

related with the following syndromes: progressive familial intrahepatic cholestasis type 3, low phospholipid-associated cholelithiasis (LPAC) syndrome, intrahepatic cholestasis of pregnancy, transient neonatal cholestasis, drug-induced liver disease, as well as fibrosis and cirrhosis in adults.<sup>5</sup>

Our patient met 2 of the following criteria for LPAC syndrome: (1) onset of symptoms before 40 years of age; (2) chronic mild cholestasis; (3) intrahepatic hyperechoic foci, biliary sludge or microlithiasis; (4) low biliary phospholipid concentration, and (5) recurrence of biliary symptoms after cholecystectomy.<sup>6,7</sup> Once the mutation had been confirmed, the presence of complicated biliary colic and chronic mild cholestasis led us to believe that this was a compound heterozygous patient, resulting in synthesis of immature or nonfunctional – but not truncated – MDR3 protein.

It is important to assess this possible aetiology in young patients who present repeated episodes of biliary colic for several reasons: first, because genetic tests are available that can confirm the diagnosis and avoid invasive tests with their associated risks; second, because it is an entity with a high response rate to UDCA,<sup>3</sup> an inexpensive, well-tolerated drug; third, because patients can present complications, such as complicated biliary colic or progressive fibrosis<sup>5</sup>; and finally, given its genetic nature, screening can be performed in first and second degree relatives who present compatible symptoms.<sup>2</sup>

## References

1. Eferink RP, Tytgat GN. Hepatic canalicular membrane 1: the role of MDR2 P-glycoprotein in hepatobiliary lipid transport. *FASEB J*. 1997;11:19–28.
2. Poupon R, Rosmorduc O, Boëlle PY, Chrétien Y, Corpechot C, Chazouillères O, et al. Genotype-phenotype relationships in the low-phospholipid-associated cholelithiasis syndrome: a study of 156 consecutive patients. *Hepatology*. 2013;58:1105–10.
3. Jacquemin E. Progressive familial intrahepatic cholestasis. *Clin Res Hepatol Gastroenterol*. 2012;36 Suppl 1:S26–35.
4. Strubbe B, Geerts A, van Vlierberghe H, Colle I. Progressive familial intrahepatic cholestasis and benign recurrent intrahepatic cholestasis: a review. *Acta Gastroenterol Belg*. 2012;75:75–80.
5. Davit-Spraul A, Gonzales E, Baussan C, Jacquemin E. The spectrum of liver diseases related to ABCB4 gene mutations: pathophysiology and clinical aspects. *Semin Liver Dis*. 2010;30:134–46.
6. Poupon R, Arrive L, Rosmorduc O. The cholangiographic features of severe forms of ABCB4/MDR3 deficiency-associated cholangiopathy in adults. *Gastroenterol Clin Biol*. 2010;34:380–7.
7. Wang D, Afdhal N. Sleisenger and Fordtran's Gastrointestinal Liver Disease. Feldman & Friedman & Brandt. Volume 2. Sección IX. Liver Chapter 65, p. 1100–1133.

Héctor Miguel Marcos Prieto, Daniel Pérez Corte, M. Concepción Piñero Pérez\*, María Cristina Revilla Morato, Ana María Mora Soler, Rosa Virginia Acosta Materán, Andrea Jiménez Jurado, Valeria Calabuig Mazzola, Antonio Rodríguez Pérez

*Servicio de Aparato Digestivo, Complejo Asistencial Universitario de Salamanca, Hospital Clínico, Salamanca, Spain*

\* Corresponding author.

*E-mail address:* [conxipi@hotmail.com](mailto:conxipi@hotmail.com) (M.C. Piñero Pérez).

## Metastatic intestinal obstruction secondary to a primary lung tumour<sup>☆</sup>



### Obstrucción intestinal por metástasis de cáncer de pulmón

Small bowel tumours are very rare, accounting for 1%–6% of all gastrointestinal (GI) cancers. Between 1% and 4% correspond to metastatic tumours, most commonly originating from malignant melanoma, lung cancer and colon cancer.<sup>1</sup>

Metastatic involvement of the small intestine is rare from a clinical point of view, but not so much from a histopathology perspective, with an incidence of 2%–14% according to autopsy series.<sup>2</sup>

The clinical manifestation of intestinal metastases is generally due to a complication of the disease, such as perforation, obstruction or active bleeding.<sup>3</sup> Presence of any of

these is associated with poorer prognosis of the underlying disease, and requires urgent surgical treatment.

