

# ERS International Congress 2021: highlights from the Clinical Techniques, Imaging and Endoscopy Assembly

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## **Abstract**

This article summarises the highlights from the European Respiratory Society's "Clinical techniques, imaging and endoscopy" Assembly 14 presented at the virtual 2021 European Respiratory Society International Congress. Cutting-edge innovative developments in both diagnostic approaches and therapeutic strategies in patients with lung cancer, interstitial lung disease, obstructive airway disorders and infectious diseases were presented on this year's interactive congress platform. In this article, the Assembly 14 subgroups summarise the key take home messages given new research outcomes and place them in the context of the current knowledge.

## Introduction

In this article, early career members of the Assembly 14 subgroups "Interventional pulmonology", "Imaging" and "Ultrasound" review the highlights of novel diagnostic approaches and innovative therapeutic strategies in patients with various lung diseases. Results of numerous trials in the field of lung cancer, interstitial lung disease, obstructive airway disorders and infectious diseases presented at the 2021 international European Respiratory Society (ERS) International Congress are summarised.

# Group 14.01: interventional pulmonology

## Peripheral pulmonary lesions and lung cancer

Effective and minimally invasive diagnosis of peripheral pulmonary nodules (PPNs) has emerged as one key issue at the ERS congress 2021. As guidelines recommend lung cancer screening, solutions to obtaining tissue from PPNs safely and effectively are required. More than in previous years, several new techniques have been established and synergistically combined to address this emerging challenge. Additionally, localised malignant PPN ablation treatment is being studied, enhancing the importance of bronchoscopy in both diagnostic and therapeutic approaches.





Bronchoscopy-guided ablation is a novel therapeutic approach for malignant PPNs. Few pilot and model studies of radiofrequency ablation and microwave ablation (MWA) in porcine and canine models are published [1, 2] and a few first trials have described encouraging results of bronchoscopic ablation

techniques in human beings [3]. At the ERS congress 2021, Lau *et al.* [4] presented MWA of peripheral lung tumours in the NAVABLATE study as a late-breaking abstract. This prospective study used MWA in combination with navigation bronchoscopy to target PPNs <30 mm. 30 patients with lung cancer or oligometastatic disease, aged 68±10 years, with a mean nodule size of 14±5 mm, were enrolled, and 39 ablations were performed using 75–100 Watts. All ablations were technically satisfactory without serious adverse events. One patient experienced mild haemoptysis as an adverse event, possibly related to the ablation device as it occurred in the first 30 days. At 1 month, technique efficacy was 100%, and the planned 2-year follow-up regarding efficacy is in progress [4].

Robotic navigational bronchoscopy (RNB) provides increased reach, stability, dexterity and needle proximity for small PPNs. In a recent study with 131 patients, RNB provided a diagnostic yield of 82% in PPNs with a mean size of 18 mm. Nodule size remains the major determining factor in diagnostic feasibility [5]. RNB combined with cytological rapid on-site evaluation (ROSE) was found to be a successful and safe method with a pneumothorax rate of 3.7% as the sole adverse event in another recent study with 55 patients [6]. A new idea that was presented at the ERS congress 2021 was the combination of RNB with the needle-based confocal endomicroscopy (nCLE), which is able to differentiate between malignant and benign processes with a concordance with cytopathology of 95% [7]. Thereby, the combination of RNB with nCLE seems to reduce the number of near-miss events. In a late-breaking study. Manley et al. [8] combined nCLE with RNB in 17 patients with a mean nodule size of 15 mm. After identifying the nodule by RNB, nCLE was performed for correcting needle position. After confirming the correct position of the catheter within the malignant nodule using nCLE, transbronchial needle aspiration was performed. The sensitivity of nCLE was 93% for detecting malignant patterns, and diagnostic yield was 82%, without relevant adverse events. Importantly, nCLE identified near-miss events leading to reposition of sampling location in 35%. Thus, the combination of nCLE and RNB improved the diagnostic yield in very small lesions, displaying a synergistic benefit when used in diagnostic procedures.

