Methods This was a retrospective cohort study of all patients undergoing antireflux surgery from 2001 to 2009 in a tertiary centre under a single surgeon. Patients undergoing fundoplication and/or repair of paraoesophageal hernias were included. Patients underwent preoperative assessment by means of endoscopy, oesophageal manometry and 24 h oesophageal pH monitoring. Patients were grouped on the basis of their 24 h pH profile into upright refluxers (daytime increased oesophageal acid exposure) and mixed refluxers (night-time increased oesophageal acid exposure +/- daytime reflux). Primary outcomes included if the patient had stopped PPIs following surgery and the incidence of postoperative dysphagia, vomiting and reflux symptoms. Secondary outcomes included if the patients deemed the operation a success.

Results 120 patients were included, with a median age of 49 years (range 24–81) at time of surgery. 53% (n=63) were male and 93% were laparoscopic procedures. 100 patients (83.3%) had Nissen fundoplication and 13% had a combination of fundoplication and paraoesophageal hernia repair. A DeMeester score >15 was present in 97.8% (n=88) patients, with 21% (n=19) patients having upright reflux and 68.5% (n=63) having mixed reflux symptoms on pH studies. Mixed refluxers were nearly twice as likely to have significant oesophagitis (grade B+) and or Barrett's oesophagus on preoperative endoscopy (mixed reflux 57.1% vs 31% upright reflux; $p=0.051$). Those patients with mixed reflux symptoms were significantly more likely to stop PPIs postoperatively (mixed reflux 96%, n=48 vs 75% n=12, upright reflux; $p=0.01$). Further, the incidence of significant post operative dysphagia was doubled in the upright reflux group (upright reflux 53.3% vs mixed reflux 26.7%; $p=0.058$). Overall, 83.5% (n=76) deemed the operation to have been a success, 7.7% (n=7) a partial success and 8.8% (n=8) considered that surgery had failed to improve their symptoms.

Conclusion In addition to the DeMeester score for predicting outcomes in antireflux surgery, the presence of reflux only in the upright position may indicate a poorer outcome.

Competing interests None declared.

PTU-191

CIRCUMFERENTIAL RESECTION MARGIN <1 MM FOLLOWING NEOADJUVANT CHEMOTHERAPY DOES NOT ADVERSELY IMPACT SURVIVAL IN PATIENTS UNDERGOING OESOPHAGECTOMY

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Introduction The TNM 7 classification system for oesophageal cancer defines circumferential resection margin (CRM) involvement as R0 (>1 mm), R1a (<1 mm), and R1b (involved margin). The prognostic importance of CRM involvement is unclear with previous studies demonstrating conflicting results. We evaluated survival according to CRM status for patients with locally advanced oesophageal adenocarcinoma who had neoadjuvant chemotherapy prior to Ivor Lewis en bloc oesophagectomy with extended lymphadenectomy.

Methods Since 2006, the CRM has been classified as R1a, R1b or R0, assessed by two specialist upper gastrointestinal pathologists in our unit. Since 2007, NAC prior to oesophagectomy was introduced as standard practice in patients with locally advanced oesophageal adenocarcinoma. A retrospective review of prospectively collected data were performed on all patients who underwent oesophagectomy for oesophageal and gastroesophageal junctional adenocarcinoma including demographic data, NAC, final histology, 30, 60, 90 and in-hospital mortality, and survival. Patients with proximal and distal resection margin involvement were excluded. The primary

outcome was overall survival evaluated by Kaplan–Meier survival analysis.

Results 63 consecutive resections were included. 62% (n=39) were R0, 25% (n=16) were R1a and 13% (n=8) were R1b. 2-year overall survival was 56%, 63% and 13% respectively, with a statistically significant reduction in survival between R1b and R1a/R0 ($p<0.05$). Local recurrence rates were similar for R0 (8%), R1a (19%) and R1b (13%). The rate of distant metastases was higher in R1b (75%) compared to R0 (19%) and R1a (20.5%). Univariate analysis demonstrated that poor survival was associated with R1b but not R1a or R0.

