Accuracy of lung ultrasonography in the hands of non-imaging specialists to diagnose and assess the severity of community-acquired pneumonia in adults: a systematic review

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ABSTRACT

Objectives We aimed to systematically review the published literature regarding adults with clinical suspicion of pneumonia that compares the accuracy of lung ultrasonography (LUS) performed by non-imaging specialists to other reference standards in diagnosing and evaluating the severity of community-acquired pneumonia. Moreover, we aimed to describe LUS training and the speciality of the physician performing LUS, time spent on the LUS procedure and potential harms to patients.

Materials and methods We searched MEDLINE, Embase, CINAHL, Web of Science and Cochrane Central Register of Controlled Trials up until May 2019. We included studies that used LUS to diagnose pneumonia, but also confirmed pneumonia by other means. Publications were excluded if LUS was performed by a sonographer or radiologist (imaging specialists) or performed on other indications than suspicion of pneumonia. Two review authors screened and selected articles, extracted data and assessed quality using Quality Assessment of Diagnostic Accuracy Studies 2.

Results We included 17 studies. The sensitivity of LUS to diagnose pneumonia ranged from 0.68 to 1.00; however, in 14 studies, sensitivity was ≥0.91. Specificities varied from 0.57 to 1.00. We found no obvious differences between studies with low and high diagnostic accuracy. The non-imaging specialists were emergency physicians, internal medicine physicians, intensivists or ‘speciality not described’. Five studies described LUS training, which varied from a 1-hour course to fully credentialed ultrasound education. In general, the methodological quality of studies was good, though, some studies had a high risk of bias.

Conclusion We found significant heterogeneity across studies. In the majority of studies, LUS in the hands of the non-imaging specialists demonstrated high sensitivities and specificities in diagnosing pneumonia. However, due to problems with methodology and heterogeneity there is a need for larger studies with uniform and clearly established criteria for diagnosis and blinding.

PROSPERO registration number Prospectively registered in PROSPERO (CRD42017057804).

INTRODUCTION

Community-acquired pneumonia (CAP) is a frequent and serious health concern, leading to increased morbidity and mortality if not detected and treated properly.1,2 CAP accounts for 2.5% of all patient contacts in Danish general practice3 and globally causes countless hospital admissions, laboratory tests and imaging procedures.4

Today, the typical imaging procedures for diagnosing pneumonia are CT scan of the chest and chest X-ray (CXR), with CT considered the gold standard.5 However, far from all patients have these imaging procedures performed due to high radiation dose, high costs and low availability.6

An alternative mode of imaging is lung ultrasonography (LUS). The advantages of LUS are absence of radiation, high availability and low cost.7 Moreover, LUS can be performed as a bedside point-of-care test to supplement the physician’s clinical
examination. Numerous reviews and meta-analyses indicate that LUS has excellent accuracy for the diagnosis of pneumonia in adults.

None of the existing literature, however, differentiates between LUS operators despite the fact that ultrasound generally is considered a highly user-dependent imaging modality. To our knowledge, no previous review has focussed solely on the accuracy of LUS in the hands of physicians working clinically.

The aim of this study was to systematically review the published literature regarding adults with clinical suspicion of pneumonia that compares the accuracy of LUS performed by physicians working clinically (non-imaging specialists) to other reference standards in diagnosing and evaluating the severity of CAP. Moreover, to describe LUS training and the speciality of the physician performing LUS, time spent on the LUS procedure and potential harms to patients.

METHODS
Data sources and search strategy
This review was prospectively registered in PROSPERO. We followed the Cochrane guideline for conducting a systematic literature review, and the Preferred Reporting Items for Systematic Reviews and Meta-analyses (PRISMA) guideline for reporting the results. The literature search was conducted by a medical librarian and JJS in February 2017 and updated in May 2019. We searched the following databases: MEDLINE and Embase via Ovid, CINAHL via Ebsco, Web of Science and Cochrane Central Register of Controlled Trials.

