

años tenían infección por *P. aeruginosa* (17% intermitente y 28,3% crónica) con una mediana de edad de primoinfección de 5,2 años<sup>15</sup>. Si comparamos los resultados de estas publicaciones con los nuestros, podemos observar que, aunque en todos ellos se objetiva una disminución de la prevalencia de infección crónica por *P. aeruginosa* en los pacientes DCN, nuestros resultados muestran una menor prevalencia. Esto podría ser debido a un seguimiento muy estrecho, con visitas y cultivos de secreciones respiratorias mensuales, con tratamientos precoces e intensivos de la primoinfección por *P. aeruginosa* y también porque hay mayor cantidad de pacientes del grupo FR.

En conclusión, observamos un cambio en la historia natural de la infección bronquial por *P. aeruginosa* en la fibrosis quística tras la implantación del cribado neonatal, con una disminución significativa ( $p < 0,001$ ) de la prevalencia de infección crónica por dicho patógeno en estos pacientes.

### Financiación

Los autores no han recibido ayudas de financiación.

### Conflicto de intereses

Los autores declaramos que no hay conflicto de intereses.

### Bibliografía

1. Palser S, Smith S, Nash EF, Agarwal A, Smyth AR. Treatments for preventing recurrence of infection with *Pseudomonas aeruginosa* in people with cystic fibrosis. *Cochrane Database Syst Rev*. 2019;12:CD012300.
2. Cantón R, Fernández A, Gómez E, Campo R, Meseguer MA. Infección bronquial crónica: el problema de *Pseudomonas aeruginosa*. *Arch Bronconeumol*. 2011;47 Supl 6:8-13.
3. Milczewska J, Wolkowicz T, Zacharczuk K, Mierzejewska E, Kwiatkowska M, Walicka-Serzysko K, et al. Clinical outcomes for cystic fibrosis patients with *Pseudomonas aeruginosa* cross-infections. *Pediatr Pulmonol*. 2020;55:161-8.
4. Parkins MD, Somayaji R, Waters VJ. Epidemiology, biology, and impact of clonal *Pseudomonas aeruginosa* infections in cystic fibrosis. *Clin Microbiol Rev*. 2018;31(4). E000019-18.

5. Cantón R, Maíz L, Escribano A, Oliveira C, Olicer A, Asensio O, et al. Consenso español para la prevención y el tratamiento de la infección bronquial por *Pseudomonas aeruginosa* en el paciente con fibrosis quística. *Arch Bronconeumol*. 2015;51:140-50.
6. Lund-Palau H, Turnbull AR, Bush A, Bardin E, Cameron L, Soren O. *Pseudomonas aeruginosa* infection in cystic fibrosis: Pathophysiological mechanisms and therapeutic approaches. *Expert Rev Respir Med*. 2016;10:685-97.
7. Malhotra S, Hayes DJ, Wozniak DJ. Cystic fibrosis and *Pseudomonas aeruginosa*: The host-microbe interface. *Clin Microbiol Rev*. 2019;32(3).
8. Gartner S, Cobos N. Cribado neonatal para la fibrosis quística. *An Pediatr (Barc)*. 2009;71:481-2.
9. Cantón R, Cobos N, de Gracia J, Baquero F, Honorato J, Gartner S, et al., en representación del Grupo Español de Consenso del Tratamiento Antimicrobiano en el Paciente con Fibrosis Quística. Consenso del tratamiento antimicrobiano en el paciente con fibrosis quística. *Arch Bronconeumol*. 2005;41 Suppl 1:1-25.
10. Gartner S, Moreno A, Cobos N. Tratamiento de la enfermedad respiratoria en la fibrosis quística. En: Cobos N, Pérez-Yarza EG, editores. *Tratado de neumología pediátrica*. Madrid: Ergon; 2008. p.849-66.
11. Flume PA, O'Sullivan BP, Robinson KA, Goss CH, Mogayzel PJ, Willey Courand DB, et al. Cystic fibrosis pulmonary guidelines. *Am J Respir Crit Care Med*. 2007;176:957-69.
12. Proesmans M, Balinska-Miskiewicz W, Dupont L, Bossuyt X, Verhaegen J, Hoiby N, et al. Evaluating the 'Leeds criteria' for *Pseudomonas aeruginosa* infection in a cystic fibrosis centre. *Eur Respir J*. 2006;27:937-43.
13. Mak DY, Sykes J, Stephenson AL, Lands LC. The benefits of newborn screening for cystic fibrosis: The Canadian experience. *J Cyst Fibros*. 2016;15:302-8.
14. Schlüter DK, Southern KW, Dryden C, Diggle P, Taylor-Robinson D. Impact of newborn screening on outcomes and social inequalities in cystic fibrosis: A UK CF registry-based study. *Thorax*. 2020;75:123-31.
15. Annual Data Report 2018. Cystic Fibrosis Foundation Patient Registry; 2018.

Roser Ayats Vidal\*, Montserrat Bosque García, Miguel García González y Óscar Asensio de la Cruz

Unidad de Fibrosis Quística y Unidad de Neumología, Alergia e Inmunología pediátrica, Servicio de Medicina Pediátrica, Parc Taulí, Hospital Universitario de Sabadell, Sabadell (Barcelona), España

\* Autor para correspondencia.

