

### **E3. What is Hot in breast cancer radiation oncology in 2012?**

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#### **Partial Breast Irradiation**

Most patients with early breast cancer can be treated with breast-conserving therapy (BCT) consisting of a lumpectomy and axillary staging followed by whole-breast irradiation [1,2]. The addition of a boost to the primary tumour bed further reduces the local relapse rate, with a relevant benefit in patients with risk factors [3].

Because of the inconvenience linked to protracted radiation therapy (RT) fractionation schedules, many women living in remote areas cannot benefit from BCT [4]. Two approaches have been used to diminish this drawback, the first being hypofractionation, and the second accelerated partial-breast irradiation (APBI) [5,6]. The latter is built on the finding that the recurrence pattern after BCT is generally confined to the vicinity of the primary tumour and on the expectation of a lower rate of adverse effects when large doses of RT are delivered in a short time to a part of the breast only. Over recent years, several techniques of APBI have been developed and tested [7,8]. From the recent and ongoing prospective randomised clinical trials, the only results published are those of a small trial with multicatheter brachytherapy also including a conventionally fractionated external electron beam technique with a median follow-up of 66 months, and the results of a trial using a dedicated 50 kV applicator to deliver a single intraoperative dose to the lumpectomy cavity after a median follow-up of only slightly more than 2 years [9,10]. In the presence of the available (mostly non-randomised) evidence showing that APBI can be both safe and effective, both the American Society for Radiation Oncology (ASTRO) and Groupe Européen de Curiethérapie – European Society for Radiotherapy and Oncology (GEC-ESTRO) published consensus guidelines describing patient categories that might be suitable for APBI [11,12]. However, as historically required levels of evidence are still unavailable, we should continue to support participation in clinical trials comparing APBI with conventional BCT.

on a complete set of imaging data is used to prepare sophisticated image-guided individualised treatments to adequately cover the target volumes while simultaneously limiting the dose to the normal structures. Quality assurance tools and guidelines are progressively being applied in daily clinical practice.

Among the first to evaluate the influence of dose homogenisation on cosmetic outcome after BCT were Donovan and colleagues who randomised patients between standard RT and RT using three-dimensional dose optimisation techniques based on either physical compensators or step-and-shoot multileaf collimator (MLC) segmented fields. The improved dose homogeneity with a decrease in the breast volume receiving >105% of the prescribed dose was associated with a decreased change in breast appearance during follow-up as scored by photographic and clinical assessment [13]. Several technical solutions depending on the target volumes to be irradiated (breast; boost; lymph-node areas) are proposed, also demonstrating that an optimal compromise between dose coverage, homogeneity, and sparing normal tissues does not always require the use of the most advanced techniques [14,15,16]. Depending on the individual patient's anatomy, position and treatment technique should be individualised to obtain optimal dose coverage and organ sparing [17,18]. A significant reduction in the volume of irradiated lung and heart tissue can be obtained with the use of breath-hold and gating techniques [19,20].

All these improvements have led to a decrease in the dose to normal structures, which is expected to further increase the net benefit for the patients [21]. A prerequisite for modern RT planning starts with adequate volume delineation, for which there still remains a large source of variation and uncertainty [22]. Although it remains difficult to prove that optimised RT techniques have a direct effect in themselves, we should accept that they have contributed to the substantial improvement in the outcome of breast cancer patients in terms of disease control, survival, and quality of life [23].

#### **Modern Radiotherapy Techniques**

Preparation and delivery of RT has changed considerably over the years. Nowadays, fully virtual simulation based

#### **Conflict of interest statement**

The authors declare that they do not have any conflict of interest related to the work presented in this abstract.