The search terms ‘ultrasonography’ and ‘pneumonia’ were used in combination and with thesaurus terms (online supplementary e-appendix 1). Reference lists of included articles and identified reviews were evaluated manually for further eligible studies. Patients or the public were not involved in our research. All data relevant to the study are included in the article or uploaded as online supplementary information.

Eligibility and selection of studies
Studies were eligible if a full-text paper with original data was available, the paper described the use of LUS for diagnosing CAP in adults (≥18 years) and the diagnosis of CAP was confirmed by other means, for example, other imaging. Hence, we included all diagnostic accuracy studies that used any reference standard other than LUS. Studies were excluded if not published in English, Danish, Norwegian, or Swedish, if LUS was performed on other indications than suspicion of pneumonia, if LUS was performed by an imaging specialist or if the pneumonia was considered to be ventilator-associated or nosocomial. We defined an imaging specialist as a sonographer or radiologist, and a non-imaging specialist as a physician working clinically.

Two review authors (JJS and PSH or MPH) independently screened the titles and abstracts of all studies identified. Any disagreements were resolved by consensus or by consulting other review authors (CAA and MBJ).

Two review authors (JJS and PSH or MPH) independently extracted data using an adapted version of the Cochrane data exaction template (online supplementary e-appendix 2). We contacted study authors when information about the physician performing the LUS was incomplete or missing, or if important data could not be derived directly from the published study.

Methodological assessment
Methodological quality of the selected studies was evaluated according to the Quality Assessment of Diagnostic Accuracy Studies 2 (QUADAS-2). Two reviewers (JJS and PSH or MPH) independently performed the assessment of methodological quality. Any disagreements were resolved by a third review author (CAA).

Patient and public involvement
No patient involved.

RESULTS
The database search identified 7285 individual, non-duplicate articles, and one potential article was identified through the reference lists (figure 1). Twelve studies had little or no information about the physician performing LUS and we contacted the corresponding authors of these studies. Based on additional information provided by the study authors, two studies were excluded and two studies were included. No elaboration was available for the remaining eight studies. They were thoroughly assessed and four were included, as they clearly described the scanning physicians as a non-imaging specialist physician working clinically. The remaining four studies were excluded.

One study included both patients with CAP and nosocomial pneumonia. However, data on the CAP subgroup was obtained by correspondence with the study authors.

In total, 17 studies describing LUS in the hands of the non-imaging specialist to diagnose CAP in adults were included (figure 1).

Study characteristics
The studies were published between 1996 and 2019; 16 were prospective diagnostic accuracy cohort studies, and one was a retrospective study (online supplementary e-table 1).

The majority of studies included patients admitted to hospital, although one multi-centre study enrolled both hospitalised patients and outpatients (table 1). The studies included between 11 and 356 adult patients with a mean age from 34.0 to 84.8 years, of whom between 47% and 93% were men. Two studies included only patients aged ≥65 years.

The signs and symptoms of pneumonia described in the American Thoracic Society guidelines (cough, pleuritic
pain, sputum production, fever and dyspnoea) were used as inclusion criteria in nine studies, and six studies based inclusion on comparable, but not identical, criteria. The remaining two studies only included patients with respiratory complaints like cough, dyspnoea, chest pain or haemoptysis leading to a chest CT being ordered.

Definition of pneumonia based on LUS varied across studies. Still, presence of subpleural or alveolar consolidation or a tissue-like lesion was part of the definition in all studies except one, in which no definition was described. The physicians performing and interpreting LUS were generally blinded to the reference standard; however, in four studies, this matter was unclear.

Diagnostic accuracy of LUS

Diagnostic accuracy is presented in Table 1. The sensitivity of LUS to diagnose CAP ranged from 0.68 (95% CI, 0.52 to 0.81) to 1.00 (95% CI, 0.95 to 1.00); in 14 of the 17 studies, it was ≥0.91. The specificity could be calculated in 13 of the studies. It varied from 0.57 (95% CI, 0.34 to 0.78) to 1.00 (95% CI, 0.92 to 1.00), but in seven studies, it was ≥0.94. We found no systematic differences between studies with low and high diagnostic accuracy in terms of study setting, participant training or experience, or choice of reference standard. Inter-observer agreement was reported in two studies with κ-values of 0.83 and 0.90.