For patients with suspected lung cancer, fast and accurate tissue diagnosis is important for therapeutic decisions and prognostication. A laser-based real-time visualisation of tissue with higher harmonic generation (HHG) microscopy seems to be an innovative tool that provides the ability to identify pathological features in small bronchoscopic biopsies without the need for tissue processing. One study presented at ERS congress 2021 reported about 29 biopsies from 11 patients that were imaged by using the HHG microscope within a few minutes after bronchoscopic sampling and compared results to routine pathological practice [9]. The HHG microscope was able to identify pathological hallmarks so that in future the use of HHG microscopes within the endoscopy units may reduce the number of biopsy samples required and may reduce endoscopy time.

The use of cryoprobes that provide larger tissue samples is well established for diagnosis of thoracic malignancy. Thus, the diagnostic yield of endobronchial biopsies of endoluminal lesions and of transbronchial cryobiopsies of periphery pulmonary nodules was found to be 90% and 93%, respectively [10, 11]. When added to conventional sampling methods, cryobiopsy improved the diagnostic success rate in 8.6% of patients, and with low adverse events (1.2% severe bleeding and 0.8% pneumothorax events) [12]. At ERS congress 2021, Gonuguntla and co-authors [13] described endobronchial ultrasound-guided transbronchial Cryo Nodal Biopsy (EBUS-TBCNB) as a novel approach for mediastinal lymph node sampling in four patients with mediastinal lymphadenopathy. These first EBUS-TBCNB revealed encouraging results, but further studies are necessary.

Another topic that was discussed at the congress was the complication rate of bronchoscopic procedures in patients with severe respiratory failure. As an alternative diagnostic approach, transoesophageal bronchoscopic ultrasound-guided fine needle aspiration (EUS-B-FNA) was described in 16 patients with severe respiratory insufficiency and central lesions suspicious for lung cancer [14]. EUS-B-FNA showed a diagnostic yield of 81% with no severe side-effects, thus demonstrating an overall acceptable safety profile in this patient group with high morbidity.

# Take home messages

- RNB is a promising tool to reach small peripheral pulmonary lesions.
- nCLE is a safe and effective diagnostic modality in addition to navigation bronchoscopy, causing sampling area readjustment in 35% of cases.
- MWA therapy seems to be a safe, tissue-sparing alternative therapy for malignant lung nodules smaller than 30 mm, but further trials are required.
- EBUS-TBCNB is currently being explored as a novel approach for mediastinal lymph node sampling.

## Interstitial lung disease

Transbronchial cryobiopsy (TBCB) has become a relevant tool to obtain histology in patients with interstitial lung disease (ILD). In 2020, the COLDICE trial showcased TBCB as a safe alternative to surgical diagnostic biopsy, as there was high histopathological agreement between these two approaches [15, 16]. In the COLDICE cohort, the main factor predicting histopathological agreement was the number of TBCB samples. In a multicentre study by Hetzel *et al.* [17] reported at the ERS congress 2021, 127 ILD patients underwent TBCB with  $4.8\pm1.3$  biopsies, total area of  $51\pm29$  mm<sup>2</sup> and freezing time of  $\sim$ 7 s. The authors described a significant correlation of the total area of all histological specimens per patient and total area per biopsy with the confidence level of histological diagnosis. Contrary to the findings in the COLDICE trial, the number of biopsies did not have a significant impact on the yield.

Another approach for performing TBCB in ILD patients was presented in the FROSTBITE study, which evaluated the feasibility and safety of a novel smaller sheath cryoprobe and demonstrated an acceptable risk profile with low complication rates [18].

However, lung biopsies and also computed tomography scans are often indeterminate in early disease of ILD patients and thus stimulating the search for further diagnostic tools. Use of polarisation-sensitive optical coherence tomography (OCT) may help to distinguish between different ILD patterns. OCT, using infrared light, has the ability to assess larger lung volumes than biopsies, with higher visual acuity than computed tomography through lengthwise recording of the wall lining of multiple bronchial trees, showing loss of aerated space and increase of collagen with a  $10 \, \mu m$  resolution and  $2-3 \, mm$  depth. In a study from Amsterdam, collagen present in OCT showed 87% concordance with biopsies in patients with different ILDs and controls [19]. OCT showed a 100% sensitivity and specificity for diagnosis of usual interstitial pneumonia (UIP) in another recent study with 27 patients [20].

# Take home messages

- TBCB has become implemented as a standardised diagnostic tool in ILD with an acceptable safety profile. Larger sample size improves diagnostic confidence regardless of sample number.
- OCT assesses collagen deposition and loss of aerated space in large lung volumes, with an 87% concordance with biopsies and may help to distinguish different ILD patterns.

