



New smart polymers that can accurately measure lentivirus were synthesized and evaluated in just 4 months.

## Development of VSV-G Smart Polymers for Lentivirus Quantification: from *in silico* modelling to application

E. Daniels\*, K. McBain<sup>†</sup>, J. Czulak\*, A. Peon\*, A. Thomson\*.

\*Tozaro, Bedford, United Kingdom

<sup>†</sup>Sartorius, Royston, United Kingdom

### *In silico* Modelling

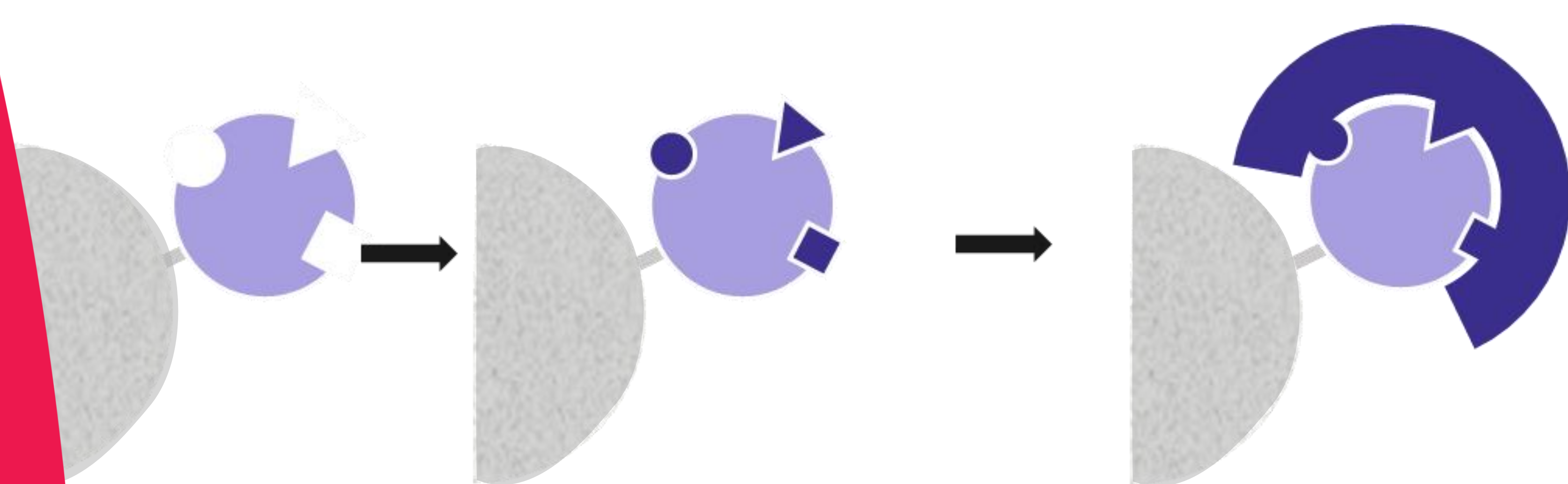
