A rare case of recurrent acute pancreatitis after EUS-FNA in a patient with pancreatic intraepithelial neoplasia type 2

Un raro caso de pancreatitis aguda recurrente después de EUS-FNA en un paciente con neoplasia intraepitelial tipo 2

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ABSTRACT

Pancreatic intraepithelial neoplasia (PanIN) is considered a pre-malignant lesion difficult to identify by imaging methods. EUS-FNA is an effective technique to obtain material for histopathological study of pancreatic cystic tumors, but it is not free of adverse events. We report a case of a 56 years old patient, with chronic abdominal pain (early 1994). MRI showed pancreatic cystic images. The etiologic diagnosis was doubtful and EUS-FNA was performed. Immediately after a FNA, patient had an episode of acute pancreatitis, requiring hospitalization. During one year after FNA he had five episodes of AP. A new EUS suspected of PanIN, which was confirmed by surgery. After surgery the patient is well and has no more episodes of AP. Although the risk of AP, EUS-FNA should be performed to determine the best treatment for these patients with chronic abdominal pain who have cystic changes of pancreatic gland.

Keywords: Pancreatic ducts; Pancreatitis; Pancreatic neoplasms; Endosonography; Fine needle aspiration (source: MeSH NLM).

RESUMEN

La neoplasia intraepithelial pancreática (PanIN) es considerada una lesión premaligna con dificultad diagnostica mediante métodos imagenlógicos. La EUS-FNA es una técnica efectiva para obtener material para el estudio histopatológico de tumores quísticos pancreáticos, pero no está libre de efectos adversos. Nosotros reportamos un caso de un paciente de 56 años de edad, con dolor abdominal cronico (desde 1994). En la resonancia abdominal se observó un quiste pancreático. El diagnostico etiológico fue dudoso y EUS-FNA fue realizada, después de la cual el paciente presento un cuadro de pancreatitis, requiriendo hospitalización. Durante el periodo de un año posterior a la FNA, éste presento cinco episodios de pancreatitis aguda. Un nuevo estudio ecoendoscopico dio la sospecha de PanIN, la cual se confirmó con la realización de la cirugía. Después de la cirugía quedo asintomático y sin presentar nuevos episodios de pancreatitis aguda hasta el momento. Aunque exista riego de pancreatitis aguda, la EUS-FNA debe realizarse para determinar el mejor tratamiento para pacientes con dolor abdominal crónico que tienen cambios quísticos de la glándula pancreática.

Palabras clave: Conductos pancreáticos; Pancreáticos; Pancreático; Endosonografía; Biopsia con aguja fina (fuente: DeCS BIREME).

INTRODUCTION

Endosonography-guided fine needle aspiration (EUS-FNA) is effective technique to obtain material for pathological studies in patients with pancreatic cystic lesions ⁽¹⁾. Because low risk of developing acute pancreatitis (AP) after procedure, it has been used in place of brushing guided by endoscopic retrograde cholangiopancreatography (ERCP) ⁽²⁾. In addition, it provides superior results for etiological diagnosis of these tumors, compared to percutaneous biopsy techniques guided by abdominal ultrasound (US) or helical computerized tomography (CT), which gives effectiveness for its use ^(3,4). It is safe with a low rate of complications, ranging from 0.3 to 5% ⁽⁵⁾.

Perforation, hemorrhage, AP and a case of tumor cells seeding ⁽⁶⁾ at needle's route has been described after this technique. An important fact is that even with introduction of fine needle aspiration and guided biopsy technique; a study showed there is no statistical difference between diagnostic EUS and EUS-FNA related to episodes of AP after aspiration which is rare, up to 2% ⁽⁷⁾.

There are few publications and scarce data to demonstrate the occurrence of AP after EUS-FNA in pancreatic cystic lesions tumors (8-10). One study suggests that patients with a history of AP might be more likely to develop it (7). In any case the risk factors for the AP after EUS-FNA are unknown, and the frequency and severity of episodes vary

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Figure 1. MRI showed cystic images located in the body of the pancreas, the largest of them measuring 1.2 cm, accompanied by other smaller cystic images.

from one center to another and must be related to practitioner's experience (11,12).

