


Application of confocal endomicroscopy in the diagnostic elucidation of pancreatic cyst

 Rodrigo C. Scomparin¹
Luiza H. Bento¹
Clelma P. Batista¹
Bruno C. Martins¹
Luciano Lenz¹
Elisa R. Baba¹
Adriana Vaz Safatle-Ribeiro¹
Fauze Maluf-Filho¹

1. Cancer Institute of the State of São Paulo, São Paulo/SP, Brasil

<http://dx.doi.org/10.1590/1806-9282.64.10.885>

KEYWORDS: *Pancreatic cyst. Pancreatic neoplasms/diagnostics. Echo-endoscopy/methods. Confocal Microscopy/methods*

INTRODUCTION

Treatment of patients with pancreatic cysts is challenging and follow-up remains controversial. Some guidelines allow the standardization of the treatment of these patients, especially in cases of intraductal papillary mucinous neoplasia (IPMN).

New technologies have been developed and applied for diagnostic assistance and therapeutic management. Confocal laser endomicroscopy (CLE) represents a technique in which a probe is used, promoting the *in-vivo* microscopic image in real time of the tissue studied¹. The CLE technique through EUS-guided needle aspiration CLE – nCLE has been shown to be promising, especially in IPMN cases.

The purpose of this study is to report the applica-

tion of nCLE technology in a patient with pancreatic cyst, already with a previous diagnosis of IPMN at follow-up.

CLINICAL CASE

Patient, female, 74 years old, married, from São Paulo, hypertensive, diabetic, on atenolol, enalapril and glycated. She presented with nonspecific abdominal pain and cystic lesion in the pancreas, with communication with the main pancreatic duct, compatible with IPMN and in follow-up since 2011. Until 2016, she underwent an annual follow-up with magnetic resonance imaging showing cystic lesion with localized lobulated contours in the pancreatic body,

DATE OF SUBMISSION: 27-Jan-2018
DATE OF ACCEPTANCE: 16-Feb-2018
CORRESPONDING AUTHOR: Rodrigo Corsato Scomparin
Instituto do Câncer do estado de São Paulo
Av. Dr. Arnaldo, 251 - Cerqueira César, São Paulo - SP, 01246-000
E-mail: r.scomparin@outlook.com

measuring approximately 27 mm, in communication with the main pancreatic duct, which had a gauge of 23 mm. In 2017, there was a slight increase in its dimensions to 30 mm (Figure 1). Echoendoscopic evaluation with echo-guided puncture and nCLE were performed in search of signs of malignancy.

In September 2017, echoendoscopy showed an anechoic image of 30 mm x 16 mm, oval, with regular contours, precise limits, without nodules or solid components in the interior, with posterior acoustic reinforcement, in communication with the main pancreatic duct. An echo-guided puncture with 19G needle was performed and nCLE was carried out, observing the coffee beans aspect (pancreas), digitiform projections, and no signs of malignancy were found (Figures 2 and 3). The aspect was suggestive of IPMN of the intestinal type. At the end of the examination, the cyst was completely emptied, recovering 3 ml of fluid and translucent liquid. An antibiotic prophylaxis was performed for seven days with ciprofloxacin. The dosage of markers was amylase 111770 U/L, lipase 482960 U/L, CEA 8.3 ng/mL and CA 19.9 58.3 ng/mL. Cytological analysis was negative for neoplasia.

There were no complications related to the procedure. The multidisciplinary team decided on clinical follow-up.

DISCUSSION

In recent years, the incidental finding of pancreatic cysts in imaging studies has been increasing, probably due to equipment sophistication, the development of new technologies and a larger number of imaging studies carried out^{2,3}. The prevalence of pancreatic cystic lesions in imaging studies may vary between 3% and 19%⁴⁻⁶. At autopsies, the number is even higher, 24%²⁻⁵. Pancreatic cysts may be congenital, inflammatory or neoplastic. It is estimated that

less than half of the cases are of intra-ductal papillary mucinous neoplasia (IPMN), of which only a small share will develop into invasive carcinoma^{2,4}.

Several tests can be used to detect and evaluate pancreatic cysts, such as abdominal ultrasonography, computed tomography, nuclear magnetic resonance and echoendoscopy⁷. New technologies have been developed, such as CLE, promoting the microscopic image *in vivo* and in real time, with magnification in about a thousand times¹. The CLE technique with a probe inserted into the EUS-guided needle-based CLE (nCLE) is promising, especially in IPMN cases.

In a multicentre study¹, the diagnostic capacity of nCLE was evaluated in 29 cases of pancreatic cystic lesions, with clinical information omitted. The rates of sensitivity, specificity, accuracy and intra-observer and inter-observer agreement were evaluated. The results for cystic lesions were 95%, 94% and 95%, respectively, with Kappa agreement of 0.81 for intra-observer and 0.86 for inter-observer. The authors considered the agreement in both cases to be almost perfect. The results of this study demonstrate that the nCLE technique in cases of pancreatic cystic lesions can be very useful in the conduct and follow-up, as was the case of the patient reported.