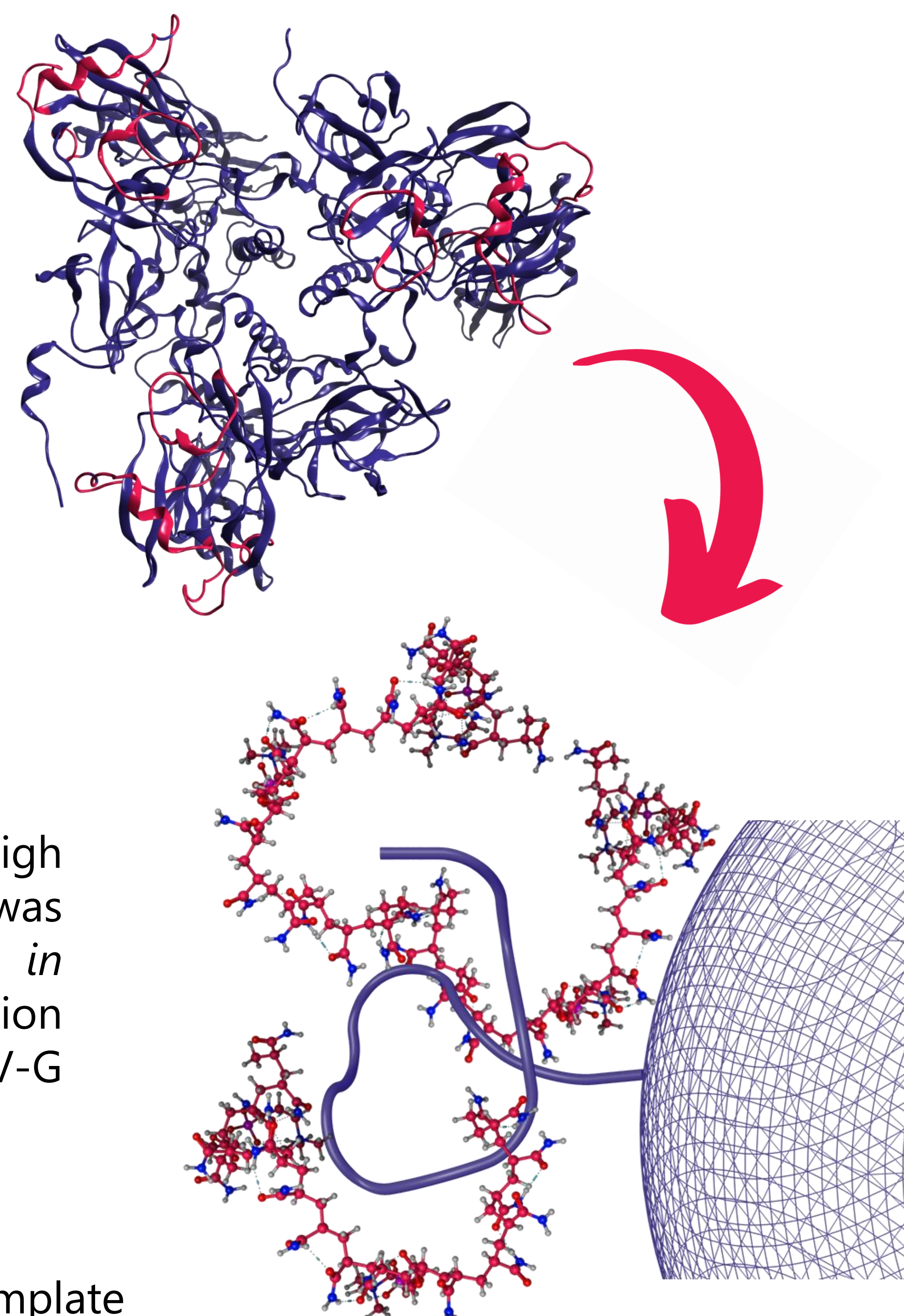
Computational modelling was used to assess the surface of VSV-G, an envelope protein of Lentivirus (LV). **7 epitopes** were selected for use as templates in the development of smart polymers. Subsequently, each epitope was screened against a library of **>500 monomers** (the building blocks of smart polymers). Monomers that interacted favorably with the epitopes were used in the next step of polymer development.