We present 2 cases of patients with primary lung cancer who presented intestinal obstruction due to metastasis.

The first case, a 78-year-old man, smoker of 40 pack-years, was diagnosed with squamous cell lung cancer T2aNOM1 (hepatic) and treated with chemo- and radiotherapy. Nine months later, he presented to the emergency department for symptoms of bowel obstruction. Abdominal X-ray and computed tomography (CT) revealed dilatation of loops to the mid-jejunum with change in calibre secondary to adhesion or internal hernia. The patient underwent surgery, during which an obstruction was found in the jejunum as a result of a stenosing tumour. Complete resection and anastomosis were performed. Histopathology classified the tumour as squamous cell carcinoma metastasis. The patient was discharged on postoperative day 8. He continued with chemotherapy but died due to disease progression 20 months after the surgery.

The second case, a 49-year-old man, smoker of 20 pack-years, was seen in the emergency department for abdominal pain. Abdominal CT (Fig. 1) showed a mass in the right iliac fossa with signs of necrosis and regional lymphadenopathies. Colonoscopy was performed due to suspected caecal cancer, with no evidence of lesions. However,

<sup>☆</sup> Please cite this article as: Romera-Barba E, Castañer-Ramón-Llín J, Navarro-García I, Carrillo López MJ, Sánchez Pérez A, Vazquez-Rojas JL. Obstrucción intestinal por metástasis de cáncer de pulmón. *Gastroenterol Hepatol*. 2016;39:466–468.



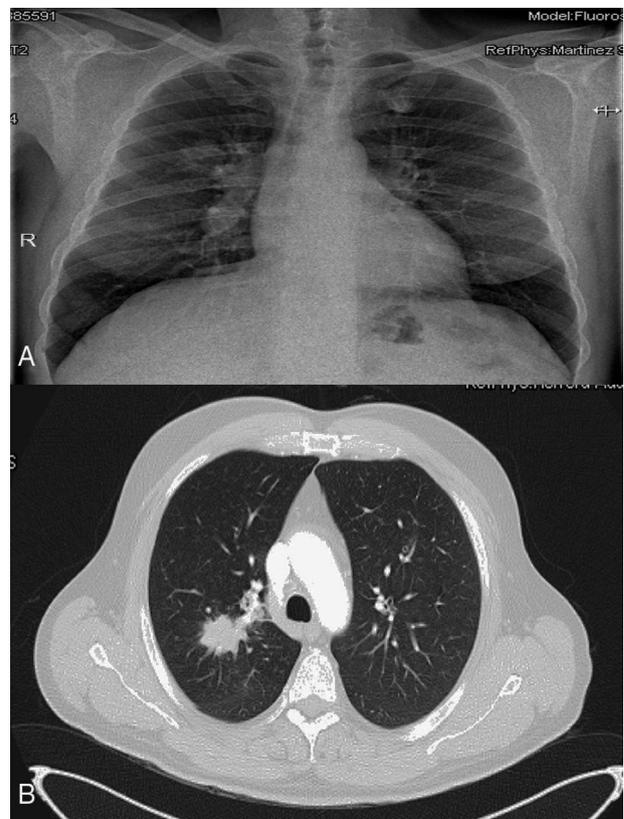
**Figure 1** Abdominal computed tomography, with oral and intravenous contrast. A large mass measuring  $10.2 \times 8.8$  cm can be seen in the right iliac fossa, affecting the terminal ileum and adjacent distal ileal loops; it also appears to encompass the appendix, with lateral displacement of the caecum. The inside of the mass shows fluid and oral contrast content, with air bubbles, probably due to necrosis and ulceration.

a preoperative chest X-ray identified a poorly defined nodular image in the right upper lobe (Fig. 2A). A chest CT scan and bronchoscopy performed to complete the study showed a mass in the right upper lobe, with right hilar lymphadenopathies (Fig. 2B). Biopsy of the pulmonary and abdominal lesions identified small cell lung cancer. The patient was diagnosed with stage IV small cell lung cancer and treated initially with cisplatin-etoposide chemotherapy. After receiving the first cycle, he required urgent surgery due to a bowel obstruction secondary to the metastasis in the right iliac fossa affecting the terminal ileum. Complete resection of the tumour and ileocolic anastomosis were performed. The patient was discharged on day 7 post-surgery and recommenced chemotherapy treatment. Despite completing 6 cycles, disease progression was detected on the follow-up CT scan. Now, 9 months after diagnosis, he is receiving second-line chemotherapy, with stable disease.

