

Preliminary Results Simulating Direct and Indirect Neutron-Induced DNA Damage With **Repair Mechanisms**

Nicolas Desjardins¹ and John Kildea²

¹Dept. of Physics, McGill University, Montreal, QC, H4A 3J1, Canada ²Medical Physics Unit, McGill University, Montreal, QC, H4A 3J1, Canada

INTRODUCTION

- High-energy radiotherapy patients (>10 MeV) and astronauts in deep space are subject to neutron radiation which can lead to carcinogenesis [1-2].
- The risks associated with neutron radiation are **energy** dependent [3-4].
- Previous studies have used Monte-Carlo simulations to study this energy dependence but **none included repair**

OBJECTIVES

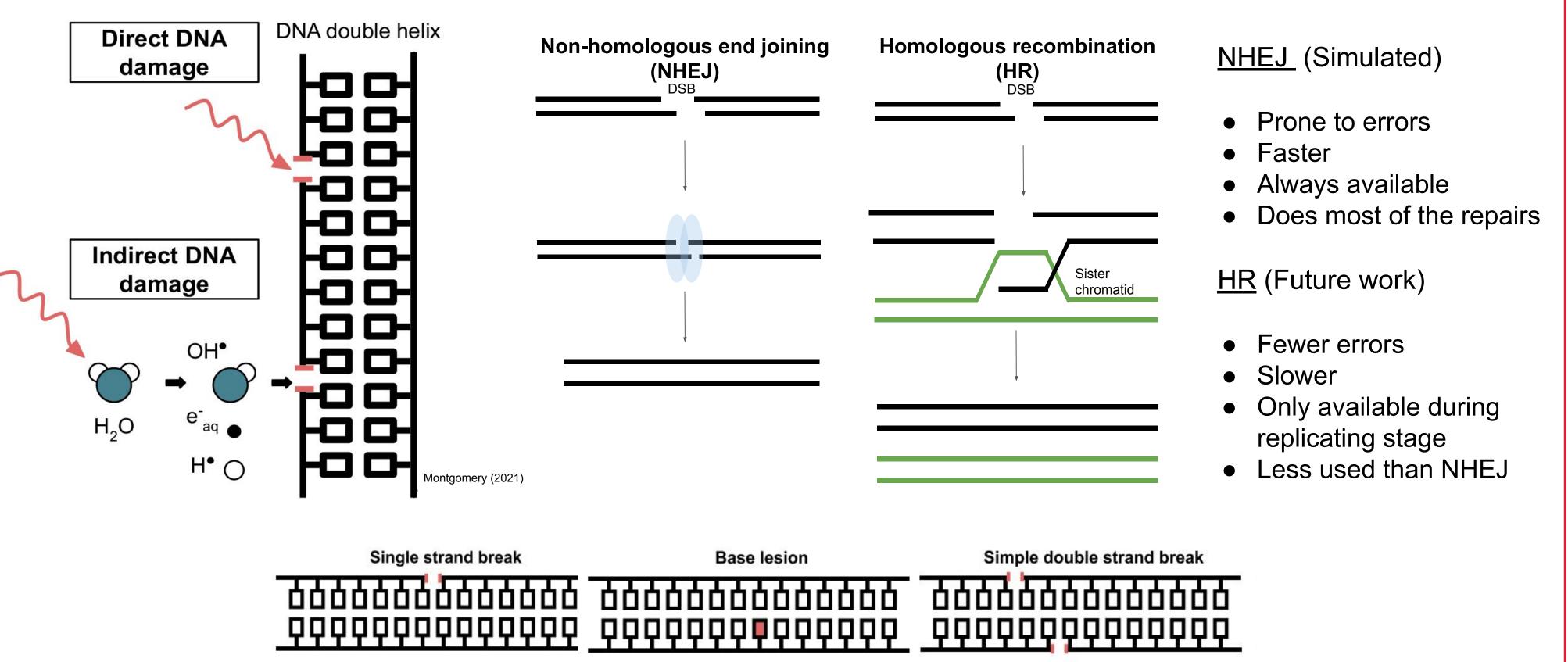
- Overall goal: Better understand how neutrons cause cancer.
- <u>Specific goal</u>: Estimate the neutron relative biological effectiveness by simulating DNA damage and repair mechanisms.

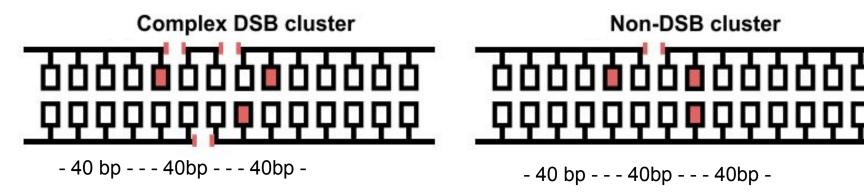
The relative biological effectiveness tells us how the effects of a given radiation compare to those of a reference radiation, which is typically 250 keV photons.



