrovascular accident'/exp OR 'cerebral apoplexy'/exp) AND ('aspiration pneumonia'/exp OR 'pneumonia'/exp) AND ('dysphagia'/exp OR 'swallowing disorder'/exp OR 'swallowing dysfunction'/exp) AND ('incidence'/exp); Pubmed: (stroke or cerebral apoplexy or cerebrovascular accident) and (dysphagia or swallowing disorder) and (pneumonia or aspiration pneumonia or pulmonary infection) and incidence; CNKI: (SU= swallowing disorder OR SU= dysphagia) AND (SU= stroke OR SU= cerebral apoplexy OR= cerebrovascular accident) AND (SU= pneumonia OR SU= pulmonary infection) AND (SU= incidence); The search strategy of Wanfang database, cqvip and CBM were similar to CNKI with respective medical subject headings.

### Study selection

The literature was screened in the following logical three-stage: title and abstract screening, full text screening, and extraction. Two authors (Qiuping Ye and Jiahui Hu) independently screened the retrieved studies. When the full text was not immediately available, it would be obtained through correspondence with the authors. No additions were requested other than those contained in published manuscripts.

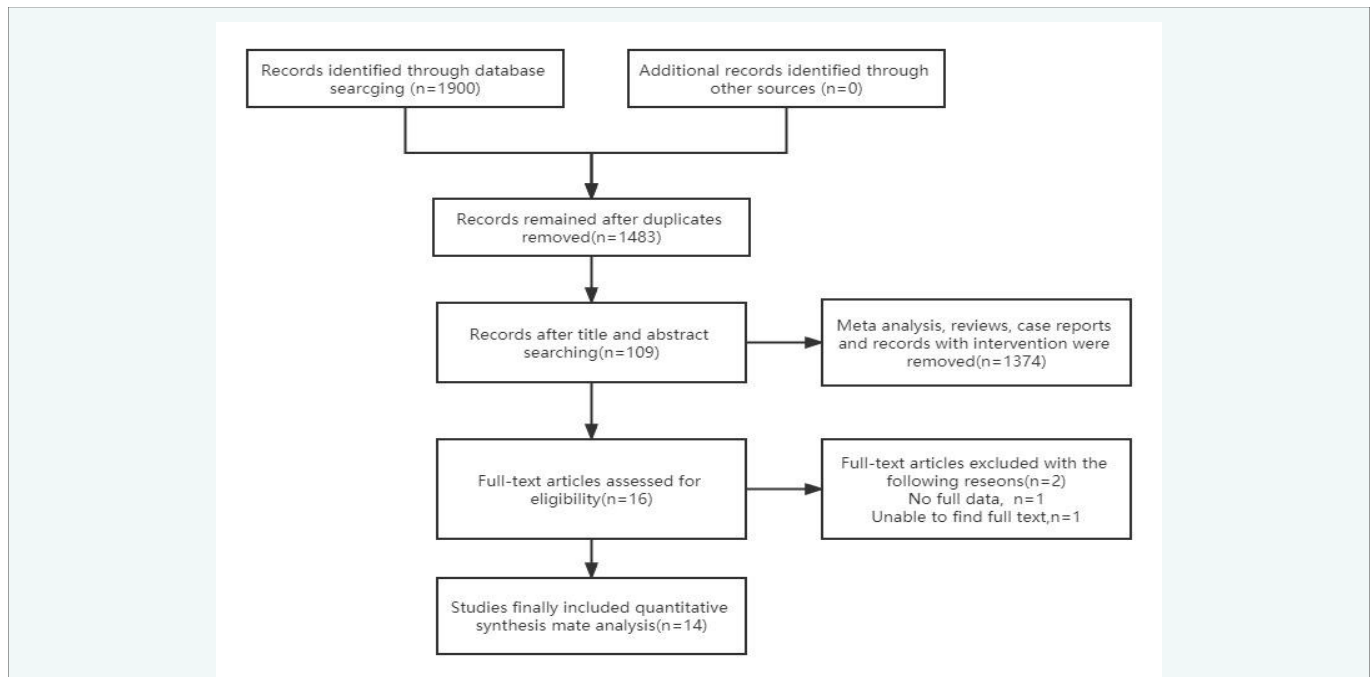
### Data extraction

Data extraction was carried out in parallel by two reviewers (Qiuping Ye and Jiahui Hu). The study details (author, year of publication), study characteristics (design, setting) and participant data (population size, male/female ratio, mean age at diagnosis, incidence rate for diseases and the corresponding data of amount) were extracted. The data extraction process was shown in Figure 1. Conflicts in the process of study selection could be discussed and determined by the third author (Yong Dai).

### Risk of bias assessment

The heterogeneity assessment was performed using the  $I^2$  statistic.  $I^2 < 50\%$  was thought to represent low heterogeneity,  $I^2 > 50\%$  was thought to represent moderate heterogeneity, and  $I^2 > 75\%$  represented high heterogeneity. Publication bias within the studies was evaluated with a funnel plot using a random effects model.

The quality of the included studies was evaluated using the Newcastle-Ottawa Scale<sup>11</sup> that was adapted to fit this review. They were assessed using the following terms: selection (representative of the exposure cohort, determination of exposure, definition of controls, and whether the study population is large enough), comparability (comparability of cases and controls on the basis of the design or analysis). The maximum score of available points was five.



