

MEDICAL INSIGHTS

Transbronchial cryobiopsy for the diagnosis of interstitial lung diseases (ILDs)

Current procedural guidance for interventional bronchoscopists according to the literature

Background

Data supporting transbronchial cryobiopsy (TBCB) for diagnosis of ILDs continues to mount. The technique represents a reasonable alternative to surgical lung biopsy (SLB) contributing to the multidisciplinary discussion (MDD) and is adopted in an increasing number of centers around the globe^{1,3}.

Mortality rates for TBCB are low with a pooled estimate of 0.5 %³. Acute exacerbation rates and length of hospital stay were also reported to be lower compared to SLB¹. Although SLB carries an in-hospital mortality of up to 16 % if conducted non-electively, it is still commonly used for sampling in ILD-patients². 80 % of SLBs could be avoided by performing a TBCB¹.

Challenges and goals

For idiopathic pulmonary fibrosis (IPF) and fibrotic hypersensitivity pneumonitis, recommendations have been made for TBCB over SLB^{6,7}. A lack of procedural standardization and high degree of interobserver variability between different institutions make it difficult to recommend TBCB over SLB in general¹. To this end, common questions on the application of Erbe flexible cryoprobes as an indispensable part of TBCB need to be addressed.

Method

Avasarala et al. conducted research on MEDLINE and Cochrane library comprising articles from 2009 to 2020 on cryobiopsy. 263 publications were selected for an evidence-based narrative review to address common procedural questions¹.

Results and key findings

Diagnostic yield

Pooled estimate for the diagnostic yield for TBCB independently from the MDD was 82.5 % but should be viewed with caution, as the MDD represents the gold standard^{1,3}. Nevertheless, TBCB could add diagnostic confidence of between +31 % and +34 % to clinical, radiographic and bronchoalveolar lavage data^{1,4}.

Procedural environment

Physicians should be trained on the procedure and management of severe complications (severe bleeding & tension pneumothorax). It is suggested that TBCB is performed in an operating room or endoscopy suite under general anesthesia. Interventional radiology, thoracic surgery and critical care unit should be available for emergent situations¹.

Contraindications

- → Rapid clinical decline (can indicate an impending exacerbation, e.g. development of unexplained patchy ground-glass opacities)¹
- → Pulmonary hypertension (> 50 mmHg relative contraindication, risk of damage to pathologic vasculature)¹
- → Uncorrected bleeding diathesis (iatrogenic or pathologies, aspirine medication is considered a relative contraindication)¹
- \rightarrow Compromised pulmonary function (see Fig. 1)¹

Cryobiopsy collection

TBCB can be performed by the use of a flexible bronchoscope through an endotracheal tube or a rigid bronchoscope¹. A bronchial blocker is placed for bleeding control on segmental level^{1,3}. The cryoprobe is retracted 1 cm after resistance of the visceral pleura is felt¹. Fluoroscopy for probe placement helps preventing a pneumothorax¹. Depending on the size of the cryoprobe, freezing is activated for 3 to 6 seconds before en-bloc extraction¹. A freeze test in water before the procedure can support adjustment of the activation time¹.

Management of complications

Pneumothorax occurs in approximately 10% of patients³. A chest drainage insertion kit, drainage apparatus and fluoroscopy/ultrasound to aid placement should be available. Further, supplies to secure and connect a chest tube are needed¹.

Severe bleeding occurs in 0.3 % of patients¹. A larger endotracheal tube (8.5 mm diameter or larger) or rigid bronchoscope are necessary to secure the airway along with a bronchial blocker¹.

Histopathological considerations

The fragile specimen should be carefully thawed into saline without scraping or squeezing¹. It is embedded in paraffin with an orientation to maximize the surface area¹.



Thresholds in pulmonary function representing contraindications

Implications

Avasarala et al. conclude that TBCB is a minimally invasive, effective and safe biopsy technique. TBCB outcomes could be improved by further technological harmonization¹.

Friction in the 2.0 mm working channel hampering tactile identification of the pleura, and a small remaining cross section of the working channel available for suction to control bleeding were discussed¹. With respect to prevention of a pneumothorax, Avasarala et al. make a weak recommendation for the flexible reuseable 1.9 mm cryoprobe (recommendation grade 2B)¹.

The new flexible single-use 1.7 mm cryoprobe did not show a significantly different sample size when compared to the flexible reuseable 1.9 mm cryoprobe⁵. Additionally, the smaller cryoprobe provides less friction and better suction through the working channel¹. To reduce the risk of a pneumothorax, retraction of 1 cm after contact with the visceral pleura is strongly recommended¹. The radiopaque probe tip of the 1.7 mm flexible single-use cryoprobe has a length of approximately 6 mm and can serve for distance estimation under fluoroscopy¹. When the angle of the fluorosocpy beam is not perpendicular to the probe, care should be taken in distance estimation. To prevent coughing and dislocation of the cryoprobe, Avasarala et al. recommend the use of general anesthesia¹. In previous literature, deep sedation has been deemed applicable, too⁸.



