Introduction

Meta-analysis of published literature suggests that up to 14% of upper gastrointestinal cancer (UGIC) subjects had a negative oesophagogastroduodenoscopy (OGD) up to 3 years prior to diagnosis. We have examined how often UGIC is missed at OGD in our organisation and associated risk factors.

Methods

Computerised OGD records from Sandwell General and City Hospitals between 1999 and 2007 were retrieved and submitted to the West Midlands Cancer Intelligence Unit (WMCIU) for UGIC registrations linkage. Subjects undergoing OGD 3 months to 5 years before diagnosis were identified as potentially missed UGIC cases and those with no OGD 3 months to 5 years before diagnosis served as controls. The influence of age, gender, indication, endoscopist specialty, trainee involvement, sedation use, number of biopsies taken, and histology of UGIC on missed UGIC were examined by logistic regression analysis.

Results

36577 OGD records were submitted to WMCIU for cancer linkage and 524 UGIC were linked. 419 control UGIC. 388 (93%) diagnosed at initial OGD; 31 (7%) diagnosed at repeat OGD appropriately (within 90 days, with previous benign history). 105 (20%) potentially missed UGIC cases: 39 (37%) oesophageal cancer (OC) and 66 (63%) gastric cancer (GC); 42 (40%) had OGD 3 months to 1 year prior to diagnosis; 33 (31%) had OGD 1–3 years prior and 30 (29%) had OGD 3–5 years prior. Furthermore, 30% of missed UGIC cases had OGD where abnormality at site of UGIC was seen but not biopsied or benign histology from an inadequate number of biopsies (< 4) within 3 months to 1 year prior to diagnosis.

Lack of alarm symptoms (2.51, 95%CI 1.58–4.00, p = 0.001) and female gender (1.79, 1.16–2.79, p = 0.009) were associated with missed UGIC. The number of biopsies taken was significantly lower in the missed UGIC group than in the controls (2.1 ± 0.2 vs 5.4 ± 0.1, p < 0.05).

In subjects with OC, mid-oesophageal OC appeared more likely to be missed than lower-oesophageal OC (2.04, 0.99–4.23, p = 0.05). Oesophageal squamous cell carcinoma was much more likely to be missed than oesophageal adenocarcinoma (4.47, 1.83–10.65, p < 0.001). In GC subjects, there was no association between missed UGIC and tumour site (1.10, 0.61–1.97, p = 0.5) or histology subtype (1.10, 0.46–2.67, p = 0.8).

Age (1.5, −0.8–3.8, p = 0.2), endoscopist specialty (1.39, 0.70–2.76, p = 0.34), trainee involvement (1.2, 0.78–1.86, p = 0.59) and sedation use (0.98, 0.64–1.51, p = 0.9) were not associated with increased risk of missing UGIC.

Conclusion

Missing UGIC at OGD was seen in 14.3% of subjects within 3 years of diagnosis. It was associated with lack of alarm symptoms, female gender, oesophageal squamous cell carcinoma and an insufficient number of biopsies from recognised abnormalities.

Disclosure of Interest

None Declared

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Introduction

The Autologous Stem Cell Transplantation International Crohn’s Disease (ASTIC) Trial is a randomised controlled trial co-sponsored by ECCO and EBMT and funded by the Broad Foundation that investigates immunoaablation and hemopoietic stem cell transplantation (HSCT) in Crohn’s disease (CD) over 1 year: all patients will have reached this endpoint by April 2013.

Methods

Patients with impaired quality of life due to active CD, despite ≥ 3 immunosuppressive agents all underwent mobilisation (iv cyclophosphamide 4 g/m2 over 2 days then filgrastim10µg/kg/ day) before randomisation to immediate (1 month) or delayed (15 months) HSCT. The conditioning regimen was iv cyclophosphamide 50mg/kg per day for 4 days, anti-thymocyte globulin 2.5 mg/kg/day and methyl prednisolone 1mg/kg on days 3–5. The bone marrow was reconstituted by infusion of an unselected graft of 3–8×10⁶/kg CD34+ stem cells. Clinical (CDAI), endoscopic (SES-CD) quality of life and safety data are compared 1 year after mobilisation alone or after mobilisation and HSCT.

Results

As of Jan 2013, data are available on 34/45 patients. Following mobilisation and HSCT, the CDAI fell from 317 (median, IQR 244–407) to 157 (71–246, n = 17) vs 351 (313–446) and 298 (220–370, n = 17) with mobilisation alone. The aggregate lower GI SES-CD score was 13.0 (8.5–24.5) before and 3.0 (1.5–10.0) after HSCT compared to 13.0 (6.5–15.5) before and 6.5 (3.5–17.8) after mobilisation alone. Over the whole study to end 2012 there were 62 SAEs in 19 patients randomised to early transplantation (3.3 per patient) and 58 in 18 patients randomised to delayed transplantation (3.2 per patient). One patient died following HSCT. Final results of the study will be available to be presented for the first time at BSG 2013.

Conclusion

Immunoaablation and HSCT appears to be an effective treatment for CD that may substantially reduce endoscopic evidence of disease but incurs significant toxicity. The final results of the trial will allow a rational evaluation of the effectiveness and safety of HSCT to be discussed at BSG 2013.

Disclosure of Interest

None Declared
are few explanations for the wide variation in inter-individual TPMT activities in the wild-type range. Bioavailability of the cofactor S-adenosylmethionine (SAM) and RBC age may play a role. The aim of this study was to determine if gender or anaemia influences RBC TPMT activity.

**Methods** We analysed a retrospective cohort of 6,496 RBC TPMT samples (n = 3804 females, n = 2692 males) measured in the PRL since 2004 and correlated enzyme activity with gender and haemoglobin concentrations.

**Results** A greater portion of females exhibited intermediate TPMT activity (13.46%) as compared to males (11.07%). The mean TPMT activity was also significantly lower in females (32.46 pmol/mg Hb/h) versus males (34.13 pmol/mg Hb/h; p = 0.0001, 95% CI 0.7950–1.589). When separated by low, intermediate or normal haemoglobin concentrations, this relationship only remained in patients with normal TPMT activity. TPMT activity was significantly higher in female patients with an Hb < 12g/dl (n = 250, mean TPMT 35.06 pmol/mg Hb/h) versus females with an Hb > 12g/dl (n = 2192, mean TPMT 32.46 pmol/mg Hb/h; p = 0.0001). Similarly TPMT activity was significantly higher in male patients with an Hb < 10g/dl (n = 125, mean TPMT 38.14) versus males with an Hb > 12g/dl (n = 1901, mean TPMT 35.76; p = 0.0001).

**Conclusion** TPMT activity in the wild-type range is lower in females than males, suggesting a post-translational influence on TPMT activity related to gender. Lower levels of SAM have been reported in females, which may explain this observation. Re-appraisal of the concordance between TPMT genotype and phenotype, adjusting for gender is therefore indicated. The finding of higher TPMT activity with anaemia may be due to a younger red cell population in this group. The difference in TPMT activities between patients with or without anaemia is clinically relevant, particularly where the TPMT activity is around the cut-off between intermediate (10–25 pmol/mg Hb/h) and normal (≥26 pmol/mg Hb/h) ranges.

**Disclosure of Interest** None Declared

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**REFERENCES**