

Lung Ultrasound in Diagnosis of Acute Respiratory Failure: BLUE Protocol Based Evaluation

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ABSTRACT

Introduction: Acute respiratory failure is a distressing situation for the patient, demanding dynamic evaluation and interventions. Lung ultrasound is a bedside technique, very promising in this setting. Aims/Objectives: To evaluate the diagnostic accuracy of the lung ultrasound algorithm (BLUE protocol) in ICU patients admitted with acute respiratory failure.

Methodology: This study was conducted in the department of respiratory medicine in consensus emergency medicine in Pariyaram Medical College, Kannur over a period of 1 year investigating 111 consecutive adult patients who presented with acute respiratory failure. A double blinded screening lung ultrasonography was performed according to the blue protocol, and was compared with final diagnosis. Uncertain diagnoses and rare causes were excluded. Three items were assessed: artefacts (horizontal A lines or vertical B lines indicating interstitial syndrome), lung sliding, and alveolar consolidation and/or pleural effusion.

Results: 111 patients were evaluated (63.2% male), mean age was 62 years (SD \pm 12.9). Anterior absent lung sliding plus A lines plus lung point indicated pneumothorax with 68% sensitivity, 100 % specificity and an accuracy of 94.1%. A normal anterior profile plus deep venous thrombosis indicated pulmonary embolism with 50% sensitivity, 96.6% specificity

and 97% accuracy. Multiple anterior diffuse B lines with lung sliding indicated pulmonary oedema with 90.5% sensitivity, 97.5% specificity and 96% accuracy. C profile and A/B profile showed > 80% accuracy in pneumonia.

Conclusion: Lung ultrasound, as proposed in the BLUE protocol, has good accuracy, and seems reproducible and useful in acute respiratory failure.

Key Words: Lung Ultrasound, Acute Respiratory Failure, BLUE Protocol.

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INTRODUCTION

Acute respiratory failure is a distressing situation, demanding dynamic evaluation and interventions. It is frequently encountered in intensive care units (ICUs), remains a major cause of morbidity and mortality.^{1,2} Furthermore, there are often situations of severe respiratory distress in which an urgent diagnosis is required within minutes to direct potentially life-saving therapy.^{3,4}

Management of such critically ill patients requires imaging techniques, which are essential for optimizing diagnostic and therapeutic procedures. To date, chest imaging has relied on bedside chest radiography and lung computed tomography (CT).⁵ To perform a lung CT scan, however, requires transportation to the department of radiology, a risky procedure necessitating the presence of trained physicians and sophisticated cardio-respiratory monitoring.⁶ In addition, helical multi-detector row CT exposes the patient to a substantial radiation dose, which limits the repeatability of the procedure.⁷ General and cardiac ultrasound can be easily performed at the bedside by physicians working in the intensive care unit (ICU) and may provide accurate

information with diagnostic and therapeutic relevance. It has become an attractive diagnostic tool in a growing number of situations, including evaluation of cardiovascular status, acute abdominal disease such as peritoneal collections, hepatobiliary tract obstruction, acalculous acute cholecystitis and diagnosis of deep venous thrombosis.⁸ Lung ultrasound is an imaging modality which is easily available at bedside, is real time and free of radiation hazards in comparison to conventional imaging. Lichtenstein and colleagues proposed an algorithm approach with a diagnostic accuracy of 90.5%, named Bedside Lung Ultrasound in Emergency (BLUE protocol).⁹ Chest ultrasonography, enables a quick bedside examination of the patient, the patient does not absorb any ionizing radiations, and the scan can be performed by the emergency physician (EP), who can immediately integrate the findings with the clinical data.¹⁰

The aim of the current study was to evaluate the diagnostic accuracy of the lung ultrasound algorithm (BLUE protocol) in ICU patients admitted with acute respiratory failure.

MATERIALS AND METHODS

The present study was done in the intensive care units in the respiratory medicine department, Pariyaram Medical College, Kannur, Kerala. It was a hospital based observational study (double blinded). The period of study was one year. We prospectively recruited 111 adult patients admitted for acute respiratory failure (ARF) to the ICU whose final diagnosis was established in the hospitalization report using standardized tests by the pulmonologists (at discharge).

