

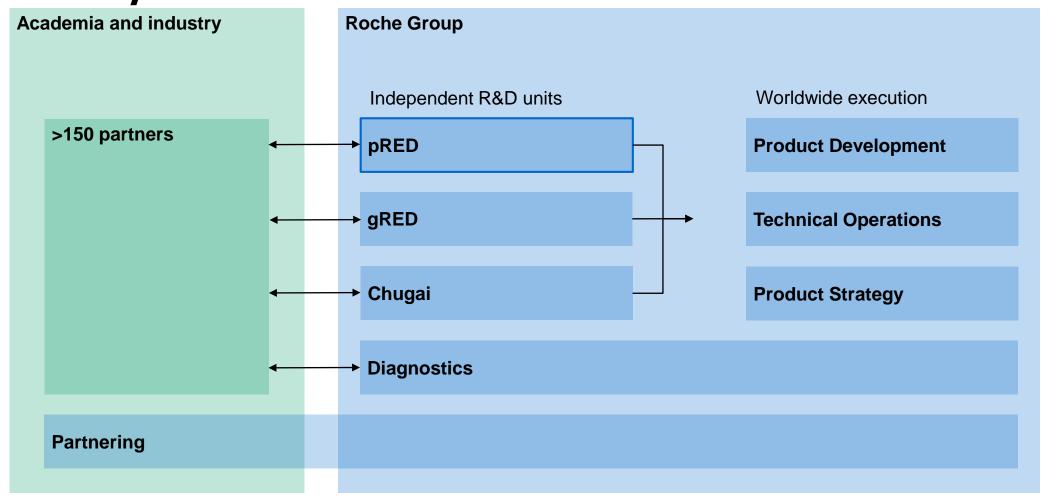
The CAIMAN project

Making most of our data with D360

Marc Pompiati, LMR, Roche Innovation Center Munich



The Roche Group organisational structure Roche pRED is one of three independent research & development units

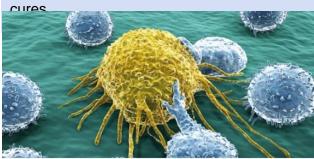


The Roche pRED disease areas



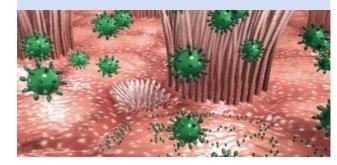
Oncology

Combining cancer immunotherapy and tumour targeted therapy approaches to seek



Infectious diseases

Developing targeted treatments for life-threatening infectious diseases



Immunology & inflammation

Developing differentiated medicines for people with immune and inflammatory

diseases

Ophthalmology Restoring sight



Neuroscience

Developing medicines for serious neurological diseases



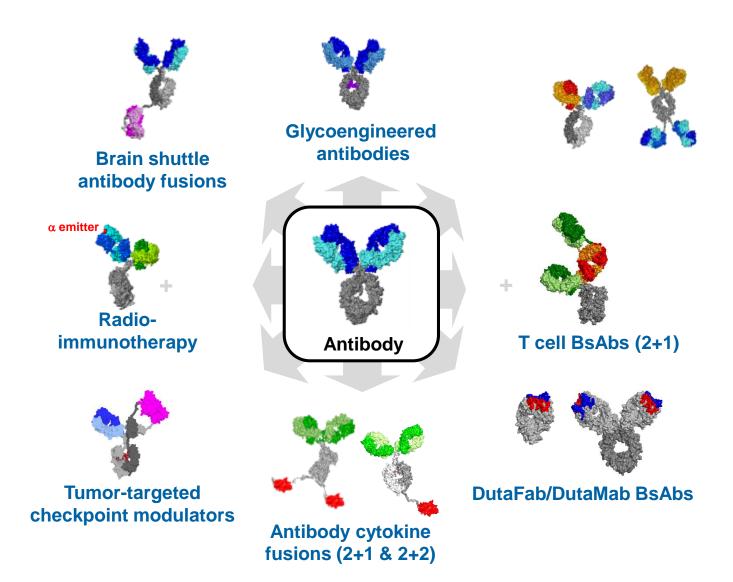
Monogenic rare diseases

Tackling inherited genetic disorders





Roche pRED's modular biologics Advancing therapeutic antibody sciences









- Digitalization is creating exciting new opportunities.
- To be equally successful in this new environment, Roche pRED depends on making better use of our research data, which continue to increase exponentially.
- The CAIMAN project is part of the Lab of the future initiative that addresses digitalization and supports the overall pRED digitalization goals.



