

Role of contrast harmonic endoscopic ultrasonography in other pancreatic solid lesions: Neuroendocrine tumors, autoimmune pancreatitis and metastases

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ABSTRACT

Contrast harmonic endoscopic ultrasonography (CH-EUS) is a new technique which allows the dynamic study of the microvascularization of a target tissue. Its application is validated for the diagnosis of pancreatic adenocarcinoma but remains unclear for other solid pancreatic tumors (neuroendocrine tumors [NETs], autoimmune pancreatitis [AIP], metastases). The purpose of this review is to outline the potential role of the CH-EUS in these indications. NETs are typically iso/hyperenhanced at CH-EUS, and a heterogeneous enhancement seems a good predictor of malignancy in neuroendocrine pancreatic tumor. AIP is often iso/hyperenhanced at CH-EUS. Quantitative analysis of time-intensity parameters is promising for the distinction between pancreatic adenocarcinoma and AIP. The appearance of pancreatic metastases at CH-EUS is various depending on the origin of the primary tumor. Data from the literature remain to this day weak to determine the role of the CH-EUS in the management of rare solid tumor of the pancreas (NETs, AIP, and metastases). Specific studies are expected to further clarify the impact of this procedure in this field.

Key words: Autoimmune pancreatitis, contrast harmonic endoscopic ultrasonography, neuroendocrine tumor, pancreatic metastasis

INTRODUCTION

Contrast harmonic endoscopic ultrasonography (CH-EUS) is a new technique which allows the dynamic study of the microvascularization of a target tissue. Its application in the characterization of solid tumors of the pancreas is validated for the diagnosis of pancreatic adenocarcinoma.

Its role is less clear for other solid pancreatic tumors (neuroendocrine tumors [NETs], autoimmune pancreatitis [AIP], metastases).

The purpose of this review is to outline the potential role of the CH-EUS in these indications.

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CONTRAST HARMONIC ENDOSCOPIC ULTRASONOGRAPHY IN PANCREATIC NEUROENDOCRINE TUMORS

Positive diagnosis

Pancreatic NETs (PNETs) represent between 5% and 10% of all pancreatic solid tumors. They are typically richly vascularized and have an early arterial enhancement in cross-sectional imaging. This behavior can be demonstrated in CH-EUS and enables a differential diagnosis with other pancreatic solid masses [Figure 1]. However, no study has specifically addressed PNETs up to now. In the study from Kitano *et al.*,^[1] 95% of PNETs were iso/hyperenhanced ($n = 18/19$). In work from Gincul *et al.*,^[2] 100% were iso/hyperenhanced ($n = 10/10$). In another work from Yamashita *et al.*,^[3] 100% ($n = 8/8$) were iso/hyperenhanced at early arterial phase.

Prediction of malignancy

One study^[4] assessed the value of Doppler-EUS sensitized with a second-generation ultrasound contrast agent injection in predicting malignancy of PNETs. Forty-one tumors were evaluated. Heterogeneity after injection of contrast had a diagnostic accuracy of 90.2% with a sensitivity of 90.5% and a specificity of 90%. A study submitted as an abstract at DDW 2015 assessing 92 PNETs had promising results with diagnostic values >90% for the prediction of malignancy in cases of a heterogeneous enhancement [Figures 2-4].

Detection of small neuroendocrine tumors

There is currently no data to conclude about the potential diagnostic value of CH-EUS in detecting some small PNETs such as insulinomas in comparison to conventional EUS.

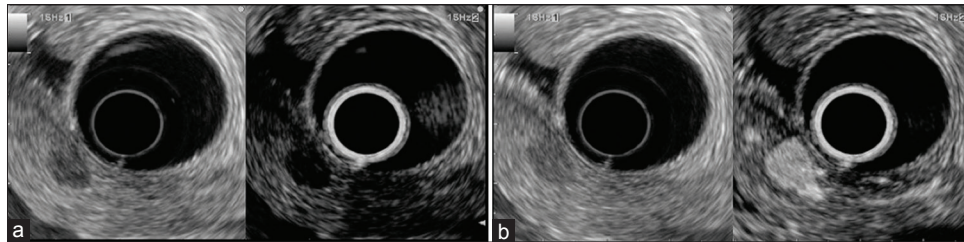


Figure 1. Typical benign G1 neuroendocrine tumor with early homogeneous strong hyperenhancement. (a) Image immediately after injection. (b) Image 20 s after injection

