



# Beyond diagnosis: a narrative review of the evolving therapeutic role of medical thoracoscopy in the management of pleural diseases

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**Background and Objective:** Medical thoracoscopy (MT) is an endoscopic technique performed by interventional pulmonologists with a favorable safety profile and few contraindications, providing diagnostic and therapeutic intervention in a single sitting. This narrative review was designed to summarize the therapeutic role of MT based on the latest results from the available literature.

**Methods:** Pertinent literature published in English, relative to human studies, between 2010–2022 was searched in Medline/PubMed and Cochrane databases. Publications regarded as relevant were considered for inclusion in this review; additional references were added based on the authors' knowledge and judgment. The review considered population studies, meta-analyses, case series, and case reports.

**Key Content and Findings:** MT has mostly been described and is currently used globally in the diagnostic approach to exudative pleural effusion of undetermined origin. Carefully evaluating the literature, it is clear that there is initial evidence to support the use of MT in the therapeutic approach of malignant pleural effusion, pneumothorax, empyema, and less frequently hemothorax and foreign body retrieval.

**Conclusions:** MT is an effective procedure for treating the clinical entities presented in this document; it must be carried out in selected patients, managed in centers with high procedural expertise. Further evidence is needed to assess the optimal indications and appropriate patients' profiles for therapeutic MT. The endpoints of length of hospital stay, surgical referral, complications and mortality will have to be considered in future studies to validate it as a therapeutic intervention to be applied globally.

**Keywords:** Thoracoscopy; talc; poudrage; pneumothorax; empyema

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## Introduction

Medical thoracoscopy (MT) is a minimally invasive procedure that allows for direct visualization of the pleural space and intrathoracic structures while obtaining image-guided biopsies and performing therapeutic interventions. It was first described in 1910 when Jacobaeus utilized rigid cystoscopes to evaluate the pleural space (1-3). In current practice, thoracoscopy represents one of the main endoscopic procedures, along with bronchoscopy in interventional pulmonology (4-6). Thoracoscopy can be broadly divided into minimally invasive MT/pleuroscopy and surgical thoracoscopy, also known as video-assisted thoracic surgery (VATS). VATS is performed under general anesthesia with selective bronchial intubation and is performed in an operating room (OR); on the other hand, MT can be performed under local anesthesia, mild or moderate sedation (non-operating room anesthesia; NORA) in an endoscopy suite (7). The most common use of MT is in the setting of an undiagnosed exudative pleural effusion to confirm or confute a diagnosis of malignant pleural effusion (MPE) (8-10); however, additional conditional indications of MT include pleural effusions of undetermined etiology, biopsies of pleural anomalies detected on chest computed tomography (CT), chemical and mechanical pleurodesis, and mechanical adhesiolysis (9,10). The additional benefits of performing a MT instead of just placing a chest drain are the possibility of carrying out a direct endoscopic evaluation of the pleural cavity, performing adhesiolysis and carrying out a biopsy sampling if necessary. The decision to perform MT rather than VATS should be carefully made based on patient-related factors, the underlying pathology and the operator's experience in MT.

MT is a safe procedure with low mortality, major and minor complications reported in the current literature (10-12). Its application has also been described in patients admitted to the intensive care unit, where, in highly selected cases, it may represent a valuable therapeutic alternative in critically ill patients (13,14).

The increasing adoption of MT worldwide (15,16) is due to interventional pulmonologist's efforts in offering less invasive procedures to reduce complications and length of hospital stay (LOS) with an overall better cost-effectiveness profile (17). We present this article in accordance with the Narrative Review reporting checklist (available at <https://jtd.amegroups.com/article/view/10.21037/jtd-23-1745/rc>).

## Rationale and knowledge gap

MT is an endoscopic technique with well-defined procedural standards (18). However, few randomized controlled trials (RCTs) compare MT with other procedures focusing on diagnostic yield and treatment success in pleural disease (19-23). The latest guidelines also elaborate on the role of MT in pleural disease diagnosis and treatment (24). In diagnosing MPE, its use aimed at bioptical diagnosis is considered the gold standard (24). In the context of pleurodesis for symptomatic MPE, talc slurry or talc poudrage (TP) may be equally offered to patients in order to control fluid accumulation and reduce the need for repeated invasive procedures (24). The guidelines do not currently support using MT in pneumothorax, favouring a surgical approach where necessary (24). Although in daily clinical practice MT is often used as first approach in pleural infection (PI), due to lack of supporting evidence, latest guidelines do not recommend MT as a viable treatment option for patients with infection of pleural space (24). On the contrary, in the remaining literature and daily clinical practice, MT is also applied in other contexts, including pneumothorax, PI, retained hemothorax (RH), and retrieval of intrapleural foreign bodies (FBs) (25-28).

## Methods

### Objective

This narrative review aims to summarize current literature, performing an overview of novel clinical contexts in which the use of MT has the potential to bring additional therapeutic alternatives to patient care. This manuscript is written as per SANRA quality scale (29). A comprehensive and systematic online literature search via Medline/PubMed and Cochrane database for the period January 2010 to December 2022 was performed for articles published using the keywords “medical thoracoscopy”, “local anesthetic thoracoscopy”, and “thoracoscopy”. The search strategy is summarized in *Table 1*.

