

SciLifeLab possibilities in Drug Discovery & Development

Per I Arvidsson

Director SciLifeLab Drug Discovery & Development

SciLifeLab as a government-funded national infrastructure

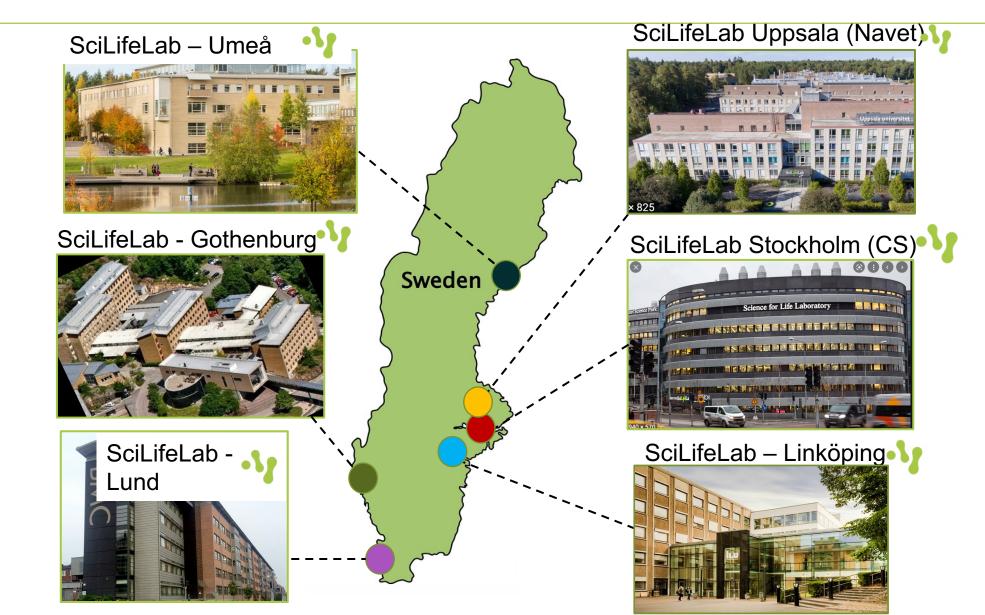




Max-IV Next-generation synchrotron **ESS** European Spallation Source SciLifeLab Molecular Life Sciences

New SciLifeLab national sites – Est. 2021





Many faces of SciLifeLab





Infrastructure

10 technology platforms, >40 units >1400 users/yr, 3000 projects/yr

- ~ 500 tech experts
- Bioinformatics
- Cellular and Molecular Imaging
- · Chemical Biology and Genome Engineering
- **Clinical Genomics**
- Clinical Proteomics and Immunology
- Genomics
- Drug Discovery and Development
- Integrated Structural Biology
- Metabolomics
- Spatial and Single Cell Biology •



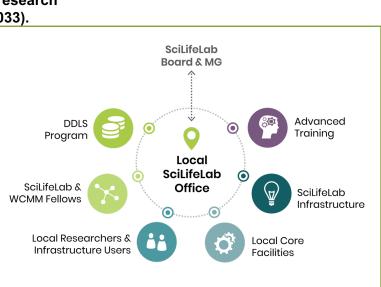
Research environment

- ~ 1500 scientists
- 189 affiliated research groups
 - o KI 32
 - o KTH 47
 - o SU 27
 - o UU 83
 - o Other univs
- Recruitment of 35 SciLifeLab Fellows
- 7 Research Community Programs
- 16 Technology Development Projects
- COVID-19 research program

