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Short Report

Frequency of occurrence and characteristics of primary pancreatic lymphoma during endoscopic ultrasound guided fine needle aspiration: A retrospective study



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ABSTRACT

Background: Primary pancreatic lymphoma is a rare tumour of the pancreas. Data on the role of endoscopic ultrasound guided fine needle aspiration for its diagnosis are scant.

Aim: To identify the frequency of occurrence, sonographic characteristics and cytological findings that are predictive of primary pancreatic lymphoma.

Methods: Pancreatic lymphoma cases were identified by retrospective review of solid pancreatic masses over 10-year period.

Results: 12/2397 (0.5%) lesions were identified. Patients were predominantly white (92%) and male (58%). Mean largest dimension was 47.5 mm and 83.3% were located in the head. The mass appeared heterogeneous in 75% and peripancreatic lymphadenopathy was noted in 58%. None of the patients showed features of chronic pancreatitis or pancreatic ductal dilation. Rapid onsite analysis revealed atypical lymphocytes in 92%. Flow cytometry confirmed diagnosis in 75% of cases.

Conclusions: Primary pancreatic lymphoma is encountered in 0.5% of patients undergoing endoscopic ultrasound guided fine needle aspiration. A large heterogeneous mass, in the absence of chronic pancreatitis or pancreatic duct dilation that reveals atypical lymphocytes on fine needle aspiration is suggestive.

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1. Introduction

In the United States, it is estimated that 45,220 cases of pancreatic cancer will be diagnosed in 2013 with an estimated death in 38,460 individuals [1] more commonly from adenocarcinoma. However, solid pancreatic masses identified on imaging vary from adenocarcinoma in 85% to mass forming autoimmune pancreatitis, primary pancreatic lymphoma (PPL), neuroendocrine tumours and complex pancreatic cyst neoplasms. While the latter conditions are rare, treatment outcomes are vastly different.

Pancreatic involvement in lymphoma can be due to peripancreatic lymph node mass causing direct invasion of the parenchyma;

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intrinsic pancreatic mass lesion with or without discrete peripancreatic lymphadenopathy or secondary extranodal deposit [2]. Widely accepted criteria for diagnosis of PPL are (a) mass involving the pancreas with or without loco-regional lymph nodes, (b) absence of superficial/mediastinal lymph nodes, (c) absence of hepatic or splenic involvement and (d) normal leucocyte count [3]. Although previous history of haematological malignancy, young age, presence of B symptoms (fever, night sweats and weight loss), large size tumour, low CA19-9 level and absence of jaundice or diabetes mellitus are shown to indicate haematological malignancy in the pancreas [4], tissue diagnosis is imperative for appropriate treatment given the differing outcomes of therapy for these conditions

EUS-FNA with its very high sensitivity and specificity has taken a central role in the management algorithm of solid pancreatic masses [5]. The objective of this study is to identify the frequency of occurrence, endosonographic characteristics and cytological findings of PPL in patients undergoing endoscopic ultrasound guided fine needle aspiration (EUS-FNA) of solid pancreatic masses.

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2. Materials and methods

This study was executed by reviewing the Institutional Review Board approved database of all patients who underwent EUS-FNA of solid pancreatic masses over a period of 10 years. The data extracted included findings at EUS, rapid on site cytopathology evaluation (ROSE) and final diagnosis. Criteria for inclusion in the study were solid pancreatic masses and a final diagnosis of PPL. Excluded from analysis were pancreatic or retroperitoneal lymph node masses that were adherent or adjacent to the pancreas.

EUS was performed under moderate sedation after administration of intravenous meperidine, midazolam and diazepam using a linear array echoendoscope (Olympus UCT140, Olympus America Corp, Center Valley, PA). When a mass was identified, its size, echo features, involvement of adjacent vasculature, and presence of peripancreatic lymph nodes was documented. Fine needle aspiration (FNA) of the mass was performed with a standard FNA needle (Echotip, Cook Endoscopy, Winston-Salem, NC, USA; ExpectTM, Boston Scientific Corporation, Natick, MA, USA), the needle size was determined by endosonographer preference. ROSE was undertaken of air-dried smears with Diff-Quick staining (Dade Diagnostics, Miami, FL, USA) and additional FNA passes were made for flow cytometry or other ancillary studies as determined by the onsite cytopathologist. The number of passes required to establish onsite preliminary assessment was documented. Patients were then recovered in the endoscopy unit and discharged when stable. Follow-up telephone call was made 24 h after the procedure to assess for delayed complications.