## Equipment

In order to perform the procedure, standard equipment should be available (1,2) (*Figure 1*). There are two types of thoroscopes, the rigid model being the most widespread and the novel flexi-rigid thoracoscope, produced only by Olympus Corporation (Tokyo, Japan). Each device has its characteristics, and the choice of a specific model is

**Table 1** Search strategy summary

Items	Specification
Date of search	01 February 2023
Databases and other sources searched	Medline/PubMed and Cochrane
Search terms used	“medical thoracoscopy” OR “local anesthetic thoracoscopy” OR “thoracoscopy”
Timeframe	From January 2010 to December 2022
Inclusion and exclusion criteria	Inclusion criteria: human-based studies (study type: original article, research article, full paper available), English language Exclusion criteria: animal-based studies
Selection process	The selection process was conducted by A.F. and N.C. independently, then removing duplicated results. Consensus on additional papers consideration and review of the final list of references included was performed by all authors



**Figure 1** Preparation of the operating trolley for rigid medical thoracoscopy. Items: scalpel, scope forceps, zero-degree scope, 30-degree scope, light source, kidney basin, sterile drapes, scissors with blunt tip, Klemmer forceps, suction tube, needle holder, suture thread, trocar, cotton, material for collection of biopsy specimens, gauze.

based on operator comfort, level of experience, and type of procedure to be carried out (see *Table 2*), with the aim to maximize both therapeutic results and diagnostic yields. In this regard, the rigid thoracoscope is the instrument most chosen by operators, with an external diameter between 6 and 10 mm and a large working channel, compared to the 3 mm one equipped on the flexi-rigid thoracoscope (8). A greater working channel allows for the insertion of bigger tools than the standard flexible biopsy forceps and the cryoprobe utilized within the bronchoscopy field and flexi-rigid instrumentation. The mechanical advantage of rigid forceps favors obtaining larger biopsy samples, even

from very dense lesions, such as fibrous plaques, and solid malignant lesions, such as sarcomatoid mesothelioma, which may require greater tearing force (9,31). In addition, when using rigid thorascopes, it is possible to insert different instruments through the working channel, facilitating the work of debridement of the pleural space by using rigid biopsy forceps and management of complications such as controlling intra-procedural hemorrhage (30-32). Lastly, the rigid thoracoscope may offer more durability with less frequent need for maintenance and repair compared to flexi-rigid instrumentation when preserved appropriately (8).

On the other hand, the flexi-rigid thoracoscope combines the flexibility of bronchoscopes with the stability of rigid thorascopes, thus providing increased maneuverability and flexibility, which facilitates the complete exploration of the pleural space available and homogenous insufflation of talc within the pleural space (10). Its resemblance to the familiar bronchoscope increases confidence among interventional pulmonologists approaching the procedure for the first time. Furthermore, the external smaller diameter reduces the incision size site and, consequently, post-procedural chest wall pain (9,10,33).

In a recent RCT, Bansal and colleagues compared a mini-thoracoscopy set-up with a flexirigid one, demonstrating no significant difference in diagnostic yield between the two devices (33). There is still no literature data on comparison between the various devices in the therapeutic setting of pleural diseases.

### Contraindications

Absolute contraindications to performing MT include an

**Table 2** Different properties of the rigid and flexi-rigid thoroscopes

Type of thoracoscope	Learning curve length	Ease and completeness of exploration of the pleural space	Biopsy size	Possibility of talc instillation under direct vision	Suitability for mechanical lysis of adhesions
Rigid	+	++	+++	+++	+++
Flexi-rigid	+++	+++	+ <sup>†</sup>	+++	+

<sup>†</sup>, increasable by using a cryoprobe (30). +, not favourable; ++, adequately favourable; +++, optimal.

**Table 3** Levels of sedation and anaesthesia

Level of sedation	Responsiveness	Airway	Ventilation
Minimal sedation	Normal response	Unaffected	Unaffected
Moderate sedation	Response to verbal stimuli	No intervention required	Adequate
Deep sedation	No response to verbal stimuli	Intervention may be required	May be inadequate
General anaesthesia	Unarousable	Must be supported via airway device	Inadequate, must be artificially supported

Modified from Kochhar *et al.* 2016 (35).

uncorrectable bleeding disorder, significant pulmonary hypertension, cardiovascular instability, acute uncorrectable type 1 respiratory failure, ongoing type 2 respiratory failure, and absence of the possibility of generating an adequate pleural space (10,18,24). Conditions that may require changing the entry site into the pleural cavity are the presence of cutaneous infection, tract metastasis or rib fracture around the port insertion site.

Of note, if the procedure is therapeutic, such as in the setting of pleural effusion or pneumothorax, and clinical improvement may be expected, MT may be performed even in the presence of hypercapnia, considering adequate post-procedural assistance and monitoring (7,34).

## Sedation

MT is performed in an endoscopy suite without the mandatory necessity of an anesthesiologist. It is typically performed under local anesthesia with conscious, moderate, or deep sedation (see *Table 3*) (35-41). The sedation and analgesic components should be carefully managed during the operative and postoperative phases since pain is a common symptom during pleural and chest wall manipulation. Pain may be elicited by the cutaneous incision, when the thoracoscope applies pressure on the ribs, while the operator samples the parietal pleura, and when initiating talc insufflation (1,42,43). With the increase of patient's age and in case of infiltration of the pleura by malignant diseases, the pleura itself tends to become less

tender (8,9).

When performing MT, local anesthesia, e.g., lidocaine 1%, is typically injected at the entry site to anesthetize the skin and the parietal pleura while being careful not to exceed the 4.5 mg/kg dose (300 mg maximum total dose) in order to avoid systemic toxicity (1,24,36).

Koulelidis *et al.* performed a RCT comparing MT with local lidocaine alone and with intravenous midazolam, enrolling 80 patients, finding no significant difference in hypoxemia measured by peripheral oxygen saturation SpO<sub>2</sub> at the beginning of the procedure, at 15 minutes, and at the end of the procedure. However, midazolam was associated with reduced periprocedural and post-procedural cough and pain (44).

