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**Introduction** Squamous Cell Cancer (SCC) of the oesophagus has a poor prognosis with 5-year survival at 10%. Squamous high grade dysplasia (HGD) is the precursor lesion to SCC. Risk of progression to SCC with Squamous HGD can be 65% at 5 years. Radiofrequency ablation (RFA) is a minimally invasive ablation technique with proven efficacy for early neoplasia in Barrett's Oesophagus.

**Methods** Prospective multicenter registry of patients undergoing RFA for Squamous HGD and early carcinoma in situ (CIS) from eight UK centers. Nodular lesions were removed by endoscopic mucosal resection before RFA. Treatment consisted of a single ablation at 12 J/cm<sup>2</sup>. Patients were followed-up 3 monthly and repeat biopsies taken. Those with residual dysplasia underwent RFA 3 months later until 12 months where they were assessed for treatment success or failure.

**Results** 25 patients have undergone ablation for squamous HGD/CIS. We report on 17 patients to have completed protocol. Mean length of dysplastic epithelium ablated was 5 cm (1–14). Mean time to protocol completion was 8.7 months. CR-HGD was seen in 59% of patients and CR-D in 47% at end of protocol biopsy with mean of 1.4 RFA treatments (1–3). All those with successful outcomes remain free of dysplasia at most recent biopsy, median follow-up 10.6 months (2–36) from first treatment. At protocol completion, six of 17 patients (35%) had progressed to invasive cancer and referred for surgery or chemo-radiotherapy. Three patients (18%) required dilatations for oesophageal structuring after first treatment. Two of these patients have required serial dilatations thereafter with an average of four dilatations per patient.

**Conclusion** Squamous HGD and CIS are very aggressive pathologies as evidenced by the fact a third of patients progressed to invasive disease despite RFA. The role of RFA in these patients remains unclear. In our series 47% of patients responded to RFA & have reassuringly remained free of dysplasia at last follow-up. These figures are lower than limited published data to date but in our series an emphasis was placed on restaging carefully after each treatment to assess for progression. As our experience grows with confidence in identifying these lesions more accurately and increasing the frequency and number of ablations administered over the protocol period, dysplasia reversal rates will be expected to increase.

Competing interests None declared.

PWE-028

HALO RADIOFREQUENCY ABLATION FOR HIGH GRADE DYSPLASIA AND EARLY MUCOSAL NEOPLASIA ARISING IN BARRETT'S OESOPHAGUS: INTERIM RESULTS FORM THE UK HALO RADIOFREQUENCY ABLATION REGISTRY

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Introduction Barrett's oesophagus (BE) is the pre-cursor to oesophageal adenocarcinmoa (OAC). High grade dysplasia (HGD) and early mucosal neoplasia in BE has historically been treated with surgery. Recently there is a shift towards minimally invasive endotherapy with endoscopic mucosal resection (EMR) and Radiofrequency ablation (RFA).

**Methods** Prospective registry from 14 UK centers to audit RFA outcomes in patients with HGD and early neoplasia in BE. Prior to RFA, any visible lesions were first removed by EMR. Patients then underwent RFA 3 monthly until all visible BE was ablated or cancer developed. Biopsies were taken at the end of this protocol.

Results 216 patients have completed protocol, mean age 68.6 years (40-90), 81% male. Mean time to protocol end 11.3 months (IQR 8-14.3), median 2 ablations and mean of 2.4 (2-6) during protocol with mean 1.4 circumferential ablations and 1.2 focal ablations performed during protocol. Mean length BE segment ablated is 5.8 cm (1-20). CR-HGD was achieved in 83% patients at protocol end biopsy. CR-D was 76% and CR-BE 50% at this point. CR-D was more likely in short segment BE (<5 cm) at protocol end (82% vs 54%, p<0.0001, Fisher's exact test). Patients who required EMR during RFA protocol were less likely to achieve CR-D than those who had RFA alone (52% vs 79%, p=0.002, fishers exact test). 3.7% patients progressed to invasive cancer at protocol end. Complications include one perforation and 1% incidence of superficial tears. 37 patients have at least 12 months or more follow-up after successful completion of protocol (range 12-42), median 16.2 months. Durability in these is excellent with 95% dysplasia free at most recent biopsy.