Some guidelines have attempted to standardize the treatment of patients with IPMN, such as the one published in 2012 by the International Association of Pancreatology⁸. According to it, patients with secondary ductus IPMN with no worrisome features can be followed. Through nCLE, it was possible to infer the histological type of IPMN, that is, intestinal⁹. This histological type generally presents a more favourable evolution when compared to the oncocytic, pancreatobiliary and gastric types⁸.

In a Swedish study¹⁰, published in 2017, the survival and the risk of progression of IPMN under surveillance were analysed in a group of 395 patients

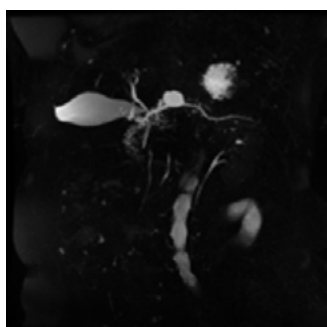


FIGURE 1



FIGURE 2

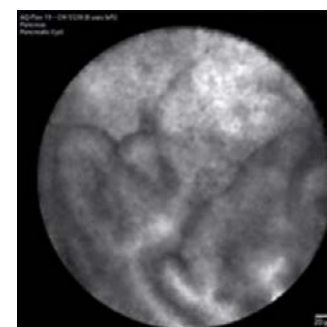


FIGURE 3

between January 2008 and December 2013. The authors found that the main pancreatic duct with ≥ 6 mm and cephalic location, including the uncinate process, are associated with an increased risk of progression, especially after five years of diagnosis. According to the guideline of the International Association of Pancreatology⁸, there is no evidence to support that surveillance can be spaced or discontinued even in the face of morphological stability of the lesion. The patient in question maintained morphological stability for five years. The current recommendation is that surveillance should be continued at shorter intervals, especially in the period after five years, which is known when the risk of progression

increases. However, the routine use of nCLE may suggest a new guidance regarding the follow-up interval. Prospective studies with adequate casuistry can prove this assertion.

CONCLUSION

Treatment of patients with incidental pancreatic cysts is based on clinical, radiological and biochemical findings. With the advent of echoendoscopy and nCLE, it became possible to aggregate information on the microscopic characteristic of cyst lining, which may be useful in the management and therapeutic strategy of this group of patients.

PALAVRAS-CHAVE: *Cisto pancreático. Pancreatic Neoplasms/diagnóstico. Endossonografia/métodos. Microscopia confocal/métodos.*

REFERENCES

1. Krishna SG, Brugge WR, Dewitt JM, Kongkam P, Napoleon B, Robles-Medrandia C, et al. Needle-based confocal laser endomicroscopy for the diagnosis of pancreatic cystic lesions: an international external interobserver and intraobserver study (with videos). *Gastrointest Endosc.* 2017;86(4):644-54.
2. Laffan TA, Horton KM, Klein AP, Berlanstein B, Siegelman SS, Kawamoto S, et al. Prevalence of unsuspected pancreatic cysts on MDCT. *AJR Am J Roentgenol.* 2008;191(3):802-7.
3. Jong K, Nio CY, Hermans JJ, Dijkgraaf MG, Gouma DJ, van Eijck CH, et al. High prevalence of pancreatic cysts detected by screening magnetic resonance imaging examinations. *Clin Gastroenterol Hepatol.* 2010;8(9):806-11.
4. Zhang XM, Mitchell DG, Dohke M, Holland GA, Parker L. Pancreatic cysts: depiction on single-shot fast spin-echo MR images. *Radiology.* 2002;223(2):547-53.
5. Kimura W, Nagai H, Kuroda A, Muto T, Esaki Y. Analysis of small cystic lesions of the pancreas. *Int J Pancreatol.* 1995;18(3):197-206.
6. Canto MI, Hruban RH, Fishman EK, Kamel IR, Schulick R, Zhang Z, et al. Frequent detection of pancreatic lesions in asymptomatic high-risk individuals. *Gastroenterology.* 2012;142(4):796-804.
7. Del Chiaro M, Segersvärd R, Lohr M, Verbeke C. Early detection and prevention of pancreatic cancer: is it possible today? *World J Gastroenterol.* 2014;20(34):12118-31.
8. Tanaka M, Fernández-del Castillo C, Adsay V, Chari S, Falconi M, Jang JY, et al; International Association of Pancreatology. International consensus guidelines 2012 for the management of IPMN and MCN of the pancreas. *Pancreatol.* 2012;12(3):183-97.
9. Giovannini M, Caillol F, Lemaistre A, Monges G, Napoleon B, Pujol B. Endoscopic ultrasound guided confocal microscopy: atlas of cystic pancreatic lesions. *Endosc Ultrasound.* 2014;3(Suppl 1):S19-21.
10. Del Chiaro M, Ateeb Z, Hansson MR, Rangelova E, Segersvärd R, Kartalis N, et al. Survival analysis and risk for progression of intraductal papillary mucinous neoplasia of the pancreas (IPMN) under surveillance: a single-institution experience. *Ann Surg Oncol.* 2017;24(4):1120-6.

