

## Tridimensional cellular models of prostate cancer for the evaluation of copper-64 chloride as a theranostic agent



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### BACKGROUND

Despite medical advances that contributed to the decline of prostate's cancer (PCa) mortality, it still remains incurable in advanced stages.



Important to develop new approaches to ensure <u>early diagnosis</u> and <u>treatment</u> of chemoresistant tumours.

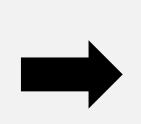


Radiopharmaceuticals have the potential to be used in <u>diagnosis</u>, <u>therapy</u> or simultaneously for both purposes – <u>theranostics</u>.

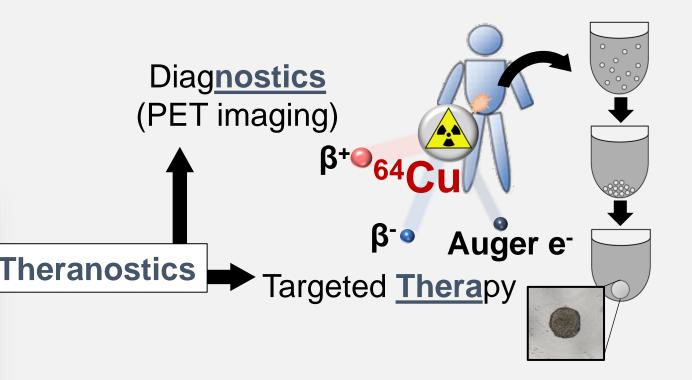
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**▼** 2.8 MBq

Time (days)



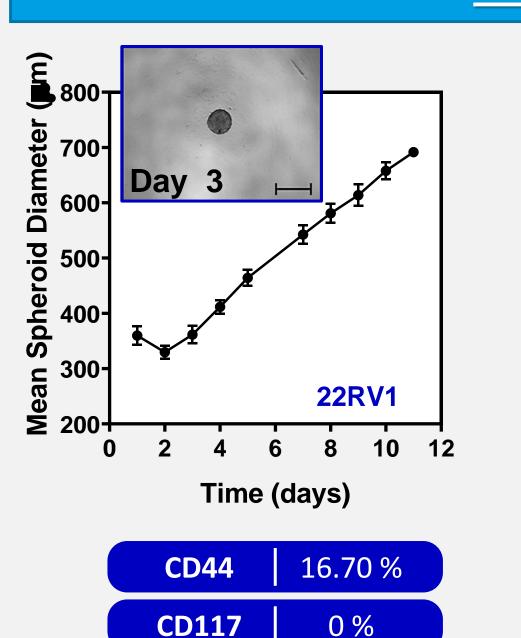
64CuCl<sub>2</sub> has the potential to induce damage in monolayer cultured PCa cell lines, while bearing minimal side effects in non-tumoral cells. [1]

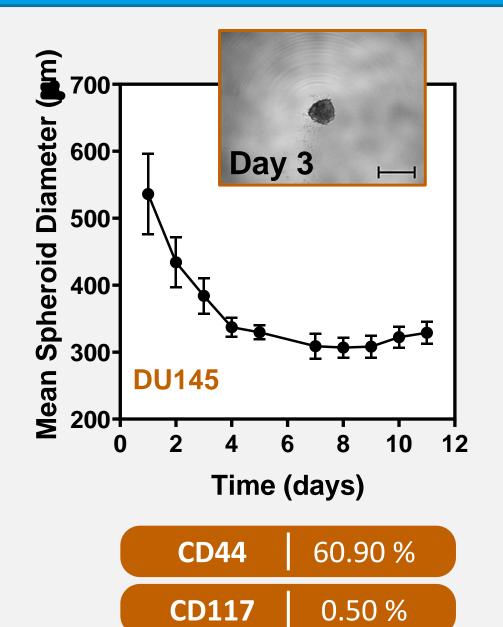


Need to use more sophisticated culture models, such as <u>multicellular tumor spheroids</u>, that better replicate the metabolic and proliferative gradients of tumors, with enriched populations of <u>cancer stem cells (CSCs)</u>, involved in treatment failure and cancer relapse. <sup>[2]</sup>

