A B S T R A C T

The authors address the value of endoscopic retrograde cholangiopancreatography, ultrasonography, computed tomography, magnetic resonance imaging and endoscopic ultrasound for the diagnosis of cystic and solid neoplasms of the pancreas, demonstrating that each of them is of great importance to undoubtedly increase the diagnostic accuracy of the biliopancreatic system diseases. The best method for each of several tumors is then determined.

Key words: Diagnostic imaging. Neoplasms. Neoplasms, cystic, mucinous, and serous. Pancreas.

INTRODUCTION

In the 1960s the biliopancreatic system was considered a restricted area for an approach with diagnostic imaging methods. In the early 1970s the Endoscopic Retrograde Cholangiopancreatography (ERCP) was introduced and the accurate diagnosis, etiology and location began to be determined with a high level of accuracy1.

Abdominal ultrasound (US), introduced in the late 1970s, increased identification of diseases that affected the biliopancreatic tree in a less invasive way2. With the use of in real time, sectorial and high resolution systems for US, the gallbladder, the extrahepatic biliary tree and the pancreatic gland could be better studied3.

Also in the 1970s computed tomography (CT) was introduced in daily clinical practice and revolutionized medical diagnosis by images. The rapid acceptance of the method and its real diagnostic capacity gave the Nobel Prize for medicine to its creators, Godfrey Hounsfield and Alan Cormack4.

The principle of magnetic resonance imaging (MRI) is known since the 1940s, but only in the 1970s the first medical images were obtained through MRI5.

As if all these were not enough, the endoscopic ultrasound experimentally begun almost at the same time. Also known as endosonography or ecoendoscopy (EE), it is a technique that allows the placement of a transducer ultrasound in all points accessible to classic endoscopy. Thus, organs and regions less accessible to other methods of image, such as the pancreas, the distal portion of the biliary tree and the papilla are studied with unparalleled accuracy6.

These methods have undergone modifications over time and their high problem-solving capacity has shown that each of them has a fundamental role to undoubtedly increase the diagnostic accuracy of the biliopancreatic system’s diseases.

The purpose of this revision is to clearly and succinctly demonstrate the actual role of each of these exams for each of the various cystic and solid tumors of the pancreas, comparing them and determining which method is best for each of the several lesions.

This article will be presented in two parts: the first will address the methods of image in cystic Neoplasms; the second will address the study of methods of image in solid Neoplasms of the pancreas.

Cystic Neoplasms
Pancreatic cystic neoplasms are rare tumors, contributing with only 10% to 15% of all pancreatic cysts and 1% of pancreatic cancers. These tumors were classified as cystic neoplasms, composed by mucinous cystadenoma, cystadenocarcinoma, mucinous papillary intraductal neoplasm and, finally, the microcystic adenoma, also known as serous cystadenoma.
These clinical subtypes are very important, since the microcystic adenomas are benign and asymptomatic and therefore do not require any kind of treatment. While all the others are presently considered to be premalignant lesions of the pancreatic carcinoma.

**Mucinous Cystic Neoplasm**

All mucinous cystic neoplasms should be treated surgically, for it is believed that the mucinous cystadenomas coexist with mucinous cystadenocarcinomas or evolve into them.

**Mucinous Cystadenoma**

With characteristically macrocystic morphology, they are unilocular cystic lesions, with well differentiated walls from the rest of the pancreatic parenchyma and may be divided into multiple compartments by thin septa with or without a thick content that corresponds to mucin. The presence of localized thickening, irregularity of the wall or solid component suggests malignant degeneration to cystadenocarcinoma. At US and EE these cystic neoplasms seem similar to pseudocysts. They contain cysts with diameters larger than 2 cm, anechoic, with posterior enhancement and internal septations. The septa are thin and, to the extent that gain is increased, the cystic areas get filled with echoes. In 10 to 18%, calcifications are present in the wall and are seen as echogenic areas with shadows. These cysts may not be defined as benign or malignant.