Acute respiratory failure was defined based on the classical clinical and biological criteria of admission to the ICU. Patients were diagnosed with acute respiratory failure clinically according to symptoms and signs of hypoxemia with or without hypercapnia, and biologically by arterial blood gases indicating very low oxygen tension (PO2 less than 60 mmHg) or hypoxic index less than 300 with or without hypercapnia (CO2 more than 45 mmHg) and low PH.¹¹

All patients underwent standard medical care including History, clinical examination, routine lab investigations, Plain X-ray chest and CT chest or other imaging techniques (when needed) and other diagnostic tools when needed (as pleura aspirate examination and fibro-optic bronchoscopy).The diagnosis of the underlying etiology of acute respiratory failure was established by exhausting all investigations and Standardized tests by the pulmonologists and not including lung U/S. Thoracic ultrasound examination was done for all patients by trained emergency

medicine physician who did not participate in the patients' management by using Micromaxx ultrasound sonosite machine. The ultrasound was performed to all patients on admission without interrupting their management. Ultrasound was performed with both deep (2.5 MHZ) and superficial (5 MHZ) probes in a semi-recumbent position, or supine if intubated.

Regarding ultrasound examination, signs with dual answers (Absent or present) were assessed, as follows: artifact analysis (A or B lines), lung sliding, abnormal ultrasonic shadow (alveolar consolidation or pleural effusion). A lateral sub-posterior search for postero lateral alveolar and/or pleural syndrome (PLAPS) was essential. Other ultrasonic signs such as lungpoint sign and sinusoid signs were reported when they were identified (Table 1).^{9,12,13} In patients suspected to have pulmonary embolism, deep venous thrombosis was sought using the same probe.

Visualization of anatomic echoic intraluminal thrombosis or absence of compressibility was considered as a positive finding. The signs observed in each disease were methodically collected; then the ultrasound data were compared with the diagnosis established by the pulmonologist. Uncertain diagnoses and rare causes were excluded: patients with non-respiratory as or rare causes of acute respiratory failure and patients given multiple final diagnoses at the end of hospitalization.

Statistical Analysis

Data were collected, tabulated, and then analyzed using the SPSS software package version 16.

Table 1: Description of common chest ultrasonic signs

Ultrasonic sign	Ultrasonic description
Sliding sign	Dynamic transverse twinkling movement visible at the pleural line and synchronized with respiration
A lines	Horizontal lines parallel to the pleural line recurring at regular intervals (normal sign)
B lines	Vertical pleural based lines (normal sign if less than three lines per view)
Consolidation	Hypoechoic shadow
Pleural effusion	Appears mostly as an anechoic, homogeneous space between parietal and visceral pleura
Lung point	Sudden, on-off visualization of a lung pattern
Sinusoids sign	Inspiratory centrifugal shifting of the visceral pleura with decrease in apparent thickness



Figure1: Ultra Sound Machine

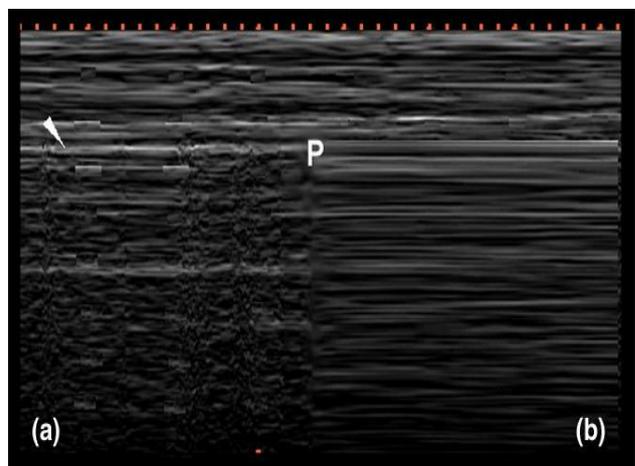


Figure 2: Lung Point In M-Mode

RESULTS

This study included 111 patients with acute respiratory failure. 10 patients were excluded of the analysis because they had rare/uncertain diagnosis. Out of 101 patients, 64 (63.4%) were

male and 37 females (36.6%). Their mean age was 62 years (SD ± 12.9). Average time taken per person was 10-15 min. At inclusion in the study, patients had a mean PaO2 of 51.47 ±8.43 mmHg and a mean PaCO2 of 52.10 ±24 mmHg.