We describe a case of pancreatic intraepithelial neoplasia (PanIN) with an recurrent episodes of AP after EUS-FNA. Based on scientific evidence the authors discuss its occurrence and risk factors for AP.

CASE REPORT

Caucasian male patient, 56 years old, complained of abdominal pain since 1994. Reported colonic diverticular disease for 15 years and nephrolithiasis. He smoked for over 20 years and said to be a social drinker. In 2001 presented high intensity abdominal pain and underwent CT which revealed hypodense areas in the body of pancreas. Medical treatment was performed with improvement in clinical status. Two years later, he had a new episode similar to previous. Another abdominal CT scan showed diverticulitis, swelling and changes in fat surrounding pancreatic tail. Magnetic resonance imaging of abdomen (MRI) revealed small cystic images in the body, along with dilation and tortuosity of main pancreatic duct (MPD) and branch duct (Figure 1). EUS-FNA was indicated to evaluate

the cystic lesion and obtain material for pathological examination.

In the body we saw MPD dilation (33 mm). We noted numerous cystic areas, communicating with MPD, and the largest one reached 17 x 8 mm (Figure 2a and 2b). The lesion together (macrocystic and microcystic areas) had 26 mm. The diagnosis of EUS associated with the appearance of fluid aspirated reinforced the suspicion of branch-duct intraductal papillary mucinous tumor of the pancreas. The pathological examination was negative for malignancy, with rare leukocytes, scanty degenerate red blood cells and amorphous proteinaceous material.

Two hours after procedure, patient had intensive abdominal cramps in mesogastrium, radiating to the back, accompanied by nausea and vomiting refractory to analgesics and antispasmodics. Deep abdominal palpation revealed rigidity and severe pain in mesogastrium. Immediately, he was admitted and laboratory tests revealed: high levels of amylase (3x normal) and lipase (3x normal). Plain X-ray in upright position and both hemidiaphragm were normal. The clinical picture associated with laboratory tests showed AP after EUS-FNA. A symptomatic medical treatment was imposed along with antibiotics. Patient remained under medical care until painful symptoms disappear, and was discharged in good condition after 5 days of stay. In July of the same year he presented abdominal pain radiation to the back and located in LIF. CT showed increased pancreatic volume with small hypodense lesions without contrast enhancing. The largest one measured 18 mm. Amylase and lipase were normal. CEA- 1.62 mg / ml and CA 19-9- 12 U / ml, also were normal.

One year after presented four episodes of abdominal pain accompanied by increased amylase and lipase (2x normal). This situation led the doctor to request a new EUS. The examination revealed a cyst 20 mm length



Figure 2. Endoscopic ultrasound image (a and b). Note the small cystic areas, located in the pancreatic body. See anechoic area without wall, appearing to be secondary to pancreatic duct dilation. Note (yellow arrow) communicating with the MPD (red arrow).



Figure 3. EUS showed a macrocystic area measuring 20 X 11 mm (a) as a component of the entire lesion which measures 40 X 28 mm in major axes (b).

and other smaller cysts. It measured 40 x 28 mm in its major axes, with dilatation of secondary ducts (Figure 3a and 3b). The other regions were normal.

Based on EUS results patient was referred for surgery, undergoing subtotal pancreatectomy without complications and with good outcome. Macroscopic analysis of surgical specimen showed a segment of pancreatic body measuring 90 x 50 x 15 mm, with a nodular, soft and slightly purplish and bosselated lesion measuring 35 x 15 x 15 mm. This lesion was adjacent to MPD, which was dilated. Final diagnosis was pancreatic intraepithelial neoplasia (PanIN type II) with ductal adenomatous papillary hyperplasia, without atypical changes. Patient had good outcome and now, after 12 years is doing well.

DISCUSSION

EUS-FNA is being increasingly used for diagnosis of different pancreatic cystic lesions (3). The risks associated with this procedure are low (5,8). The technique is safe with low rate of complications, ranging from 0.3% to 5% (11). These complication rates are equivalent to those of upper digestive endoscopy (13) and colonoscopy (14). Perforation, hemorrhage, and AP are described as well one case of tumor cells dissemination after EUS-FNA for pancreatic tumor (6). O'Toole et al. (15) evaluated the complications of EUS-FNA in 322 patients. After 345 punctures they observed overall complication rate in 1.6%, without any case of serious complication or death.