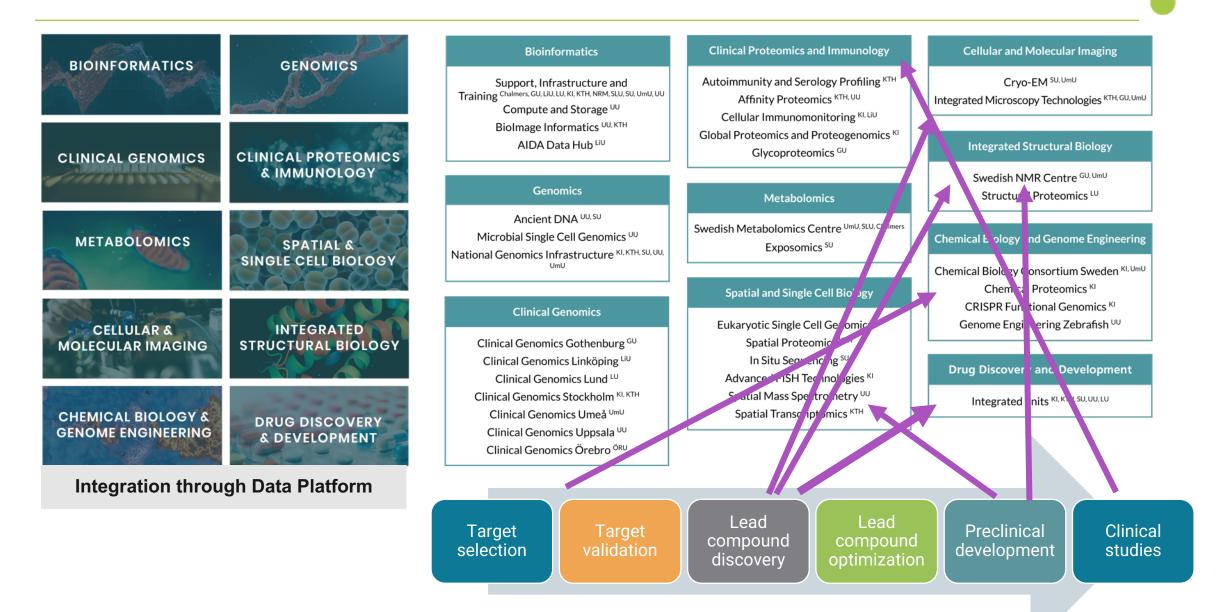
Data-driven life science

A new national 12-year 3.1 BSEK research program, funded by KAW (2021-2033).

- ~ 500 data scientists
- Data support (FAIR)
- Data analysis & AI
- 4 research areas
- Recruitment of DDLS Fellows
- Research and training •
- Collaborations with WASP •
- Industrial program



SciLifeLab infrastructure from 2021 - platforms and units





Swedish NMR Centra – New DNP-NMR



Astras nya utrustning – gör jobbet en miljon gånger snabbare

EkonomiDen ser ut som kombinerad värmepanna och kylskåp. Men den nya magnetutrustningen som Astra Zeneca och Göteborgs universitet nu investerar 40 miljoner kronor i kan analysera molekylstrukturer upp till en miljon gånger snabbare än dagens instrument.

– Den är unik för Skandinavien och kommer att hjälpa mycket forskning – från medicin till batteriteknik, säger Göran Karlsson på Göteborgs universitet.



https://www.gu.se/en/nmr/research-areas/material-science



Unit of Chemical Proteomics: Interactions with the Industry Sector

Types of interaction

- A. Projects for Industry (most interactions)
- B. Partnership with several industries in EU and Canadian calls for funding
- A. Industry projects from a total of 6 different users from Industry:
 - 3 Swedish SMEs,
 - one EU big Pharma
 - o one EU SME
 - o one non-EU SME

Major questions answered

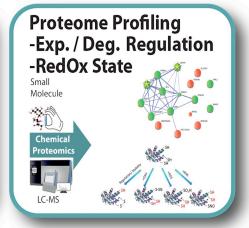
- TARGET DECONVOLUTION using MS-based proteomics, including off-targets
- MECHANISM OF ACTION ELUCIDATION of drugs / candidate drugs using MS-based proteomics
- Antigen-antibody EPITOPE MAPPING

