

Enhancing prostate cancer recurrence prediction using an atlas-based tumour control probability model

Kazi Ridita Mahtaba¹

Martin Ebert², Jeremy Booth³, Leyla Moghaddasi³, Rob Finnegan⁴, Yutong Zhao⁵, Burhan Javed⁶, George Hruby³ and Annette Haworth¹

¹ Institute of Medical Physics, The University of Sydney

² Sir Charles Gairdner Hospital

³ Northern Sydney Cancer Centre, Royal North Shore Hospital

⁴ Northern Sydney Cancer Centre

⁵ School of Physics, Mathematics and Computing, University of Western Australia

⁶ Genesis Care, Rockhampton Hospital

Enhancing prostate cancer recurrence prediction using an atlas-based tumour control probability model

Kazi Ridita Mahtaba¹, Martin A. Ebert^{2,3,4}, Jeremy Booth^{1,5}, Leyla Moghaddasi⁵, Robert Finnegan⁵, Yutong Zhao², Burhan Javed⁶, George Hruby⁵, Annette Haworth¹

¹Institute of Medical Physics, The University of Sydney, Camperdown, New South Wales, Australia

²School of Physics, Mathematics and Computing, University of Western Australia, Crawley, Western Australia, Australia

³Department of Radiation Oncology, Sir Charles Gairdner Hospital, Perth, Western Australia, Australia

⁴Centre for Advanced Technologies in Cancer Research, Perth, Western Australia, Australia

⁵Northern Sydney Cancer Centre, Royal North Shore Hospital, St Leonards, New South Wales, Australia

⁶Genesis Care, Rockhampton Hospital, Rockhampton City, Queensland, Australia

Abstract:

Aim:

This study aimed to evaluate an atlas-based tumour control probability (TCP) model^{1,2} for predicting prostate cancer (PCa) recurrence by integrating patient-specific primary radiotherapy and histopathology data. The goal was to investigate segment-wise adjustments to TCP model parameters to derive realistic TCP estimates and improve recurrence prediction.

Method:

Nine patients with available histopathology reports and primary radiotherapy dose-fractionation data were selected from an ethics-approved study (NCT03073278) on re-irradiation of locally recurrent prostate cancer. A population-based biological atlas³, consisting of voxelised cell density and tumour probability models, was deformably registered to each patient's prostate and segmented based on individual histopathology. Radiosensitivity parameters, including a single PCa grade-independent α/β ratio, four Gleason Pattern (GP)-dependent α values, and nine Gleason Score (GS)-dependent α/β ratios, were derived from a separate cohort using numerical optimization². The TCP model calculated overall TCP and

generated voxel-wise TCP maps. Three segment-wise parameter adjustment approaches (cell density only, with GP-dependent α , and with GS-dependent α/β) were compared to a baseline model without adjustments, with recurrent tumour contours overlaid on TCP maps to assess alignment with lower TCP regions.

Results:

The combination of segment-wise cell density with GS-dependent α/β adjustments demonstrated the strongest predictive performance (Figure1). All nine recurrent PCa patients showed a significant overall TCP reduction, and in seven patients, regions of low TCP corresponded with the relapsed gross tumour volume. Voxel-level histogram analysis and statistically significant volume-weighted TCP differences between tumour and non-tumour regions (Wilcoxon signed-rank test, $p = 0.003$) further supported this outcome. In contrast, adjustments based on GP-dependent α with cell density failed to effectively predict recurrence, while cell density alone provided only moderate improvement.

Conclusion:

The atlas-based TCP model, incorporating patient-specific histopathology, showed potential to enable personalized, biologically-based dose prescriptions.

