



Diagnosis of Pneumothorax by Radiography and Ultrasonography

A Meta-analysis

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Objective: This study compares, by meta-analysis, the use of anterior-posterior chest radiography (CR) with transthoracic ultrasonography for the diagnosis of pneumothorax.

Methods: English-language articles on the performance of CR and ultrasonography in the diagnosis of a pneumothorax were selected. In eligible studies, data were recalculated, and the forest plots and summary receiver operating characteristic (sROC) curves were analyzed.

Results: Pooled sensitivity and specificity were 0.88 and 0.99, respectively, for ultrasonography, and 0.52 and 1.00, respectively, for CR. For ultrasonography performed by clinicians other than radiologists, pooled sensitivity and specificity were 0.89 and 0.99, respectively. The sROC areas under the curve were compared, and no significant differences between ultrasonography and CR were found. Meta-regression analysis implied that the operator is strongly associated with accuracy (relative diagnostic OR, 0.21; 95% CI, 0.05-0.96; $P = .0455$).

Conclusions: The meta-analysis indicated that bedside ultrasonography performed by clinicians had higher sensitivity and similar specificity compared with CR in the diagnosis of pneumothorax, but the accuracy of ultrasonography in the diagnosis of pneumothorax depended on the skill of the operators.

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Abbreviations: AUC = area under curve; CR = chest radiography; DOR = diagnostic OR; PNx = pneumothorax; QUADAS = the quality of diagnostic accuracy studies; sROC = summary receiver operating characteristic

Pneumothorax (PNx) frequently occurs in the ED and ICU, especially in patients with trauma and those who are ventilated. Tension PNx is a very serious condition that can potentially lead to cardiac arrest and requires early diagnosis and urgent treatment. A small or medium PNx generally is not life-threatening, but delays in diagnosis and treatment may result in progression of respiratory and circulatory compromise in unstable patients. The diagnosis of PNx generally is confirmed by chest radiography (CR), but CR has been demonstrated to be an insensitive and

unreliable examination.¹⁻⁴ Kirkpatrick and colleagues⁵ evaluated the use of anterior-posterior supine CR with CT scanning for the diagnosis of PNx. In their study of 225 trauma patients, the sensitivity of CR

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was only 20.9%. CT scanning is considered the gold standard for detection of PNx. Sometimes, however, it is impractical to transfer a critically ill patient for

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CT scanning. The high doses of radiation in CT scanning also cannot be neglected.

Ultrasonography was first used in the diagnosis of PNx in humans in 1987.⁶ In recent years, some characteristic signs have been identified for the diagnosis of PNx with ultrasonography, such as lung sliding,⁷ comet tail artifacts,⁸ the A line sign⁸ and lung point.⁹ Because lung ultrasonography can be performed easily and quickly at the bedside by intensivists, pneumologists, and emergency physicians, it can be used in the diagnosis of PNx in ventilated patients,^{7,10} in trauma patients,¹¹⁻¹³ and after lung biopsy.^{14,15} The accuracy of ultrasonography in the detection of PNx varies across studies and is associated with the operator's experience. In the study by Sartori et al,¹⁶ the sensitivity and specificity were 100% for transthoracic ultrasonography to detect PNx in 285 patients after lung biopsy. However, in another study, the sensitivity of ultrasonography was 58.9% and specificity, 99.1%. Slater and colleagues¹⁷ concluded that sometimes ultrasonography only could exclude but not confidently be used to diagnose PNx without the use of other imaging modalities.

Static and dynamic ultrasonography features of PNx have been identified in a number of studies, but the contemporary diagnostic performance of ultrasonography in the detection of PNx has not been well characterized. Should we use the thoracic sonographic examination in addition to the standard focused abdominal sonography for trauma examination in the ED, which is designated the extended focused abdominal sonography for trauma?²⁵ We undertook a meta-analysis of the published literature to compare the accuracy of ultrasonography and CR in the diagnosis of PNx.