### HPLC Screening

Each epitope template was immobilized onto porous beads and used as the solid support in high performance liquid chromatography (HPLC). Each epitope-functionalized solid phase was screened against **91 linear polymers**, built using the favorable binders identified from *in silico* modelling. If a polymer showed a strong interaction to the template, its HPLC retention time changed. The compositions of strongly binding polymers were used to design new VSV-G smart polymers, and the remaining data was used to improve our *in silico* predictive model.

### Smart Polymer Synthesis

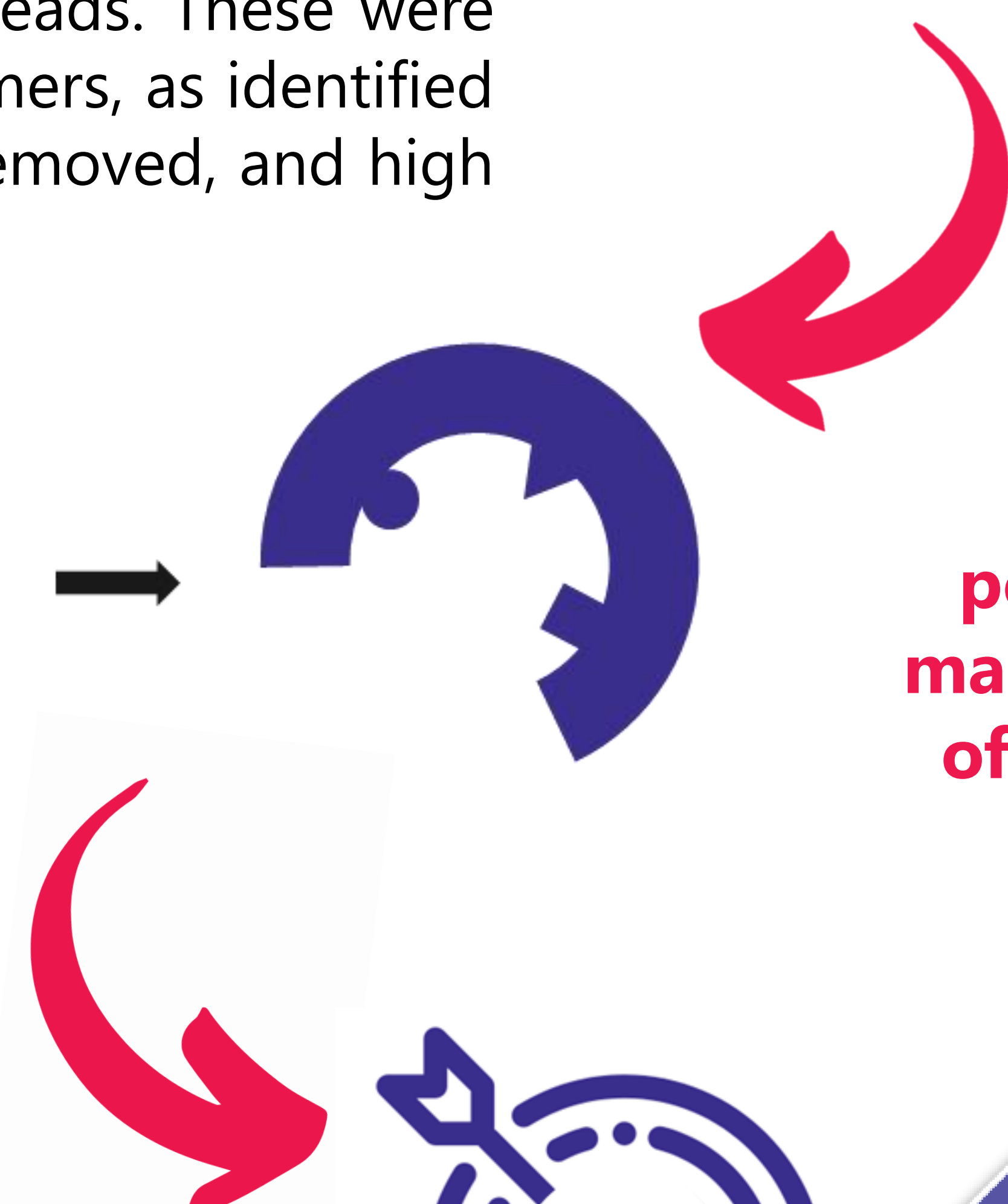
Polymers were made using our patented solid phase imprinting technique. Each template (epitope) identified by *in silico* modelling was immobilized onto the glass beads. These were then exposed to a polymerization solution including crosslinkers and monomers, as identified by HPLC screening. After polymerization, the templated glass beads were removed, and high affinity nanoparticles (smart polymers) were collected.



### Evaluation in Application

Each VSV-G smart polymer was tested for affinity to VSV-G protein in SPR. **19 hits** were identified, giving a **~38% hit rate**! So far, 3 of those hits have been further tested against intact lentivirus using the Octet BLI platform. All **3 smart polymers worked in application** – they could be immobilized onto BLI biosensors and showed affinity towards LV across a  $10^5$  to  $10^8$  TU/mL range.

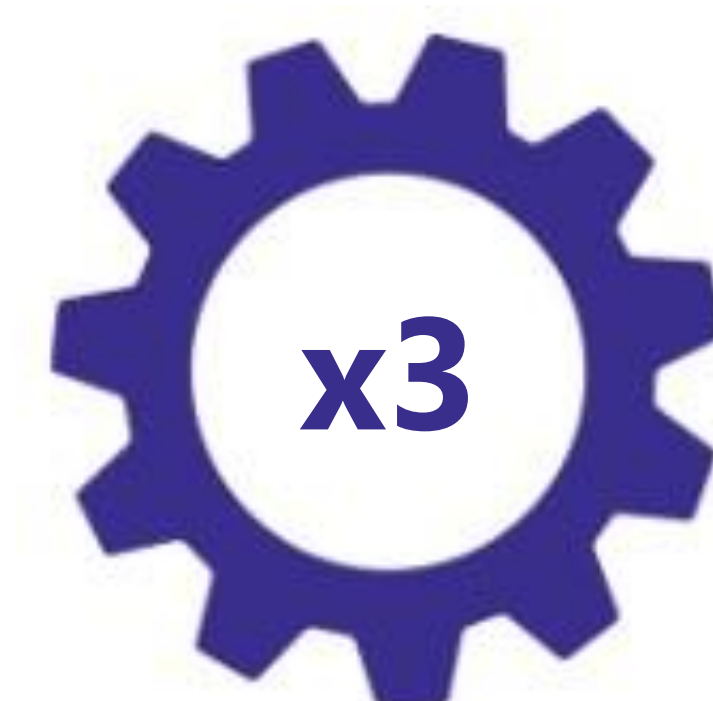
3 new prototypes for the detection of in-tact lentivirus were made and evaluated in just **4 months**, showing that our **smart polymer development process** can be used for **rapid generation of new reagents** to support proprietary gene therapy manufacturing processes.



**>50 smart polymers were made in 4 months of hands-on lab time!**



**19 Hits against VSV-G in SPR**



**3 x Prototypes against LV in BLI**