Table 2: Distribution of the sample according to age

Age	Count	Percent
<=50	21	20.8
51 – 60	23	22.8
61 – 70	29	28.7
>70	28	27.7
Mean ± SD	62 ± 12.9	

Table 3: Distribution of the sample according to sex

Sex	Count	Percent
Male	64	63.4
Female	37	36.6

According to history, clinical examination and investigations (not including ultrasound examination), the final etiological diagnosis of acute respiratory failure in the 111 studied patients was: pneumonia 22 cases (CAP 15 cases and HAP 5 cases, aspiration pneumonia 2 cases), acute exacerbation of chronic obstructive pulmonary disease (AECOPD) 21 cases, pulmonary edema 21 cases, pulmonary embolism (PE) 6 cases, massive effusion/empyema 5 cases, bronchial asthma (BA) 12 cases, pneumothorax 19 cases, idiopathic pulmonary fibrosis (IPF) 3 cases and no/multiple diagnosis in 2 cases. (Table 4)

US Examination

To facilitate the comparison of the ultrasound data to the final etiological diagnosis obtained by the conventional diagnostic tools in the ICU, the characteristic ultrasound signs were collected together and interpreted as different ultrasound profiles according to the previously designated nomenclatures.^{9,13} The A profile designates anterior predominant bilateral A lines associated with

lung sliding (with possible focalized B lines).The A' profile is an A profile with abolished lung sliding and without lung point. The B profile designates anterior predominant bilateral B lines associated with lung sliding (with possible focalized A lines). The B' profile is a B profile with abolished lung sliding. The A/B profile designates anterior predominant B lines on one side and predominant A lines on the other. The C profile designates anterior alveolar consolidation(s). PLAPS represents posterior and/or lateral alveolar and/or pleural syndrome. All these definitions are based on the patient being supine or semi recumbent. The normal profile associates the A profile without PLAPS (regardless of posterior A or B lines).

The ultrasound profiles of different final underlying etiologies of ARF: (Table 5)

Pneumonia was presented by: A (normal) profile in 9% of patients, AB profile in 23% of patients, A+ PLAPS profile in 23% of patients ,B' profile in 14% and C profile in 32% of patients.

Acute exacerbation of chronic obstructive pulmonary disease (AECOPD) was presented by: A (normal) profile in 66.6% of patients, A' profile in 19.1%, B profile + PLAPS in 9.5% and A+ PLAPS profile in 4.7% of patients. Pulmonary edema was presented by: A (normal) profile in 9.5% of patients, and B + PLAPS profile in 90.5% of patients.

Pulmonary embolism (PE) was presented by: A (normal) profile in 33.3%, C profile in 17% of patients and A+ PLAPS + DVT profile in 50% of patients.

Bronchial asthma (BA) was presented by: A (normal) profile in 91.7% of patients and A +PLAPS profile in 8.3% of patients.

Pneumothorax was presented by: A 'profile in 31.6% of patients and lung point profile in 68.4% of patients.

Rare diagnosis, unknown diagnosis and multiple diagnosis cases were omitted.

Table 4: Final Diagnoses and Methods of Diagnosis

DIAGNOSIS	METHODS
For all patients	History, clinical examination, radiography read by radiologists, CT when available
Pulmonary oedema n= 21	Favourable clinical progression under treatment
Pneumonia n=22	Evaluation of cardiac function using echocardiography, functional tests, and American Heart Association recommendations
COPD n=21	Clinical profile, x-ray chest finding, microorganism isolated (blood, sputum or by invasive tests), recovery with antibiotics. Included were infectious, aspiration, community, or hospital-acquired pneumonia.
Acute asthma n=12	Diagnosed by clinical, radiological and spirometric evidence of irreversible obstructive airway disease.
Pulmonary embolism n=6	History, spirometry, responds to bronchodilator treatment
Pneumothorax n=19	CT pulmonary angiogram
	X-ray chest, CT thorax (if needed)
Excluded patients	
Rare n = 8	Chronic diffuse interstitial disease (n = 3), effusion/empyema (n =5),
No final diagnosis n = 1	Unknown diagnosis at the end of hospitalization
Several final diagnoses n=1	Pulmonary oedema plus pneumonia