MATERIALS AND METHODS

Study Design and Data Sources

A literature review and meta-analysis were conducted. Original articles published in English up to the end of October 2010 were searched in Medline, EMBASE, and the Cochrane Library. We used combinations of the following key words to identify all original articles in which ultrasonography, CR, or both were used in diagnosing PNx: ("ultrasound" or "sonography" or "ultrasonography" or "radiography" or "chest film" or "chest radiograph") and ("pneumothorax" or "aerothorax" or "aeropleura") and ("sensitivity" and "specificity"). New links displayed beside the abstracts were followed and retrieved. Bibliographies of retrieved articles were searched independently and checked for additional studies. No attempt was made to include unpublished data.

Study Selection

We selected articles for analysis that included the following criteria: (1) evaluation of the diagnostic performance of ultrasonography, CR, or both for the detection of PNx; (2) comparison of imaging results with a gold standard (ie, CT scan or composite

standard that included clinical presentation and documentation of the escape or aspiration of intrapleural air at the time of drainage); (3) reporting of results in sufficient detail to allow reconstruction of contingency tables of the raw data (ie, true-positive, true-negative, false-positive, and false-negative results); and (4) having diagnostic criteria for abnormal test results (eg, on ultrasonography, the disappearance of lung sliding; on CR, the appearance of air within the pleural space). Two of the authors (W. D. and Y. S.) independently reviewed the articles and ascertained the criteria for inclusion in the pooled data analysis, with disagreements resolved by discussion. Articles with the same authors were carefully investigated, and some were excluded to avoid duplicate data analysis.

Quality of Study Reports

The quality of diagnostic accuracy studies (QUADAS) tool¹⁸ (e-Table 1) was applied in our analysis to assess the quality of the studies included. The 14-item QUADAS tool assesses study design-related issues and the validity of the study results. Each item may be scored "yes" if reported, "no" if not reported, or "unclear" if no adequate information is available in the article to make an accurate judgment. We considered the quality items 1 (about the spectrum of patients), 4 (about the time period between reference standard and index test), 12 (whether the same clinical data were available when test results were interpreted as would be available when the test is used in practice), and 13 (whether uninterpretable/intermediate test results were reported) not relevant for our analysis; thus, only the remaining 10 items were applied.

Data Extraction

In many of the studies included, hemithorax was used as the study unit for interpretation of the results instead of patient number. Because the diagnosis of PNx in one lateral hemithorax has no relationship with the other side, and usually both hemithoraces must be examined to exclude PNx in one patient, we reconstructed some results as the number of hemithoraces. If there was no specific description, we recalculated one patient as two hemithoraces. For those postbiopsy, one biopsy specimen in one patient was counted as one hemithorax.

Data Analysis

We analyzed the forest plots and summary receiver operating characteristic (sROC) curves with freeware Meta-DiSc, version 1.4 software (http://www.hrc.es/investigacion/metadisc_en.htm; Ramon y Cajal Hospital; Madrid, Spain). The Spearman correlation coefficient between the logit of sensitivity and the logit of 1-specificity was calculated to test the threshold/cutoff effect. Meta-DiSc allows users to test for heterogeneity (other than threshold effect) among various studies by statistical tests, including χ^2 and Cochran Q . A low P value suggests the presence of heterogeneity beyond what would be expected by chance alone. In addition to these heterogeneity statistics, Meta-DiSc computes the inconsistency index (I^2), which has been proposed as a measure to quantify the amount of heterogeneity.

RESULTS

From the literature search, we retrieved 20 English-language articles eligible for analysis. The characteristics of the eligible articles are shown in Table 1.

