From lineal to cyclic peptidomimetics; Retained binding motif and improved physico-chemical features

**Standardized Approach**

- PDB: 5W5S (cis amide)
- Post-filtering
- MD & MM/PBSA
- Repeated N conformations

**Iproteos Macrocyclization Approach**

- IPRO Docking
- IPRO Cyclogen
- IPRO Permeability
- Synthesis & evaluation
  - Preserve binding
  - Higher permeability
  - Enhanced stability

**Results**

- **IPRO Docking**
  - Score: -8.9 kcal/mol
  - RMSD: 0.89 Å

- **IPRO Cyclogen**
  - Binding motif blocked when linker introduced

- **IPRO Permeability**
  - Ligand-based approach to predict permeability