# Obstructive lung disease

In patients with severe asthma, bronchial thermoplasty improves asthma control, quality of life and exacerbation frequency for up to 10 years, with few adverse effects such as mild-to-moderate bronchiectasis in 7% after treatment [21]. In the randomised controlled TASMA trial, airway smooth muscle reduction was confirmed as a mechanism of action with long-term altered airway remodelling [22]. One trial presented at the ERS congress 2021 evaluated the use of OCT to assess airway remodelling following bronchial thermoplasty in 15 patients with severe asthma treated in the TASMA trial [22, 23]. Following bronchial thermoplasty, a reduction in segmented high-intensity OCT area was measured in the proximal larger airways reflecting changes in airway extracellular matrix content [23]. As an alternative diagnostic tool to assess airway pathology in patients with asthma, the feasibility, safety and efficacy of mucosal cryobiopsies were compared to endobronchial forceps biopsies in 48 patients. Mucosal cryobiopsies were safe and yielded biopsies that were larger and better preserved compared with forceps biopsies [24]. The cryotechnique seems to be a promising tool for future *in vivo* studies of airway pathology.

In patients with advanced COPD and emphysema, surgical or endoscopic lung volume reduction is an established guideline-endorsed treatment option. As an endoscopic approach, the implantation of one-way valves presents a reversible, symptom-improving and disease-modifying therapeutic option for patients with complete interlobar fissures [25]. Data from the Lung Emphysema Registry (LE-R) in Germany confirmed that surgical lung volume reduction as well as endoscopic valve therapy led to significant improvements in various outcome parameters [26]. Another bronchoscopic method to achieve lung volume reduction and improvement of elastic recoil is endoscopic coil implantation. In a 2016 published randomised controlled trial, coil implantation was shown to lead to an improvement of median exercise tolerance though this was only modest and of uncertain clinical importance [27]. At ERS congress 2021, authors presented results from one study that evaluated an improved coil-like device, Lung Volume Reduction Reverser (LVR-R System). In this pilot study, 15 COPD patients were treated bronchoscopically with the LVR-R System. 6 months following intervention, no significant improvement in lung function parameters, exercise capacity or quality of life questionnaires was seen. However, a significant complication rate, mainly due to pneumonias, was noted [28].

## Take home messages

- Clinical applications of OCT to visualise extracellular matrix composition and collagen deposition in patients with severe asthma are emerging.
- Extracellular matrix content is reduced in OCT measurements after bronchial thermoplasty.
- Data from the "Lung Emphysema Registry" emphasises the efficacy of surgical lung volume reduction and endoscopic valve therapy in patients with advanced emphysema.

## Pleural disease

Malignant effusions are accompanied by considerable symptoms and morbidity. Recently, early definite interventional treatment was shown to be beneficial in malignancies both responsive and unresponsive to anticancer therapy [29]. Talc pleurodesis and indwelling pleural catheters (IPC) lead to relief of symptoms but require up to 6 days of inpatient stay with a pleurodesis rate of 51–92% [30–33]. To reduce inpatient stay to a median of 2 days with comparable pleurodesis success rates, the SIMPLE trial used ultrasonographic controls to predict pleurodesis success early by demonstrating cessation of lung sliding [34]. At the ERS congress 2021, a retrospective study presented a pragmatic approach in managing MPE by combining thoracoscopy, talc poudrage and insertion of IPC into a single day case procedure. Admission was required in only 13% of 45 patients, because of air leak and unrelated pneumonia, while mean rate of pleurodesis success was 78% [35].

Empyema was another topic presented at the ERS congress 2021. Intrapleural enzyme therapy (IET) with tissue plasminogen activator and deoxyribonuclease has been established as a therapeutic option in pleural infection. Complications of infectious pleural effusion treatments with IET were assessed by the RETROLYSIS study [36]. In 1833 patients, pleural bleeding occurred in only 4.1%, independent of IET dose. Bleeding risk was doubled with concurrent systemic anticoagulation treatment.

Persistent air leak presents a challenge in the clinical routine. ZHANG *et al.* [37] reported a combination of targeted body positioning with thrombin injection and suction drainage to assist pleural adherence in 20 predominantly male patients after spontaneous pneumothorax or bullous rupture. Air leak remission was seen after 1.3±1 days, with 5% recurrence, and without serious adverse events [37].