The studies by Liu et al and Amatya et al were the two studies of highest methodological quality (Table 3). Both studies compared LUS to CT (Table 1) and LUS was performed by emergency physicians whose prior experience and training was described (Table 2). However, they differed with regards to procedure and characteristics of LUS in terms of areas examined and definition of pneumonia on LUS (Table 2). They found sensitivities of, respectively, 0.95 (95% CI, 0.89 to 0.98) and 0.91 (95% CI, 0.78 to 0.98) and specificities of 0.99 (95% CI, 0.92 to 1.00) and 0.61 (95% CI, 0.36 to 0.83).

None of the studies compared sonographic findings to clinical outcomes. Three studies assessed the severity of pneumonia in patients with either CURB-65 or Pneumonia Outcome Research Team score, but these were not compared with LUS findings. Bourcier et al stratified their results according to onset of symptoms of pneumonia (<24 hours versus >24 hours). They found that LUS (sensitivity of 0.97) was significantly more effective than CXR (sensitivity of 0.30) in diagnosing pneumonia when time from clinical onset was <24 hours.

Speciality and training of non-imaging specialists

Information about speciality, experience and training of physicians performing LUS is presented in Table 2. LUS was performed by emergency physicians, internal medicine physicians and by intensivists, while four studies did not declare the specific speciality of the non-imaging specialists.

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LUS varied from 1 week in the emergency department to more than 10 years of clinical experience.

Five studies described a LUS training programme for the participating physicians. Three studies provided a reference for an established educational programme, whereas the remaining studies described training specifically designed for their study. All training programmes included both theoretical and practical sessions. A large variation in the extent of the training programmes was noted, ranging from a few hours at a course facility to completion of a European Federation of Societies for Ultrasound in Medicine and Biology (EFSUMB) level one qualification. Four studies reported the time spent performing LUS, which was overall <10 min.

### Potential harms to patients

Twelve studies reported false positive results from LUS, and 14 studies described false negative results (table 3). Corradi et al reported a high number of false negative results as they found 14 (22%) false negative hemithorax LUS examinations. However, five of these were reported in patients with bilateral pneumonia, in whom LUS examination only detected pneumonia in one hemithorax. Moreover, Corradi et al described that LUS-positive pneumonia were larger in diameter (81±55 mm) and close to the pleural line (1±3 mm). Likewise, more studies described false negative results that were mainly seen in patients with small consolidations where pneumonia did not reach the pleura.

Parlamento et al reported two incidental findings of subpleural consolidations in patients without pneumonia. In both cases, LUS findings were verified by chest CT scan and confirmed to be, respectively, an atelectasis caused by a large pleural effusion and a case of pulmonary embolism.
Table 2  Speciality of non-specialists, experience and training in LUS