**Figure 1** PRISMA flow diagram for study screening and selection for Meta-analysis of pneumonia after PSD was showed.

## Data synthesis

The distribution of the incidence rate and the 95% confidence intervals obtained from the studies were examined using a forest plot. A random effects model was used to pool the incidence rate among these data. Subgroup analysis was used to explore the effect of different types of study, including the prospective study or retrospective study, and the incidence rate of hospital-acquired pneumonia or follow-up acquired pneumonia. The meta-analysis was performed using Stata 16.

## Results

The study selection process was shown in a PRISMA flow diagram in Figure 1. 1900 studies were retrieved according to the initial search strategy. 417 studies were excluded after repeated checking and 1482 studies remained for the next-stage screening. After selecting the title and abstract, meta-analysis, review, case report, and studies that included intervention were removed, including 1374 studies, leaving 109 studies for the full text screening.

Following the full text screening, 93 studies without dysphagia were excluded, two studies had incomplete data. Finally, 14 studies met the inclusion criteria for the meta-analysis. Table 1 provided an overview of the characteristics of the 14 studies. And the quality of the included studies was shown in Table 2. There were two articles with four asterisks (\*) and four had three asterisks (\*), which might have relatively high quality, while others had only two asterisks (\*) with probably low quality.

Among the 14 studies, hospital-acquired pneumonia was reported as the result in 3 studies<sup>12</sup>, while the others were

pneumonia. The follow-up acquired pneumonia was shown to be the outcome in two studies<sup>13,14</sup>. Furthermore, 6 studies<sup>15</sup> were the perspective study and 8 studies<sup>16</sup> were the retrospective study, among which the incidence rate of pneumonia after PSD was compared with that of non-dysphagia in 7 studies.

The total meta-analysis for the incidence rate of pneumonia was shown in the forest plot in Figure 2. The random effects meta-analysis demonstrated that the pooled incidence rate of pneumonia was 39% (95%CI 0.25-0.36) (Figure 2). The overall effect was 11.046 with high heterogeneity ( $I^2=95.7\%$ ). From the confidence intervals, we might speculate that the percentage of variation between studies was due to heterogeneity rather than chance. Combined with the funnel plot that distributed asymmetry (Figure 6), the existence of publication bias was indicated.

For comparison of the incidence rate between the dysphagia group and the non-dysphagia group, the meta-analysis with 7 results was carried out. And the incidence rate of pneumonia in the dysphagia group was 4.39 times higher than in the non-dysphagia group, with medium heterogeneity  $I^2=60.5\%$  (Figure 3).