More than half of lung cancers present distant metastases at the time of diagnosis,<sup>4</sup> most often in the lymph nodes, liver, adrenal glands, brain and bone. The most common GI site for metastasis is the oesophagus, due to local infiltration. Intestinal metastases are rare, although more common than initially thought. They are most often located—in order of frequency—in the jejunum, ileum and duodenum.<sup>3</sup>

The most common histology of the primary lung tumour in our setting is adenocarcinoma and squamous cell carcinoma, in similar proportions (23.5% and 22.7%, respectively),<sup>1,3</sup> followed by large cell carcinoma (20.6%) and small cell carcinoma (19.6%).<sup>3</sup>

They are usually asymptomatic or generate non-specific symptoms such as anorexia and abdominal distension or pain, so clinical symptoms are sometimes difficult to distinguish from the effects of chemotherapy when the patient is already on treatment for their lung disease. Progression of the lesions leads to acute abdomen due to perforation, obstruction or haemorrhage.<sup>5</sup> Manifestation prior to diagnosis of the primary tumour is unusual,<sup>4,5</sup> although this



**Figure 2** (A) Chest X-ray: mass in right upper lobe. (B) Computed tomography with oral and intravenous contrast, where a poorly defined mass with spiculated margins can be seen (measuring  $50 \times 27$  mm in axial plane), extending from the right hilar region with obstruction of the right superior lobe bronchus and posterior segmental bronchus, associated with right hilar lymphadenopathies.

occurred in one of our cases where the lung cancer was diagnosed during the preoperative study of the abdominal mass, considered as the primary tumour.

Diagnosis is usually made during surgery. Imaging techniques such as oesophago-gastro-duodenal transit, endoscopic capsule or CT can aid in the preoperative diagnosis, although they have low sensitivity in the detection of small intestinal lesions. Systematic use of positron emission tomography-computed tomography (PET-CT) in lung cancer staging could help diagnose these small lesions, although there is still no conclusive evidence of its sensitivity. Definitive diagnosis is usually obtained by histopathology studies with the support of immunohistochemistry techniques for TTF-1, CDX2, CK7 and CK20, which can differentiate between primary and metastatic adenocarcinoma.<sup>5,6</sup>

The treatment of choice is resection of the affected area, with palliative intent. Intestinal metastases represent an advanced stage of the disease; they are generally associated with the presence of metastases in other locations, and indicate a poor prognosis, with maximum survival of 4 months.<sup>1,5</sup> However, in certain patient series (prior surgery of the primary tumour, single metastases) where the small bowel metastases have been resected, patients have presented longer survival (6–8 months); cases of prolonged

survival have also been reported, as happened in one of our patients.<sup>5,6</sup>

Advances in the systemic treatment of advanced stage lung cancer have managed to prolong survival in these patients, and consequently increase the likelihood of these types of complications. This heightens the importance of diagnosing intestinal metastases before their clinical expression, which could result in a relatively early diagnosis of the intestinal metastasis. Early surgical management of these lesions can help increase the survival of certain patients (low tumour burden, good general health) and prevent serious associated complications. The routine use of PET-CT in lung cancer staging will likely continue to increase the diagnosis of asymptomatic intestinal metastases, so the above should be taken into account in therapeutic decisions.

Thus, when non-specific bowel symptoms, anaemia or GI bleeding are found in a patient with a history of primary lung cancer, the possibility of intestinal metastatic disease should be considered as a possible cause. The appropriate diagnostic tests should then be performed, which can improve the prognosis by permitting complete resection of the lesion while the patient is in a better clinical condition, and preventing subsequent complications.<sup>7</sup>

### Conflict of interests

The authors declare that they have no conflict of interests.