The most common agents used for sedation include a combination of midazolam and fentanyl. However, there is currently no consensus regarding the choice of sedative agents while performing MT regarding safety and reducing the risk of respiratory side effects such as hypoxemia and cardiovascular events (12,44).

Different prospective and pilot studies evaluating propofol (45,46) and dexmedetomidine have been published (40). Tschopp *et al.* published a prospective cohort study evaluating pulmonologist-administered propofol for MT in 53 patients. They achieved a high safety profile, with only four patients requiring pharmacological intervention due to the appearance of side effects but with a rapid discharge of all patients from the recovery unit after a median dose of 145 mg of propofol (41). A noninferiority trial by

Grendelmeier *et al.* compared propofol with midazolam, reporting a higher incidence of hypoxemia and hypotension when using the former (46). In a pilot RCT, Sirohiya *et al.* compared dexmedetomidine with midazolam sedation in 60 patients. Post-procedural satisfaction was greater in patients sedated with dexmedetomidine, and patients treated with midazolam had a higher proportion of needs for concomitant administration of fentanyl (40).

While most cases of MT can be performed with conscious sedation, deep sedation is preferred for uncooperative patients, children, foreseeably prolonged or technically advanced procedures, and patients with demonstrated allergy to the active ingredients for local anesthesia (1,47).

More recently, the literature explored the possibility of nerve blocks as intercostal and erector spinae (ESP) blocks (48,49). McPherson *et al.* published a pilot study on ESP blocks for day-case MT. They enrolled five patients, associating propofol and remifentanyl analgesation with 20–35 mL of 0.25% 1-bupivacaine injection. They concluded that ESP blocks provide a sufficient level of intraprocedural analgesia. However, 78% of patients required oral analgesia on day zero post-discharge, and 55% required oral analgesia on postoperative day one (49).

## Monitoring

Patient monitoring should be continuous until 15 minutes post-procedure (18). Afterward, parameters monitoring should be done every 30 minutes for 1 hour, followed by four hourly observations (18). Current guidelines do not suggest monitoring hypoventilation phenomena; therefore, skin carbon dioxide tension measurement is optional (50). Tschopp *et al.* demonstrated the possibility of adjusting the depth of sedation guided by the Bispectral index (BIS). In their study, the level of sedation was individually optimized by titrating the propofol infusion according to BIS and clinical assessment (41).

## Complications

MT is a safe procedure with an overall reported mortality of 0.34%, which nears 0% when performing a diagnostic procedure (10). Mortality in MT has primarily been attributed to using non-graded talc for TP in MPE, which is associated with respiratory failure and acute respiratory distress syndrome (ARDS), resulting in a 0.69% mortality rate. However, this problem has been resolved with widespread implementation of graded talc (51–54).

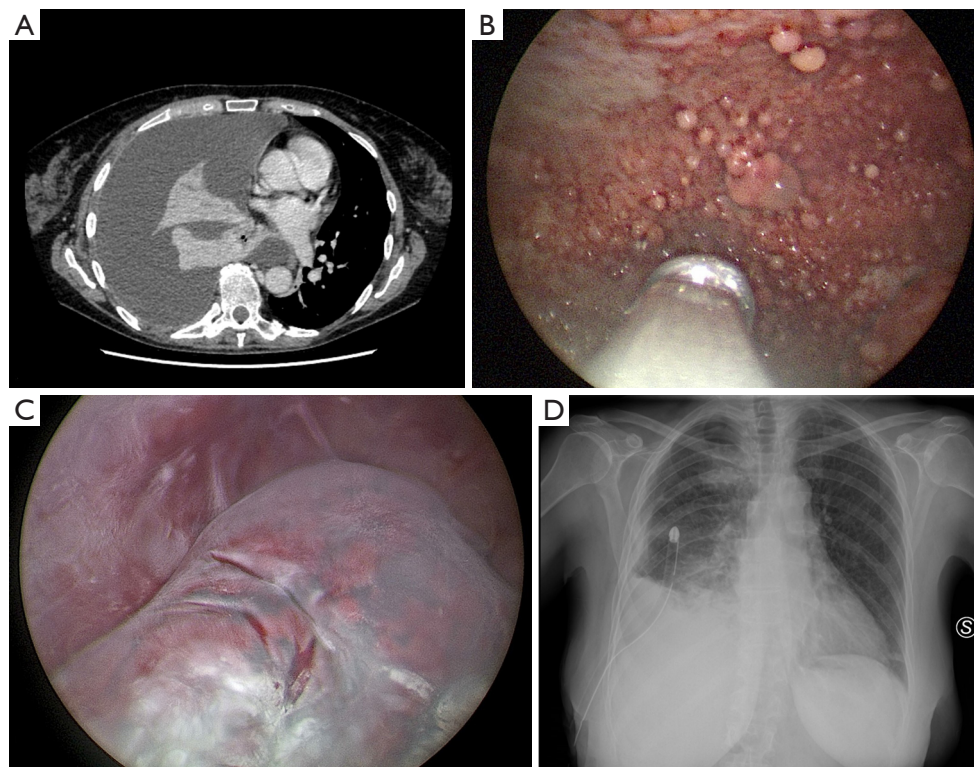
Complications in MT can be divided in major and minor adverse events. Significant complications include empyema, hemorrhage, prolonged air leak, periprocedural pneumothorax, pneumonia, and port site tumor growth, and have a cumulative rate of 1.8% (52,53,55). Minor complications include subcutaneous emphysema, minor hemorrhage, operative skin site infection, procedural hypotension, atrial fibrillation, and increased body temperature, with a cumulative rate of 7.3% (10,55). While these rates refer to MT performed typically in an endoscopy suite, similar numbers were reported in its use in the intensive care unit and when performed at the patient's bedside (14).

