

only target tumours is to be evaluated. As a consequence, the ability to make meaningful inter-patient, inter-tumour or inter-time-point comparisons can be challenging. This abstract details the **Neo-tAnGo** group's methodological approaches to these challenges.

**Neo-tAnGo** is a national 800-patient breast cancer trial assessing neo-adjuvant chemotherapy in patients with T2 tumours or above ( $>2$  cm). A secondary endpoint is radiological tumour response, assessing treatment effect in terms of change in tumour size. Mammograms and/or ultrasounds and/or MRI scans are undertaken at baseline and up to two breast and one axillary lesion are identified as target lesions. Scans are then repeated after 4 and 8 cycles of neo-adjuvant chemotherapy. At each time-point and for each target lesion, the scan type(s) and largest diameter(s) observed (maximum of two) are reported, along with a clinical judgement of response.

The frequency of scan type(s) undertaken, number of dimensions reported, resulting incidence of valid comparisons, clinical response categorisation, statistical approaches adopted and general observance of the RECIST criteria will be reported on the 550 patients who have completed chemotherapy.

#### **O-111 Functional evaluation of stem cells in breast tumours**

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**Introduction and Aim:** Side population (SP) cells have been discovered in many tissues, several cancer cell lines and solid tumours including brain and breast. These SP cells exhibit many features of stem cells. Their role in normal breast development and breast cancer is uncertain and we hypothesise that SP cells have independent functional status in relation to transporter mechanisms compared to the main population (MP) cells.

**Methods:** SP cells were identified using dual wavelength flow cytometry combined with Hoechst 33342 dye efflux. SP cells exclude Hoechst dye via an ABCG2 half transporter, ABCG2. We examined 6 breast cancer cell lines for the presence of SP cells using immunocytochemistry for stem cell and breast cancer cell markers (ABCG2). In addition we exposed SP and MP cells to mitoxantrone and determined the effect on cell viability.

**Results:** SP cells were found only in MCF-7 and MFM-223 cell lines. Both populations were found to express stem cell and typical breast cancer cell markers. However, it was only the population of SP cells which were capable of pumping out Hoechst through the ABCG2 transporter mechanism. Real-Time PCR arrays for cancer drug resistance and metabolism genes showed an upregulation of Cytochrome P450 transporters, ABCB1 and PPARG in SP cells when compared to MP cells.

**Conclusion:** Our results show a differential expression of cancer drug resistance and metabolism genes between SP cells and MP cells suggesting that the former may have an important role in chemotherapy resistance within the tumour microenvironment.

#### **O-112 The Cambridge Breast Intensity Modulated Radiotherapy Trial: dosimetry results for 1089 patients**

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**Introduction:** 2D radiotherapy (RT) breast plans can lead to substantial dose inhomogeneities, which may cause increased normal tissue toxicity. We report the dosimetry results to date of our NCRN-adopted randomised

trial comparing standard 2D RT with IMRT (intensity-modulated RT).

**Methods:** Following 3D imaging, a standard plan was produced for all patients. Plans were classified as having significant dose inhomogeneities if they exceed the upper limit of ICRU 50 ( $>107\%$  of prescribed dose). Those patients with satisfactory dose homogeneity were treated with standard RT. Patients with significant dose inhomogeneities were randomised to standard breast RT or IMRT. The intervention group were re-planned with forward-planned IMRT.

**Results:** 317/1089 (29%) had acceptable dose homogeneity with standard 2D RT. The mean difference in breast volume between randomised and non-randomised patients was  $594 \text{ cm}^3$  ( $p < 0.0001$ , 95% CI  $526\text{--}662 \text{ cm}^3$ ). However, there was considerable overlap in the range of breast volumes between the 2 groups. The mean improvement in volumes  $>107\%$  for IMRT plans was  $34.3 \text{ cm}^3$  ( $p < 0.0001$ , 95% CI  $25.7\text{--}43.0 \text{ cm}^3$ ). The mean improvement in volumes  $<95\%$  for IMRT plans was  $47.4 \text{ cm}^3$  ( $p = 0.0001$ , 95% CI  $32.1\text{--}62.7 \text{ cm}^3$ ).

**Conclusion:** This trial, which closes to recruitment in June 2007, will quantify the clinical benefit of breast IMRT, in a patient group who consume 30% of RT resources. It will also provide DNA samples linked with high quality clinical outcome data, for a translational study investigating individual patient variation in normal tissue toxicity. This will bring us closer to the ultimate aim of individualised RT based on patient's genetics.

#### **O-113 Impact of a higher radiation dose on local control and survival in breast-conserving therapy of early breast cancer: 10-year results of the Randomized Boost Versus No Boost EORTC 22881-10882 Trial**

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**Purpose:** To investigate the long-term impact of a boost dose on local control, fibrosis, and overall survival for patients with stage I–II breast cancer undergoing breast-conserving therapy.

**Patients and Methods:** A total of 5,318 patients with microscopically complete excision followed by whole-breast irradiation of 50 Gy were randomly assigned to a boost dose of 16 Gy or no boost dose (median follow-up: 10.8 years, median age 55 years).

**Results:** Local recurrence was reported as the first treatment failure in 278 patients with no boost versus 165 patients with boost at 10 years, the cumulative incidence of local recurrence was 10.2% versus 6.2% for the no boost and the boost group, respectively. The hazard ratio of local recurrence was 0.59 in favor of the boost, with no statistically significant interaction per age group. The absolute risk reduction at 10 years per age group was the largest in patients  $\leq 40$  years of age: 23.9% to 13.5%. Severe fibrosis was statistically significantly increased in the boost group (4.4% versus 1.6%). Survival at 10 years was 82% in both arms.

**Conclusion:** A boost dose of 16 Gy led to improved local control in all age groups, more severe fibrosis, but no difference in survival.