CT (Figures 1A and 1B) and MRI (Figure 2) have important roles in the diagnosis of this type of lesion, as well as its differentiation from pseudocysts. The differentiation is also based on clinical data, because the image aspects are sometimes superimposable. Many authors believe that if there is no history of acute pancreatitis with possibility of evolution to a pseudocyst, one must consider the a cystic lesion as potentially neoplastic and approach it surgically. Evaluation with these exams also includes staging of the lesion, searching for hepatic metastasis and peritoneal implants. Despite their current skills, the complete clinical characterization of such a lesion, when incidentally found, is around 25% 30%.

The fine needle aspiration (FNA) guided by EE of this neoplasia is relatively easy and the cytological analysis of the aspirated liquid may show the presence of columnar epithelial cells (benign or malignant) and mucin. The most serious complication of a pancreatic cyst puncture is contamination and abscess formation, which may be avoided by completely emptying the cyst and with antibiotic prophylaxis (Figure 3).

**Cystadenocarcinoma**

Some researchers identified four sonographic patterns associated with this type of cyst: 1) anechoic mass with posterior enhancement and irregular margins; 2) anechoic mass with inner homogeneous echoes that

Figure 1 - a) tomographic aspect of a mucinous cystadenoma, with thickening and irregularity in the cyst wall, b) thin, well-shaped septa forming part of the radiological picture of a mucinous cystadenoma.

Figure 2 - MRI of a mucinous cystadenoma.
are laminated in supine position and moving with lateral decubitus; 3) anechoic mass, with internal regular vegetations that protrude into the lumen without showing movements; and 4) completely echogenic and heterogeneous mass\textsuperscript{12}. The modern equipment of CT (Figure 4) and MRI with associated Magnetic Resonance Cholangiopancreatography (MRCP) (Figure 5) provide detailed information of pancreatic cysts, like: presence of septa, size, location and communication with the main pancreatic duct (MPD). In rare cases it may provide data on the presence of nodules or vegetation in the interior of the cysts. The critical analysis of these factors is important to differentiate a mucinous adenoma (MAC) cyst from a mucinous papillary intraductal neoplasia (MPIN)\textsuperscript{13,14}.

EE-FNA is relatively simple. With a single needle passage, the tip is placed in the center of the cyst and all content is aspirated. Once aspirated between 2-3 ml of clear liquid, the tip of the needle may be advanced through the wall or through a thin septum. This technique improves liquid outflow (Figure 6). Eventually, the viscosity of the liquid may impede the aspiration of its contents. The use of a needle of large caliber associated with prolonged aspiration usually provides enough material for analysis. Occasionally, bleeding may occur at the location of the puncture and this rarely happens inside the cyst. Should this happen, it tends to stop when the cyst fills with blood\textsuperscript{15}.

**Mucinous Papillary Intraductal Neoplasia**

The MPIN consists not only of a distended MPD, but also peripheral cystic lesions, nodules or vegetations, with mass effect. It is difficult to differentiate the MPIN from chronic pancreatitis (PC) based on the dilatation of the MPD. If there are not parenchymal lesions that suggest chronic pancreatitis, the MPD should be aspirated. Sometimes a focal lesion adhered to the wall may be seen in the MPD of patients with MPIN. These nodules, when carefully studied and aspirated, may render diagnosis of focal malignancy. Cystic lesions associated to MPIN have a wide range of presentations and may simulate serious cystadenomas (SCA) or microcysts. The large unilocular cystadenomas are commonly found in advanced cases of MPIN and must be aspirated because of the possibility of an early stage malignant tumor. The lesions that have mass effect look similar to pancreatic adenocarcinoma and must be aspirated to the diagnosis of malignancy or of a focal chronic pancreatitis nodule.

At US these lesions are difficult to distinguish. Radiological methods demonstrate focal or diffuse ductal dilatation and sometimes filling failure images may be observed within the MPD, corresponding to the tumor or impacted mucin. As these tumors are small, the predominant radiological aspect may be similar to that observed in chronic pancreatitis with ductal dilatation and advanced atrophy. It is important to highlight that parenchymatous calcifications are usually not seen in cases of MPIN. They may occur only in extremely advanced cases (Figure 7).