Remaining cross section of a 2.0 mm working channel depending on the diameter of the inserted cryoprobe

Products



The Erbe portfolio includes a flexible single-use cryoprobe with a diameter of 1.7 mm which can be used equivalently to the flexible re-useable 1.9 mm cryoprobe⁵. These are operated with the ERBECRYO[®] 2. Depending on the size of the cryoprobe, freezing times between 3 and 6 seconds were reported¹.

References

1. Avasarala SK, Wells AU, Colby TV, Maldonado F. Transbronchial Cryobiopsy in Interstitial Lung Diseases: State-of-the-Art Review for the Interventional Pulmonologist. J Bronchology Interv Pulmonol. 2021 Jan 1;28(1):81-92. doi: 10.1097/ LBR.000000000000716. PMID: 32960830.

2. Hutchinson JP, Fogarty AW, McKeever TM, Hubbard RB. In-Hospital Mortality after Surgical Lung Biopsy for Interstitial Lung Disease in the United States. 2000 to 2011. Am J Respir Crit Care Med. 2016 May 15;193(10):1161-7. doi: 10.1164/rccm.201508-16320C. PMID: 26646481.

3. Maldonado F, Danoff SK, Wells AU, Colby TV, Ryu JH, Liberman M, Wahidi MM, Frazer L, Hetzel J, Rickman OB, Herth FJF, Poletti V, Yarmus LB. Transbronchial Cryobiopsy for the Diagnosis of Interstitial Lung Diseases: CHEST Guideline and Expert Panel Report. Chest. 2020 Apr;157(4):1030-1042. doi: 10.1016/j. chest.2019.10.048. Epub 2019 Nov 27. PMID: 31783014.

4. Hetzel J et al. Transbronchial cryobiopsy increases diagnostic confidence in interstitial lung disease: a prospective multicentre trial. Eur Respir J. 2020 Dec 10;56(6):1901520. doi: 10.1183/13993003.01520-2019. PMID: 32817003.

5. Hetzel J et al. Evaluation of Efficacy of a New Cryoprobe for Transbronchial Cryobiopsy: A Randomized, Controlled in vivo Animal Study. Respiration. 2020;99(3):248-256. doi: 10.1159/000506017.

6. Behr J et al. S2K Guideline for Diagnosis of Idiopathic Pulmonary Fibrosis. Respiration. 2021 Jan 22:1-34. doi: 10.1159/000512315.

7. Raghu G et al. Diagnosis of Hypersensitivity Pneumonitis in Adults. An Official ATS/JRS/ALAT Clinical Practice Guideline. Am J Respir Crit Care Med. 2020 Aug 1;202(3):e36-e69. doi: 10.1164/rccm.202005-2032ST. Erratum in: Am J Respir Crit Care Med. 2021 Jan 1;203(1):150-151.

8. Hetzel J et al. Transbronchial Cryobiopsies for the Diagnosis of Diffuse Parenchymal Lung Diseases: Expert Statement from the Cryobiopsy Working Group on Safety and Utility and a Call for Standardization of the Procedure. Respiration. 2018;95(3):188-200. doi: 10.1159/000484055. Epub 2018 Jan 9. PMID: 29316560.

Prof. Felix Herth (Thoraxklinik, University of Heidelberg) demonstrating a transbronchial cryobiopsy for ILD diagnosis





CRYO.ERBE-MED.COM

medical-videos.com

Important information

We have prepared this document with care. Nonetheless, we cannot completely rule out errors in this document.

The information, recommendations and other data ("Information") contained in this document reflect our state of knowledge and the state of science and technology at the time of preparing the document. The information is of a general nature, non-binding and serves solely for general information purposes and does not represent instructions for use or notes on application.

The information and recommendations contained in this document do not constitute any legal obligations on Erbe Elektromedizin GmbH or their associated companies ("Erbe") or any other claims against Erbe. The information does not represent a guarantee or other quality statement; these require an express contractual arrangement with Erbe in individual cases.

Erbe shall not be liable for any type of damage resulting from following information given in this document, regardless of the legal reason for liability.

Every user of an Erbe product is responsible for checking the respective Erbe product for its properties as well as the suitability for the intended type of application or intended purpose in advance. The suitable type of application of the respective Erbe product is given by the user manual and the notes on use for the corresponding Erbe product. The user is obliged to check whether the existing user manual and the notes on use correspond with the status for the specific Erbe product. The devices may only be used according to the user manual and the notes on use.

The information on setting values, application sites, duration of application and the use of the respective Erbe product is based on the clinical experience of physicians independent from Erbe. They represent guidelines which need to be checked by the user for their suitability for the actual planned application. Depending on the circumstances of an actual application case, it may be necessary to deviate from the information provided. The user is responsible for checking this in each case when using an Erbe product. We wish to point out that science and technology is constantly subject to new developments arising from research and clinical experience. For this reason, it may be necessary for the user to deviate from the information provided in this document.

This document contains information about Erbe products which may possibly not be approved in a specific country. The user of the respective Erbe product is obliged to inform him/herself as to whether the Erbe product he/she is using is legally approved in his/her country and/or if legal requirements or restrictions for use possibly exist and to what extent.

This document is not intended for users in the USA.

Erbe Elektromedizin GmbH Waldhoernlestrasse 17 72072 Tuebingen Germany Phone +49 7071 755-0 info@erbe-med.com erbe-med.com medical-videos.com