2.1. Statistical analysis

The baseline characteristics of patients were summarized as median, interquartile range for continuous variables such as age, size and number of passes with the FNA needle. Categorical variables such as gender, race, clinical symptoms, were on the other hand expressed as frequencies and proportions. Procedure details including echo characteristics; pancreatic parenchymal appearance; calibre of pancreatic duct; presence or absence of peripancreatic lymphadenopathy was summarized. The frequency of occurrence of primary pancreatic lymphoma was determined with the denominator of the total number of EUS-FNA of solid pancreatic lesions and 95% confidence interval of population proportion was then calculated. The datasets were compiled using Microsoft Excel and SAS software, version 93 (SAS institute, Cary, NC, USA) was used to perform the analysis.

3. Results

3.1. Patient demographics and clinical details

Of 2397 patients who underwent EUS-FNA of solid pancreatic masses over the study period, 21 were identified to have a diagnosis of lymphoma. In 9/21 patients, there was ambiguity regarding the origin of the tumour mass at EUS, these were classified as secondary pancreatic lymphoma and excluded from the analysis. The remaining 12 patients were categorized to have PPL and constituted the study cohort. The frequency of PPL occurrence was 0.5% [95% CI 0.22–0.78]. Majority of patients were male (58.3%) and Caucasian (91.7%) with a median age of 65.5 years (IQR = 41–89). Clinical presentation included abdominal pain (91.7%), jaundice (25%), weight loss (16.7%), nausea and vomiting (8.3%) and B-grade symptoms (8.3%). One patient had a prior diagnosis of AIDS. All patients had identifiable mass lesions on CT scan.

Table 1 Endosonographic features, fine needle aspiration and final flow cytometry analysis of the study group (N = 12).

Maximum size (mm) Number of passes	Mean 47.5, SD: 21 Median = 4, IQR = 1–6
Location Head Body Tail	10(83.4%) 1(8.3%) 1(8.3%)
Vascular invasion Yes None	5 (41.7%) 7 (58.3%)
EUS features Heterogeneous Hypoechoic	9 (75%) 3 (25%)
Peripancreatic LN Yes No	7(58.3%) 5(41.7%)
Dilated pancreatic duct Yes None	0 12 (100%)
ROSE Numerous lymphocytes Atypical lymphocytes Necrosis	9 (75%) 2 (16.7%) 1 (83%)
Flow cytometry Conclusive Inconclusive	9 (75%) 3 (25%)
Adequacy of material Yes No	9 (75%) 3 (25%)
Repeat EUS Yes No	3 (25%) 9 (75%)
Open biopsy Yes No	3 (25%) 9 (75%)

LN: lymph node; EUS: endoscopic ultrasound; ROSE: rapid on-site cytopathology diagnosis; SD: standard deviation; IQR: inter-quartile range.

3.2. EUS features

The mean largest dimension of the mass was $47.5 \, \mathrm{mm} \, (\mathrm{SD} = 21)$ and more than 80% were located in the head region with one each in the body and tail of pancreas. Echo features were heterogeneous 75% while the rest were uniformly hypoechoic. The margins were ill defined in all patients and vascular invasion was noted in 41.7%. The rest of the pancreatic parenchyma was unremarkable without features of chronic pancreatitis. The main pancreatic duct was not dilated in any patient and peripancreatic lymphadenopathy was noted in 58.3% (Table 1).

3.3. Fine needle aspiration

FNA was performed with 22G needle in 833% and with 25G in others. One patient underwent EUS-FNA using a 19G needle following a failed procedure using a 22G needle. Median number of passes was four (IQR 1–6). Rapid onsite cytopathology revealed numerous atypical lymphocytes (Figs. 1 and 2) in 91.7% with one other showing necrosis. Flow cytometry confirmed lymphoma in 9 patients (75%). Three patients underwent repeat EUS-FNA with flow cytometry that was inconclusive; surgical biopsy established a definitive diagnosis of lymphoma in these patients. No complications were encountered with EUS-FNA.

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