To figure out the reason for high heterogeneity among the meta-analysis, a subgroup analysis was then performed according to the types of study (prospective study or retrospective study). The pooled results in the prospective study showed that the incidence rate of pneumonia was 37% (95% CI 0.22-0.52) with high heterogeneity ( $I^2=94.3\%$ ) (Figure 4a). The overall effect was 5.622. For the retrospective study, the pooled incidence rate of pneumonia was 26% (95% CI 0.20-0.32) with a general effect of



**Table 1:** The characteristics in the 14 studies.

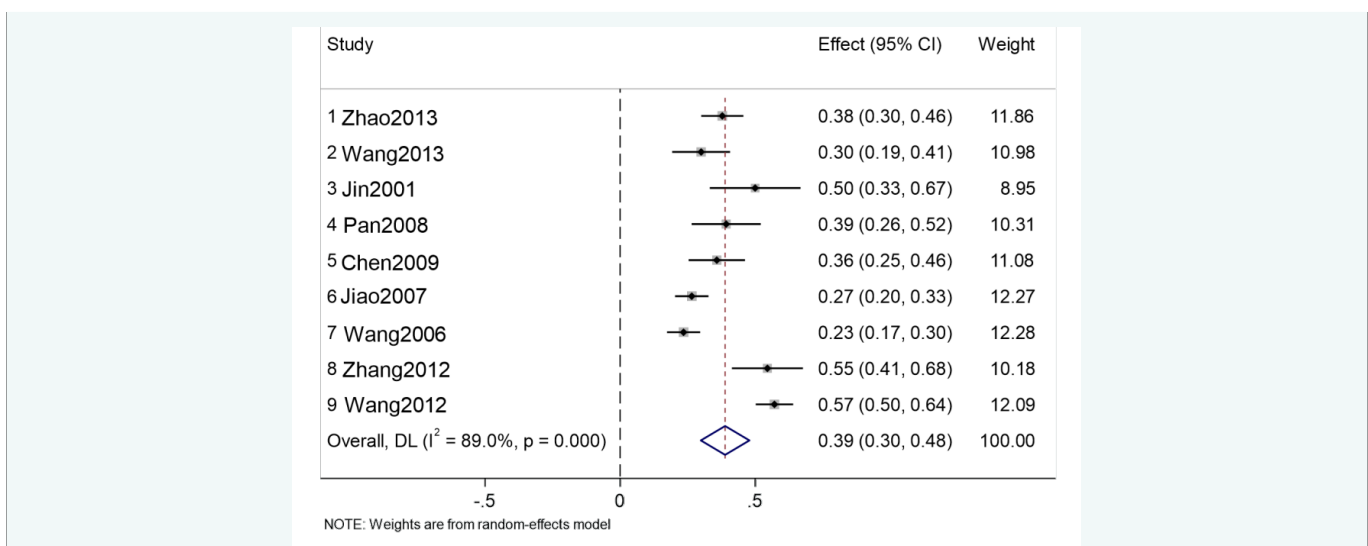
Study	Year	Male/ Female	Age	Evaluation method of swallowing	Types of stroke	Medical-history	Incidence in groups
Zhao	2013	104/44	67~83	Kubota drinking water	TACI and PACI/ POCI	Not mentioned	56/148
Ho	2018	2921/ 2111	18~80	VFSS/ FEES	Not mentioned	HTN/CAD/ cardiac dysrhythmia/ hyperlipidemia/ valvular heart disease/ renal disease/ diabetes and COPD	945/5032
Matsumura	2014	29/47	74.7± 8.4	Fujima Ichiro swallowing efficacy evaluation criteria	Ischemia/ cerebral hemorrhage	Vascular impairment/ Respiratory disease/ diabetes/ Hypertension/ heart disease	10/76
Wang	2013	75/52	65~95	SSA	Ischemia/ cerebral hemorrhage	Diabetes/Hypertension/atrial fibrillation/smokers drinking	D, 21/70 N-D, 5/57
Jin	2001	69/21	36~96	Kubota drinking water	Ischemia/ cerebral hemorrhage	Not mentioned	D, 17/34 N-D, 7/56
Masiero	2008	32/35	72.9± 12.2	FEES	Ischemia/ cerebral hemorrhage	Diabetes/ Hypertension/ COPD/ smokers	9/67
Pan	2008	120/72	36~84	Kubota drinking water	Ischemia/ cerebral hemorrhage	Hypertension/ coronary disease/ hyperlipidemia	D, 22/56 N-D, 8/136
Chen	2009	123/95	32~81	Kubota drinking water	Ischemia/ cerebral hemorrhage	Not mentioned	D, 28/81 N-D, 10/137
Jiao	2007	277/ 103	43~82	Kubota drinking water	Ischemia/ cerebral hemorrhage/ subarachnoid hemorrhage	Not mentioned	D, 52/196 N-D, 19/184
Wang	2006	191/ 155	36~90	Kubota drinking water	Ischemia/ cerebral hemorrhage/ subarachnoid hemorrhage	Not mentioned	D, 43/183 N-D, 8/173
Zhang	2012	39/16	45~90	MMASA	Ischemia/ cerebral hemorrhage	Hypertension/ coronary disease/ stroke/ Heart disease/ smoking/drinking	30/55
Brogan	2014	292/241	71± 14.9	clinical bedside evaluation of Swallowing	Not mentioned	Not mentioned	52/312
Al-Khaled	2016	1336/ 1747	76± 12	dysphagia screening	Not mentioned	Diabetes/ Hypertension/ Hypercholesterolemia/ Atrial fibrillation	917/3083
Wang	2012	Not mentioned	≤85	SSA	Not mentioned	Not mentioned	D, 36/184 N-D, 17/187