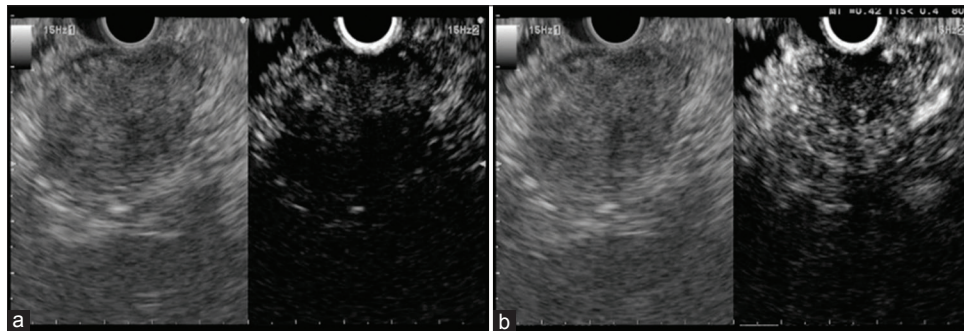


Figure 2. G1 malignant neuroendocrine tumor with heterogeneous enhancement. (a) Image immediately after injection. (b) Image 20 s after injection

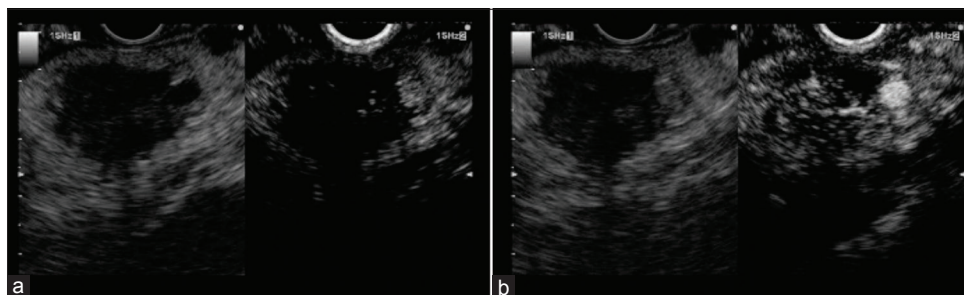


Figure 3. G2 malignant neuroendocrine tumor with heterogeneous enhancement. (a) Image immediately after injection. (b) Image 20 s after injection

CONTRAST HARMONIC ENDOSCOPIC ULTRASONOGRAPHY IN AUTOIMMUNE PANCREATITIS

The differential diagnosis between pancreatic adenocarcinoma and pseudotumoral forms of pancreatitis such as AIP is difficult, the negativity of EUS fine needle aspiration does not rule out the malignancy with certainty because of insufficient negative predictive value.

In two works,^[2,3] AIP was iso/hyperenhanced in more than 90% of cases [Figure 5]. Two studies assessed^[5,6] the use of a quantitative tool for analyzing the dynamic of enhancement to establish the differential diagnosis between AIP and pancreatic cancer. In the first, the intensity reduction rate at 1 min in comparison with the peak-intensity had the best diagnostic value, AIP having a significantly lower rate of reduction than

pancreatic cancer. In the latter work, the maximum gain of intensity was significantly higher in AIP than in pancreatic cancer.

CONTRAST HARMONIC ENDOSCOPIC ULTRASONOGRAPHY IN PANCREATIC METASTASES

Only one study^[7] specifically focused on the appearance of pancreatic metastases at CH-EUS. Of 11 lesions, 6 appeared hypoenhanced and 5 were iso/hyperenhanced depending on the origin of the primary tumor. In accordance with this study, in my experience, pancreatic metastases from adenocarcinoma (e.g., colon, breast) were hypoenhanced [Figure 6]. Metastases from renal cell carcinoma, lymphoma [Figures 7 and 8], and melanoma were iso/hyperenhanced. Notably, when lesions become larger, they tend to be heterogeneous with hypoenhanced areas.

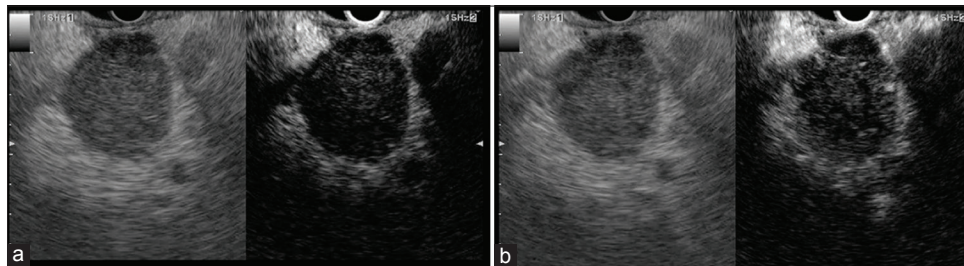


Figure 4. G3 malignant neuroendocrine tumor with almost no enhancement. (a) Image immediately after injection. (b) Image 20 s after injection