| Study                | Number of and speciality of physicians performing LUS | Prior experience in LUS or ultrasonography in general | Description of training in LUS                                                                 | Time consumption on LUS |
|----------------------|------------------------------------------------------|-------------------------------------------------------|-------------------------------------------------------------------------------------------------|-------------------------|
| Amatya et al18       | Four emergency resident physicians                   | One week of performing LUS in the ED.                  | One hour lecture on LUS. Five pre-enrolment LUS scans and interpretation reviewed by expert sonographer. | 7 min 9 s (SD 1 min 57 s) |
| Corradi et al18      | One intensivist with PhD in US*                        | More than 10 years of experience in LUS*               | –                                                                                                | –                       |
| Fares et al22        | A single physician                                    | –                                                     | –                                                                                                | –                       |
| Karimi et al39       | Trained emergency residents under supervision of the attending emergency specialist in charge | –                                                     | –                                                                                                | –                       |
| Liu et al31          | Three emergency physicians                           | At least 50 cases of LUS examination                  | Twenty-eight hours course based on US emergency medicine guidelines issued by the American College of Emergency Physicians in 2001 | –                       |
| Nazerian et al32     | Four internal medicine and emergency medicine attending physicians. Four resident physicians (two internal medicine and two emergency medicine) | Attending physicians; at least 5 years of experience in POC-US Resident physicians; at least 1 year of training in emergency US | –                                                                                                | –                       |
| Taghizadieh et al35  | One emergency specialist                              | –                                                     | –                                                                                                | –                       |
| Parlamento et al34   | One emergency physician                               | Thirty years of experience in general and cardiac US and 10 years of training in LUS | –                                                                                                | <5 min                  |
| Reissig et al39      | Experienced physicians (number and speciality not described) | At least 100 chest US procedures done prior to study | Three hours of didactic and 3 hours of hands-on thoracic US taught by an experienced radiology specialist to learn the diagnostic criteria of alveolar consolidation | <10 min                 |
| Unluer et al37       | Three attending emergency physicians                  | –                                                     | –                                                                                                | –                       |
| Benci et al17        | Physicians (number and speciality not described)      | Considerable experience in US techniques               | –                                                                                                | –                       |
| Bitar et al29        | Intensivist (number not described)                    | –                                                     | –                                                                                                | –                       |
| Bourcier et al30     | Five emergency physicians                             | –                                                     | Two days of theoretical formation alternating with practical ultrasounds sessions in groups of three people | –                       |
| Cipollini et al37    | Internal medicine specialist*                          | More than 1 year of bedside US experience*            | –                                                                                                | –                       |
| Cortellaro et al19   | One expert operator                                   | –                                                     | –                                                                                                | <5 min                  |
| Pagano et al33       | Five trained emergency physicians                     | More than 2 years of experience in LUS                | –                                                                                                | –                       |
| Ticinesi et al36     | Three internal and emergency medicine physicians      | More than 1 year of bedside US experience             | Level one of training completed according to the guidelines by the EFSUMB                       | –                       |

*Data collected by correspondence with author.
- not described; EFSUMB, European Federation of Societies for Ultrasound in Medicine and Biology; LUS, lung ultrasonography; POC-US, point-of-care ultrasonography; US, ultrasonography.

**DISCUSSION**

To our knowledge, this is the first systematic review to focus specifically on LUS to diagnose CAP in adults in the hands of non-imaging specialists physicians working clinically. These non-imaging specialists were emergency physicians, internal medicine physicians, intensivists or unclassified physicians and obtained LUS sensitivities and specificities that were typically above 0.90. We found no overall difference in diagnostic accuracy when compared with study setting or the physicians’ speciality, experience or training. Importantly, the variation in sensitivitiy and specificity was found across reference standards. No study compared sonographic findings to the severity of pneumonia. Only a few studies described LUS training of the non-imaging specialists, and these training programmes varied from short lectures to fully accredited ultrasound education.
We highlighted the results of Liu et al and Amatya et al due to the quality of the studies, but the studies were not completely comparable in other parameters. Both studies found high and comparable sensitivities of 0.95 (95% CI, 0.89 to 0.98) and 0.91 (95% CI, 0.78 to 0.98), respectively. However, in Amatya et al, LUS specificity was 0.61 (95% CI, 0.36 to 0.83) and significantly lower than the specificity in Liu et al of 0.99 (95% CI, 0.92 to 1.00). According to Amatya et al, this was due to a higher prevalence of pulmonary co-morbidities which resulted in false positive LUS results. Low specificity may lead to over-diagnosis of pneumonia and inappropriate use of antibiotics.