The details of the quality assessment are shown in e-Table 1. The eligible studies achieved most of the quality items. All 20 studies passed QUADAS items

Table 1—Characteristics of Eligible Studies

Study	Origin	Design	Modality ^a	Patient Type	No. ^a	Us Operator	Diagn Crit ^b	Us				CR			
								TP	FP	FN	TN	TP	FP	FN	TN
Lichtenstein and Menni ⁷	France	NR	Us, CR	Critically-ill	148 ^c	NR	1 ^d	41 (100)	6	0	101 (94.4)	40 (93)	0	3	41 (100)
Hill et al ¹⁹	United States	Retrospect	CR	Trauma	1,684 ^e	NR	24 ^{d,e,f}	41 (100)	5	0	138 (96.5)	107 (65.6)	0	56	1,521 (100)
Lichtenstein et al ⁸	France	Prospect	Us	Critically-ill	184	NR						6 (46.2)	0	7	28 (100)
Goodman et al ²⁰	United Kingdom	Prospect	CR	Postbiopsy	41		31 ^{d,e,f}	66 (100)	11	0	222 (95.3)	60 (85.7)	0	10	58 (100)
Lichtenstein et al ¹⁰	France	Prospect	Us, CR	Critically-ill	299	Intensivist						13 (52)	0	12	1,051 (100)
Holmes et al ²¹	United States	Prospect	CR	Trauma	1,076 ^c							4 (36.4)	0	7	43 (100)
Rowan et al ²²	Canada	Prospect	Us, CR	Trauma	54 ^e	Radiologist	2 ^f	11 (100)	1	0	42 (97.7)				
Kirkpatrick et al ¹⁵	Canada	NR	Us, CR	Trauma	266	Trauma surgeon	2 ^f	21 (48.8)	3	22	220 (98.7)	9 (20.9)	1	34	222 (99.6)
Blaivas et al ¹¹	United States	Prospect	Us, CR	Trauma	352 ^c	Emergency physician	1 ^d	52 (98.1)	1	1	298 (99.7)	40 (75.5)	0	13	299 (100)
Reissig and Kroegel ²³	Germany	Prospect	Us, CR	Postbiopsy	53	Pneumologist	31 ^{d,e,f}	4 (100)	0	0	49 (100)	3 (75)	0	1	49 (100)
Chung et al ²⁴	Korea	NR	Us, CR	Postbiopsy	97	Radiologist	NR	28 ^g (80)	3.75 ^g	7 ^g	58.25 ^g (94)	16.5 ^g (47.1)	3.75 ^g	18.5 ^g	58.25 ^g (94)
Ball et al ¹	Canada	Retrospect	CR	Trauma	676 ^c							46 (44.7)	0	57	573 (100)
Garofalo et al ²⁵	Italy	NR	Us, CR	Postbiopsy	184	NR	3 ^f	44 (95.7)	0	2	138 (100)	19 (41.3)	0	27	138 (100)
Zhang et al ¹³	China	Prospect	Us, CR	Trauma	270 ^e	Emergency physician	3 ^f	28 (87.5)	3	4	235 (98.7)	8 ^h (27.6)	0 ^h	21 ^h	106 ^h (100)
Soldati et al ²⁶	Italy	Prospect	Us, CR	Trauma	372 ^c	Emergency physician	3 ^f	55 (98.2)	0	1	316 (100)	30 (53.6)	0	26	316 (100)
Sartori et al ¹⁶	Italy	Prospect	Us, CR	Postbiopsy	285	NR	3 ^f	8 (100)	0	0	277 (100)	7 (87.5)	0	1	277 (100)
Soldati et al ¹²	Italy	Prospect	Us, CR	Trauma	218	Emergency physician	3	23 (92)	1	2	192 (99.5)	13 (52)	0	12	193 (100)
Ball et al ²⁷	United States	Prospect	CR	Trauma	810 ^e							26 (24.3)	0	81	703 (100)
Brook et al ²⁸	Israel	Prospect	Us, CR	Trauma	338	Radiologist	2 ^f	20 (46.5)	3	23	292 (99)	7 (16.3)	0	36	295 (100)
Galbois et al ²⁹	France	Prospect	Us, CR	Postdrainage	162	Intensivist	3	33 (100)	1	0	128 (99.2)	20 (60.6)	0	13	129 (100)

Data are presented as No. (%), representing the sensitivity (TP) and the specificity (TN). CR = chest radiography; Diagn Crit = diagnostic criteria; FN = false negative; FP = false positive; NR = not reported; Prospect = prospective study; Retrospect = retrospective study; TN = true negative; TP = true positive; Us = ultrasound.