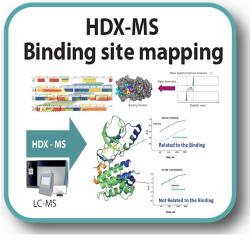
Provided service

- ✓ PISA (Gaetani M. et al 2019) for deep, high throughput and proteome-wide thermal stability/solubility profiling, also with orthogonal proteomics approaches integrated with it.
- $\checkmark\,$ Whole pipeline from cells to data analysis provided.
- ✓ Hydrogen / Deuterium Exchange Mass Spectrometry (HDX-MS) for binding site mapping

Web page: <u>https://ki.se/en/mbb/chemical-proteomics-core-facility</u> or <u>https://www.scilifelab.se/facilities/chemical-proteomics</u>

<section-header>





Chemical Proteomics. Massimiliano Gaetani. e-mail: massimiliano.gaetani@ki.se



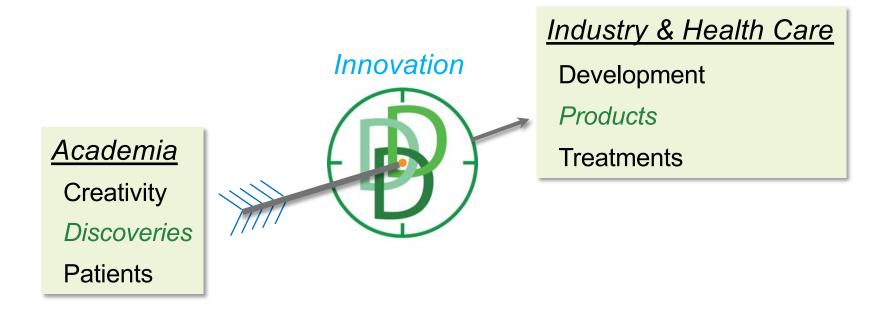
SciLifeLab DDD Mission



Separately funded capability within SciLifeLab with a specific task from the Government

Turn Academic Discoveries into Innovations

Provide State-of-the-art Drug Discovery & Development knowledge in Sweden

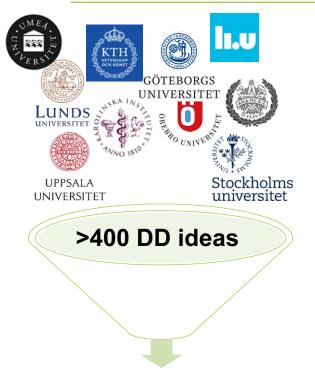


Drug Discovery with the best of both Worlds!



SciLifeLab DDD Objective

"Turn academic discoveries into innovations"



Validated DD programs

Exits: 2 Programs & 40 Service / y

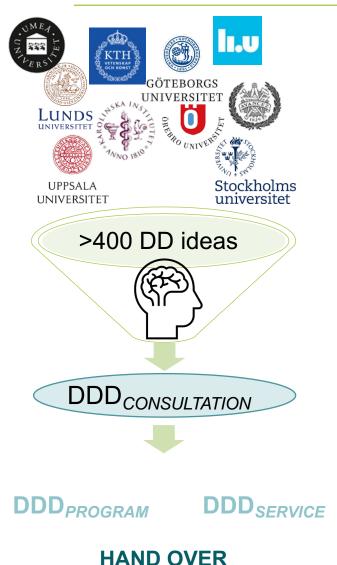
- 3 Clinical
- 4 Internationally partnered
- 5 Swedish biotechs (3 listed)



SciLifeLab DDD Capabilities



"Provide State-of-the-art Drug Discovery & Development knowledge in Sweden"



Small molecules

- DECL DNA encoded chemical libraries
- Al-enabled IT system
- Targeted Protein Degradation
- Massive virtual screening

Antibody therapeutics

- SciLifeLibs phage display libraries
- Cell therapies
- Bispecific mAbs
- New modalities
 - Conjugates
- Oligonucleotides

🖌 SciLifeLab

CBGE, Integrated Structural Biology, Cellular and Molecular Imaging, Metabolomics, Genomics, Bioinformatics, DataCenter, Etc.



