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Conformational epitopes-based lipopeptide vaccines against Schistosoma mansoni and Necator americanus

Mariusz Skwarczynski<sup>1</sup>, Abdullah A H Ahmad Fuaad<sup>1</sup>, Mark S Pearson<sup>2</sup>, Saranya Chandrudu<sup>1</sup>, Guangzho Zhao<sup>1</sup>, Darren A Pickering<sup>2</sup>, Alex C Loukas<sup>2</sup> and Istvan Toth<sup>1</sup>

<sup>1</sup>The University of Queensland, Australia <sup>2</sup>James Cook University, Australia

Hookworms (including *Necator americanus*) infect more people than HIV and malaria combined, predominantly in third world countries. Human blood fluke (*Schistosoma mansoni*) also infects millions of people worldwide. Both infections are one of the most important health problems in developing countries. Treatment of diseases related to the infections with chemotherapy have limited efficacy and re-infections after treatment are common. The development of vaccines for these infections could substantially reduce the global disability associated with these parasites. Two proteins crucial for pathogens feeding were chosen as antigens for vaccine design: hemoglobin digestion cascade protease Na-APR-1 from hookworm and cathepsin D hemoglobinase (Sm-CatD) from *Schistosoma*. In the case of both antigens high-yield production of them (eukaryotic proteins) was problematic. This prompts us to propose peptide-based strategy for the vaccines development. Thus, appropriate B-cell epitopes have been identified and conjugates were formulated into nanoparticles under aqueous conditions. Several series of vaccine candidates were tested in mice model. The lead candidates induced robust neutralizing humoral immune responses which strongly depended on conformational properties of the peptide epitopes incorporated into conjugates. The use of potentially toxic adjuvants was eliminated and vaccine candidates showed generally good safety profile. These findings are particularly encouraging for the development of single vaccine which can target both helminthiases.

m.skwarczynski@uq.edu.au

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