^aIn some articles, only parts of the data that fulfilled the inclusion criteria were analyzed.
^bDiagnostic criteria of pneumothorax by ultrasonography contains the absence of lung sliding sign, absence of comet tail sign, and the presence of lung sliding sign; 1 = the absence of lung sliding sign; 2 = the absence of both lung sliding sign and comet tail sign; 3 = the absence of both lung sliding sign and comet tail sign and the seek of lung point.

^cThe data were reconstructed.

^dSelected for the subgroup analyses: diagnosing pneumothorax by the absence of lung sliding sign.

^eSelected for the subgroup analyses: diagnosing pneumothorax by the absence of comet tail sign.

^fSelected for the subgroup analyses: diagnosing pneumothorax by the absence of both lung sliding sign and comet tail sign.

^gThe data are the mean value of the four observers' numbers.

^hThere was not sufficient detail in the results to recalculate the data per patient as opposed to per hemithorax; thus, original data of the article were used.

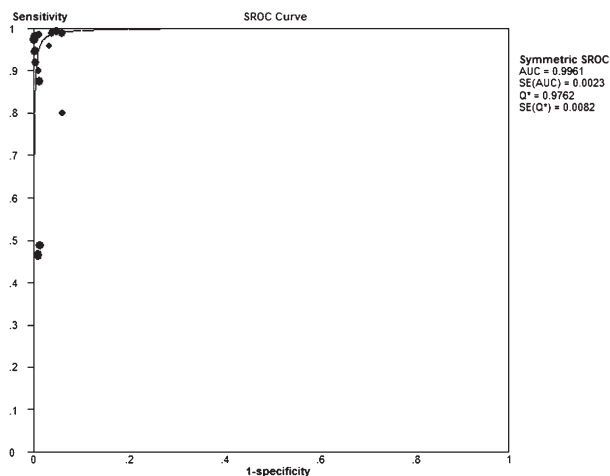
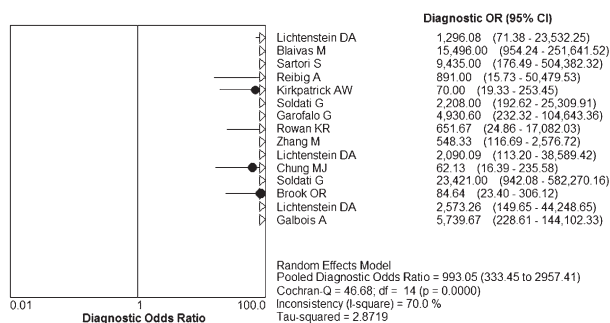
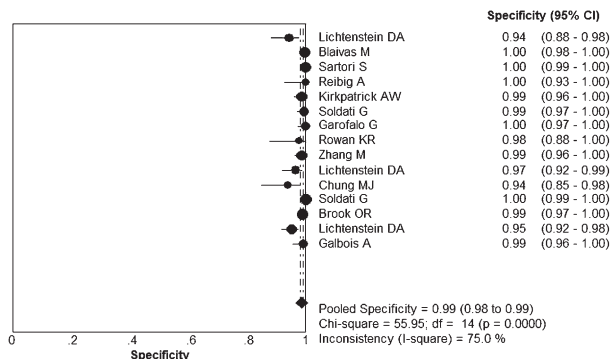
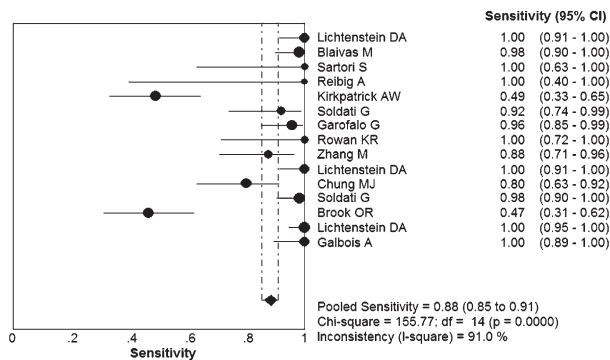