Total anterior circulation infarcts, TACI; Partial anterior circulation infarcts, PACI; Posterior circulation infarcts, POCI; Modified Mann assessment of swallowing ability, MMASA; Hypertension, HTN; Coronary artery disease, CAD; Chronic obstructive pulmonary disease, COPD; Standardized swallowing assessment, SSA; Dysphagia, D; Non-Dysphagia, N-D



**Table 2:** The quality of included studies.

Evaluation projects Studies	Representativeness				Comparability	Results		
	Representative of the exposure cohort	Whether the study population is large enough	Ascertainment of exposure	Demonstration that outcome of interest was not present at start of study	Comparability of cohorts on the basis of the design or analysis	Result assessment	Is the follow-up time long enough	Adedance of cohort follow-up
Zhao 2013	*				*			
Ho 2018	*				*			
Matsumura 2014	*				*			
Wang 2013	*				*			
Jin 2001	*				*			
Masiero 2008	*				*		*	*
Pan 2008	*							
Chen 2009	*				*			
Jiao 2007	*				*			
Wang 2006	*							
Zhang 2012	*				*			
Brogan 2014	*				*		*	*
Al-Khaled 2016	*				*		*	
Wang 2012	*				*		*	



**Figure 2** Forest plot for the pooled meta analysis of the incidence rate of pneumonia after PSD, with 95% confidence intervals.

8.01 (Figure 4b). However, the heterogeneity was high both in the prospective study and in the retrospective study.

Furthermore, the combined incidence rate of hospital acquired pneumonia was 0.26 (95% CI 0.22-0.30) without heterogeneity (I<sup>2</sup>=0) (Figure 5), which could indicate that the type of pneumonia is also a source of heterogeneity.

## Discussion

The main finding of our study indicated that the incidence rate

of pneumonia after PSD was 39% (95% CI 0.3, 0.48). Importantly, the incidence rate of pneumonia in the dysphagia group was 4.39 times higher than that in the non-dysphagia group, indicating that dysphagia was an important factor influencing pneumonia. Furthermore, the incidence rate for the prospective study was 35% (95% CI 0.23, 0.48), while there was 26% (95% CI 0.20, 0.32) in the retrospective study.