Procedural risks are influenced by the underlying pathology with visceral pleural tears more frequently reported when obtaining biopsy samples of honey-comb lung in end-stage pulmonary fibrosis and lung laceration during trocar insertion in the presence of extensive pleural adhesions (53). Precautions may be considered to reduce the risk of complications, such as postponing MT in the presence of severe cough, maintenance of chest tube until no air leakage is detected, and gradual lung re-expansion to prevent re-expansion pulmonary edema. In addition, in the setting of suspected mesothelioma, it is possible to administer radiation therapy to the incision area to prevent port site tumor growth (56,57).

## Pleurodesis

### MPE

The most common application of MT is in the diagnostic and therapeutic pathway of MPE, which typically presents unilaterally as an exudative effusion (58–60). In these patients, the initially performed procedure is often an ultrasound-guided thoracentesis for pleural fluid analysis (59,61–63). The general diagnostic yield rate for pleural fluid cytology is about 60%, with better results for primary adenocarcinoma of the lung, breast, and ovarian neoplasms (64). However, mesothelioma and squamous cell lung cancer are typically less exfoliating, resulting in a lower diagnostic yield (64–66). When negative cytology results are obtained, diagnosis may be attempted by ultrasound-guided or CT-guided pleural sampling (67). However, a diagnosis of malignancy will be commonly followed by a medical or surgical additional procedure such as pleurodesis to improve symptoms and reduce the risk of MPE recurrence. Current emphasis is placed on obtaining a timely diagnosis while



**Figure 2** Clinical case of massive MPE managed with medical thoracoscopy and talc poudrage. (A) Baseline chest CT. (B) Pathological endoscopic appearance. (C) Talc deposition. (D) Post-procedural result on chest X-ray. MPE, malignant pleural effusion; CT, computed tomography.



**Figure 3** Talc poudrage by insufflation.

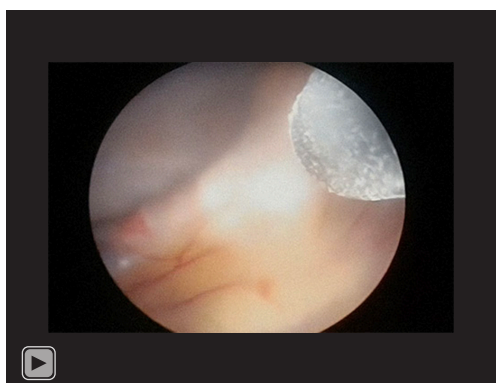
minimizing manipulation of the pleural space and the chest wall tract (24). In this regard, MT maximizes diagnostic yield allowing both diagnostic and therapeutic interventions to be performed during a single session, using a single thoracic port.

TP is the primary therapeutic procedure carried out during MT (Figure 2). The procedural workflow has been described in different scientific works that propose different methods of talc deposition during MT (Figure 3, Table 4) (70). Upon completion, uniform distribution of the talc can be confirmed by direct visualization during thoracoscopy, and a chest tube is placed inside the pleural space to aspirate residual air and allow lung re-expansion and pleural adhesion (1). Only one study has been performed directly comparing MT and TP with VATS in the setting of MPE: a retrospective study by McDonald *et al.* found that MT with TP was associated with a significantly lower hospital length of stay (0 vs. 3 days,  $P < 0.001$ ) and per-procedure cost (17).

When a diagnosis of malignancy is made in symptomatic pleural effusion, further intervention is generally required

**Table 4** Talc delivery systems for talc poudrage described in humans

Delivery system	Technique	Description	Manufacturer	Talc dose	Advantages
Through the thoracoscope	Talc insufflation	Instillation via dedicated operating channel with vials loaded with talc	Storz	3–4 g	Direct observation and orientation in talc deposition
Through other devices	Talc insufflation	Deposition using a cannula connected to an insufflator	Novatech	3–4 g	Does not require a large trocar lumen
	Talc spray	Instillation via a vaporizer/spray	Novatech, Sciarra Laboratories Inc., experimental models (68,69)	3–5 g	High level of vaporization and distribution of talc in the pleural cavity

**Video 1** Talc deposition under direct endoscopic vision.

either with chest drain insertion and medical pleurodesis or by either talc slurry or TP by MT. A RCT on 330 patients comparing TP with talc slurry yielded no statistically significant difference in terms of pleurodesis failure at 90 days (22% and 24%, respectively) defined by a need for insertion of chest tube, indwelling pleural catheter or use of MT for persistent pleural effusion (21). These findings align with a previous prospective randomized trial by Dresler *et al.* comparing TP with talc slurry, which found no significant difference in success rate at 30 days (78% *vs.* 71%, respectively) (71). However, the results from Dresler *et al.* suggested that TP may have higher success rates in subgroups of patients with primary lung or breast cancer compared to talc slurry (82% *vs.* 67%, respectively). In addition, the distribution of talc powder is more uniform at the parietal and visceral pleura levels with TP compared to talc slurry and can also be carried out under direct endoscopic vision (see *Video 1*).