FIGURE 1. Forest sensitivity, specificity, diagnostic OR, and the sROC of ultrasonography. Inconsistency (I^2) describes the percentage of total variation across studies that is due to heterogeneity rather than to chance. I^2 can be readily calculated from basic results obtained from a typical meta-analysis as $I^2 = 100\% \times (Q - df)/Q$. Cochran Q is computed by summing the squared deviations of each study's estimate from the overall meta-analytic estimate. The

2, 3, 8, and 9. The sample of 17 studies (85%) achieved verification using the standard of diagnosis (item 5). QUADAS item 6 (patients received the same reference standard regardless of the index test result) was reported in 75% of the studies. Item 7 (the reference standard was independent of the index test) was achieved in 95% of the studies. Thirteen studies (65%) reported on blinding in the results of the reference test (item 10), whereas six (30%) reported on blinding in the index test results (item 11). Of the nine studies withdrawn from the study, all had an explanation (item 14).

The pooled sensitivity, specificity, diagnostic OR (DOR), and curves for detection of PNx with ultrasonography and CR are shown in Figures 1 and 2, respectively. Pooled sensitivity and specificity were 0.88 (0.85-0.91) and 0.99 (0.98-0.99), respectively, for ultrasonography and 0.52 (0.49-0.55) and 1.00 (1.00-1.00), respectively, for CR. Pooled DOR was 993.05 (333.45-2,957.41), and sROC area under the curve (AUC) was 0.9961 (SE, 0.0023) for ultrasonography. For CR, the DOR was 304.81 (121.94-761.90), and sROC AUC, 0.9435 (SE, 0.0531).

The Spearman correlation coefficient between the log of sensitivity and log of 1-specificity was 0.136 ($P = .629$) for ultrasonography and 0.069 ($P = .778$) for CR. The significant χ^2 P values, shown in the forest plots for each test, implied that there were causes of variations other than a cutoff effect. Possible sources of heterogeneity across the studies were explored using meta-regression analysis with the following covariates as predictor variables: study design (prospective vs retrospective), type of patient (eg, critically ill, trauma), blinded test or not, ultrasonography diagnostic criteria, and operator. Results suggest that the operator is strongly associated with accuracy (relative DOR, 0.21; 95% CI, 0.05-0.96; $P = .0455$) (Table 2). Subgroup analyses based on the ultrasonography operator (clinicians other than radiologists) were performed. In the χ^2 test, pooled sensitivity was 79.93 ($P = .0000$), and pooled specificity was 26.71 ($P = .0004$). The Cochran Q was 25.02 ($P = .0008$) for DOR, which implied that heterogeneity resulted from factors other than the way a study was designed. We considered that the differences between the operators (their skill, experience, knowledge of chest ultrasonography, etc) resulted in this heterogeneity, but there were no sufficient details in the studies for us to make a classification of the operators' skills. The CR and ultrasonography modalities could be judged by their AUC.

τ^2 statistic is a method for random-effects analysis, testing the heterogeneity other than threshold effect. AUC = area under the curve; df = degrees of freedom; sROC = summary receiver operating characteristic.

