

## Development of therapeutic cancer nanovaccine candidates

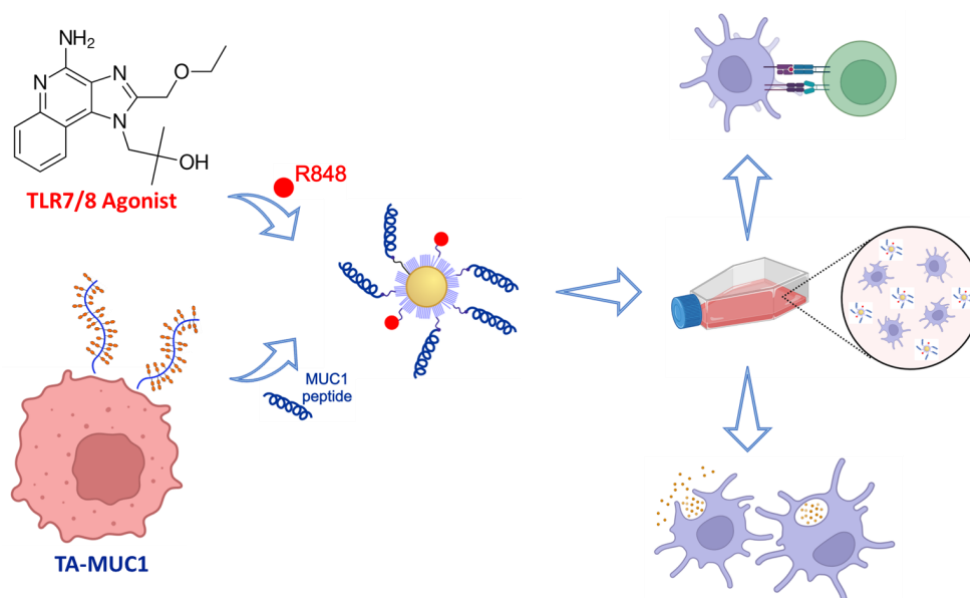
Ander Eguskiza, Ana Rita Ribeiro, Maria Morbidelli, Gilda Compagnone, Giulia Climani, Emanuele Papini, Jutta Horejs-Hoeck, Roberto Fiammengo

Department of Biotechnology, University of Verona, Strada le Grazie, 15, 37134, Verona, Italy

ander.eguskizabilbao@univr.it

Therapeutic cancer vaccines hold significant promise for achieving long-lasting remission, or even cure, in various forms of advanced stage and/or refractory cancers. Nanoparticle-based formulations are expected to play a key role in this field. In fact, nanoparticles (NPs) may considerably improve tumor-associated self-antigens or neoantigens presentation to the immune system. NPs (typically 10–100 nm), are effectively internalized by antigen presenting cells (APCs) and reach draining lymph nodes, allowing a more effective stimulation of the immune system. Additionally, the organic coating of the NPs may be programmed to adjuvate the immune response if a suitable protein corona forms upon *in vivo* administration of the vaccine formulation.

This project focuses on gold NPs (AuNPs) coated with poly(ethylene glycol) (PEG) as carriers for tumor-associated self-antigens derived from MUC1, an overexpressed and altered mucin found in different cancer types, due to compelling preliminary results showing good immunogenicity of such vaccine candidates *in vivo*. To increase the potency of the vaccine, a TLR-7/8 agonist (R848), was conjugated to the surface of the vaccine. Immune cell activation and nanovaccine internalization by antigen presenting cells was studied *in vitro*.



- [1] Compañón, I., Guerreiro, A., Mangini, V., Castro-López, J., Escudero-Casao, M., Avenzoza, A., Busto, J., Castellón, S., Jiménez-Barbero, J., Asensio, J., Jiménez-Osés, G., Boutureira, O., Peregrina, J., Hurtado-Guerrero, R., Fiammengo, R., Bernardes, G. and Corzana, F. Structure-Based Design of Potent Tumor-Associated Antigens: Modulation of Peptide Presentation by Single-Atom O/S or O/Se Substitutions at the Glycosidic Linkage. *Journal of the American Chemical Society*, **2019**, 141(9), pp.4063-4072.
- [2] Hemmi, H., Kaisho, T., Takeuchi, O. et al. Small anti-viral compounds activate immune cells via the TLR7 MyD88-dependent signaling pathway. *Nat Immunol* **2002** 3, 196–200