However, there was high heterogeneity in the pooled meta-analysis, so were the subgroup meta-analysis in the prospective

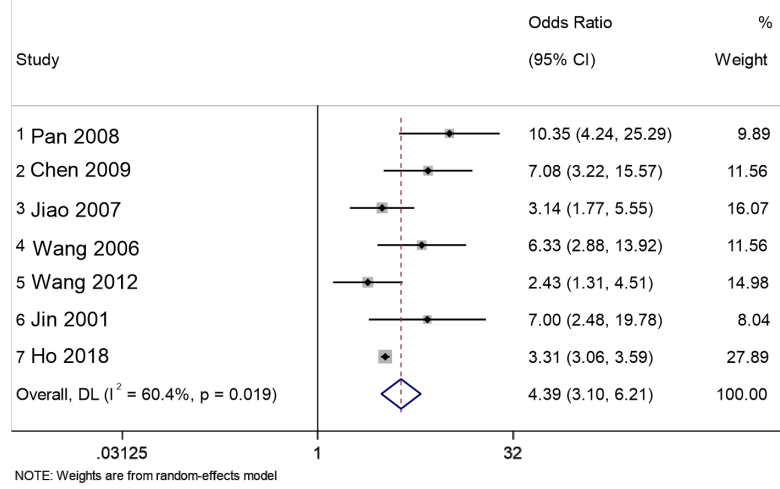


Figure 3 Forest plot for the pooled meta analysis of the incidence rate of pneumonia between dysphagia group and non-dysphagia group, with 95% confidence intervals.

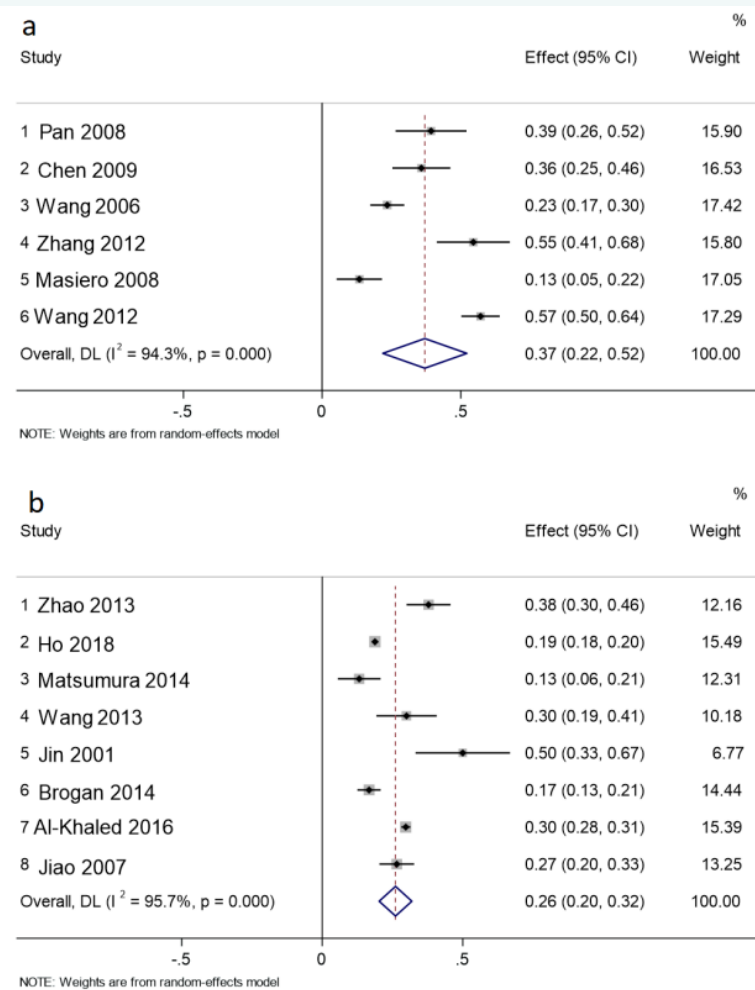
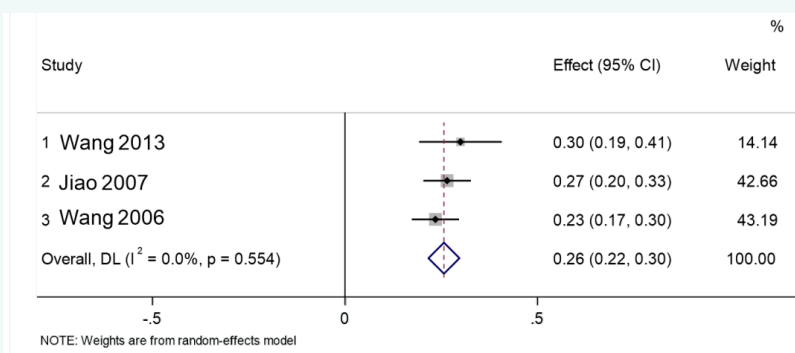


