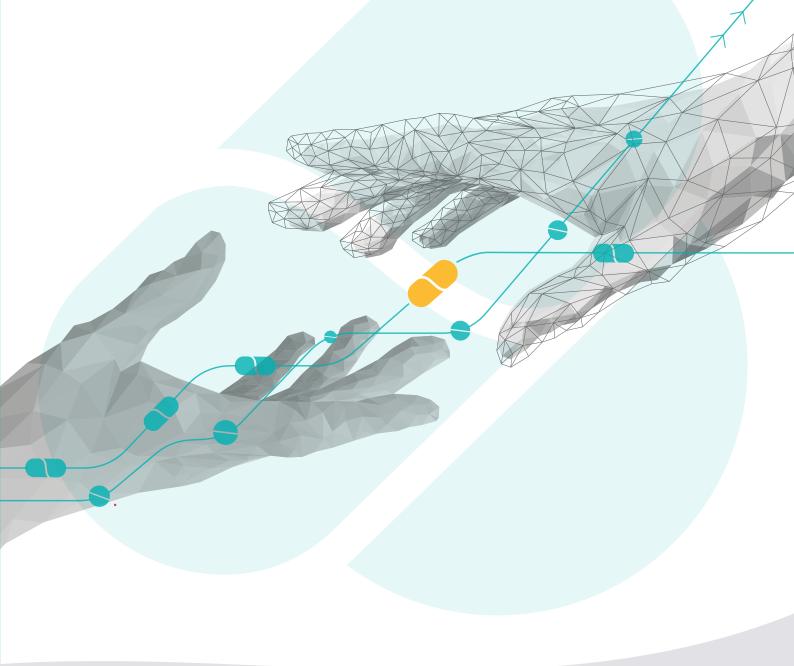


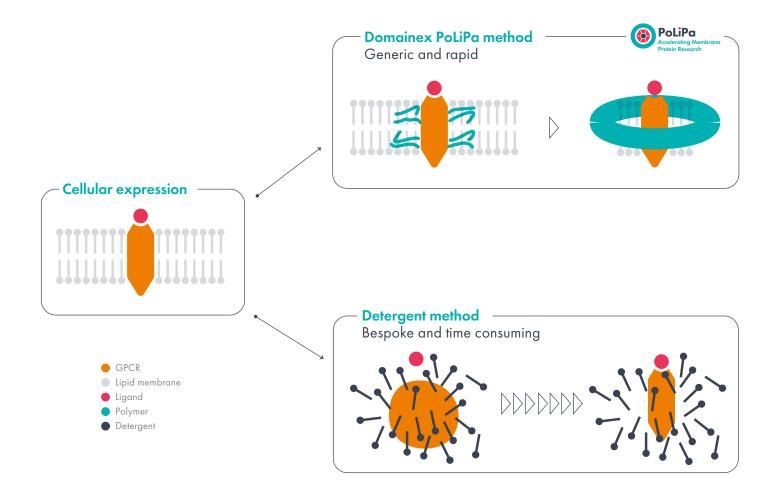
Innovative Polymer Lipid Particle (PoLiPa) Technology





Combining High Quality Functional Membrane Protein Preparations with LC-MS Read-outs

Domainex has established a generic platform to generate any purified membrane protein without the need for thermostabilising mutations or detergents. This was achieved using Polymer Lipid Particle (PoLiPa) technology that can stabilise membrane proteins by encapsulating the target protein in a polymer that encloses a small disc of the native cell membrane lipids. Once isolated, we could use sensitive, label-free LC-MS detection technology to provide full pharmacological characterisation of known ligands and test compounds.



Advantages of soluble PoLiPa-GPCRs

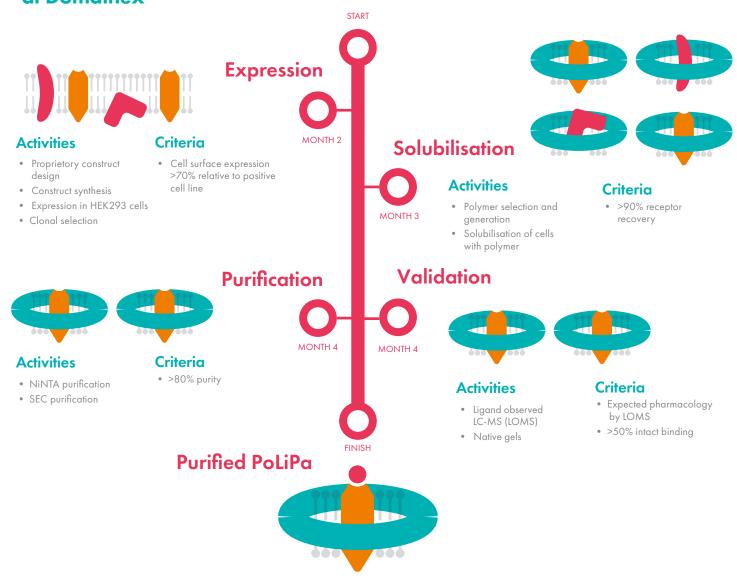
Enables rapid and generic access to pure samples of GPCRs

- Mutagenesis
- Detergents
- Generation of pharmacologically intact membrane targets
- Preparations stable over several months
- ✓ Versatile applications

Applications

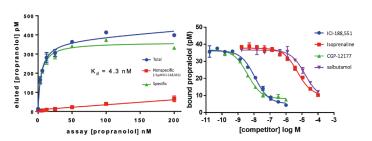
- FBDD
- SBDD
- Biophysical characterisation
- Structural determination
- DNA-encoded library screening
- Biologic hit ID
- Orphan receptor profiling

Generation of PoLiPa-GPCRs at Domainex



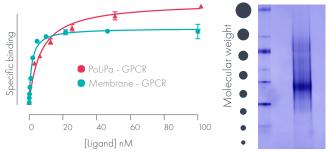
Example Data

Example 1: BETA-2-Adrenergic Receptor (\$2AR)



Left: PoLiPa-β2AR propranolol saturation binding. **Right**: Competition experiment with four known agonists/antagonist. PoLiPa-β2AR demonstrated expected pharmacology.

Example 2: Neurotensin Receptor 1 (NTSR1)



Left: Saturation binding analysis (LC-MS) of a known NTSR1 antagonist to isolated membranes expressing NTSR1 or PoLiPa-purified NTSR1. Kd values are comparable between the two preparations suggesting that a similar pharmacologically intact protein is presented in each system. **Right:** Native-PAGE gel with Coomassie staining over PoLiPa-NTSR1.

About Domainex

scientists. We provide integrated services, from disease target selection to candidate drug nomination. We have a very strong reputation for contributing innovative ideas, undertaking high-quality experiments and for generating intellectual property on behalf of our clients. We strive to build strong, dynamic relationships and work with our clients to provide customised services.

How Can Domainex Help Your Drug Discovery Project?

you to advance your drug discovery projects towards drug development effectively and efficiently. We provide customised programmes to address your specific needs at each stage of the pre-clinical drug discovery process. We draw from a wealth of expertise built up over the last 20 years across a wide range of drug targets and therapeutic areas. From our sites within Europe's leading bioscience hub at Cambridge, UK and with access to the very latest cutting-edge technologies, we are able to help you realise your goals and enrich your discovery pipeline.

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in Domainex





Chemistry & Bioanalytical Centre of Excellence

Churchill Building Chesterford Research Park Little Chesterford CB 10 1XL



Biology Centre of Excellence

Sigma Building, 40 South Street Unity Campus Cambridae CB22 3FW