Abstract PTU-191 Figure 1 Kaplan–Meier overall survival according to CRM.

Conclusion R1a and R0 CRM are associated with equivalent recurrence (local and distant) and survival rates to R0 in patients with oesophageal and GOJ adenocarcinoma following NAC and oesophagectomy. R1a CRM involvement does not adversely survival in patients with oesophageal and GOJ adenocarcinoma following NAC.

Competing interests None declared.

REFERENCES

1. **Pultrum B**, et al. A critical appraisal of circumferential resection margins in esophageal carcinoma. *Ann Surg Oncol* 2010;**17**:812–20.
2. **Sujendran V**, et al. Effect of neoadjuvant chemotherapy on circumferential margin positivity and its impact on prognosis in patients with resectable oesophageal cancer. *Br J Surg* 2008;**95**:191–4.

PTU-192

TIME: PROSPECTIVE STUDY COMBINING ENDOSCOPIC TRIMODAL IMAGING AND MOLECULAR ENDPOINTS TO RISK STRATIFY BARRETT'S OESOPHAGUS

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Introduction Biomarkers have been proposed to improve risk stratification in Barrett's oesophagus (BO), however molecular heterogeneity of BO can hamper detection of molecular changes in random biopsies. Use of Autofluorescence Imaging (AFI) within endoscopic Trimodal Imaging (ETMI) can improve dysplasia detection, but has high false positive rate. Aims of study were (a) validate biomarkers previously published in separate patient cohorts in single study (b) assess whether AFI can increase detection of biomarkers (c) combine ETMI and biomarkers to improve risk stratification of patients with BO.

Methods Prospective European multicentre study. Each patient underwent ETMI with targeted biopsies on AFI positive (AFI+) areas and one AFI negative (AFI-) area, as well as random

quadrantic biopsies. DNA content abnormalities (aneuploidy/tetraploidy); loss of heterozygosity (LOH) at 9p and 17p loci; RUNX3, HPP1 and p16 methylation; immunohistochemistry (IHC) for p53 and Cyclin A were tested on targeted biopsies. Each biomarker was correlated with the dysplasia and AFI status.

Results 111 patients with 210 biopsy areas were included in the analysis (AFI+, n=120; AFI-, n=90). Univariate per-biopsy analysis showed that all biomarkers correlated with dysplasia ($p<0.05$), with exception of 9p LOH. Multivariate analysis showed that aneuploidy, p53 IHC and Cyclin A (3 biomarker panel) were independently associated with dysplasia with an AUC=0.93 (95% CI 0.88 to 0.98) for any dysplasia and AUC=0.95 (95% CI 0.89 to 1) for HGD/early cancer (EC). AFI positivity significantly correlated with aneuploidy, p16 methylation, cyclin A and p53 staining ($p<0.05$). After excluding dysplastic areas, aneuploidy ($p=0.03$) and p53 ($p=0.04$) staining retained statistical correlation with AFI positivity. Analysis of the 3 biomarker panel in patients with dysplasia showed significant biomarker enrichment in AFI+ compared to AFI- areas ($p=0.001$). Finally, 3 biomarker panel was used to predict prevalent dysplasia. Using a cut-off of ≥ 2 biomarkers, the panel when applied to AFI+ areas alone, showed sensitivity and specificity of 88% and 90% respectively for diagnosis of HGD/EC, and 64% and 96% respectively for diagnosis of any dysplasia, compared to overall histology.

Conclusion AFI increases detection rate for molecular biomarkers. A panel of 3 molecular biomarkers on a small number of AFI targeted biopsies can efficiently predict the dysplasia status and potentially inform therapeutic management of patients with BE.

Competing interests None declared.

PTU-193 GASTRO-OESOPHAGEAL REFLUX (GORD) AND COELIAC DISEASE: A BIDIRECTIONAL STUDY

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Introduction There are limited data assessing the relationship between coeliac disease and GORD. We aimed to establish the prevalence and nature of GORD symptoms in patients with coeliac disease, and the prevalence of undetected coeliac disease in those presenting to endoscopy with GORD symptoms.