- Contractual ability
- Partnerships
 - EUbOpen (IMI)
 - ENABLE2 (VR)
 - Conception (IMI)
 - InnoPharma (Vinnova)
 - Nevermore Covid (KAW)
 - ...

Validated DD programs

Exits: 2 Programs & 40 Service / y

- 3 Clinical
- 4 Internationally partnered
- 5 Swedish biotechs (3 listed)





SciLifeLab DDD Platform – 2022

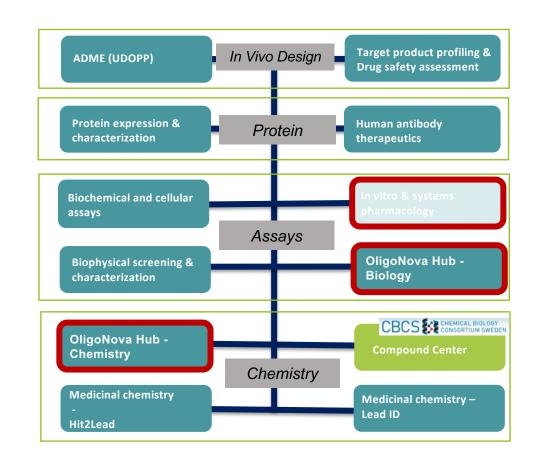
- 40+ Industrially experienced scientists
- 11 Platform Scientific Directors
 Area expertize and network
- National Platform Steering Group

 Project prioritization, budget and strategy

Håkan Billig (GU, Chair), Lars Lannfelt (UU), Maria Jenmalm (LiU), Lars Ny (GU), Anna Sandström (AZ), Outi Vaarala (Orion Corporation, FI), Tomas Lundquist (Pretzel Therapeutics)

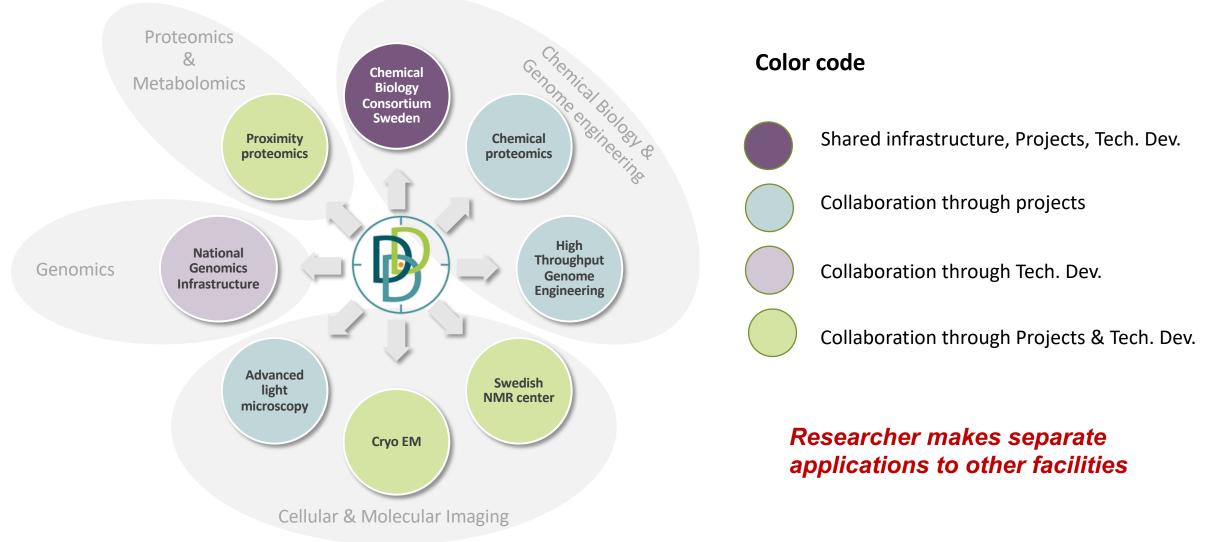
 International Platform Advisory Board – Long-term strategic input