# Take home messages

- To minimise interventional burden for a quality-of-life intervention, shifting malignant effusion pleurodesis to day-case treatment showed acceptable efficacy and safety, with a low admission rate.
- In pleural empyema and septated pleural effusion, IET resulted in 4% pleural bleed risk independently of IET dose. Full anticoagulation doubled risk to 8%.

# Group 14.02: imaging

## Coronavirus disease 2019

Coronavirus disease 2019 (COVID-19), a highly infectious respiratory disease caused by SARS-CoV-2, has completely changed our lives. Over the past 2 years much has been learned about its clinical and radiological manifestations. Imaging, especially with computed tomography (CT), has established its role not only in diagnosing and assessing disease extent, but has also shown promise in assessing short-term outcomes. In a late-breaking abstract presented at ERS congress 2021, the prognostic value of chest CT findings in older and multimorbid COVID-19 patients was evaluated [38]. In this study, which included 380 hospitalised patients aged 60 years or older (mean 78 years), ground-glass opacities (GGO) (85%) and consolidations (41%) were the most common CT findings, but pleural effusion (23%) and pulmonary nodules (12%) were more prevalent than previously reported in younger cohorts [39, 40]. Patients were also grouped in five different clusters according to the prevalence of GGO and pleural effusions. At multivariate analysis, only cluster "LH" (i.e., low GGO, high pleural effusion prevalence) was an independent predictor for mortality. This study suggests that pleural effusions may be an important prognostic factor in older patients with COVID-19.

Although the radiological signs of the disease have been extensively reported in the literature, the long-term health consequences remain largely unknown. This year, Tomassetti *et al.* [41] presented an interim analysis from the ongoing PCOILS study (Prevalence and Characteristics of Post-COvid-19 Interstitial Lung Syndrome). In this multicentre prospective study, 550 patients underwent high-resolution chest CT (HRCT) 6 months after being hospitalised due to COVID-19. Out of 550 subjects, 212 (38.6%) showed no CT abnormalities, 147 (26.7%) had interstitial abnormalities affecting <5% of the lung and 191 (34.7%) had interstitial abnormalities affecting volume. All patients presenting with

interstitial abnormalities affecting >5% of the lung at 6 months then undergo follow-up CT at 12 and 18 months. In this subgroup of patients, 133 (24.2%) had fibrotic interstitial abnormalities, which were diffuse in 98 cases (17.8%), and from these, the majority of patients (61, 11%) had a CT pattern resembling fibrotic nonspecific interstitial pneumonia/organising pneumonia; 31 (5.8%) had a pattern which was indeterminate for UIP and only six patients (1.1%) had a UIP pattern (definite UIP or probable UIP). Also, 58 patients (10.5%) had interstitial abnormalities without fibrotic-like features. The potential progression of the observed fibrotic changes has to be further elucidated in subsequent analysis.

# Take home messages

- Pleural effusions may be an important poor prognostic factor in older patients hospitalised with COVID-19.
- At 6-month follow-up after hospitalisation from COVID-19 infection, almost 25% of patients had fibrotic interstitial lung abnormalities on a follow-up CT.

## Interstitial lung diseases

ILDs encompass a variety of disorders with a wide range of causes, clinical manifestations and radiological and pathological features. Although HRCT plays an absolutely central role in the diagnostic process of these entities, chest radiograph is commonly the first indicator of the presence of an ILD. At ERS congress 2021, Nishikiori *et al.* [42] presented a validation study of an artificial intelligence (AI) software to detect chronic fibrosing ILDs in chest radiographs. In 1280 consecutive patients who had chest radiography and chest CT on the same day, 352 patients (27.5%) had chronic fibrosing ILD, 618 (48.3%) had abnormal radiological findings other than fibrosing ILD and 310 (24.2%) had normal chest radiographs. Area under receiver operating characteristics curve was 0.872 for cystic fibrosis (CF)-ILD patients *versus* other radiological abnormalities and 0.946 for CF-ILD *versus* no abnormal findings.

Recent evidence suggests that small airways disease may be a feature of idiopathic pulmonary fibrosis (IPF) [43]. In a late-breaking abstract presented at the congress, a multistage convolutional neural network to perform airway segmentation from CT volume was applied to 31 IPF patients [44]. This software showed that in comparison with "stable IPF" patients, "progressive" disease is characterised by larger increases in median airway sizes and larger declines in lung volumes.