Correo electrónico: rayatsv@tauli.cat (R. Ayats Vidal).

<https://doi.org/10.1016/j.arbres.2020.03.024>

0300-2896/© 2020 SEPAR. Publicado por Elsevier España, S.L.U. Todos los derechos reservados.

## Acute Respiratory Failure Caused by Pulmonary Lymphangitic Carcinomatosis in a Patient With Lung Adenocarcinoma at Initial Diagnosis



### Insuficiencia respiratoria aguda causada por linfangitis carcinomatosa pulmonar en un paciente con adenocarcinoma de pulmón desde el diagnóstico inicial

Dear Editor:

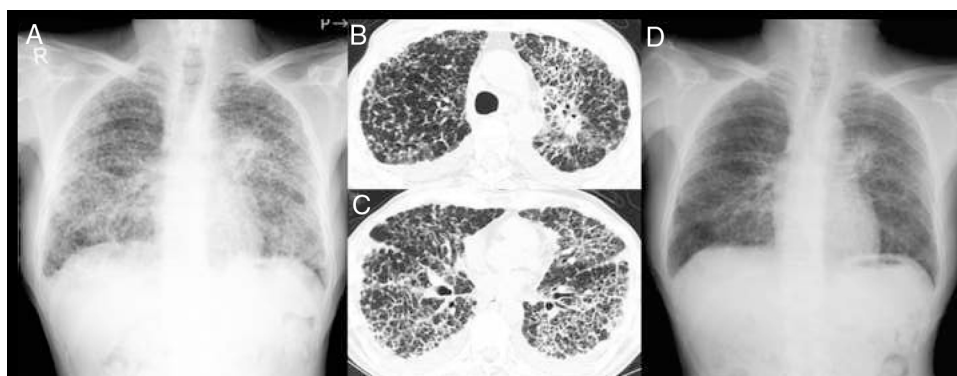
Pulmonary lymphangitic carcinomatosis is associated with poor prognosis and tissue collection is difficult depending on the respiratory condition in such patients.<sup>1</sup> Liquid biopsy could be highly effective and help in addressing the problem of poor prognosis in lymphangitic carcinomatosis patients. Here, we report a case of lung adenocarcinoma with acute severe respiratory failure caused by pulmonary lymphangitic carcinomatosis at initial diagnosis, who could receive appropriate therapy by using liquid biopsy.

A 70-year-old man with a 30 pack-year history of smoking was referred to our hospital with a 1-month history of slight fever and dyspnea on exertion. Initial diagnosis at a previous hospital had indicated bacterial pneumonia accompanied by interstitial pneumonia, which had been treated with an antibacterial drug. Sputum cytology had revealed adenocarcinoma and the patient was referred to our hospital.

On admission, the patient had bilateral inspiratory crackles on chest auscultation. Laboratory findings showed severe hypoxemia and elevated levels of Krebs von den Lungen-6 (KL-6, 2638 U/mL) and carcinoembryonic antigen (CEA, 6.9 ng/mL). Chest X-ray showed a mass shadow in the left hilar region and bilateral diffuse interstitial opacities (Fig. 1A). Computed tomography (CT) imaging revealed a mass in the left upper pulmonary lobe, bilateral diffuse beaded thickening of the intralobular septum, and minor bilateral pleural effusions (Fig. 1 B and C). Performing bronchoscopy was considered difficult because high flow of oxygen (reservoir mask: 6L/min) was required for severe respiratory failure.

Adenocarcinoma was confirmed by cytological examination of the pleural effusions obtained by left pleural puncture, and epidermal growth factor receptor (EGFR) exon 19 deletion was found in the plasma EGFR gene mutation test at the time of hospitalization. The patient was confirmed for EGFR mutation-positive lung adenocarcinoma accompanied by carcinomatosis lymphangitis and pleurisy, and treatment with osimertinib (80 mg/day) was initiated on the fourth day of admission.

Although respiratory failure exacerbated and noninvasive positive pressure ventilation (NPPV) was required temporarily, respiratory failure resolved and NPPV treatment was withdrawn after two weeks. After four weeks of treatment, the patient experienced symptomatic relief as evidenced by marked improvement in the patient's X-ray (Fig. 1D). EGFR exon 19 deletion was also detected in



**Fig. 1.** Chest X-ray showing a mass shadow in the left hilar region and bilateral diffuse interstitial opacities (A). Computed tomography revealing a mass in the left upper pulmonary lobe (B), pulmonary carcinomatous lymphangitis, and minor bilateral pleural effusions (C). Chest X-ray showing marked improvement after four weeks of treatment (D).

the pleural effusion samples. The patient was returned to the previous hospital. Currently, the patient is in partial remission following 12 months of treatment with osimertinib.

