N staging were significant different between arm A and C (P = 0.023), arm A and B (P = 0.029). Surgical decision-making in arm A was more accurate than that in arm C (96.2% vs. 80.6%, P = 0.001). Pathological T stage (P < 0.001), N stage (P < 0.001), TNM stage (P < 0.001), serum level of SAA (P = 0.002) and tumor height (P = 0.030) were significantly associated with final surgical procedures.

Conclusions: MPE is a powerful strategy in preoperative staging and more accurate than other available strategies in surgical decision-making for rectal cancer. The final surgical procedures are associated with pathological T, N, TNM stages, which MPE could dependably provide for surgical practice.

6019 POSTER

Phase II XERT trial: Neoadjuvant cetuximab, capecitabine and radiotherapy (RT) in locally advanced resectable rectal cancer

V. Velenik¹, J. Ocvirk², I. Oblak¹, F. Anderluh¹. ¹Oncology Institute, Radiotherapy, Ljubljana, Slovenia; ²Oncology Institute, Medical Oncology, Ljubljana, Slovenia

Background: Preoperative chemoradiotherapy (CRT) with capecitabine is a widely accepted treatment for locally advanced rectal cancer. Tumor response may be further enhanced by the radiosensitizing effect of the EGFR-targeting monoclonal antibody, cetuximab. This prospective, non-randomized, open-label phase II study evaluated the efficacy and safety of cetuximab combined with capecitabine and concurrent RT for locally advanced resectable rectal cancer.

Materials and Methods: Enrolled patients (pts) with MRI-confirmed stage II/III rectal cancer received capecitabine 1250 mg/m² twice daily for 2 wks, followed by IV cetuximab 400 mg/m² at wk 3, then cetuximab 250 mg/m²/wk and capecitabine $825 \, \text{mg/m}^2$ twice daily (including weekends during RT). RT was started at wk 4 at a 45 Gy dose (25×1.8 Gy, 3D conformal technique). Total mesorectal excision was scheduled 4–6 wks after CRT completion with tumor regression grades (TRG) assessed using the Dworak system. The primary endpoint was complete pathologic response (pCR; TRG 4).

Results: A total of 37 pts were evaluable for efficacy and safety; 81% male; median age 55 (range 33-72) yrs. Four pts (10.8%) had T4N2 tumors, 15 pts (40.5%) T3N2, 1 pt (2.7%) T2N2, 13 pts (35.1%) T3N1, 1 pt (2.7%) T2N1, and 3 pts (8.1%) T3N0. The median tumor distance from the anal verge was 6 (range 1–11) cm. A pCR (TRG 4) was reported in 3 pts (8.1%), TRG 3 in 7 pts (18.9%), and T-, N-, and overall downstaging rates were 56.8%, 81.1%, and 73.0%, respectively. The total sphincter preservation rates were 75.7% and 53.0% in 17 pts whose tumors were located ≤5 cm from the anal verge. All pts received 45 Gy RT. Dose reduction or treatment interruption was required for 9 pts (24.3%) due to hypersensitivity reaction (n = 4), grade 3 diarrhea (n = 4), and grade 3 hepatotoxicity (n = 1). Other grade 3 toxicities included dermatitis (n = 6, 16.2%), infection and anorexia (each n = 1, 2.7%). Eleven pts (29.7%) experienced non-fatal perioperative complications; 6 of whom had wound healing problems. One pt (2.7%) with anastomotic leakage and abdominal abscess and 1 pt (2.7%) with incarceration of transversostoma required reoperation, and 34 pts (91.9%) received postoperative chemotherapy. One pt died from sepsis after colonic necrosis and perforation.

Conclusions: Preoperative CRT with cetuximab and capecitabine is safe and feasible. While the pCR rate was in the range previously reported for CRT with capecitabine, a high pathologic downstaging rate was achieved.

6020 POSTER

Large variation between hospital types and pathology laboratories in lymph node evaluation in colon cancer in the Netherlands and its impact on survival, a national population-based study

M. Elferink¹, S. Siesling¹, O. Visser², H.J. Rutten³, J.H.J.M. van Krieken⁴, R.A.E.M. Tollenaar⁵, <u>V.E.P.P. Lemmens⁶</u>. ¹Comprehensive Cancer Centre North East, Department of Research, Enschede, The Netherlands; ²Comprehensive Cancer Centre Amsterdam, Department of Research, Amsterdam, The Netherlands; ³Catharina Hospital, Department of Surgery, Eindhoven, The Netherlands; ⁴Radboud University Nijmegen Medical Centre, Department of Pathology, Nijmegen, The Netherlands; ⁵Leiden University Medical Centre, Department of Surgery, Leiden, The Netherlands; ⁶Comprehensive Cancer Centre South, Department of Research, Eindhoven, The Netherlands

Background: Adequate lymph node evaluation is important for staging and subsequent planning of treatment in patients with colon cancer. Adjuvant chemotherapy should be considered for patients with lymph nodes metastasis. A large variation in the number of evaluated lymph nodes exists. The aim of this study was to describe the influence of patient and tumour characteristics, hospital type and pathology laboratory on adequacy of

nodal examination, and to determine its relationship with stage distribution and survival.