Table 5: Ultrasound profile in relation to final etiological diagnosis

Profile	Frequency in all cases	Differential diagnosis	In relation to final diagnosis	
			Frequency	%
A	31	Pulmonary oedema	2	9.5
		AECOPD	14	66.6
		Asthma	11	91.7
		PE	2	33.3
		Pneumonia	2	9
A'	10	Pneumothorax	6	31.6
		AECOPD	4	19.1
AB	5	Pneumonia	5	23
A + PLAPS	7	AECOPD	1	4.7
		Asthma	1	8.3
		Pneumonia	5	23
B	0			
B + PLAPS	21	Pulmonary oedema	19	90.5
		AECOPD	2	9.5
B'	3	Pneumonia	3	14
C	8	PE	1	17
		pneumonia	7	32
A + PLAPS + DVT	3	PE	3	50
Lung point	13	Pneumothorax	13	68.4
TOTAL	101		101	

Table 6: Accuracy of USG profile

Final diagnosis	USG Profile	Sensitivity (%)	Specificity (%)	Positive Predictive value (%)	Negative Predictive value (%)	Accuracy (%)
COPD + asthma	A Profile	75.76	91.2	80.7	88.6	86.1
Pneumonia	A/B Profile	22.7	100	100	82.3	83.2
	C Profile	31.8	98.7	87.5	83.9	84.2
	Lung point Profile	68.4	100	100	93.2	94.1
Pulmonary embolism	A' profile	31.6	95.1	60	85.7	77.2
	A Profile with Plaps + DVT	50	96.9	50.0	96.9	97.0
Pulmonary edema	B Profile with Plaps	90.5	97.5	90.5	97.5	96.0
	A profile	9.5	63.75	6.5	72.86	52.5

Accuracy of ultrasound diagnosis

As some profiles were presented in more than one aetiology, the sensitivity, specificity, positive predictive value and negative predictive value were calculated to determine the accuracy of each profile to diagnose each aetiology of acute respiratory failure.

Characteristic profiles that produced specificities >90% for each of the final etiologic diagnosis are presented in Table 6.

DISCUSSION

The key inference from our study was the significant diagnostic accuracy of applying chest ultrasound in comparison to the standard approach that included clinical, radiologic and biologic data. Moreover, the greater diagnostic ability of the ultrasound

approach was demonstrated when each main etiologic entity of acute respiratory failure was independently analysed.

The main diagnostic profiles were as follows:

The AB profile (asymmetric anterior interstitial syndrome) and the C profile (anterior consolidation) indicated pneumonia. Since pneumonia has numerous causes, several pathologic and radiologic presentations, and can be found in a wide variety of locations, it generates, several profiles, mainly asymmetry (from left to right, explaining the A/B profile, from posterior to anterior, explaining the A-profile plus PLAPS), and anterior consolidations (C-profile).^{9,14} The AB and the C profiles were the profiles which had a high specificity for pneumonia in our study, 100% and 98% respectively, which is consistent with the study by Rasha Dadit et al.¹⁵

COPD and asthma are bronchial diseases assumed to yield a normal lung surface. They are diagnosed mainly by their clinical presentations and are not usually accompanied by significant radiologic presentations that can be echoed by ultrasound, explaining the predominance of the A profile (i.e., normal lung surface) in these patients.⁹ Our study showed less sensitivity and specificity compared to the Lichtenstein et al's study.

Alveolar interstitial syndrome (AIS) constitutes a group of diseases that is caused by an increase in lung fluid and/or a reduction in its air content. The major causes of AIS are pulmonary oedema, interstitial pneumonia and pulmonary fibrosis. The sonographic appearance of AIS is vertical artefacts called B lines. When compared with CT scan images, B-lines correspond to oedema of the interlobular septa. When diffuse B-lines are visualized, this gives the B profile and suggests a diagnosis of AIS. The causes of diffuse AIS presented to the ICUs in our study were pulmonary oedema and interstitial pulmonary fibrosis. The B profile (with or without PLAPS due to gravitational filling of dependent alveoli) characterizes pulmonary oedema with high accuracy of 96%.^{16,17} Lung ultrasound for the diagnosis of pulmonary embolism (PE) is a relatively new concept. There are several limitations of lung ultrasound to diagnose PE. Only two-thirds of the lung area is accessible to ultrasound examination because the remainder is covered by bony structures. However, almost 80% of lesions are located in the lower lobes, which are accessible to examination.¹⁸ Only thromboembolic lesions extending to the pleura can be detected. However, it has been demonstrated that central and peripheral lesions occur concurrently in 80% of cases.¹⁹ Furthermore, the criteria used to diagnose PE are not standard. However, Lichtenstein proposed simplified criteria to diagnose PE. According to his Bedside Lung Ultrasound in Emergency (BLUE)