LMR: Lab of the future Embrace digital and automated solutions to aid discover and develop superior biologics

Landscape	Data&learning	People
We are developing ONE landscape from registration, data capture, visualisation to interpretation and learning.		



LMR: Lab of the future Embrace digital and automated solutions to aid discover and develop superior biologics

Landscape **Data&learning People** Value will increasingly be We are developing ONE landscape from registration, generated by digital and data capture, visualisation to automated solutions. interpretation and learning. Data are a central asset.



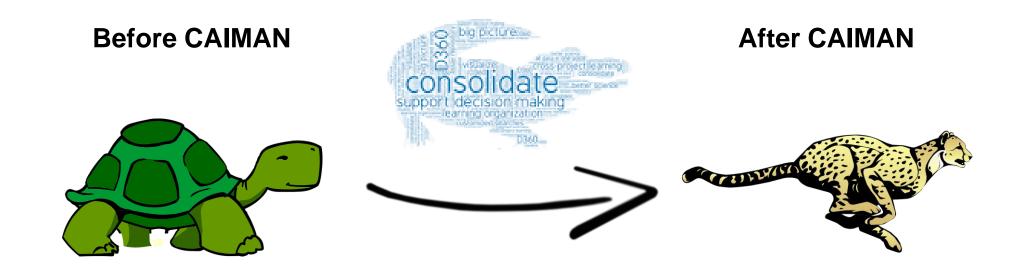
LMR: Lab of the future Embrace digital and automated solutions to aid discover and develop superior biologics

Landscape **Data&learning People** Value will increasingly be We are developing ONE Skills in biology, landscape from registration, generated by digital and biotechnology, data science and automation is needed. data capture, visualisation to automated solutions. interpretation and learning. Data are a central asset.

The CAIMAN project



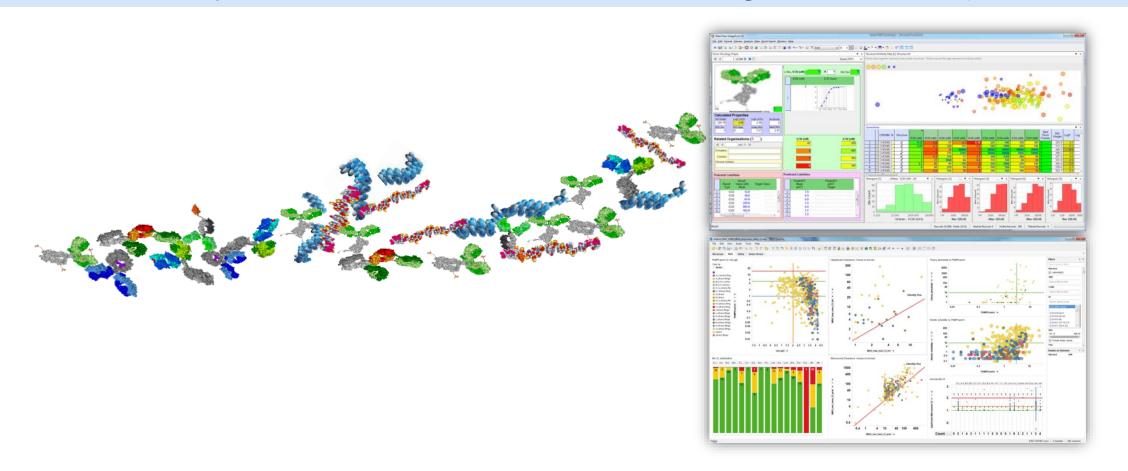
CAIMAN = Consolidated Platform for Assay, Inventory, Molecule and Analytics Data



CAIMAN Vision



A single interface for accessing and analyzing data across modalities and achieving quality data mining for project teams to increase efficiency and reduce effort for advancing portfolio projects