### References

- Garrigos L, Iglesias M, Lloreta J, Jimeno J, Arriola E. Perforación intestinal secundaria a neoplasia primaria de pulmón. *Cir Esp*. 2010;87:257–69.
- Salemis NS, Nikou E, Liatsos C, Gakis C, Karagkiouzis G, Gourgiotis S. Small bowel perforation secondary to metastatic non-small cell lung cancer. A rare entity with a dismal prognosis. *J Gastrointest Cancer*. 2012;43:391–5.
- Garwood RA, Sawyer MD, Ledesma EJ, Foley E, Claridge JA. A case and review of bowel perforation secondary to metastatic lung cancer. *Am Surg*. 2005;71:110–6.
- Yang CJ, Hwang JJ, Kang WY, Chong IW, Wang TH, Sheu CC, et al. Gastrointestinal metastasis of primary lung carcinoma: clinical presentations and outcome. *Lung Cancer*. 2006;54: 319–23.
- Song Y, Li M, Shan J, Ye X, Tang S, Fang X, et al. Acute small bowel obstruction: a rare initial presentation for the metastasis of the large-cell carcinoma of the lung. *World J Surg Oncol*. 2012; 10:26.
- Rossi G, Marchioni A, Romagnani E, Bertolini F, Longo L, Cavazza A, et al. Primary lung cancer presenting with gastrointestinal tract involvement: clinicopathologic and immunohistochemical features in a series of 18 consecutive cases. *J Thorac Oncol*. 2007;2:115–20.
- Lee PC, Lo C, Lin MT, Liang JT, Lin BR. Role of surgical intervention in managing gastrointestinal metastases from lung cancer. *World J Gastroenterol*. 2011;17:4314–20.

Elena Romera-Barba\*, Juan Castañer-Ramón-Llín, Inmaculada Navarro-García, María Jose Carrillo López, Ainhoa Sánchez Pérez, José Luis Vazquez-Rojas

*Servicio de Cirugía General, Hospital General Universitario Santa Lucía, Cartagena, Murcia, Spain*

\* Corresponding author.

E-mail address: [percentila@hotmail.com](mailto:percentila@hotmail.com)

(E. Romera-Barba).

## Peripancreatic lymph node micrometastases from a mammary carcinoma in a patient with pancreatic ductal adenocarcinoma<sup>☆</sup>



### Micrometástasis en ganglios peripancreáticos por carcinoma mamario en paciente con adenocarcinoma ductal pancreático

Lymph node staging in pancreatic carcinomas is of great prognostic and therapeutic importance.<sup>1</sup> In the large majority of patients in whom peripancreatic lymph node metastases are detected, the primary origin of the tumor is the pancreas.<sup>1–6</sup> Nevertheless, in patients with a history of extra-pancreatic neoplasms, it is essential to rule out

the possibility of these pancreatic lymph node metastases originating from structures other than the pancreas.<sup>2–6</sup> We describe an unusual case of micrometastasis in peripancreatic lymph nodes due to breast cancer in a patient who underwent resection for pancreatic adenocarcinoma. The patient was a 72-year-old woman with a history of breast cancer in 2000, estrogen receptor (ER) positive and HER2 negative. The pathological stage was pT1b (8 mm) N1 (1/20) M0. Radio- and chemotherapy were prescribed, along with hormone therapy with tamoxifen. Some years later, in 2013, she underwent surgery for a right lung tumor (mixed acinar/lepidic adenocarcinoma, cytokeratin [CK]-7++, thyroid transcription factor [TTF]-1+++), pathological stage pT2aN2 (3/14) M0. In November 2014, a follow-up study with computed tomography (CT) and magnetic resonance cholangiography detected dilatation of the intra- and extrahepatic bile ducts with dilatation of the duct of Wirsung, with no clearly defined pancreatic lesion (Fig. 1A and B). A 20-mm lesion was found in the head of the pancreas on ultrasound endoscopy (Fig. 1C and D). Positron emission tomography (PET) was negative and CA19.9 values were elevated. Fine needle aspiration was positive for carcinoma. Cephalic duodenopancreatectomy was performed, and the macroscopic study showed a whitish, fibrous tumor in the head of the

<sup>☆</sup> Please cite this article as: Machado I, Caballero M, Martínez de Juan F, Marhuenda A, Martínez la Piedra C, Santos J, et al. Micrometástasis en ganglios peripancreáticos por carcinoma mamario en paciente con adenocarcinoma ductal pancreático. *Gastroenterol Hepatol*. 2016;39:468–471.