Pioneering Single-Dose Radiotherapy for Breast Cancer Treatment

breast cancer therapy that requires just one shot of radiotherapy is as effective as traditional radiotherapy, and avoids potential damage to nearby organs, according to a paper by University College London (UCL) experts. The results, published in the *British Journal of Cancer*, mean that eight out of 10 patients who receive the treatment, TARGIT-IORT, will not need a long course of post-operative external beam radiotherapy (EBRT) (2021; https://doi.org/10.1038/s41416-021-01440-8). These results strengthen and expand previously published outcomes.

Patients who received the treatment are less likely to go on to experience fatal cardiovascular disease such as heart attacks, lung problems, or other cancers. As well as avoiding scattered radiation from EBRT that can damage nearby vital organs, delivering TARGIT-IORT during the lumpectomy procedure seems to lower the likelihood of death if patients do go on to develop cardiovascular disease, protecting in a drug-like manner. This was the case even when EBRT was also given post-operatively, and is thought to be because the treatment changes the microenvironment in the lumpectomy wound.

The researchers say that delivering radiation immediately to the site where the tumor was can reduce the adverse effects of surgical trauma, make the site less conducive for cancer growth, and could have an abscopal effect. This is where a treatment such as radiotherapy has a positive effect on tissue away from the operation site, which is increasingly recognized as a beneficial immunological action.

Previous studies have shown that the treatment has fewer radiation-related side effects compared with conventional whole breast radiotherapy, with less pain, a superior cosmetic outcome with fewer changes to the breast as a whole, and a better quality of life.

"With TARGIT-IORT, women can have their surgery and radiation treatment for breast cancer at the same time," noted lead author Jayant Vaidya, MBBS, MS, DNB, at UCL Surgery & Interventional Science. "This reduces the amount of time spent in hospital and enables women to recover more quickly, meaning they can get back to their lives more quickly."

"With TARGIT-IORT, the majority of patients presenting with early localized breast cancer will never need any further radiotherapy. They will avoid all the side effects of whole breast radiotherapy."

—Jeffrey S. Tobias, MD, FRCP, at the University College London

TARGIT-IORT is delivered immediately after tumor removal, and under the same anesthetic, via a small ball-shaped device placed inside the breast, directly where the cancer had been. The single-dose treatment lasts for around 20-30 minutes and replaces the need for extra hospital visits in eight out of 10 cases. Further tumor subgroup analysis also found that there was a significant overall survival benefit with TARGIT-IORT in patients with grade 1 or 2 cancer.

"These new results make it clear that the TARGIT-IORT is effective in all tumor subgroups of invasive duct cancer, the most common type of breast cancer," Vaidya added. "Our new online tool can help clini-



cians make a decision about additional radiotherapy (recommended in a small proportion of cases) for each individual patient.

"The finding that fewer deaths are from the avoidance of scattered radiation and the possible abscopal effect of TARGI-IORT is important and should fuel further research, opening doors to new treatments," noted.

Study Details

For the clinical trial, which started in March 2000, 2,298 women aged 45 or over with invasive breast cancer and a tumor up to 3.5 cm in diameter were randomly assigned to receive either TARGIT-IORT during lumpectomy or post-operative EBRT.

The trial was designed and run from UCL, involving 32 hospitals and medical centers in 10 countries: the UK, France, Germany, Italy, Norway, Poland, Switzerland, the US, Canada, and Australia.

"These results are the highest level of evidence proving not only the effectiveness of TARGIT-IORT, but confirming that it avoids deaths from other causes," noted Michael Baum, MB, ChB, FRCS, ChM, Professor at UCL Surgery & Interventional Science. "The new data is biologically very interesting and the new tools will make its application in routine clinical practice much easier. I am pleased that it will benefit thousands of breast cancer patients around the world."

Jeffrey S. Tobias, MD, FRCP, Professor of Clinical Oncology at UCL and UCLH, shared his thoughts: "With TARGIT-IORT, the majority of patients presenting with early localized breast cancer will never need any further radiotherapy. They will avoid all the side effects of whole breast radiotherapy. The chance of remaining free of local recurrence (in the breast itself) is the same as with traditional treatment, but our new analysis shows that even if they do get a local relapse, it will not detract from an excellent prognosis—as good as not having a relapse—a rather different state of affairs from the more serious outlook if this were to happen after EBRT."

To date, 45,000 patients in 260 centers in 38 countries have received TARGIT-IORT. The clinicians hope that following the latest results, more patients can be offered the treatment both in the UK and around the world instead of EBRT.

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