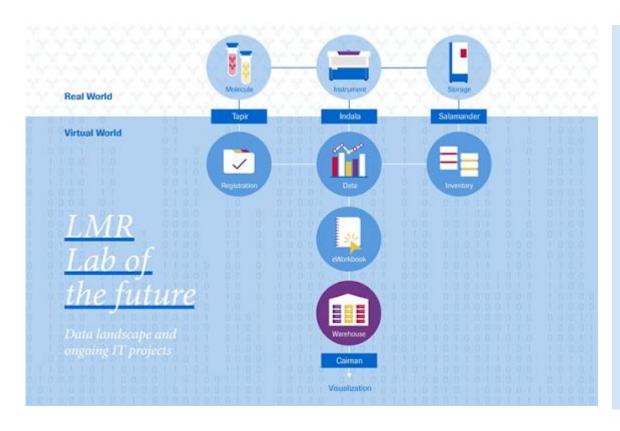
The goals of the CAIMAN project

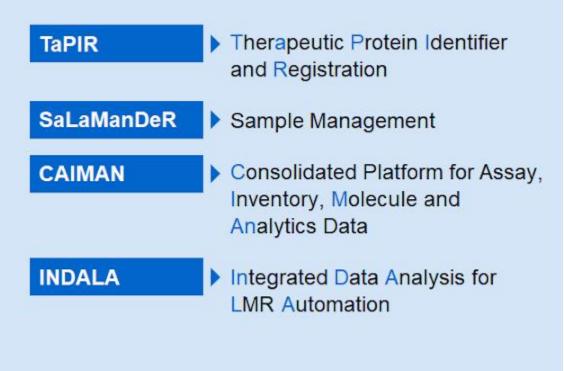


- Make access, integration and analysis of relevant and understandable data less cumbersome and time-consuming and more reliable.
- Help shift the focus from data collection and capture to data analysis by establishing automated data workflows that use existing assay data-capturing tools like BioBook and registration systems like TaPIR, and merging them with other existing project metadata into scientifically meaningful records.
- Guarantee that high-quality data are available in pRED for informed decisions by establishing a review process for assay description and data structure.





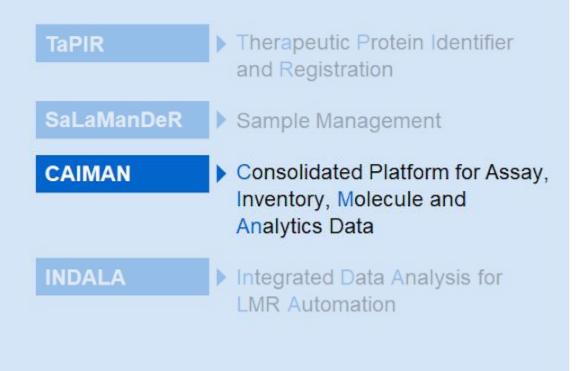














CAIMAN Definition Phase In a Nutshell – User Stories and Data define the Project



User Stories evolved into 23
Decision Points in 5 major
phases

The CAIMAN phases



- Tool Generation
- Binder Generation
- Primary Screening
- Secondary Screening
- Lead Characterization

TE Target Enabling Lead Identification

- Tool Generation
- Binder Generation
- Primary Screening
- Secondary Screening
- Lead Characterization

LO^{pre}
Lead Optimization
pre CLS

Engineering

Evaluation

Supply

Lead Characterization

Pool Selection &

LO^{post}
Lead Optimization
post CLS

- Lead Characterization
- Analytical Method Development
- Single Cell Cloning I
- Single Cell Cloning II
- Batch Cultivation
- Cell Line Stability
- PSB Clone Selection
- Supply
- GLP tox supply
- DSP Development
- USP Development