# Take home messages

- AI software can detect chronic fibrosing ILDs in chest radiographs.
- Airway measurements with AI may have a role as a surrogate marker for IPF progression.

## Lung cancer screening

Lung cancer is the leading cause of cancer-related death in the world [45]. Randomised controlled trials such as the National Lung Screening Trial (NLST) [46] or the Nederlands-Leuvens Longkanker Screenings Onderzoek (NELSON) trial [47] have shown a significant reduction in lung cancer specific mortality with low-dose chest CT screening. In the USA, previous U.S. Preventive Services Taskforce (USPSTF) guidelines recommended screening for asymptomatic adults aged 55–80 years who had a ≥30 pack-years smoking history and were smokers or had quit smoking within the previous 15 years [48]. One Australian study presented at the congress aimed to identify the proportion of newly diagnosed lung cancer patients who would have been eligible for screening according to USPSTF criteria [49]. During the study period, 175 patients were diagnosed with lung cancer and only 43% (75/175) would have qualified for screening in the preceding 12 months, capturing 54% of nonsmall cell lung cancers and 100% of small cell lung cancers. Also only 40% (70/175) of patients would have been eligible for screening in the preceding 5 years. It would be interesting to re-analyse the data according to the recently updated and wider USPSTF criteria, which now include a lower age limit (50–80 years old) and a minimum smoking history of ≥20 pack-years [50].

AUDELAN *et al.* [51] evaluated whether a deep-learning system trained to detect lung nodules from low-dose computed tomography (LDCT) could detect malignant lesions 1 year prior to diagnosis by radiologists. This system was tested in 1179 examinations from NLST in which two CT scans were available 1 year apart. Among the 177 biopsy-proven malignant nodules, the algorithm detected 172 regions of interest within 3 cm of their ground truth (97% sensitivity). Missed malignant nodules were all located close to the mediastinum. On CT images obtained 1 year prior to diagnosis, 20 nodules were not visible, and the system was able to detect 97% of the visible nodules (152/157). However, this system also provided a large number of false positive nodules (mean 12 new candidate nodules per scan).

# Take home messages

- A significant proportion of newly diagnosed cancer patients may not be eligible for lung cancer screening according to most widely used criteria.
- Deep-learning-based systems may detect malignant pulmonary nodules 1 year prior to their detection by radiologists.

## COPD

Asthma and COPD are two prevalent chronic lung diseases characterised by airflow limitation. While treatment differs between these entities, they often present with clinical similarities. HRCT is the main imaging tool for evaluation of emphysema and airways disease. One study presented at the congress aimed to determine if a machine learning approach using quantitative CT-derived features could differentiate COPD and asthma [52]. Using airway and parenchymal-related CT features, the algorithm could discriminate between asthma and COPD patients with 87% sensitivity, 71% specificity and 80% accuracy.

Another study tried to investigate the relationship between 129-xenon (129Xe) MRI and pulmonary function test (PFT) metrics in patients with asthma and/or COPD [53]. Worsening disease severity as assessed by spirometry and transfer factor of the lung for carbon monoxide ( $T_{\rm LCO}$ ) was associated with increased ventilation defect percentage (VDP) and mean acinar dimensions (LmD) and with decreased red blood cell to tissue-plasma ratio (RBC/TP), RBC/gas and TP/gas ratios. 129Xe MRI ventilation metrics also correlated moderately with spirometry, residual volume/total lung capacity,  $T_{\rm LCO}$  and lung clearance index, but not with oscillometry. Moreover, a significant proportion of patients with normal spirometry had elevated VDP due to the high sensitivity of 129Xe MRI in detecting ventilation abnormalities.

# Take home messages

- Machine learning using CT imaging-derived features may have a role in distinguishing patients with COPD and asthma.
- 129Xe MRI-derived metrics correlate moderately with COPD and asthma disease severity as assessed by PFT.

# Group 14.03: ultrasound

## Coronavirus disease 2019

The continued presence of the global SARS-CoV-19 pandemic encourages physicians to seek out ways of improving bedside evaluation, aiding in clinical assessment, diagnosis and prognostication for patients admitted with COVID-19. As predicted in the 2020 ERS congress highlights, the past year has led multiple researchers to explore the clinical utility of lung ultrasound in these patients [54].