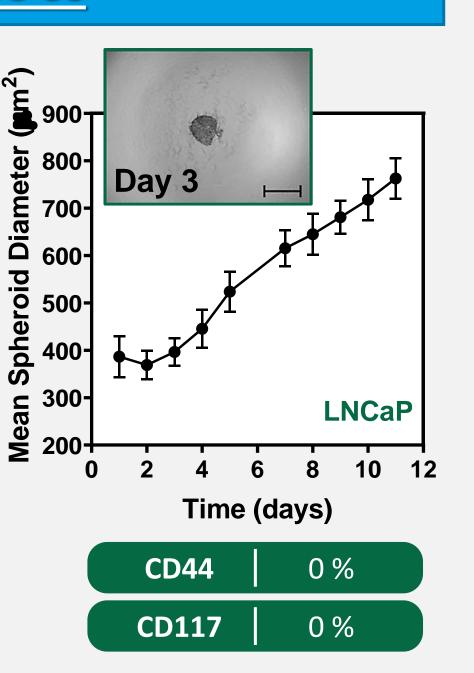
**GOAL:** To assess the theranostic potential of <sup>64</sup>CuCl<sub>2</sub> through the evaluation of the radiobiological effects of exposure of PCa spheroids from 3 cell lines (22RV1, DU145 and LNCaP) to this radionuclide

## PCa spheroids were fully established at $\underline{day\ 3}$ , with a mean diameter from 350 to 400 $\mu$ m, and had distinct populations of CSCs