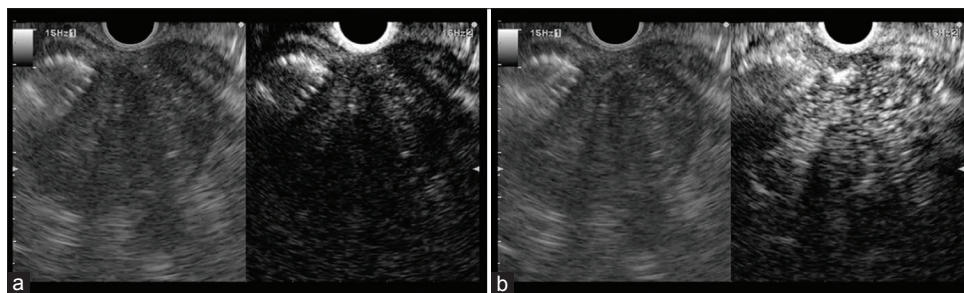


Figure 5. Typical mass-forming autoimmune pancreatitis with homogeneous intense hyperenhancement. (a) Image immediately after injection. (b) Image 20 s after injection

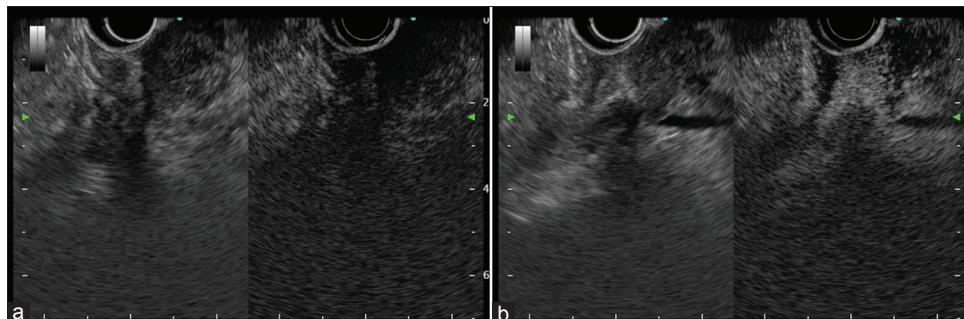


Figure 6. Metastasis from colon cancer with hypoenhancement. (a) Image immediately after injection. (b) Image 20 s after injection

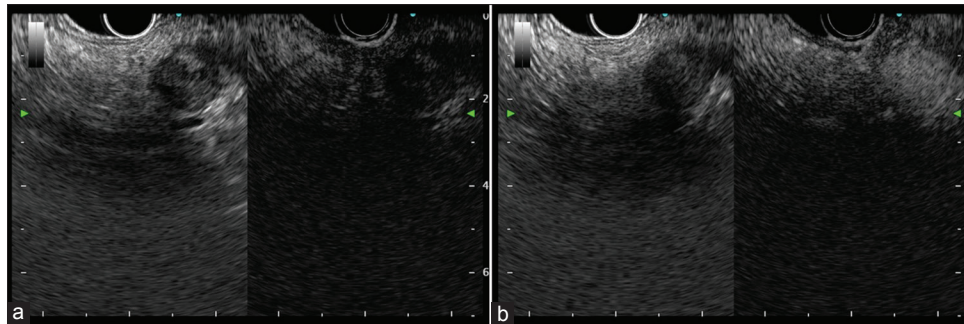


Figure 7. Metastasis from renal cell cancer with slight hyperenhancement. (a) Image immediately after injection. (b) Image 20 s after injection

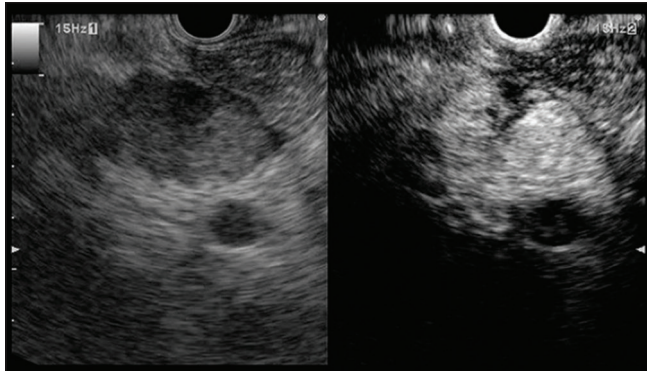


Figure 8. Metastasis from lymphoma with strong enhancement. Image 20 s after injection

CONCLUSION

Data from the literature remain to this day weak to determine the role of the CH-EUS in the management of rare solid tumor of the pancreas (NETs, AIP, and metastases). Specific studies are expected to further clarify the impact of this procedure in this field.

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Conflicts of interest

There are no conflicts of interest.

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