ALTERED FC AND FAB GLYCOSYLATION STATUS IN PATIENTS WITH IGG4-RELATED SCLEROSING CHOLANGITIS AND AUTOIMMUNE PANCREATITIS

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Introduction IgG4-related disease (IgG4-RD) is a systemic fibro-inflammatory condition characterised by an abundance of IgG4+ antibodies in the serum and tissue of involved organs. IgG glycosylation plays an important role in many chronic inflammatory and autoimmune conditions. We sought to assess the glycosylation status in patients with IgG4-RD and correlate with disease activity, damage and response to treatment.

Methods IgG Fc and Fab glycosylation status was assessed in patients with IgG4-RD involving the bile ducts (IgG4-sclerosing cholangitis, IgG4-SC) and pancreas (autoimmune pancreatitis) (n=22), disease controls with primary sclerosing cholangitis (DC n=22) and healthy controls (HC n=22). Disease activity, organ damage and response to treatment were assessed serially using the IgG4-responder index. Serum IgG and subclasses were quantified using an ELISA and nephelometry. IgG and subclass Fc glycosylation was analysed by mass spectrometry and Fab glycosylation by lectin (SNA) affinity chromatography. Statistics were performed using Prism.

Results IgG4-SC and AIP patients exhibited reduced total IgG Fc galactosylation and IgG1 Fc bisection, and increased IgG4 Fc fucosylation and IgG2/3 Fc hybrid compared with HC. There was recovery of IgG1 Fc bisection (increase) and IgG2/3 Fc hybrid (decrease) upon corticosteroid treatment. IgG Fc galactosylation and IgG2/3 Fc hybrid correlated with disease activity. IgG Fab glycosylation was higher in IgG4-RD patients, with an increase in IgG4-specific, and to a lesser extent IgG1-specific, Fab glycosylation compared to HC and DC.

Conclusions In the first study to assess glycosylation status in IgG4-RD, we demonstrated alterations in both IgG Fc and Fab glycosylation, which may play a role in pathophysiology and serve as a biomarker of disease.

WHAT IS THE YIELD AND CLINICAL UTILITY OF EUS IN PATIENTS WITH PRIOR NON-DIAGNOSTIC MRCP?

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Introduction The most effective investigation for suspected gallstones between MRCP and EUS is unclear. A 2015 Cochrane systematic review of their performance in common bile duct (CBD) stones concluded that the tests were of comparable accuracy. Conversely, a 2017 meta-analysis found EUS to be more sensitive. Any superiority of EUS may be due to better accuracy in detecting small stones. MRCP is routinely favoured as the 2nd line test following a non-diagnostic abdominal ultrasound and EUS subsequently performed as the 3rd line test when suspicion remains after a non-diagnostic MRCP. The yield and clinical utility of EUS in this setting is unclear. The aim was to identify the yield of EUS in patients with prior non-diagnostic MRCP undergoing EUS in our tertiary service.

Methods All EUS reports from 2017 were reviewed along with the electronic patient records to identify cases with prior MRCP. Indication for the procedure, symptoms, liver blood tests and interval between MRCP and EUS were recorded. Findings of sludge, microlithiasis (stones<2 mm) and discrete stones were categorised together as stones. Subsequent ERCP or cholecystectomy was identified. Yield was defined as a finding that would lead to a change in management.

Results A total of 1058 diagnostic EUS were screened of whom 253 (24%) had prior MRCP and formed the study group. Median age was 58 (16–88) years, 179 (71%) were female and 91 (36%) had a cholecystectomy. Median interval between EUS and MRCP was 5.2 (0.1–37) months. Indications for EUS were: n=76 (30%) dilated CBD, n=65 (26%) query CBD stones, n=54 (21%) unexplained acute pancreatitis (AP), n=23 (9%) right upper quadrant pain, n=17 (6.7%) abnormal LFTs, n=16 (6.3%) double duct sign and n=2 (1%) dilated PD. There was a yield from EUS in 30 (12%) patients with no significant difference between those with (n=11) or without cholecystectomy (n=19). Stones were identified in 24 cases with median size of 4 mm (range 2–8) in: CBD (n=16), cystic duct (n=1) and GB (n=7). Three had abnormal CBD without stones (calcification CBD wall, thick walled CBD, polyp), 1 patient with possible stone on MRCP had no stone seen on EUS, 1 had a pancreatic mass, and 1 had chronic pancreatitis. All patients in whom EUS findings indicated an intervention (26/30) have been referred: ERCP in 13, cholecystectomy in 9, ERCP and cholecystectomy in 3 and chemotherapy in 1.