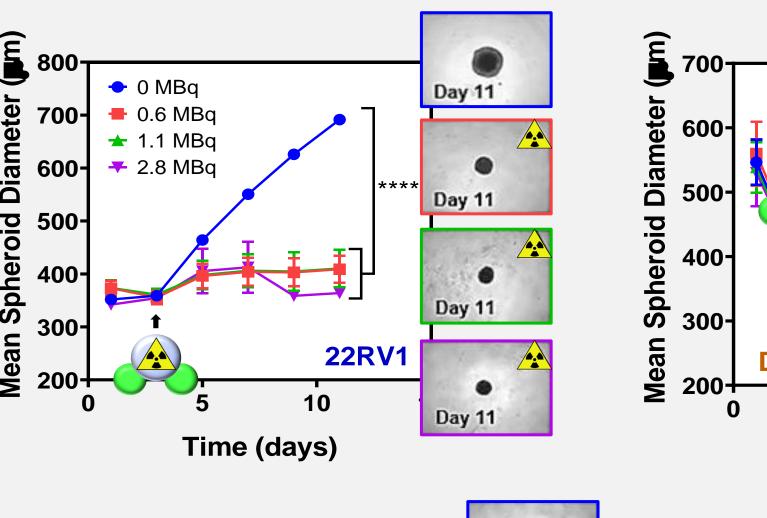


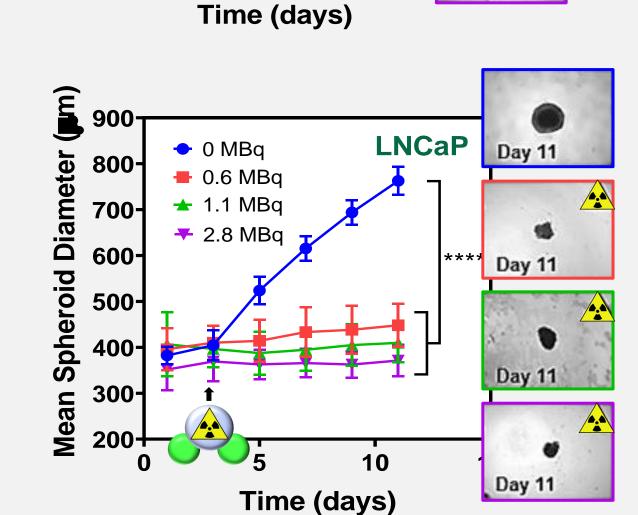


**ACKNOWLEDGMENTS** 

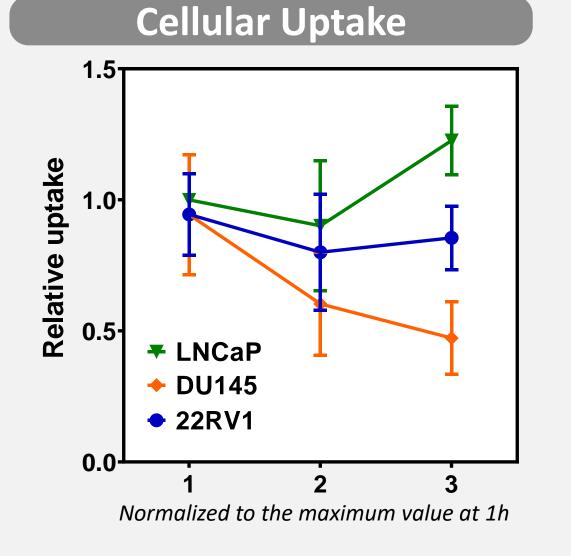


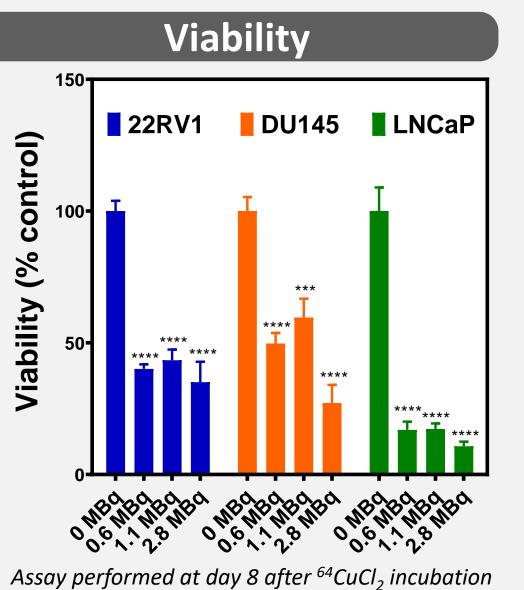
### <sup>64</sup>CuCl<sub>2</sub> significantly <u>inhibits the growth</u> of PCa spheroids



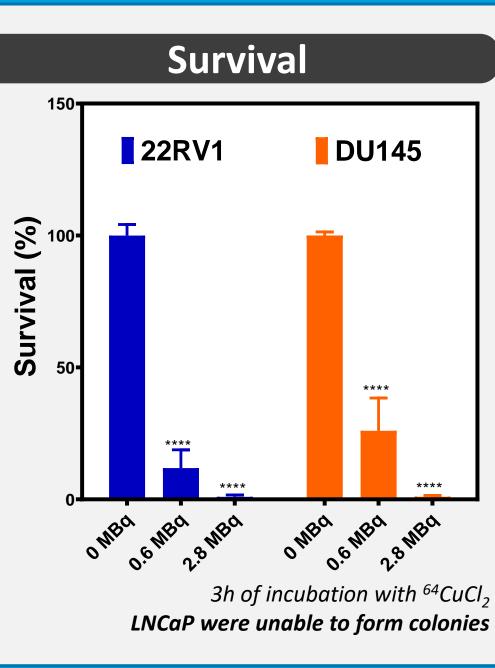


# Cellular Uptake Viability Survival 1.5 1.5 1.50





<sup>64</sup>CuCl<sub>2</sub> decreases PCa spheroids' viability and exerts an anti-



### CONCLUSIONS

- <sup>64</sup>CuCl<sub>2</sub> caused the highest cytotoxicity in LNCaP spheroids, followed by 22RV1 and DU145 spheroids. A similar trend was observed in the uptake studies: LNCaP spheroids had the highest uptake, followed by 22RV1 and DU145, as previously described for 2D cultures. <sup>[1]</sup>
- LNCaP 3D spheroids, which have the smallest population of CSCs, are more damaged by <sup>64</sup>CuCl<sub>2</sub> than spheroids from the other cell lines.
- The results here obtained confirm the high potential of 64CuCl<sub>2</sub> as a theranostic agent for PCa. [3]

#### REFERENCES

[1] Guerreiro J. F. et al. Molecules (2018). 23(11): 2944;
[2] Ishiguro, T. et al. Cancer Sci. (2017). 108, 283–289;
[3] Pinto C. I. G. et al Front. Mol. Biosci. (2020). 7:609172.

### Authors acknowledge FCT funding through UID/Multi/04349/2019 (C<sup>2</sup>TN), UIDB/04565/2020 (iBB) and PTDC/BTM-TEC/29256/2017 (FM). iBB acknowledges Lisboa 2020 for Project 007317.