

# **Response to Comment on - Long-term risks of secondary cancers for various whole and partial breast irradiation techniques** Pignol, J.P.; Hoekstra, N.

1 ignoi, j.i ., 1100itotiu, 1

## Citation

Pignol, J. P., & Hoekstra, N. (2020). Response to Comment on - Long-term risks of secondary cancers for various whole and partial breast irradiation techniques, *142*, 261-261. doi:10.1016/j.radonc.2019.11.007

Version:Publisher's VersionLicense:Creative Commons CC BY 4.0 licenseDownloaded from:https://hdl.handle.net/1887/3566422

**Note:** To cite this publication please use the final published version (if applicable).

#### Radiotherapy and Oncology 142 (2020) 261



Contents lists available at ScienceDirect

## Radiotherapy and Oncology

journal homepage: www.thegreenjournal.com



### Letter to the Editor

### Response to Comment on - Long-term risks of secondary cancers for various whole and partial breast irradiation techniques

We would like to thank and welcome the useful commentary on the article ["Long-term risks of secondary cancers for various whole and partial breast irradiation techniques" published in Radiotherapy and Oncology, volume 128, issue 3, pages 428–433], [1,2] regarding the limitations using the BEIR model [3]. It is indeed exact that the model is mainly addressing low-dose exposures and since it is a linear model it fails to incorporate a "saturation" effect beyond a threshold dose it may overestimates the rate of secondary cancers. This is well illustrated in Schneider report on the OEM model where for secondary lung cancer the model is quite linear up to 4-5 Gy then shows an inflexion with a clear saturation after 10 Gy [4]. Since all the doses we measured in the phantom were below 3 Gy, we believe the BEIR model would yield a fair estimation of secondary cancer risk. Also, we believe that the calculation was conservative as our calculated secondary cancer rates appear not overestimated but slightly underestimated. Both Grantzau meta-analysis and the EBCTCG reported secondary cancers rates after breast radiotherapy slightly higher than the one we calculated [5.6].

The OED model cited in the commentary is a powerful and sophisticated one, as it refines the BEIR model including dose distribution heterogeneity. The challenge being that it assumes all the mechanisms of response to an irradiation, at the molecular, cellular, and physiological, which occur over several decades, are fully understood and included in a mathematical model. It also assumes that each organ would be constituted by single elementary units behaving similarly.

While those models are very useful, it is critically important to acknowledge that they remain theoretical and cannot accurately calculate an absolute risk for a given irradiation situation. But they can yield rough risk estimates and enable a fair comparison between radiotherapy techniques. It is hence unsure that using another model than the BEIR VII would change the conclusions that (i) lung is the main organ at risk of secondary cancer, (ii) there is a prolong lag of time for the development of secondary cancers, (iii) the rough estimate risk of secondary radiation induced lung cancer death after 30 years largely overpass the risk of cardiac mortality so it is important to disclose this discussing a radiation treatment with a young women with highly curable disease such as DCIS and to take active measure to reduce the amount of scattered dose in the lung; and finally (iv) comparing breast irradiation techniques leads to very different risks of secondary lung cancers favoring partial breast irradiation.

#### References

- [1] The commentary.
- [2] Hoekstra N, Fleury E, Merino Lara TR, et al. Long-term risks of secondary cancer for various whole and partial breast irradiation techniques. Radiother Oncol 2018;128:428-33.
- [3] National Research Council. Health risks from exposure to low levels of ionizing radiation: BEIR VII phase 2. National Academies Press: 2006
- [4] Schneider U, Sumila M, Robotka J. Site-specific dose-response relationships for cancer induction from the combined Japanese A-bomb and Hodgkin cohorts for doses relevant to radiotherapy. Theor Biol Med Model 2011;8:27. https://doi. org/10.1186/1742-4682-8-27
- [5] Grantzau T, Overgaard J. Risk of second non-breast cancer among patients treated with and without postoperative radiotherapy for primary breast cancer: a systematic review and meta-analysis of population-based studies including 522,739 patients. Radiother Oncol 2016;121:402-13.
- [6] Taylor C, Correa C, Duane FK, et al. Estimating the risks of breast cancer radiotherapy: evidence from modern radiation doses to the lungs and heart and from previous randomized trials. J Clin Oncol 2017;35:1641-9.

Jean-Philippe Pignol\*

Department of Radiation Oncology, Dalhousie University, Nova Scotia, Canada

\* Corresponding author at: Department of Radiation Oncology, Dalhousie University 5820 University Avenue, Halifax, Nova Scotia B3H 1V7, Canada. E-mail address: Jean-Philippe.Pignol@dal.ca

Nienke Hoekstra

Department of Radiation Oncology, Erasmus MC, Rotterdam, The Netherlands Received 23 October 2018 Received in revised form 5 November 2019 Accepted 5 November 2019

Available online 27 November 2019

<sup>\*</sup> DOI of original article: https://doi.org/10.1016/j.radonc.2018.10.031