**Conclusion** This is the largest series to date of patients undergoing RFA from 14 UK centers. End of protocol CR-D is satisfactory at 76% and successful eradication appears to be durable. Patients with short segment BE are likely to respond better. Our data represent real life outcomes of integrating minimally invasive endotherapy into demanding endoscopy service commitments.

Competing interests None declared.

PWE-029

OBJECTIVE ASSESSMENT OF PHYSICAL ACTIVITY AS A MEASURE OF FUNCTIONAL RECOVERY AND QUALITY OF LIFE FOLLOWING OESOPHAGO-GASTRIC CANCER RESECTION

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**Introduction** Functional recovery following surgery is determined by the interaction between pre-operative performance, post-operative catabolism, nutritional status, and mood. Physical activity (PA) is an important domain of health-related quality-of-life (HRQL), and may be a useful objective index of recovery. We aimed to use an accelerometer-based activity metre (ActivPAL) to monitor post-operative PA in oesophago-gastric (OG) cancer patients undergoing surgery with curative intent.

**Methods** PA measures, including step count, time spent in various body positions, and energy expenditure of activity, were assessed over 7-day periods in patients undergoing oesophagectomy or gastrectomy (n=16). Nutritional status, HRQL (FAACT, FACIT-F and EORTC-QLQC30 questionnaires), and mood (HADS questionnaire) were also assessed. Time-points were pre-operatively and 1-2 weeks, 5-6 weeks, 3 months and 6 months post-operatively.

**Results** Compared with pre-operative results, PA measures were decreased by 23-89% (p<0.05) 1-2 weeks post-operatively, and were still decreased by 15-57% (p<0.05) 5-6 weeks

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post-operatively. At 3 months, all PA measures except time spent upright (p=0.009) and time spent standing (p=0.013) had recovered. Measures of PA correlated positively with physical and functional domains of HRQL, including EORTC-QLQ30 Global Health Status, FAACT Trial Outcome Index (TOI) and FACIT-TOI (p<0.001), and inversely with HADS-Depression (p<0.001).

Conclusion There is marked impairment of PA at the time of hospital discharge and a gradual recovery over 3-6 months. This carries significant implications in a disease where surgical patients may survive <2 years. PA measures are suitable outcomes for evaluating the impact of enhanced recovery programmes on functional recovery and HRQL.

Competing interests None declared.

## PWE-030 IMMUNOCYTOCHEMICAL ASSESSMENT OF INTRA-TUMOUR MICROVESSEL DENSITY IN **OESOPHAGOGASTRIC CANCER DOES NOT HAVE** PROGNOSTIC SIGNIFICANCE

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Introduction Intra-tumour microvessel density (IMD), a marker of tumour angiogenesis, correlates with metastasis and poor prognosis in many cancers. In oesophagogastric cancer however, the prognostic significance of IMD assessment remains incompletely investigated. **Methods** Patients undergoing surgery with curative intent, without pre-operative chemotherapy, were prospectively recruited between February 1999 and August 2000. Immunocytochemical staining of tumour microvessels was undertaken using anti-CD34 (QBEND 10 clone) antibodies. IMD (microvessels per mm<sup>2</sup>) was assessed using a validated "hot-spot" technique. Patients were followed-up over a 10year period using the Northern Ireland Cancer Registry. The relationship between IMD and standard clinicopathological variables was assessed using the Mann-Whitney U test. Univariate survival analysis was calculated using a Cox's proportional hazard model while survival analysis was calculated using Kaplan-Meier estimation and log rank. Results 61 patients were recruited (male=45) with a median age of 66.0 years (range 39-83). The overall 10-year survival rate was 19.7% (n=12). IMD was significantly higher in males compared to females (332.93 vs 252.44, p=0.04) and adenocarcinomas compared to squamous cell carcinomas (356.10 vs 203.66, p<0.001). On univariate survival analysis only lymphovascular invasion predicted poor prognosis (HR 2.26, 95% CI 1.01 to 5.07, p=0.05). Kaplan-Meier survival analysis demonstrated no difference in longterm survival for patients with IMD levels greater or less than the median value (738 days vs 882 days, p=0.67).