Conclusion EUS following non-diagnostic MRCP is a sizeable workload accounting for 24% of diagnostic activity in our unit with a clinically significant yield in 12% of predominantly small stones. Further prospective studies are required to ascertain the most cost-effective way to incorporate EUS into the investigation of suspected gallstone disease.
Pancreatic cysts are a frequent incidental finding on cross-sectional imaging of the abdomen. We examined the decision making and outcome of patients with a pancreatic cyst(s) discussed at the Royal Derby Hospital HPB cancer MDT and compared practice against the 2015 American Gastroenterology Association (AGA) and 2017 International Association of Pancreatologists (IAP) guidelines on the management of pancreatic cysts.

Methods A search of HPB MDT meeting reports, from January 2016 to October 2017 (n=1144) identified 88 patient (51 female) reports relating to the first discussion of a pancreatic cyst(s) discussed at the Royal Derby Hospital HPB cancer MDT and compared practice against the 2015 American Gastroenterology Association (AGA) and 2017 International Association of Pancreatologists (IAP) guidelines on the management of pancreatic cysts.

Results The median age was 72 years (range 32–87) and the median estimated 10 year survival based on the Charlson Comorbidity Index was 53% (range 0%–98%).

86% of pancreatic cysts were judged to be an incidental finding. The median cyst diameter was 19.5 mm (range 4–110 mm). 43/88 (49%) patients proceeded to endoscopic ultrasound (EUS), with 33 having a fine needle aspiration (FNA). 4/88 (5%) patients had probably malignant (C4) or malignant (C5) cytology. All 4 patients had “high risk stigmata” on their initial CT/MRI. The final outcome for most patients was no further intervention (56%) or follow-up imaging (36%), with 5 (6%) patients offered surgery.

Applying the 2017 IAP management algorithm, 13 (15%) patients had “high-risk stigmata” on CT/MRI and except where their performance status or co-morbidity precluded further investigation/treatment (5), were recommended for surgery (1) or EUS (7). Of the remaining 75 patients, 45 (60%) had no worrisome features on CT/MRI and so would not have required EUS, 21/45 (47%) of these patients in our practice underwent EUS, but none demonstrated definite mural nodules, main duct involvement or suspicious/positive cytology.

The 2015 AGA management algorithm could only be applied to those patients who had undergone initial radiological assessment with MRI (n=11). None of these patients had two positive features on MRI, indicating a need for EUS. 6/11 (55%) patients did, however, have an EUS, with none identifying positive features or concerning cytology.

Conclusions These findings suggest that a significant proportion (24%) of patients with pancreatic cysts underwent unnecessary EUS. Application of international guidelines can reduce the number of patients who require an endoscopic ultrasound.

**PTU-025** EUS FNA MICROCORE BIOPSY IS SUPERIOR TO ENDOBILIARY BIOPSY IN DIAGNOSING MALIGNANT PANCREATOCBILIARY LESIONS

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**Introduction** The ability to provide definitive and timely histological diagnosis of malignancy in suspected malignant pancreaticobiliary lesions depends on high quality tissue sampling. We conducted a study to evaluate the diagnostic utility of endobiliary biopsy versus endoscopic ultrasonography-guided fine needle aspiration (EUS FNA) microcore biopsy.

**Methods** We performed a retrospective search of our laboratory information management system for patients with clinically suspected malignant pancreaticobiliary lesions who had endobiliary and EUS FNA microcore biopsies. The haematoxylin and eosin-stained slides were retrieved and reviewed. All biopsies were categorised into whether a definitive diagnosis can be established (diagnostic) or not (non-diagnostic).

**Results** The study yielded 94 endobiliary biopsies and 78 EUS FNA microcore biopsies. 77 out of the 94 endobiliary biopsies were deemed adequate, and out of this 54 was diagnostic of malignancy (sensitivity 57%). In 11 cases, where the endobiliary biopsy was not diagnostic, subsequent EUS FNA microcore biopsies provided a malignant diagnosis in 5. 96% of EUS FNA microcore biopsies were adequate and in 62 a malignant diagnosis could be established (sensitivity 83%). Cholangiocarcinoma was the pathological diagnosis in the majority of the endobiliary biopsies and there were two metastases from lung and two neuroendocrine tumours. The main malignant diagnosis in the EUS FNA microcore biopsies was adenocarcinoma of pancreaticobiliary-type, but also included neuroendocrine tumours, solid pseudopapillary neoplasm and adenosquamous carcinoma. There were also three metastases from the colorectum, kidney and breast.