Kjetil Taskén (Prof. Oslo Cancer Cluster, NO), **Fiona Marshall** (Head of global research Merck, US), **Justin Bryans** (Head of research Lifearc, UK), **Lorenz Mayr**, (Vector Biopharma, CH), **Lovisa Afzélius**, (Flagship pioneering, US)



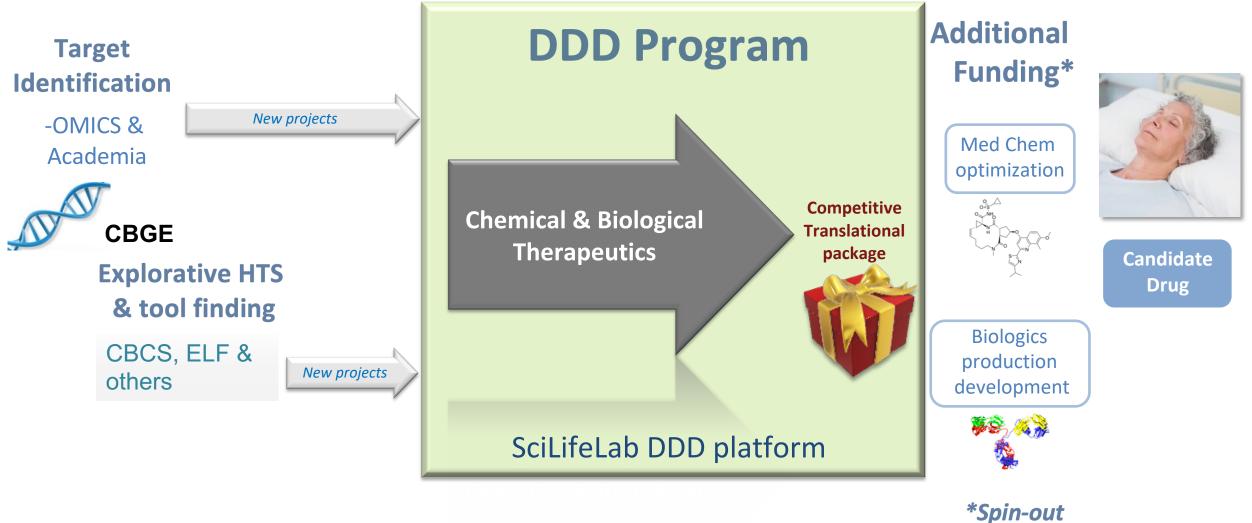


DDD within SciLifeLab ecosystem





Lead compound discovery From Exploration of Biology to Candidate Drug



Risk sharing

Preclinical development

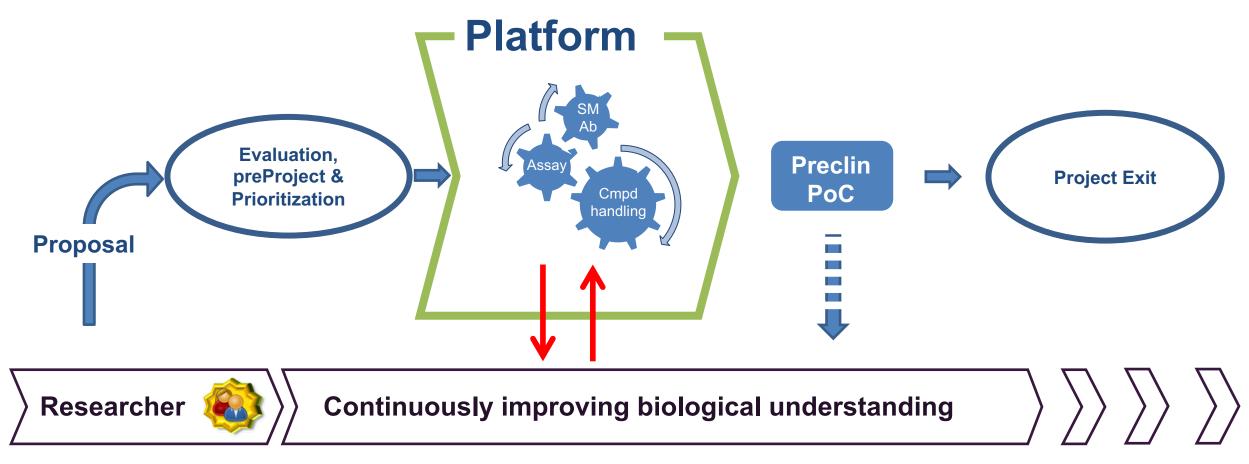
Target selection

Clinical

studies



Industry standard drug discovery infrastructure and expertise for the Swedish academic community



Researcher retains project & IP ownership

Preclinical developmer

compound discovery compour optimizati Clinical

studies

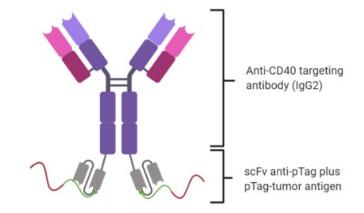
Target selection



Example



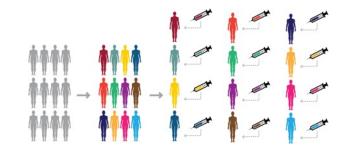
Adaptable Drug Affinity Conjugate (ADAC) Technology





pTag-tumor antigen

<u>True</u> personalized medicine by making T cells specific for the tumor - in the patient's body





Sara Mangsbo, UU New Modality – Technology platform for Oncology and infection National partnering 2021

STRIKE Pharma

Project Concept developed by Sara Mangsbo, Uppsala University

- Targeted peptide delivery
- Internalization and <u>cross-presentation</u> (harnessing the strength of the CD40 pathway)
- T cell priming and expansion in vivo
- Flexibility of cargo delivery neoantigens an opportunity
- No toxicity, safe and simple SC administration
- Fast and cost effective production of individualized part of the drug

DDD contributions

- Design of bifunctional antibody
- Library design