protocol, in the severely dyspnoeic patient not in shock, a normal lung ultrasound (presence of A-lines throughout) plus signs of deep vein thrombosis (DVT) is 81% sensitive and 99% specific for PE.⁹ Similar results were found in our study where, in pulmonary embolism where A-profile associated with deep vein thrombosis presented a sensitivity of 50% and specificity of 96%, suggesting that the search for venous thrombosis should be associated with the lung analysis.

In pneumothorax, we found that, absent lung sliding (A' profile) and the lung point, had a specificity of 95% and 100% with a sensitivity of 31% and 68%, respectively. The confirmation of lung sliding, which is the back-and forth movement of the bright echogenic parietal and visceral pleura occurring during respiration, has a 100% negative predictive value for the absence of pneumothorax.²⁰ It is important to recognize that, although the presence of lung sliding effectively rules out pneumothorax, its absence does not necessarily rule it in pneumothorax. The appearance of a single B-line confirms the opposition of both pleural surfaces and effectively rules out a pneumothorax. Therefore, the absence of a single B line can safely rule in a pneumothorax.²⁰

The lung point refers to the point on the chest wall where the visceral pleura has been separated from the parietal pleura and, therefore, defines where the pneumothorax begins. Visualization of the lung point is 100% specific for the diagnosis of pneumothorax.²¹

Our study has got 94% accuracy for pneumothorax, 97% accuracy for pulmonary embolism, 96.0% for pulmonary oedema, 86% for asthma, and COPD. Table 7 Demonstrates comparison of our study with the original study done by Lichtenstein et al. and Rasha Daabis et al.^{9,15}

Table 7: Comparison with Lichtenstein et al. and Rasha Daabis et al. study

Final diagnosis	USG Profile	Sensitivity (%)			Specificity (%)			Positive Predictive value (%)			Negative Predictive value (%)		
		P	L	R	P	L	R	P	L	R	P	L	R
COPD + asthma	A Profile	75.8	89	76	91.2	97	90	80.7	93	67	88.6	95	93
Pneumonia	A/B Profile	22.7	14.5	2	100	100	100	100	100	100	82.3	71.5	52
	C Profile	31.8	21.5	31	98.7	99	92	87.5	90	79	83.9	73	58
Pneumothorax	Lung point Profile	68.4	88	80	100	100	100	100	100	100	93.2	99	99
	A' profile	31.6		20	95.1		100	60		100	85.7		96
Pulmonary embolism	A Profile with Plaps + DVT	50	81	40	96.9	99	100	50.0	94	100	96.9	98	97
Pulmonary edema	B Profile with Plaps	90.5	97	40	97.5	95	97	90.5	87	57	97.5	99	94
	A profile	9.5		30	63.75		97	6.5		50	72.86		93

P= Present study, L= Lichtenstein et al., R= Rasha Daabis et al.

Acute respiratory failure is one of the most distressing situations for the patient. The ultrasound approach with the profiles described by Lichtenstein et al can be used in such patients, as it was corresponding to the underlying physiopathology in most of the cases, and resulted in a high diagnostic accuracy, which can be further increased when combined with simple clinical data.

CONCLUSION

Study demonstrates diagnostic accuracy similar to the original publication except in COPD and Asthma(less). Lung ultrasound, as proposed in the BLUE protocol, has good accuracy, and seems reproducible and useful in acute respiratory failure. Lung ultrasound saves time and is nearly equivalent to CT in detecting

most disorders. It can be repeated at will and generates standardized, reproducible patterns with high feasibility.

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LIMITATIONS

Single centered study.

Few final diagnoses were subsequently excluded from this study.

ETHICAL APPROVAL

The study was approved by the Institutional Ethics Committee.

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