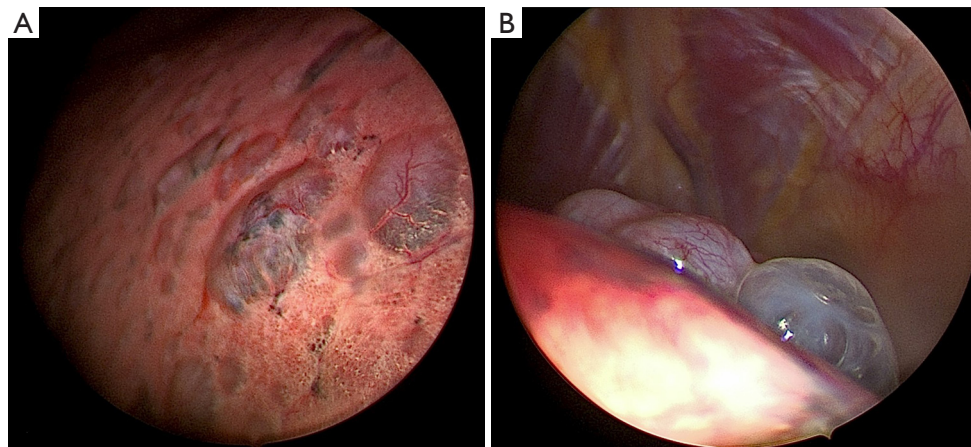
Several studies have been carried out in pre-clinical settings focusing on using talc-based foams, but the evidence on humans is still insufficient for their

application (72,73).

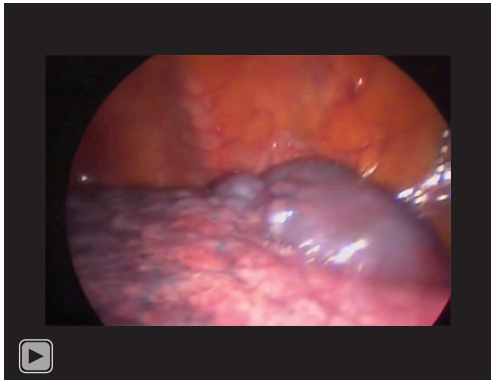
There are still few reports comparing the therapeutical efficacy between TP and indwelling tunneled pleural catheters (IPCs) in the setting of MPE pleurodesis attempts. Because IPC requires regular maintenance and MT, if not carried out as a day case procedure, is associated with a greater LOS than IPC placement alone (74–78), the debate on which of the two procedures to prefer for pleurodesis in MPE remains open. In order to propose the benefits of both treatments, Foo *et al.* published a retrospective chart-based study, enrolling 45 patients, performing TP and concomitant IPC placement into a single day-case procedure, describing pleurodesis success rates of 71.1% at 3 months and 78.8% at 6 months, with 86.7% of patients being discharged on the same day (74). Based on the current evidence, IPC should be considered an alternative or complementary treatment to TP in patients with suspected non-expandable lung (NEL) or recurrent pleural effusion (79–81). The ongoing TACTIC RCT aims to evaluate whether TP associated with IPC placement is the optimal treatment for this group of patients (22). All these data may prove vital if we consider that successful pleurodesis in MPE, as demonstrated by the systematic review of 15 pooled studies, has been associated with increased survival (3.5–5.8 months) (58).

### ***Pneumothorax***

Pneumothorax is a highly prevalent disease, classified into two main categories, namely primary spontaneous pneumothorax (PSP), in the absence of underlying lung disease, and secondary spontaneous pneumothorax (SSP), when a documented underlying lung condition exists (*Figure 4*) (82–89). Recurrence is common, ranging between 17–54% in PSP (90–93) and 13–50% in SSP within



**Figure 4** Thoracoscopic view. (A) SSP secondary to diffuse bullous lung disease (lymphangioliomyomatosis). (B) SSP associated with bullous emphysema. SSP, secondary spontaneous pneumothorax.



**Video 2** Talc poudrage in secondary spontaneous pneumothorax.

5 years, and performing a simple aspiration in the latter is unsuccessful in 15–62% of cases (94–96). In addition, for SSP, a 3.5-fold increase in mortality is associated with each secondary pneumothorax occurrence for patients affected by chronic obstructive pulmonary disease (COPD) (94,97). The current guidelines recommend different workflows for the first episodes of PSP and SSP (24).

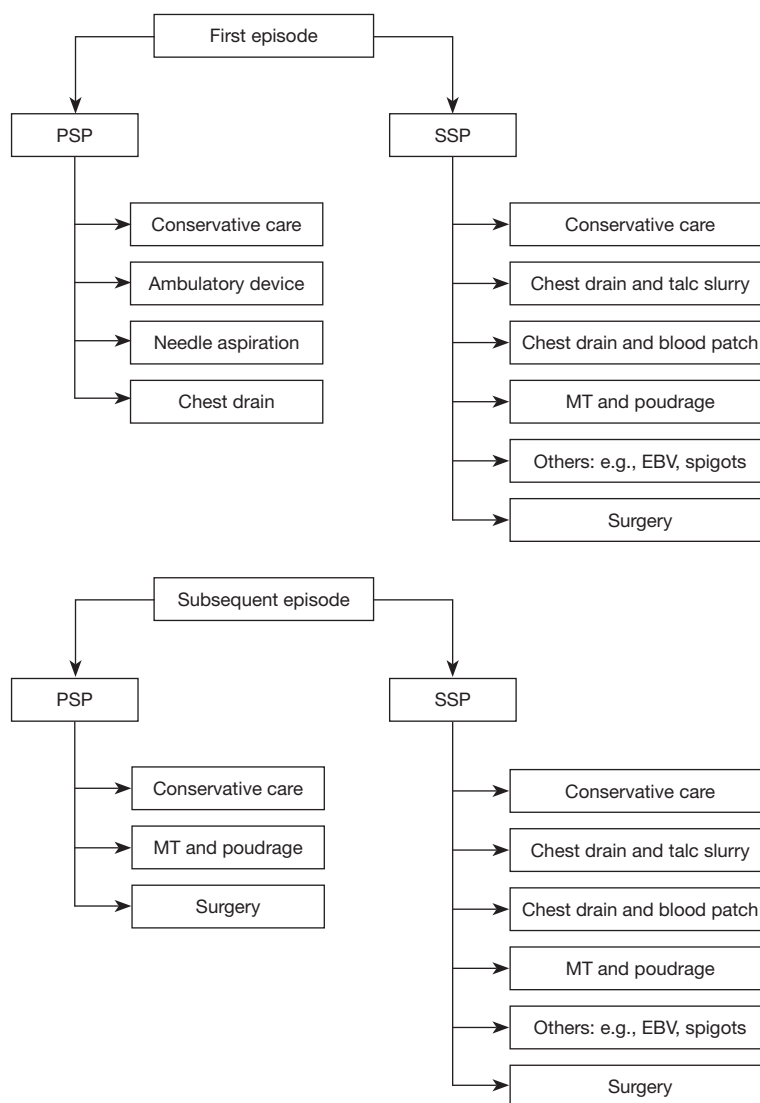
Patients with COPD who develop SSP commonly possess a significantly decreased lung function, making them poor surgical candidates; therefore, determining the best treatment for this group of patients is challenging, and MT with TP might be a practical alternative (Video 2). A case series by Lee *et al.* reported a 95% success rate for TP in SSP associated with COPD during a 3-year follow-up, with four deaths (9.8%) reported within 30 days (95). These results align with the findings described in a case series by