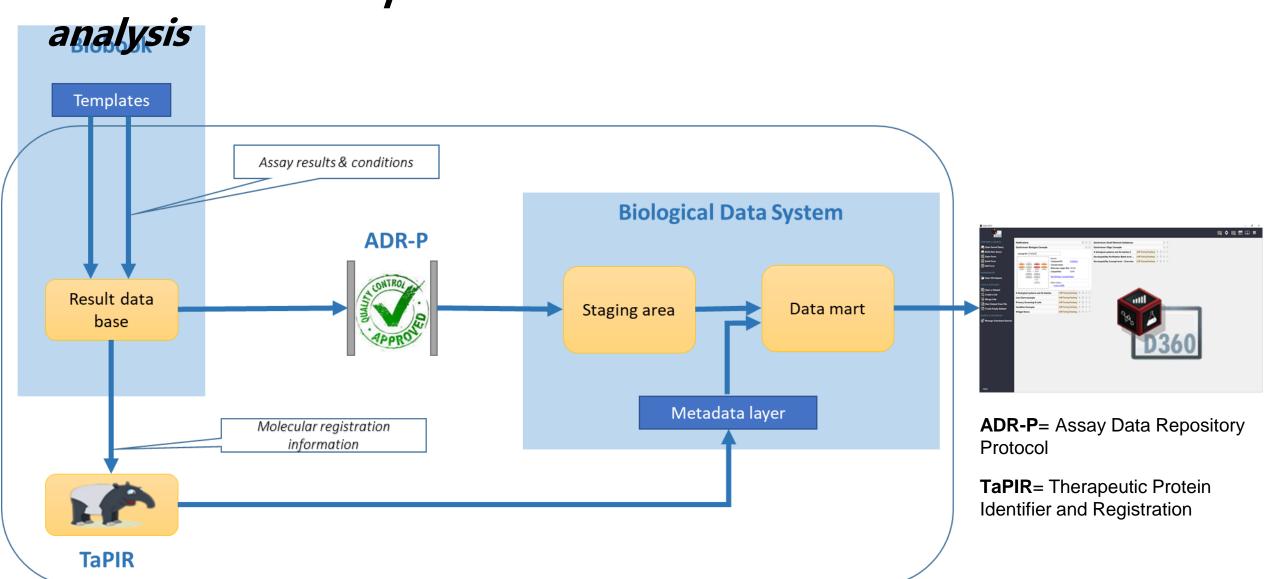
Material Supply

Process Development

Ph 0/1



The CAIMAN solution A modular landscape and foundation for advanced data



Implementation of quality controlled assay information



Assay data upload: IDBS BioBook templates

Quality control: Roche ADR-P System

Assay Data warehouse: Roche BDS System

From detailed assay description

Purification of antibodies and antibody like constructs on a liquid handling platform using OPUS robocolumns.

LIMS protocol name:
Assay owner:

Title:

Date/Site: 27.03.2018

ASSAY PURPOSE & PRINCIPLE

Protein samples are purified from cellculture supernatants before being tested in assays, because cell culture medium can disturb assay results. Another reason is that proteins are concentrated for further analysis, in order to improve signal-to-noise ratio in the assay.

A common technique is affinity chromatography. It is based on specific interaction between antibodies (or antibodylike constructs, Fabs...) and chromatography material (see Figure 1). The antibodies bind to the column and are separated from the undesired material that does not interact. Washing the stationary phase with washing buffer, undesired molecules are eluted first while the desired proteins are eluted using an acidic buffer.

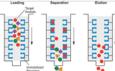
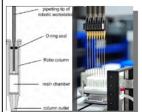


Figure 1: Principle of affinity chromatography

For the fully automated purification of antibodies in parallel, the liquid handling robot Freedom EVO from Tecan in combination with OPUS robocolumns (Replicen) (see Figure 2) is used



The robocolumns can be filled with any commercial available chromatography material. Loading of the robocolumn is performed using an 8 channel liquid handling arm (LiHa) with steel needles.

Figure 2: OPUS Robocolumn

Protein concentration is determined by UV measurement at 280 nm. Sample volume is determined by measurement at 998 nm and 900 nm.

to template

Results: PRODUCT_CONCENTRATION, PRODUCT_VOLUME,

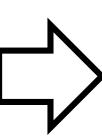
PRODUCT_AMOUNT

Conditions: LMR_PROJECT_PHASE,

CHROMATOGRAPHIC_COLUMN, PROJECT, BUFFER,

BUFFER_PH, TAPIR_BATCH

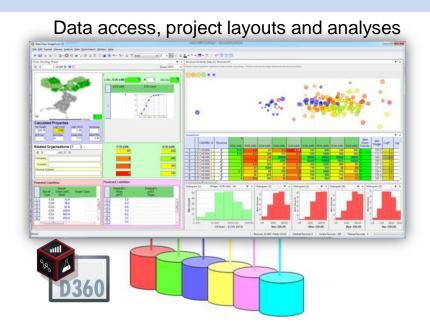


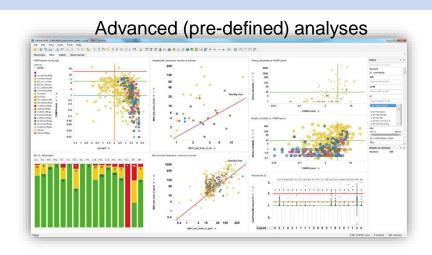


Technical & data query template coverage



- 23 data query templates (decision points) in total, 13 decision points ready
- 55 Spotfire visualizations supporting 8 decision points, 22 D360 visualizations supporting, 3 decision points
- 7 special data categories based on TaPIR data packages, 4 additional data sources integrated, 1 data source left