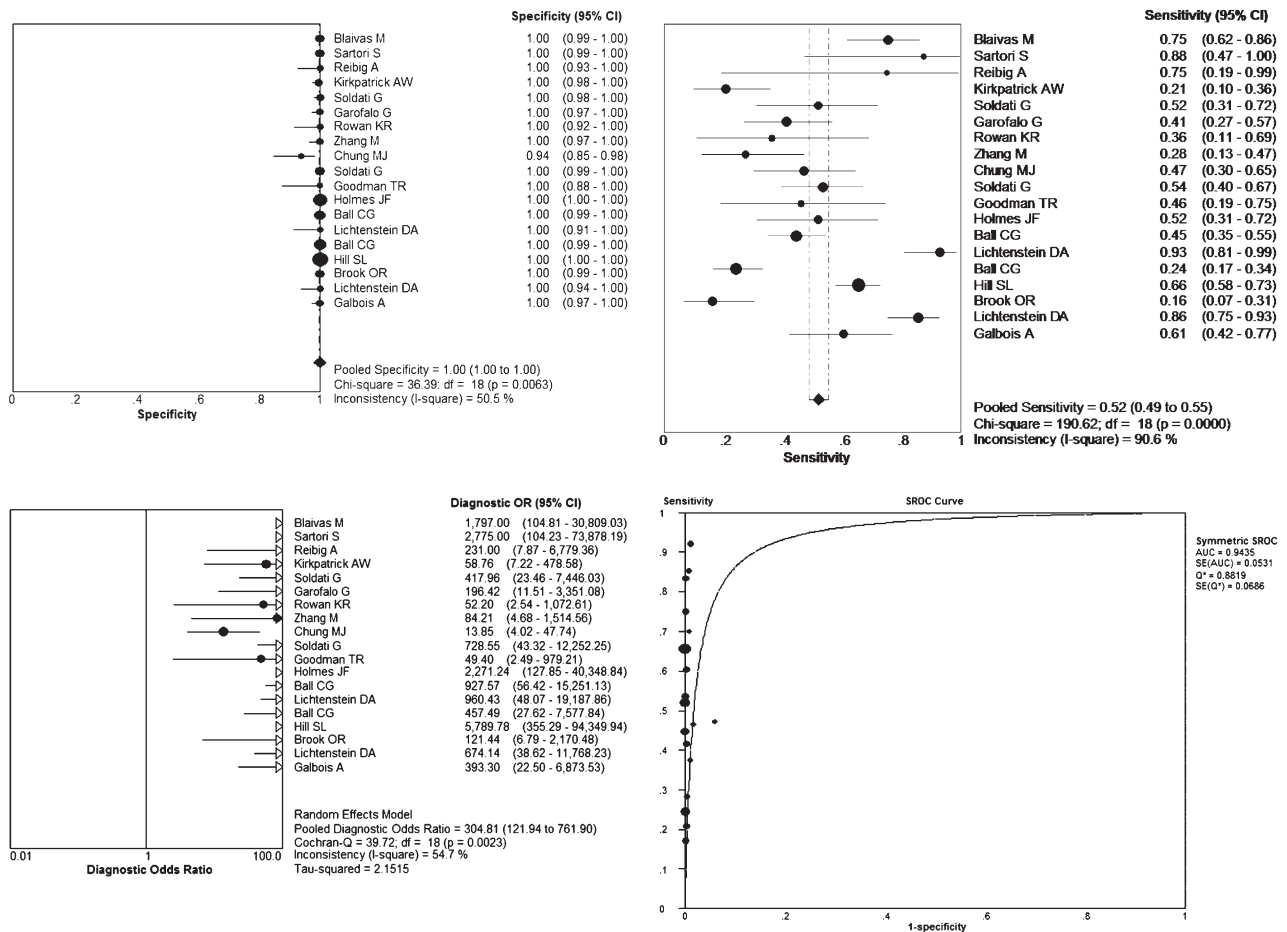


FIGURE 2. Forest sensitivity, specificity, diagnostic OR, and the sROC of chest radiography. See Figure 1 legend for explanation of the statistics and expansion of abbreviations.