- Identification, production and biophysical characterization of binders to CD40 and pTag from libraries
- Analysis of DMPK properties, bioanalysis and PKPD modelling
- Project coordination
- Coaching and networking

"DDD Industry pitch panel" – 1st attempt 2021

• Part of Vinnova funded "Innopharma project"

Objective: Assure "customer/funder"
 perspective on DDD programs

Present parties: Medivir, Oncopeptides, Eir Venture, AstraZeneca, SOBI, BII – one company had to cancel due to CDA conflict

Outcome: Valuable technical and market feedback, large interests – multiple follow up discussion, interest from all parties to renew in 2022



Ways to interact





• New assets for drug development / commercialization

COLLABORATIVE• Access to resources/instruments/technologies at DDD

- Longer term contracts based on full cost model & joint applications
- e.g. IMI programs, Vinnova, KAW Covid, ENABLE2, etc



Access to spare resources/instruments at DDD

• Non-time critical access

SERVICE

• e.g. Access to separate facility competence & instrumentation, renting of personnel, etc

Infrastructure challenges for industry collaboration



- Contractual ability to access distributed infrastructure
- University had financial restrictions for doing "contract research"
- Limited available resources "undanträngningseffekt"
- Universities are not ideal for flexible/temporary positions i.e. need longer-term contracts to hire new personnel for full cost projects
- Universities 3rd mission not prioritized to the right level, e.g. legal support
- Vinnova funding for joint technology developments with 1-2 years duration would enable us to support start-ups without infringing EU regulations for state aid!