The diagnostic accuracy of lung ultrasound for detection of peripheral changes in COVID-19 pneumonia was assessed by SMARGIASSI *et al.* [55], using a scoring system proposed by SOLADATI *et al.* [56] in which 14 thoracic zones are scored from 0 to 3 based primarily on alterations of the pleural line. 16 patients with confirmed COVID-19 pneumonia were subject to both an HRCT scan and a lung ultrasound examination within 48 h. While lung ultrasound showed encouraging results for identification of any peripheral alterations (sensitivity 92.1% and specificity 90%), the accuracy for identification of severe pneumonic changes was lower (sensitivity 70.1% and specificity 84%). Overall, these findings are impressive when considering that a recent Cochrane meta-analysis reported a pooled sensitivity and specificity of 86.4% and 54.6%, respectively, for lung ultrasound's ability to diagnose COVID-19 [57].

The utility of lung ultrasound in predicting clinical deterioration in patients admitted with COVID-19 was appraised by two studies at the ERS congress 2021. Bielosludtseva *et al.* [58] examined 20 patients with severe COVID-19 pneumonia and found that a lung ultrasound score (LUS) >18 and presence of triangular subpleural hypoechoic lesions were associated with disease progression. Utilising the same scoring system as Pierrakos *et al.* [59], a study on 18 patients found that a lung ultrasound examination within 72 h of admission to a dedicated COVID-19 ward could not predict clinical deterioration, but noted that if ultrasound is performed daily, a sudden increase in score may herald significant disease progression [60].

An international multicenter study by Pierrakos *et al.* [59] appraised the prognostic value of a global LUS-aeration score on 55 invasively ventilated patients with COVID-19. 12 thoracic zones were evaluated, each receiving a score between 0 and 3: normal aeration (0 points), B-lines covering <50% of the pleural line (1 point), B-lines covering >50% of the pleural line (2 points) or present consolidation, suggesting complete or near-complete loss of aeration (3 points). The authors concluded that a global LUS-aeration

score of  $\leq$ 13 aided in predicting successful liberation from a ventilator (p<0.01) and a score of >17 was associated with higher mortality at 28 days (p=0.01).

# Take home messages

- While some studies report promising overall diagnostic accuracy of lung ultrasound in suspected COVID-19, the authors of a recent Cochrane meta-analysis conclude that due to the disappointing specificity, the role of lung ultrasound should be to rule out COVID-19 pneumonia, rather than confirming it.
- In patients with COVID-19, ultrasound may be of value in prognostication for invasively ventilated patients and to detect peripheral pneumonia.
- Evidence is sparse for prognostic value in patients referred to dedicated COVID-19 wards not requiring
  intensive care. Study populations are generally small, and studies utilise different scoring systems,
  complicating comparison of results.

## Diaphragm

As the principal respiratory muscle, it is no surprise that ultrasound assessment of the diaphragm is receiving increasing scientific attention. Indeed, a reliable sonographic evaluation of excursion, thickness and thickening might spare the patient an invasive diaphragmatic pressure measurement [61]. While initial research on this topic was aimed at appraising diaphragmatic dysfunction in intubated patients, the proposed utility of diaphragmatic ultrasound has since expanded as highlighted by this year's congress where multiple studies appraised its value in distinct settings and conditions.

Gabrysz-Forget and colleagues [62] evaluated 111 patients with cystic fibrosis establishing a normal lower range of diaphragmatic thickening fraction at 30% and revealing that it was significantly correlated to the disease markers transthyretin, C-reactive protein, forced expiratory volume in 1 s (FEV<sub>1</sub>) and grip strength. Encouragingly, their findings suggest that the diaphragm measurement may be an even better marker of dyspnoea level than FEV<sub>1</sub> (p=0.03). Their findings establish a basis for future studies evaluating diaphragmatic activity as a marker of prognosis and efficacy of interventions in cystic fibrosis.

In patients with acute exacerbation of COPD, readmissions are common. This encouraged Kharat *et al.* [63] to examine whether diaphragmatic ultrasound was predictive of re-hospitalisation. 28 patients with exacerbation of COPD were enrolled, and the authors reported that a diaphragmatic thickening fraction above 48% at discharge was highly predictive of re-hospitalisation within 30 days (sensitivity 100% and specificity 76%).