Figure 4 Forest plot for the meta analysis of the incidence rate for pneumonia after PSD in prospective study (a) and retrospective study(b), with 95% confidence intervals.

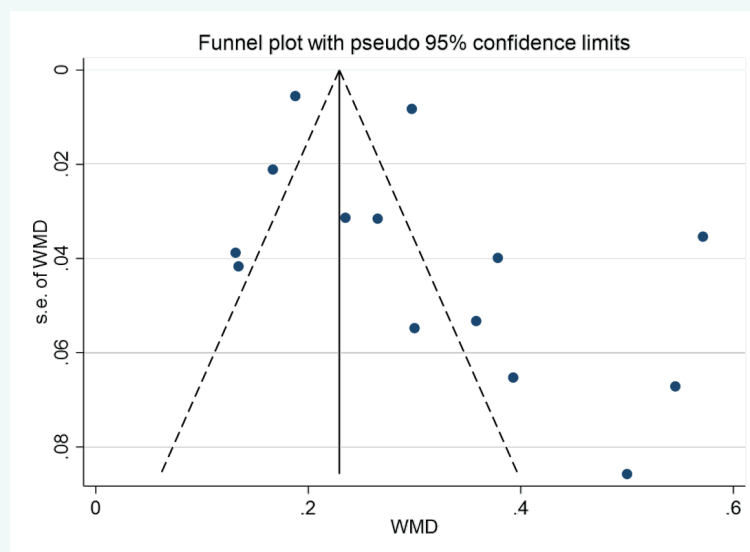


**Figure 5** Forest plot for the meta analysis of the incidence rate for pneumonia after PSD in hospital acquired pneumonia, with 95% confidence intervals.

or retrospective studies. To explore the reasons for the above results, it was suggested that the sources of these studies were different, regardless of the different methods of dysphagia evaluation or the types of study. After subgroup analysis, the heterogeneity in the subgroups remained high, indicating that the grouping of study types could not reduce the heterogeneity. Thus, the heterogeneity of the pooled meta-analysis was due to the different experiences of clinicians in evaluating dysphagia in the articles, the different places or hospitals from which patients come, the various sample sizes, etc. The funnel plot also indicated the publication bias of the included articles (Figure 6). However, there was no heterogeneity in the aggregate outcomes of hospital-acquired pneumonia, indicating that uncertainty after discharge may be another factor contributing to the high heterogeneity of these outcomes.

Furthermore, the incidence rate of pneumonia was compared between patients with and without dysphagia, with medium heterogeneity (Figure 3). In fact, dysphagia was a risk factor for

