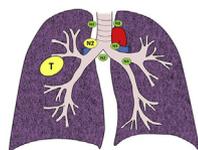


The authors reported no conflicts of interest.

The *Journal* policy requires editors and reviewers to disclose conflicts of interest and to decline handling or reviewing manuscripts for which they may have a conflict of interest. The editors and reviewers of this article have no conflicts of interest.



**SYSTEMATIC OR TARGETED SAMPLING DURING ENDOBRONCHIAL ULTRASOUND FOR MEDIASTINAL**



**STAGING IN PATIENTS WITH LUNG CANCER AND ABNORMAL MEDIASTINUM**

**To the Editor:**

In the recently published article by Sullivan and colleagues,<sup>1</sup> the authors describe the feasibility of a randomized controlled trial that aims to compare targeted with systematic endobronchial ultrasound–guided transbronchial needle aspiration (EBUS-TBNA) sampling for mediastinal staging in patients with lung cancer. The authors hypothesize that targeted EBUS-TBNA sampling is not inferior to systematic EBUS-TBNA. A targeted, selected, or “hit-and-run” EBUS-TBNA sampling is defined as a specific investigation for malignancy-suspected mediastinal nodes based on positron emission tomography-computed tomography (PET-CT) findings (fluorodeoxyglucose avid or short axis  $\geq 10$  mm).<sup>2</sup> On the contrary, a systematic EBUS-TBNA sampling is defined as a systematic investigation of all lymph nodes with a short axis  $\geq 8$  mm by EBUS, regardless of their appearance on PET-CT. This is not the first study that aims to compare both strategies. Previous studies that included patients with abnormal mediastinum on PET-CT have demonstrated the benefits of systematic over targeted sampling in terms of clinically relevant information. Compared with targeted sampling, systematic sampling enhances the sensitivity of EBUS-TBNA for the diagnosis of mediastinal nodal metastases and determines the extent of the mediastinal disease more accurately.<sup>2-4</sup>

In the study of Sullivan and colleagues,<sup>1</sup> the authors compare targeted sampling with systematic sampling based on 3 nodal features: appearance on CT, appearance on PET (fluorodeoxyglucose avidity) and appearance on EBUS, based on the Canada Lymph Node Score.<sup>5</sup> However, since all included patients present normal mediastinum on image

techniques, the choice of a systematic sampling or a targeted sampling is based only on EBUS findings (Canada Lymph Node Score). Based on their initial results, with an incidence of missed mediastinal metastases of 5.45% for targeted sampling, the authors affirm that targeted sampling is safe enough, and that there are no clinically significant differences between both strategies (although there is an increase in the detection of malignant nodes from 1.75% for targeted sampling to 7.27% for systematic sampling).

However, in our opinion, these results must be interpreted with prudence. First, because not all negative results were confirmed by surgery, and second, because all the included patients had normal mediastinum on imaging techniques. In such clinical scenario, with low prevalence of mediastinal involvement, it is well known that the sensitivity of EBUS-TBNA declines and, therefore, differences between a targeted sampling and a systematic sampling may appear lower than they are in patients with abnormal mediastinum.

We agree with the authors that it is important to establish strategies to reduce the timing of mediastinal staging in patients with lung cancer. This means avoiding redundant explorations and reducing the length of the procedures. However, mediastinal staging is a crucial step in the management of patients with lung cancer and any of these measures cannot affect its thoroughness. Systematic staging through EBUS-TBNA in patients with abnormal mediastinum on PET-CT has demonstrated to be more precise than targeted sampling and still must be recommended in such clinical scenario.

José Sanz-Santos, MD, PhD<sup>a,b</sup>

Pere Serra, MD, PhD<sup>c</sup>

Antoni Rosell, MD, PhD<sup>c,d,e,f</sup>

<sup>a</sup>Pulmonology Department

Hospital Universitari Mútua Terrassa

Terrassa, Barcelona, Spain

<sup>b</sup>University of Barcelona, School of Medicine

Barcelona, Spain

<sup>c</sup>Pulmonology Department

Hospital Universitari Germans Trias i Pujol

Badalona, Barcelona, Spain

<sup>d</sup>Universitat Autònoma de Barcelona (UAB) Cerdanyola

del Vallés

Barcelona, Spain

<sup>e</sup>Centro de investigación en red de enfermedades

respiratorias (CIBERES)

Madrid, Spain

<sup>f</sup>Institut de recerca Germans Trias I Pujol (IGTP)

Badalona, Barcelona, Spain

**References**

- Sullivan KA, Farrokhyar F, Leontiadis GI, Ptel YS, Churchill IF, Hylton DA, et al. Routine systematic sampling vs. targeted sampling during endobronchial ultrasound: a randomized feasibility trial. *J Thorac Cardiovasc Surg.* December 4, 2021 [Epub ahead of print].

Copyright © 2022 The Author(s). Published by Elsevier Inc. on behalf of The American Association for Thoracic Surgery. This is an open access article under the CC BY-NC-ND license (<http://creativecommons.org/licenses/by-nc-nd/4.0/>).

2. Crombag LMM, Dooms C, Stigi JA, Tournoy KG, Schuurbiens OCJ, Ninaber MK, et al. Systematic and combined endosonographic staging of lung cancer (SCORE study). *Eur Respir J*. 2019;E3:18008800.
3. Sanz-Santos J, Serra P, Torky M, Andreo F, Centeno C, Mendiluce L, et al. Systematic compared with targeted staging with endobronchial ultrasound in patients with lung cancer. *Ann Thorac Surg*. 2018;106:398-403.
4. Cole AJ, Harcastle N, Turgeon GA, Thomas R, Irving LB, Jennings BR, et al. Systematic endobronchial ultrasound-guided transbronchial needle aspiration improves radiotherapy planning in non-small cell lung cancer. *ERJ Open Res*. 2019;15:5:00004-2019.
5. Hylton DA, Turner S, Kidane B, Spicer J, Xie F, Farrokhyar F, et al. The Canada lymph node score for prediction of malignancy in mediastinal lymph nodes during endobronchial ultrasound. *J Thorac Cardiovasc Surg*. 2020;159:2499-507.

<https://doi.org/10.1016/j.xjon.2022.01.030>