## Take home messages

Diaphragmatic ultrasound is a promising surrogate marker of disease severity and a possible alternative
to invasive evaluation of diaphragmatic function in several pathologies, but evidence is heterogeneous
and reference values as well as standardised scanning protocols are still needed, limiting clinical
applicability.

## Ultrasound for guiding pleurodesis

Recurrent malignant pleural effusion is most often observed in relation to mesothelioma, lung or breast cancer. Even though new treatments such as IPC with or without concomitant talc pleurodesis are options, talc pleurodesis through a chest tube remains the most common definitive treatment. While successful in obtaining effusion control in four out of five instances, the procedure requires an initial hospital admission until adherence of pleural membranes is confirmed. In a recently published ambitious randomised controlled trial, Psallidas and colleagues [34] noted that the current recommendations on removal of the chest tube by the British Thoracic Society suggest awaiting daily fluid volume drainage of <250 mL and lung re-expansion confirmed radiologically; however, these recommendations are not based on strong evidence. The authors hypothesised that integration of thoracic ultrasound before and after pleurodesis could shorten the length of hospital stay without compromising pleurodesis success.

The results were eloquently summarised by the study's second author, M. Hassan, in an oral presentation at this year's ERS conference. 313 patients requiring talc pleurodesis were included and randomised 1:1 to standard care or integrated thoracic ultrasound. The length of hospital stay was significantly shorter in the thoracic ultrasound group compared to standard care (p<0.001). Furthermore, the ultrasound approach was non-inferior to standard care in relation to pleurodesis failure (34.6% *versus* 32.2% of patients).

These results raise the question of whether thoracic ultrasound should be considered a standard approach in patients undergoing pleurodesis for recurrent malignant pleural effusion.

# Take home messages

- The current recommendation for removal of a chest tube following talc pleurodesis for malignant pleural effusion is based on sparse evidence.
- Integration of thoracic ultrasound prior to and following talc pleurodesis reduces length of hospital stay and is non-inferior to standard care in terms of pleurodesis failure.

# Pulmonary embolism

Computed tomography pulmonary angiography (CTPA) is considered the diagnostic gold standard in suspected pulmonary embolism. But the procedure is time consuming, expensive and exposes the patient to radiation. Furthermore, the increasing availability of radiation-based diagnostics lowers the threshold for referral. Indeed, only 20–30% of patients referred for CTPA in Europe are diagnosed with pulmonary embolism. As such, in the hope of improving selection of patients for irradiating imaging, researchers often appraise the diagnostic accuracy of ultrasound in suspected pulmonary embolism, and this year was no exception.

A small study by ABDELKADER and colleagues [64] examined the diagnostic accuracy of ultrasonographic detection of hypoechoic rounded or wedge-shaped pleural consolidations in 30 haemodynamically stable patients with moderate to high pre-test probability of pulmonary embolism. To further increase diagnostic accuracy, the consolidations had to be hypoperfused, gauged by absence of colour Doppler signals. The authors report a sensitivity of 61% and specificity of 91.6%, hinting that this approach may be of value in ruling in suspicion of pulmonary embolism, possibly limiting the need for CTPA.

A comprehensive review and meta-analysis by Falster *et al.* [65] compiled not only studies on diagnostic accuracy of lung ultrasound, but also on deep venous, cardiac and multi-organ ultrasound. The study, analysing 70 unique publications, found that several single-organ ultrasound findings, such as detection of a deep venous thrombus, the McConnell's sign or detection of at least two hypoechoic pleural-based lesions, exhibit specificities above 95%, hinting at a high utility in confirming suspected pulmonary embolism. Furthermore, meta-analysis of a multi-organ ultrasound examination devoid of right ventricular strain or thrombi, hypoechoic pleural-based lesions and deep venous thrombi yielded a sensitivity of 90%, correlating with a high negative predictive value in patients with low or moderate pre-test probability.

## Take home messages

- >70 descriptive studies have assessed the diagnostic accuracy of ultrasound in suspected pulmonary embolism
- Several single-organ and multi-organ findings are highly predictive of pulmonary embolism while a
  normal ultrasound investigation of the lungs, heart and deep veins of the legs may be of value in
  dismissing suspicion in patients with low to moderate pre-test probability.
- No randomised controlled studies on the subject have been conducted.

Provenance: Commissioned article, peer reviewed.

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