In 3324 patients underwent radial EUS and EUS-FNA, 10 patients (0.3%) had complications related to the procedure and death in 0.06% of cases according Mortensen et al. (16). There was no statistical difference between diagnostic EUS and EUS-FNA (16).

The risk of iatrogenic AP caused by FNA ranges from 0 to 2% and increases in patients with pancreatic masses, cysts or when FNA is performed through MPD, because all these procedures consist of passing the needle directly through a normal pancreatic parenchyma (5,7-^{9,11)}. Perhaps this was the major problem, in this case, because to reach its destination the needle had to cross a reasonable amount of normal pancreatic tissue and this was done twice.

O'Toole et al reported 4 complications (1.2%) after aspiration of pancreatic cystic tumors (3 cases of AP and 1 aspiration pneumonia). The AP in this series was the result of FNA in lesions of the head or uncinate process, unlike our case, with FNA performed in the body of the pancreas. Significantly in this study, the number of punctures was not important to predict the occurrence of complications (15). Eloubeidi et al. (9) reported 1 case (0.6%) of moderate intensity AP in 158 patients underwent EUS-FNA for solid tumors of the pancreas, and the other autor reported a fatal consequence of the EUS-FNA ina a patient with pancreatic cystic lesion (12).

Other studies suggest that episodes of AP after EUS-FNA occur more frequently in patients with a history of AP (7). Some relate its occurrence in patients with benign diseases or pseudotumors (17,18). The fact is that our patient had never had AP prior to EUS-FNA presented five episodes of AP after this procedure in 1 ½ year period, despite having a benign lesion.

In several reports of fatal AP episodes after percutaneous FNA of pancreas no cancer or mass was found during autopsy (17), which ratifies the one found in our case. Unlike, recent studies demonstrate a mass or pancreatic cancer in patients with AP after EUS-FNA (6,7-9,11).

Another factor that seems important and has been discussed in the literature deals with experience of the operator, because AP was found when about 79% of echo-endoscopists had performed at least 100 FNA for pancreatic masses (11). This is not the case, because the operator (JCA), at time of this procedure, had already performed more than 600 EUS-FNA for pancreatic tumors (3).

It is not possible to determine with certainty if smaller lesions are more likely to present episodes of AP. However a study with percutaneous FNA revealed the frequency of AP is higher in lesions smaller than 30 mm compared with those larger than 45 mm (17). In our case the lesion was much smaller than 30 mm, measuring 17 mm. So perhaps this was an important factor for the occurrence of AP. Our experience confirms these data. The frequency of AP after EUS-FNA appears to be lower than in cases submitted to percutaneous FNA (11). In a retrospective study with percutaneous FNA, severe episodes of AP occurred in 3% of patients (19). All patients who developed AP related to the procedure had lesions smaller than 30 mm (19).

It is mere speculation the mechanism by which episodes of AP occurs after FNA. Perhaps is related to pancreatic juice leakage from MPD or its branch ducts during FNA (11). Perhaps MPD or branch ducts disruption caused by needle, may be responsible for AP. Anyway, in this case and in all where the lesions are smaller than 30 mm, we always take care to avoid introduce the needle into MPD, a factor we consider important in genesis of AP episodes.

Gress *et al.* ⁽⁷⁾ prospectively evaluated the frequency of AP after pancreas EUS-FNA. In this study, the levels of blood amylase and lipase were measured before and 2 hours after procedure. There was significant elevation of amylase and / or lipase in 5%, but only 2% of patients showed clinical signs of AP. Interestingly the main reason for EUS-FNA be performed for these two developed AP, was unexplained recurrent AP episode ⁽⁷⁾.

In short, it is pointed out that pancreatic EUS-FNA is safe, effective and has a low AP occurrence rate in experienced hands. On the other hand, if a patient has a history of recurrent AP, normal pancreatic parenchyma between the wall of the digestive system and the lesion to be punctured, a benign lesion less than 30 mm or MPD transfixed by inadvertently puncture, are supposedly causal factors of AP, as no clear risk factor was identified. Surely the vast majority of retrospective studies demonstrating AP after EUS-FNA are underestimated.

Conflict of interest: The authors have no potential conflicts of interest

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