

Presentation Abstracts

001

Assessment of genomic variants and overall survival in primary tumor lung adenocarcinoma and corresponding mediastinal lymph nodes obtained by EBUS- TBNA

Caroline Silvério Faria^{1,2}, Camila Machado Baldavira², Tabatha Gutierrez Prieto Martins Rocha², Viviane Rossi Figueiredo³, Ellen Caroline Toledo do Nascimento⁴, Evandro Sobroza de Mello⁵, Murilo Castro Cervato⁷, Ricardo Mingarini Terra⁶, Vera Luiza Capelozzi², Leila Antonangelo^{1,8}

¹ Laboratório de Investigação Médica LIM/03, Departamento de Patologia, Faculdade de Medicina, Universidade de São Paulo, São Paulo, SP, Brazil.

² Laboratorio de Histomorfometria e Genômica Pulmonar, Departamento de Patologia, Faculdade de Medicina, Universidade de São Paulo, São Paulo, SP, Brazil.

³ Serviço de Endoscopia Respiratória, Instituto do Câncer do Estado de São Paulo, Hospital das Clínicas, Faculdade de Medicina, Universidade de São Paulo, São Paulo, SP, Brazil.

⁴ Laboratório de Anatomia Patológica, Instituto do Coração (InCor), Hospital das Clínicas, Faculdade de Medicina, Universidade de São Paulo, São Paulo, SP, Brazil.

⁵ Laboratório de Patologia, Instituto do Câncer do Estado de São Paulo, Hospital das Clínicas, Faculdade de Medicina, Universidade de São Paulo, São Paulo, SP, Brazil.

⁶ Hospital Israelita Albert Einstein, São Paulo, SP, Brazil.

⁷ Serviço de Cirurgia Torácica Oncológica, Instituto do Câncer do Estado de São Paulo, Hospital das Clínicas, Faculdade de Medicina, Universidade de São Paulo, São Paulo, SP, Brazil.

⁸ Divisão de Laboratório Central, Departamento de Patologia, Hospital das Clínicas, Faculdade de Medicina, Universidade de São Paulo, São Paulo, SP, Brazil.

Category: Respiratory Endoscopy

DOI: 10.31744/einstein_journal/2024ABS_BTS_ST0001

Caroline Silvério Faria - <https://orcid.org/0000-0001-5272-2169>
 Camila Machado Baldavira - <https://orcid.org/0000-0002-5364-7305>
 Tabatha Gutierrez Prieto Martins Rocha - <https://orcid.org/0000-0002-2615-6922>
 Viviane Rossi Figueiredo - <https://orcid.org/0000-0002-9458-8153>
 Ellen Caroline Toledo do Nascimento - <https://orcid.org/0000-0002-2981-1419>
 Evandro Sobroza de Mello - <https://orcid.org/0000-0002-4383-2910>
 Murilo Castro Cervato - <https://orcid.org/0000-0001-8146-5798>
 Ricardo Mingarini Terra - <https://orcid.org/0000-0001-8577-8708>
 Vera Luiza Capelozzi - <https://orcid.org/0000-0001-9732-5853>
 Leila Antonangelo - <https://orcid.org/0000-0002-8634-5100>

Corresponding author

e-mail: lantonangelo@hc.fm.usp.br

Introduction: Histopathological analyses and classification by the Tumor, Node, Metastasis System (TNM) are key-elements in therapeutic decision-making for non-small cell lung cancer (NSCLC).⁽¹⁾ Endobronchial ultrasound-guided transbronchial needle aspiration (EBUS-TBNA) is effective in obtaining biopsies of hilar and mediastinal lymph nodes (MLN), for tissue collection to mediastinal staging in NSCLC.^(2,3) The assessment of clinically significant genomic alterations in surgically resected primary tumors (PT) and MLN aspirates obtained by EBUS-TBNA can provide valuable insights into the risk of recurrence, presence of occult metastases, and the identification of patients eligible for personalized therapies.⁽⁴⁾

Objectives: To analyze somatic variants of clinical relevance in genomic DNA (gDNA) extracted from formalin-fixed, paraffin-embedded (FFEP) PT tissues and MLN aspirates using next-generation sequencing (NGS), correlating the findings with patient overall survival.

Methods: The gDNA was extracted from paired samples (PT and MLN) from 32 patients with lung adenocarcinoma, resulting in 64 samples, and analyzed using a customized genetic panel consisting of 107 genes, utilized SureSelect XTHS2 kit (Agilent Technologies, Santa Clara, CA, USA). The overall

survive were assessed using Kaplan-Meier curves (IBM SPSS Statistics 25.0; Chicago, Illinois, EUA).

Results: Strong and potential clinical significance variants were detected in 72% of PT and 75% of MLN samples. These variants were identified in 14 different genes, highlighting *EGFR*, *TP53*, *KRAS*, and *ATM*, with a frequency of more than 15% and 10% in PT and MLN samples, respectively. Concerning survival analysis, no significant difference was observed in PT samples. Surprisingly, in MLN samples, significant differences were noted in the *TP53* and *ATM* genes (p-values of 0.001 and 0.035, respectively). This suggests that the identified variants may correlate with worse prognosis, recurrence or occult metastasis in the patients evaluated.

Conclusion: In summary, we identified distinct somatic variants between PT and MLN samples, with survival being negatively impacted by the presence of variants in MLN. Thus, the incorporation of genomic testing applied to MLN samples could enhance the accuracy of TNM staging, providing a more comprehensive approach to patient management.

Keywords: Cancer genomic; Mediastinal lymph node; Adenocarcinoma; EBUS-TBNA; NGS

REFERENCES

1. Barta JA, Powell CA, Wisnivesky JP. Global epidemiology of lung cancer. *Ann Glob Health*. 2019;85(1):8. Review.
2. Harangoş A, Berindan-Neagoe I, Toma L, Şimon I, Pop O, Şimon M. EBUS in optimizing non-small cell lung cancer diagnosis and treatment. *Med Pharm Rep*. 2021;94(2):176-84.
3. Martin-Deleon R, Teixido C, Lucena CM, Martinez D, Fontana A, Reyes R, et al. EBUS-TBNA Cytological Samples for Comprehensive Molecular Testing in Non-Small Cell Lung Cancer. *Cancers (Basel)*. 2021;13(9):2084.
4. Martin-Deleon R, Teixido C, Lucena CM, Martinez D, Fontana A, Reyes R, et al. EBUS-TBNA Cytological Samples for Comprehensive Molecular Testing in Non-Small Cell Lung Cancer. *Cancers (Basel)*. 2021 Apr 25;13(9):2084.

SGPP number: Not applicable.

CEP: FMUSP 3.004.983/2018.

Research funding: *Fundação de Amparo à Pesquisa do Estado de São Paulo (FAPESP): 2019/04416-3, 2018/20403-6, 2023/02755-0, 2023/01158-9; Conselho Nacional de Desenvolvimento Científico e Tecnológico (CNPq): 303735/2021-0; Coordenação de Aperfeiçoamento de Pessoal de Nível Superior (CAPES)/ Programa de Demanda Social (DS) # 88887.472497/2019-00.*