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## References

- [1] Voogd AC, Nielsen M, Peterse JL, et al. Differences in risk factors for local and distant recurrence after breast-conserving therapy or mastectomy for stage I and II breast cancer: pooled results of two large European randomized trials. *J Clin Oncol* 2001; **19**: 1688–97.
- [2] Fisher B, Anderson S, Bryant J, et al. Twenty-year follow-up of a randomized trial comparing total mastectomy, lumpectomy, and lumpectomy plus irradiation for the treatment of invasive breast cancer. *N Engl J Med* 2002; **347**: 1233–41.
- [3] Van Werkhoven E, Hart G, Van Tinteren H, et al. Nomogram to predict ipsilateral breast relapse based on pathology review from the EORTC 22881–10882 boost versus no boost trial. *Radiother Oncol* 2011; **100**: 101–7.
- [4] Athas WF, Adams-Cameron M, Hunt WC, Amir-Fazli A, Key CR. Travel distance to radiation therapy and receipt of radiotherapy following breast-conserving surgery. *J Natl Cancer Inst* 2000; **92**: 269–71.
- [5] Whelan TJ, Pignol JP, Levine MN, et al. Long-term results of hypofractionated radiation therapy for breast cancer. *N Engl J Med* 2010; **362**: 513–20.
- [6] START Trialists' Group. The UK Standardisation of Breast Radiotherapy (START) Trial B of radiotherapy hypofractionation for treatment of early breast cancer: a randomised trial. *Lancet* 2008; **371**: 1098–107.
- [7] Offersen BV, Overgaard M, Kroman N, Overgaard J. Accelerated partial breast irradiation as part of breast conserving therapy of early breast carcinoma: a systematic review. *Radiother Oncol* 2009; **90**: 1–13.
- [8] Mannino M, Yarnold J. Accelerated partial breast irradiation trials: diversity in rationale and design. *Radiother Oncol* 2009; **91**: 16–22.
- [9] Polgár C, Fodor J, Major T, et al. Breast-conserving treatment with partial or whole breast irradiation for low-risk invasive breast carcinoma – 5-year results of a randomized trial. *Int J Radiat Oncol Biol Phys* 2007; **69**: 694–702.
- [10] Vaidya JS, Joseph DJ, Tobias JS, et al. Targeted intraoperative radiotherapy versus whole breast radiotherapy for breast cancer (TARGIT-A trial): an international, prospective, randomised, non-inferiority phase 3 trial. *Lancet* 2010; **376**: 91–102.
- [11] Smith BD, Arthur DW, Buchholz TA, et al. Accelerated partial breast irradiation consensus statement from the American Society for Radiation Oncology (ASTRO). *Int J Radiat Oncol Biol Phys* 2009; **74**: 987–1001.
- [12] Polgár C, Van Limbergen E, Pötter R, et al. Patient selection for accelerated partial-breast irradiation (APBI) after breast-conserving surgery: recommendations of the Groupe Européen de Curiethérapie-European Society for Therapeutic Radiology and Oncology (GEC-ESTRO) breast cancer working group based on clinical evidence (2009). *Radiother Oncol* 2010; **94**: 264–73.
- [13] Donovan E, Bleakley N, Denholm E, Evans P, et al. Randomised trial of standard 2D radiotherapy (RT) versus intensity modulated radiotherapy (IMRT) in patients prescribed breast radiotherapy. *Radiother Oncol* 2007; **82**: 254–64.
- [14] Fournier-Bidoz N, Kirova Y, Campana F, et al. Technique alternatives for breast radiation oncology: Conventional radiation therapy to tomotherapy. *J Med Phys* 2009; **34**: 149–52.
- [15] van der Laan HP, Dolsma WV, Schilstra C, et al. Limited benefit of inversely optimised intensity modulation in breast conserving radiotherapy with simultaneously integrated boost. *Radiother Oncol* 2010; **94**: 307–12.
- [16] Schubert LK, Gondi V, Sengbusch E, et al. Dosimetric comparison of left-sided whole breast irradiation with 3DCRT, forward-planned IMRT, inverse-planned IMRT, helical tomotherapy, and topotherapy. *Radiother Oncol* 2011; **100**: 241–6.
- [17] Campana F, Kirova YM, Rosenwald JC, et al. Breast radiotherapy in the lateral decubitus position: A technique to prevent lung and heart irradiation. *Int J Radiat Oncol Biol Phys* 2005; **61**: 1348–54.
- [18] Kirby AM, Evans PM, Donovan EM, et al. Prone versus supine positioning for whole and partial-breast radiotherapy: a comparison on non-target tissue dosimetry. *Radiother Oncol* 2010; **96**: 178–84.
- [19] Korreman SS, Pedersen AN, Aarup LR, et al. Reduction of cardiac and pulmonary complication probabilities after breathing adapted radiotherapy for breast cancer. *Int J Radiat Oncol Biol Phys* 2006; **65**: 1375–80.
- [20] Topolnjak R, Sonke JJ, Nijkamp J, et al. Breast patient setup error assessment: comparison of electronic portal image devices and cone-beam computed tomography matching results. *Int J Radiat Oncol Biol Phys* 2010; **78**: 1235–43.
- [21] Taylor CW, Brønnum D, Darby SC, et al. Cardiac dose estimates from Danish and Swedish breast cancer radiotherapy during 1977–2001. *Radiother Oncol* 2011; **100**: 176–83.
- [22] Castro Pena P, Kirova YM, Campana F, et al. Anatomical, clinical and radiological delineation of target volumes in breast cancer radiotherapy planning: individual variability, questions and answers. *Br J Radiol* 2009; **82**: 595–9.
- [23] Poortmans P, Aznar M, Bartelink H. Quality Indicators for Breast Cancer: Revisiting Historical Evidence in the Context of Technology Changes. *Semin Radiat Oncol* doi:10.1016/j.semradonc.2011.09.007.