The diagnostic accuracy of LUS for diagnosing pneumonia described in this review is consistent with results from previous reviews that made no distinction between imaging specialists and physicians working clinically.8–13 Recently, Orso et al obtained a pooled sensitivity of 0.92 and a specificity of 0.93 in a review based on studies performed in emergency departments.40 Of course, the majority of LUS operators were emergency physicians, corresponding to the non-specialists in the present review. Consequently, Orso et al and this study have included many of the same studies. However, Orso et al also included studies with imaging specialists and patients with ‘acute respiratory failure’. Our review included LUS performed by non-imaging specialists from different specialties and in different settings. One study was even partly conducted in outpatient settings with non-hospitalised patients.20

| Study               | True positive LUS results, N (%) | False positive LUS results, N (%) | False negative LUS results, N (%) | True negative LUS results, N (%) | Nature of false positive LUS results                                                                 |
|---------------------|---------------------------------|----------------------------------|-----------------------------------|---------------------------------|---------------------------------------------------------------------------------------------------|
| Amatya et al46      | 40 (64.5)                       | 7 (11.3)                         | 4 (6.5)                           | 11 (17.7)                       | Three bronchiectasis, two interstitial lungs, one tuberculosis and one normal lung               |
| Corradi et al18     | 30* (46.8)                      | 1* (1.6)                         | 14* (22.0)                        | 19* (29.6)                      | –                                                                                                 |
| Fares et al22       | 28 (73.7)                       | 2 (5.3)                          | 2 (5.3)                           | 6 (15.7)                        | –                                                                                                 |
| Karimi et al23      | 263 (93.9)                      | 0 (0.0)                          | 17 (6.1)                          | 0                               | –                                                                                                 |
| Liu et al31         | 106 (59.2)                      | 1 (0.6)                          | 6 (3.4)                           | 66 (36.8)                       | –                                                                                                 |
| Nazerian et al32     | 72 (25.3)                       | 9 (3.1)                          | 15 (5.3)                          | 189 (66.3)                      | Three cancers, three parenchymal impaired ventilation not due to infection and three pulmonary fibrosis |
| Taghizadieh et al64  | 29 (96.7)                       | 1 (3.3)                          | 0                                 | 0                               | –                                                                                                 |
| Parlamento et al24   | 31 (63.3)                       | 0 (0.0)                          | 1 (2.0)                           | 17 (34.7)                       | –                                                                                                 |
| Reissig et al29     | 211 (59.3)                      | 3 (0.8)                          | 15 (4.2)                          | 127 (35.7)                      | –                                                                                                 |
| Unluer et al27      | 27 (37.5)                       | 7 (9.7)                          | 1 (1.4)                           | 37 (51.4)                       | Four pulmonary embolisms and three exacerbations of COPD                                       |
| Benci et al17       | 37 (46.3)                       | 0 (0.0)                          | 0 (0.0)                           | 43 (53.7)                       | –                                                                                                 |
| Bitar et al29       | –                               | –                                | –                                 | –                               | –                                                                                                 |
| Bourcier et al90    | 117 (81.2)                      | 9 (6.3)                          | 6 (4.2)                           | 12 (8.3)                        | Four sepsis of other origin, two pulmonary embolisms, one ARDS, one pulmonary fibrosis and one acute anaemia |
| Cipollini et al27    | 105 (82.0)                      | –                                | 23 (18.0)                         | –                               | –                                                                                                 |
| Cortellaro et al19  | 80 (66.7)                       | 2 (1.7)                          | 1 (0.8)                           | 37 (30.8)                       | One congestive heart failure and one subphrenic abscess with lung atelectasis                  |
| Pagano et al13      | 67 (63.8)                       | 13 (12.4)                        | 1 (1.0)                           | 24 (22.8)                       | Seven exacerbations of COPD, two congestive heart failure, three cancers and one pulmonary infarction |
| Ticinesi et al36    | 88 (52.1)                       | 3 (1.8)                          | 8 (4.7)                           | 70 (41.2)                       | Two pulmonary embolisms and one cancer                                                           |