Although MRCP may demonstrate ductal changes, endoscopic pancreatography is vital to the diagnosis, confirming the filling failure images, identifying the output of mucoid secretion through the duodenal papilla and enabling the access to such material and its cytological and histopathological exams (figures 8a and 8b). Cytology of material aspirated from the distended MPD or from a MPIN-associated cyst demonstrates an aspect similar to MAC, with malignant or benign columnar epithelial cells, generally associated with large amounts of mucin\textsuperscript{16,17}.

**Cystic Serous Neoplasia**

Serous cystadenomas usually display a microcystic component, with a typical “honeycomb”
endosonographic aspect, although they may also be macrocystic and unilocular, which may display area of fibrosis or a central scar. These injuries are potentially benign and their echotextural characteristics are very suggestive of this tumor type\textsuperscript{18}.

In this disease the US very accurately identifies a nodular area with multiple microcysts. Abundant fibrous stroma and small dimensions of cysts may generate a radiological aspect of a solid neoplasm. However, the careful review of all phases of the exam generally allows the identification of small cysts. The hypervascularity of numerous fibrous septa causes an intense early enhancement after intravenous administration of contrast medium at CT and MRI (Figure 9). The ERCP has no diagnostic function in this type of cystic neoplasm\textsuperscript{13,14}.

**Pseudopapillary Solid-Cystic Epithelial Neoplasia (Frantz's tumor)**

This is a pseudopapillary solid-cystic epithelial neoplasia located in the tail of the pancreas, which is often seen in young women (average age 24 years). It is an unusual, low-grade malignant tumor, curable by surgical removal. Usually the patient has no symptoms until the tumor is large. Often multiple areas of hemorrhage and cystic degeneration are found within the tumor, but mitotic figures are rarely seen. After surgical resection, prognosis is remarkably good, although duodenal and liver invasion and distant metastasis have been reported\textsuperscript{19,20}. The lesion may appear similar to the cystadenoma or cystadenocarcinoma at EE and US, as there are septa in the cystic parts, possibly due to prominent papillae projecting into the space of cystic degeneration. At US, CT and MRI a solid-cystic mass may appear. Calcifications inside the mass are not common. Although
the echographic features are indistinguishable from those of cystadenoma and cystadenocarcinoma, the asymptomatic occurrence in young women, associated with a hemorrhagic component, must lead to suspicion of such a tumor\textsuperscript{19,20}.

The radiological presentation of the solid-cystic epithelial neoplasia is dependent on the relationship between the solid and cystic components, with some cases being seen as predominantly solid, well demarcated masses and others with central areas of liquefaction (Figure 10). These, in turn, often present with hemorrhage, which may show increase of attenuation of the cystic content at CT or, more easily, the hyperintense signal in T1 and hypointense in at MRI; some of these tumors may present calcification in the periphery. EE-FNA of this type of injury has high accuracy and sensitivity\textsuperscript{21}.

\textbf{Figure 8} - \textit{a)} aspect of “fish eye” at duodenoscopy examination \textit{b)} radiological imaging during ERCP, where filling failures throughout the MPD and the cephalic portion of the secondary ducts can be seen. This aspect is of mixed type NIMP (III) (images courtesy of Prof. Dr. José Sebastião dos Santos HCFMRP-USP).

\textbf{Figure 9} - Tomographic image of a 72-year-old female patient with incidental finding of a solid-cystic area on the tail of the pancreas. EE-FNA revealed a serous microcystic cystadenoma, which was confirmed by surgery.

\textbf{Figure 10} - \textit{a)} CT image in a patient aged 12 who suffered a bike fall and had abdominal pain, \textit{b)} operative specimen of Frantz’s tumor (images courtesy of Prof. Dr. José Sebastião dos Santos HCFMRP-USP).
RESUMO

Os autores fazem uma revisão considerando o valor da colangiopancreatografia endoscópica retrógrada, da ultrassonografia, da tomografia computadorizada, da ressonância magnética e da ecoendoscopia para o diagnóstico das neoplasias císticas e sólidas do pâncreas, demonstrando que cada um deles tem grande importância para aumentar, de forma incontestável, a acurácia diagnóstica das doenças do sistema biliopancreático, determinando qual o melhor método para cada um dos diversos tumores.


REFERENCES


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