We compared the two sROC curves by the Z statistic as follows:

$$Z = \frac{|Q_1^* - Q_2^*|}{\sqrt{SE(Q_1^*)^2 + SE(Q_2^*)^2}} \quad (\text{Equation 1})$$

where Z was 1.36 ($P > .05$), which means that there was no significant difference between the two diagnostic methods for detection of PNx.

The accuracy of ultrasonography performed by clinicians other than radiologists in detecting PNx was analyzed. The pooled results are shown in e-Figure 1 (forest sensitivity, specificity, DOR, and the sROC of ultrasonography performed by clinicians other than radiologists). Pooled sensitivity and specificity were 0.90 (0.87-0.93) and 0.99 (0.98-0.99), respectively. Pooled DOR was 1,676.28 (338.03-8,312.71), and AUC was 0.9981 (SE, 0.0015). The Z statistic compared with the sROC of CR was 1.50 ($P > .05$).

Subgroup analyses based on the ultrasonography diagnostic criteria for PNx were performed, and descriptions of these studies are given in e-Table 2.

The pooled results are shown in e-Figure 2. Only two articles contained extractable data based on the ultrasonography diagnostic criterion of the presence of lung point, which was not enough for a meta-analysis. Compared with one another and with the sROC of CR, all the sROC curves showed no significant difference according to the Z statistic.

DISCUSSION

The results of the present study demonstrate superior sensitivity and similar specificity in the use of ultrasonography compared with CR for the diagnosis of PNx. Using sROC curves derived from the available published articles, we conclude that bedside ultrasonography performed by clinicians other than radiologists is as accurate as CR in detecting PNx. Although there was no statistical significance, it seemed to be more accurate for the diagnosis of PNx when both the lung sliding sign and the comet tail sign were absent in ultrasonography.

Our prior research concluded that ultrasonography allows for a significantly quicker diagnosis of PNx

Table 2—Metaregression Analysis for Possible Sources of Heterogeneity

Variance	Coeff Standard	Error	P Value	RDOR	95% CI
Inverse variance weights 1					
Cte	7.670	2.4070	.0129
S	0.322	0.2557	.2439
Design	1.091	1.0113	.3122	2.98	0.29-30.66
Patient	-1.036	0.9913	.3265	0.35	0.04-3.49
Blind	1.332	1.6944	.4546	3.79	0.08-188.46
Diagnostic					
Criteria	0.168	0.7500	.8281	1.18	0.21-6.67
Operator	-1.568	0.7330	.0648	0.21	0.04-1.13
Inverse variance weights 2					
Cte	8.070	1.3366	.0002
S	0.351	0.2276	.1578
Design	1.239	0.8480	.1779	3.45	0.51-23.52
Patient	-1.190	0.8327	.1867	0.30	0.05-2.00
Blind	1.429	1.5401	.3777	4.17	0.13-136.03
Operator	-1.613	0.6446	.0337	0.20	0.05-0.86
Inverse variance weights 3					
Cte	8.712	1.1164	.0000
S	0.253	0.1950	.2228
Design	1.549	0.7692	.0718	4.71	0.85-26.11
Patient	-0.724	0.6473	.2895	0.48	0.11-2.05
Operator	-1.836	0.5884	.0109	0.16	0.04-0.59
Inverse variance weights 4					
Cte	7.933	0.9033	.0000
S	0.182	0.2023	.3865
Design	1.286	0.8015	.1368	3.62	0.62-21.12
Operator	-1.662	0.6079	.0194	0.19	0.05-0.72
Inverse variance weights 5					
Cte	8.705	0.8616	.0000
S	0.187	0.2310	.4330
Operator	-1.550	0.6948	.0455	0.21	0.05-0.96
Inverse variance weights 6					
Cte	6.837	1.0165	.0000
S	0.295	0.2559	.2710
Design	0.874	1.0900	.4385	2.40	0.22-25.75