Tschopp *et al.* in which TP performed for persistent air leak or recurrent pneumothorax was successful in 95% of cases over a 5-year follow-up (90).

Recently, in the literature, the use of argon plasma coagulation (APC) during MT has been proposed in the setting of SSP. Guo *et al.* enrolled 70 patients with refractory pneumothorax to compare the prognosis of APC treatment performed during MT (MT/APC) with VATS and surgical pleurodesis (SP) (98). Compared with the MT/APC and SP groups, patients managed with VATS demonstrated poor short-term prognosis and higher hospitalization costs. Zhang *et al.* described a case series of patients with long-lasting SSPs treated with APC and autologous blood pleurodesis, resulting in complete resolution without further treatment in all patients considered (99).

In the context of PSP, the scientific debate is certainly more heated, given the young age of the subjects and the possibility of withstanding surgical stress. While pleurodesis is not typically performed during the first episode of PSP, a randomized study performed by Tschopp *et al.* comparing TP with intercostal tube drainage alone for the initial treatment of PSP reported superiority of TP both in terms of preventing recurrence at 5 years (5% *vs.* 34%, respectively) and increasing cost-effectiveness (100). Several systematic reviews have highlighted how a primary surgical approach to patients at the first episode of PSP seems to guarantee a lower recurrence rate than that of a primary approach consisting of chest drain placement, aspiration, or observation treatment (101–103), the latter being the lowest cost alternative (104). However, it is unclear which patients have a higher recurrence rate after the first episode





**Figure 5** Therapeutic alternatives in the treatment of the first episode of pneumothorax and subsequent episodes. MT finds a therapeutic role in both PSP and SSP. PSP, primary spontaneous pneumothorax; SSP, secondary spontaneous pneumothorax; MT, medical thoracoscopy; EBV, endobronchial valve.

and, therefore, who should immediately be a candidate for a more invasive procedure (105). The guidelines support VATS or thoracotomy as the gold standard for recurrent PSP (24). The authors' current practice (see *Figure 5*) is to consider MT as a feasible therapeutic option in specific settings, after a multidisciplinary discussion between the thoracic surgical team and the interventional pulmonology team, as an alternative or a first therapeutic proposal for relapsing PSP in patients unwilling to undergo surgery or not fit enough despite a young age (91,106).

In both PSP and SSP, differently than MPE, the

parietal pleura innervation is typically preserved; as a result, the use of talc pleurodesis is likely to evoke intense pain and autonomic nervous activity, which may require deep sedation and pain relief therapy in order to improve tolerability (95,97).

### *Chylothorax*

No further evidence has been published from the first article proposing MT as a therapeutic alternative for chylothorax (107). The authors believe MT may be

**Table 5** Stages of parapneumonic effusion

Stage/category	Pleural space appearance	Pleural fluid bacteriology	Pleural fluid chemistry	Invasive intervention needed
I	Minimal, free-flowing effusion (<10 mm on lateral decubitus)	Culture and Gram stain results unknown	pH unknown	No
II	Small to moderate free-flowing effusion (>10 mm and, <1/2 hemithorax)	Negative culture and Gram stain	pH $\geq$ 7.20	Tailored therapy
III	Large, free-flowing effusion ( $\geq$ 1/2 hemithorax), loculated effusion, or effusion with thickened parietal pleura	One positive culture or Gram stain	pH <7.20	Yes (chest drain, fibrinolytics, MT, surgery)
IV		Frank purulent appearance		Yes (chest drain, fibrinolytics, MT, surgery)

Modified from the American College of Chest Physicians guidelines on medical and surgical treatment of parapneumonic effusions (113). MT, medical thoracoscopy.

considered a pleurodesis technique if there are no surgical indications (e.g., thoracic duct ligation) (108).

### **Hepatic hydrothorax**

Patients suffering from an advanced liver disease with transudative pleural effusion are characterized by an unfavourable prognosis and are fragile from a procedural point of view. Both MT and VATS in these patients have favourable results on symptoms in the short term but are burdened by high periprocedural morbidity and mortality (109,110). There are currently no evidence-supported therapeutic roles for MT in the setting of hepatic hydrothorax outside of overlapping complications such as PI (111).