**Conclusion** Immunocytochemical analysis of IMD does not have a prognostic benefit in determining long-term survival in patients with oesophagogastric cancer.

Competing interests None declared.

# PWE-031 I

## THE POTENTIAL OF HAEM TRANSPORT PROTEINS AS THERAPEUTIC TARGETS IN THE TREATMENT OF **OESOPHAGEAL ADENOCARCINOMA**

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Introduction Epidemiological evidence supports the potential role of dietary haem iron in gastrointestinal carcinogenesis. 1 2 Oesophageal cancer cells acquire inorganic and organic iron by progressive upregulation of iron and haem transport proteins.<sup>3</sup> These alterations result in increased cellular iron loading which is likely to drive cellular proliferation.3 The effect of perturbing haem import proteins, hemopexin receptor (LRP1) and haem carrier protein (HCP1), on cancer cell iron transport, phenotype and tumour burden is unknown. The aim was to determine the in vitro and in vivo effects of depriving oesophageal cancer cells of haem iron.

Methods Stable knock-downs of LRP1 and HCP1 were created by infection with specific shRNA lenti-viral vectors. Successful knockdown was confirmed by Western blotting. The effect on cellular iron transport and cell phenotype was assessed by qRT-PCR and phenotypic experiments for viability, proliferation, migration and anchorage independent growth. Stable knock-downs of LRP1 and HCP1 were then xenografted into NOD-SCID mice as an in vivo model of oesophageal adenocarcinoma.

**Results** Perturbation of LRP1 and HCP1 caused a compensatory upregulation of inorganic iron import proteins and a decrease in iron storage capacity. Neoplastic activity was significantly impaired compared to control (proliferation, viability, colony forming and migration - all p=<0.01). Loss of active haem iron import significantly reduced xenograft tumour burden in murine models with a 70% (p=0.014) and 58% (p=0.05) reduction in average xenograft weight compared to control.

**Conclusion** Haem import significantly contributes to iron loading in oesophageal cancer cells and creates a more aggressive phenotype. Functional inhibition of haem importing proteins LRP1 and HCP1 curbs neoplastic activity and significantly reduces in vivo tumour burden in murine models. Haem iron transport proteins are potential therapeutic targets in the treatment of oesophageal adenocarcinoma.

Competing interests None declared.

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PWE-032 | ENDOSCOPIC MUCOSAL RESECTION (EMR) FOLLOWED BY ADJUVANT RADIOFREQUENCY ABLATION (RFA) CAN RESULT IN BETTER OUTCOMES COMPARED TO EMR ALONE IN PATIENTS WITH BARRETT'S EARLY **NEOPLASIA (EN). A COMPARATIVE STUDY FROM A** TERTIARY CENTRE IN THE UK

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Introduction RFA has shown efficacy in eradicating Barrett's EN (high grade dysplasia (HGD) or intra-mucosal cancer (IMC)). To our knowledge, there are no studies directly comparing outcomes in patients with EN who undergo EMR alone vs EMR followed by RFA. The aim of this study was to assess the efficacy, safety and long term outcomes of adjuvant RFA in this setting.

Methods We searched our prospective Barrett's Oesophagus EMR database for patients who had EMR of lesions harbouring EN followed by RFA for eradication of residual Barrett's mucosa between 2007 and 2008 as part of a multi-centre trial (intervention group). The control group included patients with similar lesions

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