Pulmonary lymphangitic carcinomatosis is mainly detected in patients with adenocarcinoma, especially those with lung, breast, and gastric cancers, and associated with poor prognosis.<sup>1</sup> In advanced non-small cell lung cancer (NSCLC), identifying driver gene mutations is recommended, but tissue collection is difficult depending on the general condition of the patient, such as respiratory failure, which is often a time-limited situation in such patients. Liquid biopsy is useful when tissue sampling is difficult or inadequate.<sup>2</sup> Although liquid biopsy can only currently be used in clinical practice for EGFR sensitizing and resistance mutations, various NSCLC driver genes have been detected successfully with the help of several plasma-based tests including molecular tests (such as detection of ROS-1) even before proceeding to tissue-based tests.<sup>3–6</sup>

In the present case, the time gap between the plasma EGFR mutation test and result report was only three days. This case indicates that liquid biopsy for detecting other driver gene mutations in addition to the EGFR mutation are warranted, since a therapeutic effect can be expected in the patient with driver gene mutation-positive NSCLC; this is true even in cases where tissue tests are difficult, such as in patients with lymphangitic carcinomatosis, which generally have poor prognosis.

In summary, we reported a case of EGFR mutation positive lung adenocarcinoma with acute severe respiratory failure caused by pulmonary lymphangitic carcinomatosis. Although tissue collection was difficult, the patient could receive appropriate therapy by using liquid biopsy. Liquid biopsy could help in addressing the problem of poor prognosis in lymphangitic carcinomatosis patients.

## Bibliografía

1. Klimek M. Pulmonary lymphangitis carcinomatosis: systematic review and meta-analysis of case reports, 1970–2018. *Postgrad Med.* 2019;131:309–18.
2. Rolfo C, Mack PC, Scagliotti GV, Bass P, Barlesi F, Bivona TG, et al. Liquid biopsy for advanced non-small cell lung cancer (NSCLC): a statement paper from the IASLC. *J Thorac Oncol.* 2018;13:1248–68.
3. Leighl NB, Page RD, Raymond VM, Daniel DB, Divers SG, Reckamp KL, et al. Clinical utility of comprehensive cell-free DNA analysis to identify genomic biomarkers in patients with newly diagnosed metastatic non-small cell lung cancer. *Clin Cancer Res.* 2019;25:4691–700.
4. Dagogo-Jack I, Rooney M, Nagy RJ, Lin JJ, Chin E, Ferris LA, et al. Molecular analysis of plasma from patients with ROS1-positive NSCLC. *J Thorac Oncol.* 2019;14:816–24.
5. Supplee JG, Milan MSD, Lim LP, Potts KT, Sholl LM, Oxnard GR, et al. Sensitivity of next-generation sequencing assays detecting oncogenic fusions in plasma cell-free DNA. *Lung Cancer.* 2019;134:96–9.
6. Lam VK, Tran HT, Banks KC, Lanman RB, Rinsurongkawong W, Peled N, et al. Targeted tissue and cell-free tumor DNA sequencing of advanced lung squamous-cell carcinoma reveals clinically significant prevalence of actionable alterations. *Clin Lung Cancer.* 2019;20:30–6, e33.

Akihito Okazaki<sup>a,c,\*</sup>, Kazuhiko Shibata<sup>b</sup>,  
Yasuhiko Matsuda<sup>a</sup>, Yasutaka Shiba<sup>a</sup>, Keiichi Iwasa<sup>b</sup>,  
Kazuo Kasahara<sup>c</sup>

<sup>a</sup> Department of Respiratory Medicine, Koseiren Takaoka Hospital, 5-10, Eiraku-machi, Takaoka 933-8555, Japan

<sup>b</sup> Department of Medical Oncology, Koseiren Takaoka Hospital, 5-10, Eiraku-machi, Takaoka 933-8555, Japan

<sup>c</sup> Department of Respiratory Medicine, Kanazawa University Faculty of Medicine, Institute of Medical, Pharmaceutical and Health Sciences, 13-1, Takara-machi, Kanazawa 920-8641, Japan

\* Corresponding author.

E-mail address: [akihitookazaki1017@gmail.com](mailto:akihitookazaki1017@gmail.com) (A. Okazaki).

<https://doi.org/10.1016/j.arbres.2020.03.032>

0300-2896/ © 2020 SEPAR. Published by Elsevier España, S.L.U. All rights reserved.

## Osteomielitis costal que simula un empiema pleural en una enfermedad neumocócica invasiva



### Rib Osteomyelitis Mimicking a Pleural Empyema in a Patient with Invasive Pneumococcal Disease

Estimado Director:

La enfermedad neumocócica invasiva (ENI) representa una forma grave de infección por *Streptococcus pneumoniae*, y puede

producir un amplio abanico de manifestaciones clínicas que incluyen septicemia, meningitis, artritis, osteomielitis, celulitis, endocarditis<sup>1</sup>, etc. Se trata de un cuadro grave con una morbimortalidad significativa (especialmente en niños menores de 2 años y en adultos con condiciones médicas de riesgo y/o edad avanzada) y una incidencia variable, aunque se estima que en países desarrollados es de unos 10 casos por 100.000 personas/año en la población general<sup>2</sup>. Presentamos el caso de una paciente con ENI que presentó, tras un cuadro de infección respiratoria aguda no consolidante, varias complicaciones infecciosas torácicas y extra-torácicas.