### **Lysis of septations and debridement of the pleural cavity**

#### **PI**

About 85% of parapneumonic effusions completely resolve with medical treatment alone, the remaining 15% develop empyema (112). The early stages of a parapneumonic effusion (see *Table 5*) usually do not require invasive intervention (113) stages III and IV require further procedures to reduce the risk of severe sepsis, septic shock, and mortality (114,115).

Treatment of complicated parapneumonic effusions (CPPEs) and empyema relies on pleural fluid removal and immediate definition of the next course of action. While surgery has always been a cornerstone for treating PI (116),

early pathological stages respond to chest tube drainage and administration of intrapleural fibrinolysis, improving fluid drainage and reducing surgical referral and LOS, as described in the MIST-2 trial (117). MIST-3, a prospective multicenter phase III RCT, was recently published and evaluated the outcomes of standard of care, early VATS, and intrapleural enzyme therapy (IET) in PI. The study randomized 60 patients and demonstrated that early VATS may reduce LOS, while IET was associated with reduced recovery time and earlier pain relief (118,119).

The role of MT in PI is currently supported by weak evidences. However, in clinical practice it remains a useful alternative treatment approach for PI, even if current guidelines do not recommend its use in this scenario (24). Indications of MT include mechanical clearing of septations with the thoracoscope, draining of pleural fluid, irrigation of the pleural space with saline (14,120-122) or with fibrinolytics (123,124), and insertion of chest tubes under direct endoscopic visualization (125).

While few studies are available, a retrospective case series reporting on MT in CPPE and tuberculous pleural effusion described a success rate of 97.5%, with only 1.2% of treated patients subsequently requiring surgical intervention (126). An Italian cohort described by Ravaglia *et al.* demonstrated a 75.6% success rate, with the need to proceed subsequently to surgery in 7.8% of patients (127). The same study demonstrated a success rate of 100% for MT in stage I PI, 83.3% in stage II, and 58.1% in stage III. The authors of the aforementioned paper recommended MT as the first-line treatment, especially in fragile patients, encouraging the initiation of therapeutic measures early in the disease process (127).

A recent RCT by Kheir *et al.*, with LOS as the primary endpoint, compared IET with early MT for PI. Its conclusions support MT as the first therapeutic choice, being associated with a shorter LOS than IET alone (23). In addition, a pleural biopsy obtained through MT increased the microbiological diagnostic yield by 12.5%, therefore aiding in selecting personalized rather than empiric antibiotic therapy (23).

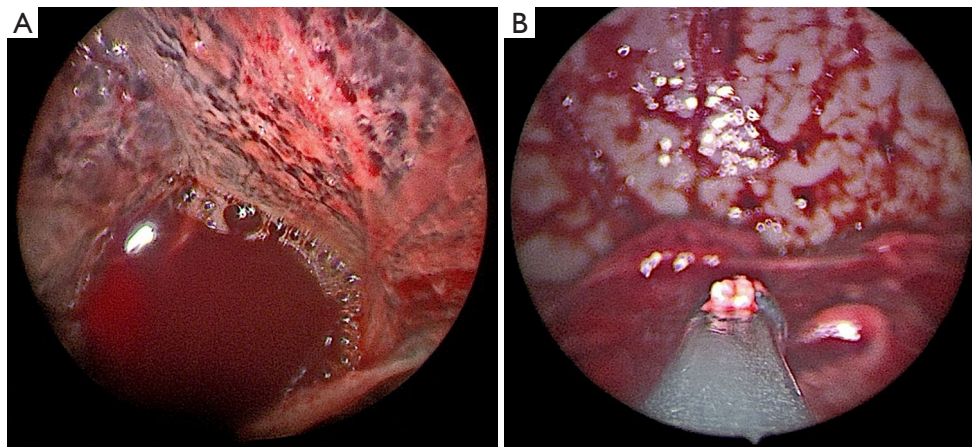
A systematic review and meta-analysis by Mondoni *et al.*, pooling the results of eight studies, reported a pooled treatment success rate of MT of 85% for PI when used as a first-line intervention or after the failure of a chest tube insertion and a pooled complication rate of 9.0% (128). The success of the treatment was further increased by carrying out subsequent fibrinolytic therapy through the chest drain positioned at the end of the MT (128). Surgical intervention remains the primary rescue therapy in case of failure of medical therapy. In addition, when lung re-expansion is not achieved with MT alone, surgery should be considered, especially if the patient remains persistently symptomatic despite adequate infection control (129). Surgical intervention should be advised in this case because lung entrapment may be determined by the development of a pleural fibrotic peel, making the decortication procedure indispensable and preferably carried out in VATS (130,131). There is a current need for large prospective randomized trials comparing MT with surgical intervention. The pending 'studying Pleuroscopy in Routine Pleural Infection Treatment' (SPIRIT) trial is worth mentioning, and its results are currently awaiting publication (132).

The clinical entity of pediatric empyema is also worth mentioning (133-136). About 5% of children with community-acquired pneumonia who require hospitalization develop empyema. The primary treatment consists of antibiotic therapy (136-138) and the placement of a chest tube in order to drain the pleural fluid (138,139). Refractory cases may require the use of fibrinolytic agents (140,141). There is currently no available data regarding the use of MT in treating PI in the pediatric setting. A recent case series by Zuccatosta *et al.* describes their experience with MT in multiloculated pediatric empyema defined by laboratory data, clinical presentation, and ultrasound imaging (142). The population consisted of six patients treated with MT; five had multiloculated empyema, with the remaining presenting with organized empyema. MT was performed in all patients with successful recovery and full lung re-expansion on chest X-rays, with microbiological diagnoses made in four cases. Despite the small sample size,

it provides promising results and inspires future studies by describing a favorable safety profile for MT while providing definitive treatment without sequelae on follow-up (142).