Methods: Data from all patients with colon carcinoma stage I-III (pT1-4NanyM0) who underwent surgical treatment, diagnosed in the period 2000–2006 were retrieved from the Netherlands Cancer Registry. Multilevel logistic analysis was performed to examine the influence of relevant factors on the number of evaluated lymph nodes. The relation between pathology laboratories and stage distribution was assessed. Cox regression analysis was performed to analyse the association between the number of examined lymph nodes, the lymph node ratio and survival.

Results: The number of examined lymph nodes was determined for 29,551 (89%) of the 33,206 tumours. The median number of evaluated lymph nodes was 8, varying from 4 to 15 lymph nodes between pathology laboratories. Median number of lymph node count was negatively associated with volume of pathology laboratory and positively associated with volume of hospital. Females, younger patients, right-sided tumours, tumours with larger depth of invasion, tumours with nodal involvement and patients treated and evaluated in a university medical centre were less likely to have 9 or less lymph nodes evaluated. After adding these factors to the multilevel model, an unexplained variation between the pathology laboratories remained. This variation led to differences in stage distribution between the pathology laboratories, correlating with the median number of evaluated lymph nodes (p < 0.001). With increasing number of evaluated lymph nodes, the risk of death decreased, both in patients with positive lymph nodes and in patients with negative lymph nodes. The risk of death increased with rising lymph node ratio in patients with lymph node

Conclusion: There was a large diversity in lymph node evaluation among patients with colon cancer, with variation between pathology laboratories, leading to differences in stage distribution and being associated with survival. These results indicate that improvement in nodal sampling is needed in many pathology laboratories.

6021 POSTER

Efficacy of chemoradiotherapy for the treatment of locally advanced squamous cell carcinoma of the rectum

M.C. Tronconi¹, R. Doci², M. Bignardi³, F. Sclafani¹, N. Personeni¹, S. Bozzarelli¹, L. Rimassa¹, M. Di Rocco⁴, A. Santoro¹, C. Carnaghi¹.

¹Istituto Clinico Humanitas, Department of Oncology and Hematology, Rozzano Milano, Italy;

²Istituto Clinico Humanitas, Surgical Oncology Unit, Rozzano Milano, Italy;

³Istituto Clinico Humanitas, Radiotherapy and Radiosurgery Unit, Rozzano Milano, Italy;

⁴Istituto Clinico Humanitas, Department of Pathology, Rozzano Milano, Italy

Summary background data: Squamous cell carcinoma (SCC) of the rectum is a rare pathologic entity, accounting for only 0.1% to 0.25% of all rectal cancers. Only 57 cases of colorectal SCC having been reported over a period of more than 60 years. From June 2006 to August 2008, six consecutive patients with squamous cell carcinoma (SCC) of the rectum were treated at the same Institution, according to protocols used for anal SCC.

Methods: All tumours were locally advanced and the clinical stage was T3N0M0 in 2 cases, T3N1M0 in 1, T4N1M0 in 2 and T3N2M1 in 1 case (lung metastases). Five patients received primary chemoradiotherapy and one received chemotherapy only due to previous pelvic irradiation. Radiotherapy was delivered to a target volume including primary tumor, internal and external iliac nodes and mesorectum. The minimum dose was 5.040 cGy; a boost dose to the primary tumor up to 5.940 cGy was given to three patients. Radiotherapy was associated with these chemotherapy schedules: 4 patients received 3 cycles of 5-fluorouracil (5-FU) (1000 mg/m²/day continuous infusion on days 1-4) and cisplatin (CDDP) (80 mg/m² on day 1) repeated every 3 weeks; 1 patient received 6 weeks of continuous infusion of 5-FU (225 mg/m² daily). One patient received 2 cycles of 5-FU (1000 mg/m²/day continuous infusion for 4 consecutive days) in combination with mitomycin-c (10 mg/m² on day 1) every 4 weeks; this patient received 2 additional cycles of chemotherapy at the end of radiotherapy because of the presence of metastases. The patient treated with chemotherapy alone received 2 preoperative cycles of 5-FU and CDDP repeated every 3 weeks. All patients concluded their treatment without a diverting enterostomy.

Results: Complete clinical response (CR) was achieved in 3 patients and partial response (PR) in 2. Disease stabilization (SD) was obtained in 1 case and no patients showed progressive disease (PD). Surgery was performed in 1 patient with PR and in 1 with SD. The patient with lung metastases received 4 courses of systemic chemotherapy. As of the last follow-up (FU) patients with CR were free of recurrence at 17 (cT4N1), 31 and 28 months (cT3N0). At a median FU of 18 months all patients are alive and all but the patient with metastasis are disease free.