RADIotherAnostic pepTidEs in Immuno-Oncology (RADIATE-IO)

Problem being solved:

According to the WHO, the global cancer burden will rise by 77% by 2050. Most of the current treatment strategies involve a combination of targeted therapy or immunotherapy with therapies that cause immunogenic cell death (ICD) such as chemotherapy and external beam radiation therapy (EBRT).

- however, chemotherapy is highly non-specific and cytotoxic; its significant side effects limiting its use in these chronically ill patients
- due to the high radiation doses and practicalities, external beam radiation therapy is limited to locoregional disease or oligometastases and cannot be used in patients with systemic disease spread
- there is a need for more effective combination therapies to obliterate tumoural heterogeneity and overcome cancer immunotherapy (CIT) resistance

Solutions:

The team aims to develop a toolbox of novel radiotheranostic peptides is anticipated to significantly improve accuracy of immunoprofiling

- developing a toolbox of novel radiotheranostic peptides that can more accurately and noninvasively immunoprofile patients to guide combination CIT use
- targeted radioligand therapy (RLT) which allows the delivery a similar effective dose of radiation to the lesion to cause ICD without the non-specific side effects of chemotherapy and EBRT but at the same time
- having a long enough path length for cytotoxic bystander effects to on neighbouring nontarget expressing tumour cells, thus overcoming intralesional heterogeneity of a specific target

Market Size:

The global theranostic market reached USD \$1.8B in 2022, and is projected to experience substantial expansion, reaching USD \$4.2B by 2031. The *in vivo* animal data generated based on the proposed project from this funding would position RADIATE-IO technology as new theranostic tools in LCa treatment, a market valued at over USD \$29.5B in 2022 with 11% CAGR.

Competition:

Duke-NUS peptide inventions integrate technologies from immunology, chemistry and *in vivo* molecular imaging for the development of a multi-modal radiotheranostic peptide, other competitors include:

- CytoSite Bio Inc (USA) has recently reported the FDA investigational new drug (IND) approval for first-in-human (FIH) trials for its flagship GZB PET imaging agent, CSB-321, for the assessment of immunotherapy efficacy
- Researchers at UCSD (USA) and Peking University that have their own GZB peptides based on similar scaffolds
- To-date no team has demonstrated the potential of a GZB peptides as a radiotheranostic in cancer or infectious disease

Team:

- <u>Ann-Marie Chacko</u>, Asst Professor, Duke NUS medical school
- Prof Chandra Verma, A*STAR Bioinformatics Institute (BII)
- A/Prof Daniel Tan, National Cancer Centre Singapore