### RH

Acute hemothorax is a medical emergency most frequently caused by trauma, and its identification by chest X-ray, chest CT, and thoracentesis requires immediate treatment, commonly with chest tube placement and antibiotic prophylaxis, VATS, or thoracotomy (143,144). Quantification of blood loss can help guide medical treatment with surgical exploration with VATS or thoracotomy performed in large effusions (>1,500 mL), ongoing blood loss (>200 mL/h), or when a conservative approach is deemed insufficient (145). Thoracotomy and VATS allow clearing of the pleural cavity, quantification of blood loss, decortication, and chest tube placement for residual fluid evacuation and lung expansion (146-148). These techniques represent the gold standard of hemothorax treatment; however, thoracotomy and VATS can be challenging in fragile patients or those with multiple comorbidities (149). Current literature reporting the use of MT in hemothorax is scarce and limited to case studies in fragile patients not deemed fit enough to undergo surgery. In light of the reduced tools to deal with acute bleeding, the main indication for MT is RH (*Figure 6*). RH is defined as a residual hematic pleural effusion larger than 500 mL after 72 h of treatment with a chest tube (150). One case report describes the use of MT in a 76-year-old male with pleural sarcomatoid carcinoma with multiple comorbidities and high risk for surgery in whom MT was used for treating hemothorax and investigating the underlying pleural pathology (151). Bioptic samples were obtained with rigid thoracoscopy followed by APC treatment of actively bleeding nodules. The case study is promising regarding underlying MT's utility in this setting and the implementation of APC for limiting active bleeding. A similar example was described by Srinivasan *et al.*, reporting the case of an 82-year-old patient treated with flexi-rigid MT with a delayed presentation of hemothorax following blunt trauma resulting in RH (27). The RH proved refractory to chest drainage but the challenge was overcome with the use of a cryoprobe which allowed the removal of retained adhesions and clots, resulting in a well-tolerated procedure with immediate relief of respiratory symptoms. The case highlights the usefulness of using bronchoscopic accessories in the pleural setting in order to achieve satisfactory results when dealing



**Figure 6** Endoscopic view of retained haemothorax. (A) View at the time of entry into the pleural cavity with evidence of adhesions. (B) View after debridement using a rigid thoracoscope with rigid forceps.



**Video 3** Endoscopic view of intrapleural foreign body: fragment of catheter for extemporaneous cardiac pacing.

with complex situations in which surgery is not an option and conservative treatment is found to be ineffective.

### Intrapleural FB recovery

Intrapleural FBs are rare and may occur after medical manipulation of the pleural space. Retrieval is typically performed through thoracotomy and VATS under general anesthesia (152).

There is a scarcity of literature describing the retrieval of FBs through MT, and most of the documents are anecdotal case reports. MT can be used as an alternative in patients with poor lung function or those who do not tolerate surgical procedures to retrieve intrapleural FBs (14,149).

A case report describes the success of rigid thoracoscopy in

the retrieval of a self-expandable metallic stent (SEMS) from the pleural cavity in a patient with right lower lobe squamous cell lung cancer that had determined endobronchial obstruction with subsequent formation of lung abscess leading to the formation of a broncho-pleural fistula and migration of the SEMS into the pleural cavity (153).

Narasimhan *et al.* described two cases in which MT was used to retrieve an aspiration needle dislodged within the pleural space during thoracentesis. In both cases, the procedure was conducted using a rigid thoracoscope under local anesthesia and conscious sedation, allowing for the removal *en bloc* of the intrapleural FB while allowing for simultaneous biopsy sampling of pleural lesions after drainage of the pleural effusion (154).

In the authors' experience, MT was used to remove a fragment of a catheter used for extemporaneous cardiac pacing from a patient who underwent major thoracic surgery and was subsequently not deemed operable (*Video 3*).

While MT is useful for retrieving FBs in fragile patients, VATS remains the gold standard, particularly for large FBs, in a condition of foreseeable significant pleural adhesions or associated hemothorax.

### Strengths and limitations of the review

The narrative review was designed to summarize findings from the literature on the identified topic. Despite the increase in prospective trials, RCTs, and meta-analyses developed in the field of MT, much of the inherent literature comprises retrospective studies, case series, and case reports. These may not be appropriately powered to

detect significant results and are likely subject to bias and procedural errors, limiting their generalizability.

## Conclusions

MT is a safe and efficient treatment with a wide range of indications for different pleural diseases. When carried out by a properly qualified operator, there are few complications even in the therapeutic setting. This procedure is one that can be performed using spontaneous ventilation and local anaesthesia with or without sedation. It should be acknowledged that there is supporting evidence that MT and VATS share common indications. Nevertheless, lung isolation and general anesthesia are frequently needed for VATS. There have been reports of VATS procedures performed under regional or epidural anesthesia and without the need for intubation (155,156), but MT continues to be the least invasive technique for visualizing the pleural space (157). The procedural boundaries may become increasingly blurred with more research on these different procedural approaches.

## Future trends

Future scientific studies should assess the optimal indication for MT regarding prognostic outcomes and cost-effectiveness. The procedure should not aim to replace surgical alternatives for the pathological entities in which it is applied, but should find its role in selected cases as a low-cost, safe, and highly effective therapeutic choice in the hands of the interventional pulmonologist. The endpoints of LOS, surgical referral, short- and long-term complications and mortality must be considered to demonstrate MT's validity as a therapeutic maneuver to be applied globally.

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