The RDOR (obtained by exponentiating the model coefficients) compared the DOR of studies of a given test that lacked a particular methodologic feature with those without the corresponding shortcomings in design. Coeff = coefficient; Cte = constant term in the equation; FPR = false-positive rate; RDOR = relative diagnostic OR; S = indicator of threshold (logit TPR + logit FPR); TPR = true-positive rate.

compared with CR and CT scanning.¹³ The research of Siström and colleagues¹⁵ showed that ultrasonography was not useful in estimating the volume of a PNx, but studies by Garofalo et al,²⁵ Soldati et al,¹² and ourselves¹³ found the opposite. Although there was no statistical significance, from our experience, we recommend that only if there is an absence of both the lung sliding sign and the comet tail sign can a diagnosis of PNx be made. The only part of normal lung visible on ultrasound is the pleura; the artifacts of normal pleura indicate the absence of a pneumothorax. Ultrasonography-based diagnosis of PNx frequently is a “rule out” test. Thus, the presence of both the lung sliding sign and the comet tail sign could rule out PNx, but absence of the lung sliding sign or comet tail sign could not confirm the existence of PNx.³⁰ In our experience, if one of these two signs is absent, the other sign must be carefully examined before a diagnosis of PNx can

be made. The lung point is a specific sign that allows PNx to be confirmed and the PNx volume to be determined,³⁰ but it is rarely found. The use of additional ultrasonography signs, such as the seashore sign and power sliding, could improve the accuracy of the ultrasonography-based diagnosis of PNx, but there were not enough data for us to analyze these separately.

Despite its simplicity, security, and portability, ultrasonography has limitations in the diagnosis of PNx. It may not be appropriate for patients with subcutaneous emphysema, adhesion of pleura, thoracic dressings, pleural calcifications, or skin injury. Slater et al¹⁷ concluded that patients with COPD commonly show signs on ultrasonography that mimic a PNx. Gillman et al³¹ found the so-called “pseudo-lung point” sign by which the diagnosis of PNx should not be made and pockets of air could be missed on ultrasound. In a study by Chung and colleagues,²⁴

the accuracy of ultrasonography depended on the skill of the operator, and the diagnostic accuracy might be lower if ultrasonography was performed by an inexperienced clinician.

The present analysis has some limitations. We did not identify unpublished studies, and no attempt was made to include articles in other languages. From a traditional viewpoint, because air stops the progression of the ultrasound beam, it might seem difficult to detect PNx with ultrasonography. Studies that concluded poor accuracy of ultrasonography or good accuracy of CR in the diagnosis of PNx might not have been published.

From the meta-analysis, the role of bedside ultrasonography in detecting PNx is very promising. It would appear to be an attractive alternative to bedside CR, especially in the emergency department, ICU, and other clinical situations where radiography is not available, such as in medical air transport and remote medical facilities. It has the potential to play a major role in the diagnosis of acute respiratory failure, effectively acting as a visual stethoscope.³²

CONCLUSIONS

Clinician-performed ultrasonography is a reliable tool in the diagnosis of PNx. It has the advantage of portability, simplicity, rapidity, and higher sensitivity and similar specificity compared with CR. Ultrasonography provides a useful adjunct for clinicians in treating patients with multiple trauma or who are ventilated, but the accuracy of ultrasonography in the diagnosis of PNx depends on the skill of the operators.

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Dr Ding: contributed to the study design; data collection, analysis, and interpretation; preparation of the manuscript; and review and approval of the final manuscript.

Dr Shen: contributed to the data collection, analysis, and interpretation; preparation of the manuscript; and review and approval of the final manuscript.

Dr Yang: contributed to the preparation of the manuscript and review and approval of the final manuscript.

Dr He: contributed to the preparation of the manuscript and review and approval of the final manuscript.

Dr Zhang: contributed the study design, preparation of the manuscript, and review and approval of the final manuscript.

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