Methods Group A were histologically proven patients with coeliac disease (n=225) who were asked to complete a validated reflux questionnaire and then compared to age/sex-matched controls (n=348). Group B were consecutive GORD patients undergoing endoscopy who had duodenal biopsies and coeliac serology taken. (n=851) Furthermore, patients with newly diagnosed coeliac disease underwent manometry and pH studies prior to commencing gluten free diet (n=33).

Results In Group A the prevalence of GORD was greater in coeliac disease (66%) than in healthy controls (50%) $p=0.0001$. Coeliac patients also report reflux of greater severity: coeliac disease OR=6.8, 95% CI=3.6 to 12.7, $p<0.001$. In Group B at endoscopy the prevalence of undetected coeliac disease was 1.66% (14/851). In Group C 31/33 were able to tolerate manometry and complete testing (2 had partial investigation before the catheter was removed) At manometry 30/33 (91%) had a normal lower oesophageal sphincter (LOS). 2/33 had a hypotensive LOS. 21/33 (64%) had normal oesophageal motility. However, 10/33 had a hypocontractile oesophagus, 1 was hypertensive and 1 showed functional oesophago-gastric junction obstruction. During manometry 6/31 (19%)

demonstrated significant reflux, 6/31 (19%) had some reflux and the final 19/31 (61%) had no reflux episodes. In these coeliac patients neither the presence of symptoms nor abnormal oesophageal study findings was related to histological grade of coeliac disease, (villous atrophy) or serology findings.

Conclusion Up to two thirds of patients with coeliac disease report reflux symptoms and one third have demonstrable abnormalities of oesophageal motility and reflux. However, in an unselected population at endoscopy, reflux symptoms are not predictive of coeliac disease.

Competing interests None declared.

PTU-194 DOES METABOLIC SYNDROME IMPACT TUMOUR PATHOLOGY IN OESOPHAGEAL ADENOCARCINOMA?

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Introduction Obesity is an established risk factor for both the increased incidence of oesophageal adenocarcinoma cancer (OAC), and adverse outcomes by increasing risk of recurrence and reducing survival in obese patients post oesophagectomy. The exact mechanism of this relationship is unclear but the pattern of fat distribution pattern is likely important. Abdominal obesity more closely reflects an increased visceral fat area and is associated with alterations in metabolic risk profile. The clustering of central obesity, hypertension, and raised plasma glucose, triglycerides and HDL cholesterol is also known as the metabolic syndrome (MetS). The processes underlying the metabolic syndrome especially insulin resistance and increased leptin, can provide a favourable growth environment for malignant cells and may have a role in cancer progression.

Methods The aim of this prospective observational study of OAC patients was to examine the incidence of MetS and its relationship to tumour pathology in an Irish population. Patients underwent a metabolic and nutritional assessment prior to initiation of treatment. Visceral fat area was measured using CT scans. MetS was defined according to the International Diabetes Federation definition.¹

Results 83 OAC patients (71 male: 12 female) were recruited with a median age of 64.6 years $\pm$ 1.0 (range 48–86). All patients underwent an oesophagectomy, with 42% (n=35) receiving neoadjuvant chemoradiotherapy. 58% of patients were either overweight or obese with a further 60% centrally obese. Males had significantly greater visceral fat area ($p=0.031$) despite no difference in total abdominal fat compared to females ($p=0.757$). The incidence of obesity may be underestimated as 41% of patients reported unintentional weight loss with 18% losing $>5\%$ of their usual body weight. MetS was diagnosed in 39% patients, which exceeds the population norms reported at 21%.² The presence of MetS was not associated with tumour length, depth of invasion, lymph node positive disease, clinical or overall pathological stage in males or females. Individual features of MetS were also not significantly related to the pathological staging of oesophageal cancer.

Conclusion We report an increased prevalence of MetS and central obesity in a cohort of Irish patients with OAC. Dysphagia and weight loss are common in the presentation of oesophageal cancer and may mask the effect of obesity and metabolic syndrome on the clinical pathological features of OAC in this cohort. Further research is needed to fully understand the underlying biological mechanisms linking obesity to oesophageal cancer.