METHODS

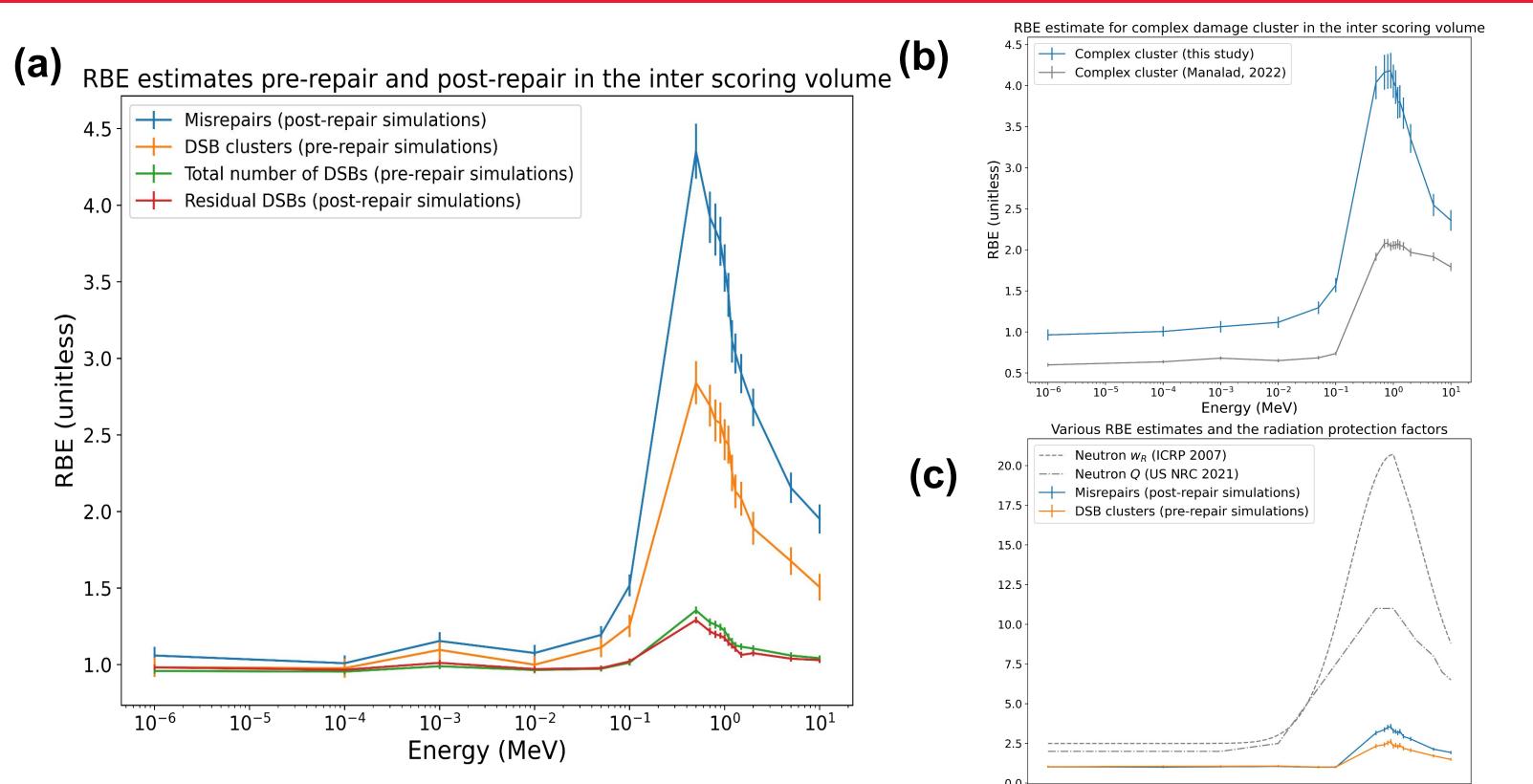
- Our team had already acquired data on the secondary particles of neutrons using Geant4 simulations [7].
- Using those data, direct and indirect damage were simulated on a DNA model by TOPAS-nBio using the TOPAS toolkit [8-9].
- Non-homologous end joining was simulated using the DaMaRiS framework adapted to TOPAS [10]. Homologous recombination will be included in future work





CURRENT RESULTS

- **Result (a):** The estimated RBE peak is more significant for mis-repairs than it is for residual DSBs post-repair. This is to be expected given that mis-repairs are believed to be caused by DSB clusters.
- **Result (b):** The pre-repair damage results from this study compared to the results obtained from a similar study using a different DNA model. The differences may be due to different physical constructors and DNA models.
- **Results (c):** The post-repair RBE estimates compared to the published neutron quality factors. The quality factors are judgment values used to evaluate the risk associated with neutron radiation.



— 10 bp —

CONCLUSION

- The addition of the NHEJ mechanism to Monte-Carlo simulations seems to increase the peak of our estimated RBE compared to DSB clusters.
- These results are preliminary and more analysis is required to conclude anything significant.
- Future work will incorporate homologous recombination.

REFERENCES

[1] Kry et al. (2017) doi: 10.1002/mp.12462 [2] Heilbronn et al. (2005) doi: 10.1093/rpd/nci033 [3] ICRP (2007) ICRP 103: The 2007 Recommendations of the ICRP

[4] US NRC (2021) Units of Radiation Dose

[5] Montgomery et al. (2021) doi: 10.1088/1361-6560/ac2998 [6] Baiocco et al. (2016) doi: 10.1038/srep34033 [7] Lund et al. (2020) doi: 10.1016/j.ejmp.2020.04.001 [8] Perl et al. (2012) doi: 10.1118/1.4758060 [9] Schuemann et al. (2018) doi: 10.1667/RR15226.1 [10] Warmenhoven et al. (2020) doi: 10.1016/j.dnarep.2019.102743 [11] Manalad et al. (2022) currently under revision

ACKNOWLEDGMENTS

Thank you to James Manalad, Felix Mathew, and the whole NICE-ROKS team!

The Neutron-Induced Carcinogenic Effects (NICE) research group led by Dr. John Kildea (kildealab.com) is funded by the Natural Sciences and Engineering Research Council of Canada (NSERC), the Canada Foundation for Innovation (CFI), and the Canadian Space Agency (CSA).