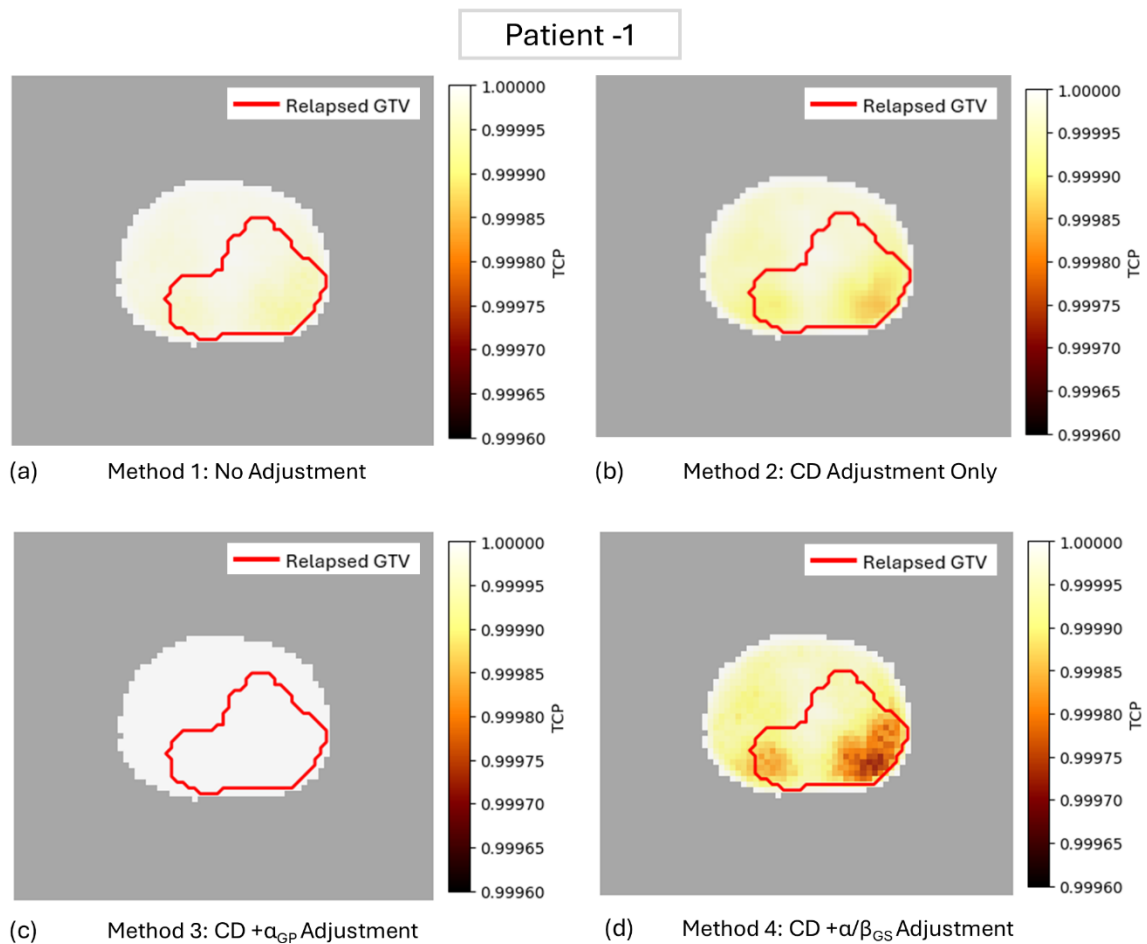


Figure 1: Axial TCP maps for Patient-1 using four methods: (a) No parameter adjustment, (b) Cell density adjustment only, (c) Cell density + GP-dependent α adjustment, and (d) Cell density + GS-dependent α/β adjustment. Warmer regions indicate lower TCP, with the red contour outlining the relapsed Gross Tumor Volume (GTV).

References:

1. Her EJ, Reynolds HM, Mears C, et al. Radiobiological parameters in a tumour control probability model for prostate cancer LDR brachytherapy. *Phys Med Biol.* 2018;63(13):135011. doi:10.1088/1361-6560/aac814
2. Zhao Y, Haworth A, Reynolds HM, et al. Patient-specific voxel-level dose prescription for prostate cancer radiotherapy considering tumor cell density and grade distribution. *Med Phys.* 2023;50(6):3746-3761. doi:10.1002/mp.16264
3. Finnegan RN, Reynolds HM, Ebert MA, et al. A statistical, voxelised model of prostate cancer for biologically optimised radiotherapy. *Phys Imaging Radiat Oncol.* 2022;21:136-145. doi:10.1016/j.phro.2022.02.011