pneumonia, as shown in a previous study<sup>17</sup>, but dysphagia was also influenced by early screening or management of swallowing. A retrospective study<sup>18</sup> explored the risk factor of aspiration pneumonia by observing the incidence rate of aspiration pneumonia in 1 year, 3 years, and 5 years, respectively, in the dysphagia and non-dysphagia groups. The results showed that the incidence rate of pneumonia ranged from 24.8% to 58.8% in the dysphagia group, while it occurred from 5.8% to 43.9% in the non-dysphagia group. Dysphagia has been considered to lead to aspiration, and about a third of these can lead to pneumonia<sup>19</sup>. Therefore, another article on acute ischemic stroke also confirmed dysphagia as a risk factor for pneumonia<sup>20</sup>. Additionally, subject to an invasive procedure, being bedridden or having a massive cerebral infarction were other risk factors for pneumonia in that investigation. Actually, old age and other concurrent diseases, such as diabetes and hypertension, were also risk factors for pneumonia<sup>21</sup>. In clinical practice, dysphagia needs multidisciplinary treatment, including swallowing treatment, adjustment of bolus types, oral nursing,



**Figure 6** Begg's funnel plot of the studies investigating the incidence rate of pneumonia after PSD included in the meta-analysis.



etc. Therefore, the reduction in the incidence rate of pneumonia is also influenced by these factors. In fact, the literature<sup>2</sup> has indicated that dysphagia was experienced in 91.7% of patients with community-acquired pneumonia, suggesting that dysphagia and pneumonia were interrelated factors.

In consideration of the influence of pneumonia in PSD patients, various treatments were used to reduce the incidence rate of pneumonia, including acupuncture<sup>22</sup>, mirror therapy<sup>23</sup> and consistency of food<sup>24,25</sup>. Those thicker foods could decrease the risk of aspiration in the process of swallowing. Furthermore, clearing bacteria in oral was another method to reduce aspiration<sup>26</sup>. Although prophylactic antibiotics have been demonstrated to decrease infections in stroke patients, it had no effect on the mortality of pneumonia<sup>27</sup>. Our study summarized the latest incidence rate of pneumonia in patients with PSD and will guide the development of clinical measures that reduced the incidence rate of pneumonia and adjusted the treatment program in time.

### Strengths and limitations

This is the first meta-analysis for the incidence rate of pneumonia in PSD patients. Although dysphagia was a risk factor for the incidence rate of pneumonia, no study had concentrated on the incidence rate of pneumonia after PSD recently. The incidence rate of pneumonia after stroke ranged from 5% to 22%<sup>5,6</sup>, but the incidence rate of pneumonia after PSD reached 39% in our analysis, which illustrated the negative effect of dysphagia on the incidence of pneumonia.

Our study has pooled studies on patients with pneumonia after PSD and conducted a meta-analysis on the incidence rate of pneumonia in PSD. But these types of study included prospective study and retrospective study, which might affect the homogeneity of the pooled meta-analysis. Additionally, those studies came from different countries, which could have a different patient populations, medical level and nursing level, even different diagnostic criteria for pneumonia, leading to heterogeneity in the final meta-analysis, as shown in Figure 6. Furthermore, the quality of the included literature was not high and there were very few studies focusing on these epidemiological data. And no recent studies were included, which was another limitation of our study.

### Conclusions

Our results have analyzed the incidence rate of pneumonia after PSD in 14 related articles and the results showed that the incidence rate of pneumonia after PSD reached 39%. Pneumonia is a risk factor for the mortality in stroke patients, reducing its incidence rate is an important work for clinical worker. In this meta-analysis, we have summarized the incidence rate of pneumonia after PSD, which might have some guiding significance for early treatment and prognosis.

### Statement of Ethics

*“An ethics statement is not applicable because this study is based exclusively on published literature.”*

### Conflict of Interest Statement

The authors declare that they have no competing interests. No competing financial or non-financial interest from the funders exist.

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### Author contributions

Zulin Dou and Qiuping Ye put forward the ideas and designed this article. Qiuping Ye, Jiahui Hu and Yong Dai screened and extracted the data. Qiuping Ye analyzed the data. Qiuping Ye and Jiahui Hu wrote the manuscript. Critical revision of the manuscript was done by Jia Qiao and Zulin Dou.

### Data Availability Statement

The data that support the findings of this study are openly available in <http://pubmed.ncbi.nlm.nih.gov>.

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