* Hemithoraxes.
- not described; ARDS, acute respiratory distress syndrome; COPD, chronic obstructive pulmonary disease; LUS, lung ultrasonography; N, number.
Importantly, the results of this particular study did not differ from the remaining studies. Hence, LUS might also be applied on non-hospitalised patients with suspected CAP, which supports the vision that LUS could be a useful tool for any clinician in the future.41

Non-imaging specialists working in primary care are first in line to see patients with CAP and general practitioners have already begun using point-of-care ultrasound.42 43

The results by Bourcier et al suggest that LUS is a better diagnostic tool for achieving an early diagnosis (≤24 hours from clinical onset) compared with CXR. The ability of LUS to accurately diagnose pneumonia early in the course of the disease may improve outcomes for patients attending primary care.45 Furthermore, improved diagnostic performance in patients with suspected CAP may reduce the need for antibiotics. On the other hand, the size of pulmonary lesions might be smaller in the early stages of disease and the results indicate that the usability of LUS to diagnose CAP is compromised by its inability to visualise pulmonary lesions that are not in contact with the pleura. However, according to Lichtenstein et al who looked for lung consolidation in intensive care patients, this occurred in only 1.5% cases of lung consolidation.44

Due to a lower prevalence and less severe disease in a general practice population, further evaluation of LUS in the diagnosis of CAP in general practice is required.

LUS is a user-dependent examination and several guidelines14 45 46 stress that diagnostic performance requires sufficient training to gain the necessary competencies. A meta-analysis by Tsou et al found a significant difference in diagnostic accuracy between LUS performed by ‘advanced’ versus ‘novice’ sonographers in the diagnosis of pneumonia in children.47 However, they defined ‘novice sonographers’ as physicians with little or no prior LUS experience or training (≤7 days); most of the non-imaging specialists in the present review would be classified as ‘advanced sonographers’ according to this definition. Though, the learning curve appears steep from paediatric data and in a randomised controlled trial by Jones et al,48 they found that substitution of CXR with LUS when evaluating children suspected of having pneumonia was feasible and safe, also in the hands of novice sonographers (≤25 examinations). Today, there are no guidelines or recommendations specifying the amount of training or level of competence needed to perform LUS.49 50

As this review has shown, however, these competencies can be reached by the non-imaging specialist physician even after a short, tailored training programme. To ensure that physicians maintain and develop skills over time and learn to incorporate LUS findings into clinical decision-making, longitudinal training elements must be incorporated into the training programmes.50

This study describes the different specialties of the non-imaging specialists and demonstrates great heterogeneity in their prior experience and training in LUS. However, sensitivities and specificities are comparable, thereby implying that LUS can be performed by physicians in various specialties, and by less experienced physicians, with comparable results to those of physicians with considerable experience in LUS.

Limitations

The aim of this study was to describe the diagnostic accuracy of LUS for diagnosing CAP when performed by physicians with considerably less ultrasound experience than imaging specialists. In four of the included studies, the speciality of the physician was not reported.17 19 20 22 These studies were included as we assessed from the clinical setting that the physicians were not radiologists or sonographers. The results from these four studies did not differ from the remaining studies. Furthermore, while some of the physicians had extensive experience with LUS,17 19 34 and their ultrasonography competencies may be compared with those of an imaging specialist, we did not find, in general, that sensitivity and specificity increased with experience. Comparison of studies was difficult due to sparse information on the non-imaging specialists’ training, their experience with LUS and the heterogeneity in the reference standards used. Due to the significant heterogeneity across studies, it was not appropriate to pool data and perform a meta-analyses.

CONCLUSION

We found significant heterogeneity across studies. In the majority of studies, LUS in the hands of the non-imaging specialists demonstrated high sensitivities and specificities in diagnosing pneumonia. However, due to problems with methodology and heterogeneity, there is a need for larger studies with uniform and clearly established criteria for diagnosis and blinding.

Contributors JJS is the guarantor of the study. JJS, PSH, MPH, MBBJ, OG and CAA contributed to the concept, design and drafting of the study. JJS, PSH and MPH conducted the systematic search strategy and the review. All authors revised critically and approved the final manuscript.

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