

Chemistry in Medicine: fluorophores and optical imaging

Marc Vendrell^{1,2}

¹ Centre for Inflammation Research, Institute for Regeneration and Repair (IRR), The University of Edinburgh, EH16 4UU Edinburgh, UK

² IRR Chemistry Hub, Edinburgh BioQuarter, The University of Edinburgh, EH16 4UU Edinburgh, UK

Fluorescent activatable probes are valuable tools for live-cell imaging because of their tunability and target specificity.¹ Over the last few years, our group has designed a collection of fluorogenic amino acids and peptides for high-resolution biological imaging and translational medicine, which was recognised with the RSC Bader Prize 2023. Our team have demonstrated that this approach can be used to generate probes to visualize infection and immune cells in human biosamples,² *in vivo*³ and in *ex vivo* human biopsies.⁴ We have designed fluorescent amino acids to: 1) be compatible with conventional solid-phase peptide synthesis, 2) maintain the biomolecular recognition features of the native peptides and 3) emit fluorescence preferentially after target binding, improving signal-to-noise ratios for imaging. Recently, we have included the smallest turn-on fluorescent amino acids for peptide-PAINT imaging and super-resolution microscopy,⁵ and to fluorogenic tags for proteins associated with immune cell function like interleukins,⁶ immunophilins and chemokines⁷ as well as nanobodies and antibodies.⁸ Finally, the talk will also briefly discuss our efforts to establish apply these fluorescent probes for clinical applications.

[1] Cheng, Z. et al. Nat. Rev. Chem. **2020**, 4, 275.

[2] a) Mendive-Tapia, L. et. al. Nat. Commun. **2016**, 7, 10940; b) Mendive-Tapia, L. et. al. Nat. Protoc. **2017**, 12, 1588; c) Mendive-Tapia et al. Angew. Chem. Int. Ed. **2022**, 61, e202117218.

[3] a) Barth N. et al. Nat. Commun. **2020**, 11, 4027; b) Kaplaneris, N. et al. Nat. Commun. **2021**, 12, 3389; c) Subiros-Funosas, R. et. al. Chem. Sci. **2020**, 11, 1368; d) Barth N. et al. Angew. Chem. Int. Ed. **2022**, 61, e20211302.

[4] Scott, J. et al. Nat. Commun. **2022**, 13, 2366.

[5] De Moliner et al. Angew. Chem. Int. Ed. **2023**, 62, e202216231.

[6] Reese et al. ACS Cent. Sci. **2024**, 10, 143.

[7] a) Bertolini et al. ACS Cent. Sci. **2024**, 10, 969; b) Bertolini et al. JACS **2024**, 146, 30565; Bertolini et al. JACS **2025**, accepted.

[8] a) Kuru et al. Nat. Commun. **2024**, 15, 7531; b) Nadal-Bufi et al. JACS **2025**, 147, 7578.