## From Hadrons to Therapy: Fundamental Physics Driving New Medical Advances

Contribution ID: 14

Type: not specified

## Production of unconventional radioisotopes, radiochemistry development and preclinical studies for cancer theranostics at TRIUMF

Tuesday 6 September 2022 16:40 (40 minutes)

Using unconventional radionuclides for cancer treatment has been gaining popularity in recent years thanks to the remarkable results from the clinical studies with 177Lu, 223Ra and 225Ac1–3. TRIUMF launched a campaign to produce 225Ac from 232Th spallation4. Benefiting from this program, several other interesting alpha-emitters are co-produced, including 213Bi, 227Th, and 212Pb, which we have developed processes to purify5. TRIUMF's ISOL facility allows the production of Tb isotopes, including 155Tb which we use as an imaging companion for 225Ac and 177Lu6.

Those unconventional radionuclides require specific chelators for efficient and stable labelling, due to their larger sizes and different chemical properties compared to conventional radiometals. We have developed several novel chelators for those radionuclides7. The novel chelators were attached to tumour targeting peptides and were subsequently labelled with radioisotopes.

Preclinical imaging and biodistribution studies with peptides targeting melanoma or neuroendocrine tumour were performed, and the results demonstrated tumour specific uptake and low background uptake, indicating the promises of the novel radionuclides and chelators. Therapy studies with 225Ac is undergoing.

REFERENCES

[1] Parker, C. et al. Alpha emitter radium-223 and survival in metastatic prostate cancer. N. Engl. J. Med.369, 213–223 (2013).

[2] Kratochwil, C. et al. 225Ac-PSMA-617 for PSMA-targeted a-radiation therapy of metastatic castrationresistant prostate cancer. J. Nucl. Med.57, 1941–1944 (2016).

[3] Strosberg, J. et al. Phase 3 Trial of 177 Lu-Dotatate for Midgut Neuroendocrine Tumors. N. Engl. J. Med.376, 125–135 (2017).

[4] Robertson, A. K. H. et al. 232Th-Spallation-Produced 225Ac with Reduced 227Ac Content. Inorg. Chem.59, 12156–12165 (2020).

[5] McNeil, B. L. et al. Production, purification, and radiolabeling of the 203Pb/212Pb theranostic pair. EJN-MMI Radiopharm. Chem.6, 6 (2021).

[6] Fiaccabrino, D. E., Kunz, P. & Radchenko, V. Potential for production of medical radionuclides with online isotope separation at the ISAC facility at TRIUMF and particular discussion of the examples of 165Er and 155Tb. Nucl. Med. Biol.94–95, 81–91 (2021).

[7] Yang, H. et al. Synthesis and evaluation of a new macrocyclic actinium-225 chelator, quality control and in vivo evaluation of 225Ac-crown- $\alpha$ MSH peptide. Chem. –A Eur. J.26, 11435–11440 (2020).

Primary author: YANG, Hua (TRIUMF Canadian Particle Accelerator, Vancouver, Canada)

**Co-authors:** MCNEIL, Brooke (TRIUMF, Vancouver, Canada); ZHANG, Chengcheng (British Columbia Cancer Research, Vancouver, Canada); BÉNARD, François (British Columbia Cancer Research, Vancouver, Canada); EN-GUDAR, Gokce (TRIUMF, Vancouver, Canada); MEERES, Harrison (TRIUMF, Vancouver, Canada); WHAR-

TON, , Luke (TRIUMF/University of British Columbia, Vancouver, Canada); SCHAFFER, Paul (TRIUMF, Vancouver, Canada); MCNEIL, Scott (TRIUMF, Vancouver, Canada); RADCHENKO, Valery (TRIUMF, Vancouver, Canada)

Presenter: YANG, Hua (TRIUMF Canadian Particle Accelerator, Vancouver, Canada)

Session Classification: Research on targeted radionuclide therapy and associated technologies

Track Classification: Targeted radionuclide therapy and associated technologies