Multiple data sources (internal and external)

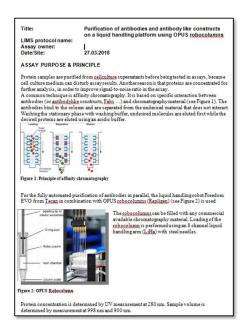
Technical & data query template coverage



631 experiments loaded to BDS/D360 PROD (128 Assay protocols, 117 LIMS protocols , 207.539 TaPIR batches, 1.184.545 data points, 16 TaPIR projects)

In total: 1085 Assay protocols created, 42 assay protocol owner

65 BioBook templates overall productive or close to productive

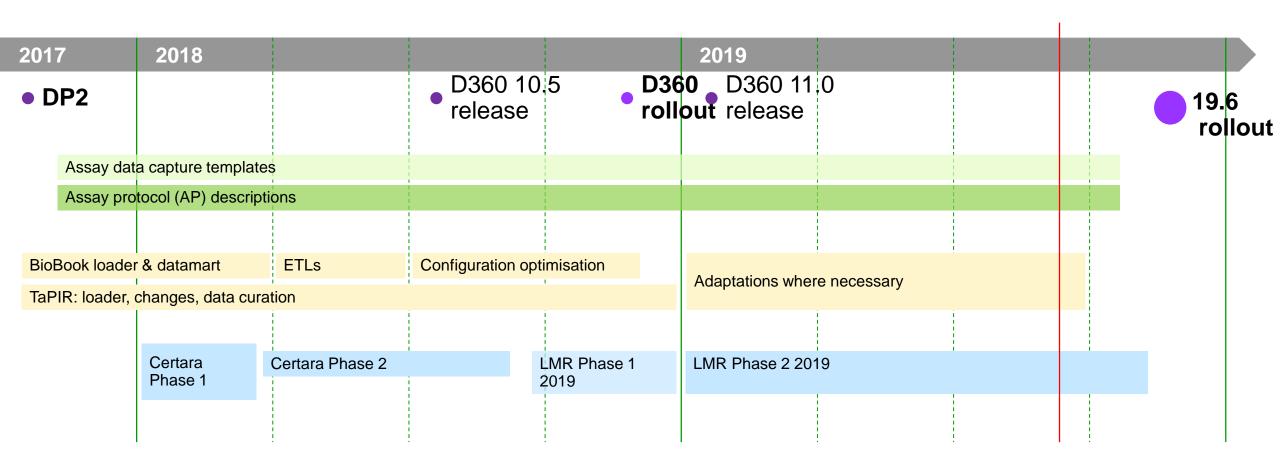


PRODUCT_A Conditions: CHROMATO	DDUCT_CONCENTRATION MOUNT LMR_PROJECT_PHASE, GRAPHIC_COLUMN, PRO TAPIR_BATCH		OLUME,
LIMS protocol reference:	IgG Micropurification Tecan/v1	ш	
Annotation scheme:	general	Protocol comment:	
Owner:	werneru	Editors: (-> RUD)	avinashn, werner
Last modified by:	rocheo	Detailed assay protocol: (-> <u>Template</u>)	Micropurification
Creation site:	PENZBERG	Data producer group:	PhysChem
NMP Project short name:	PHYSICOCHEMICAL_STUDIES $[_\to details]$	Used for multiple projects	s: No
Biosample & Target			
Biological system:	<u>in-vitro</u>	Species: Strain:	~NotApplicable
Tissue type:	~NotApplicable	Cell line: (-> RNCB)	
Subcellular fraction:	~NotApplicable	Assay target: (-> IDM)	

Timelines

Roche

CAIMAN delivery – high level activities





The CAIMAN project *Value*

The CAIMAN project helps to	By
shift focus from data collection to data analysis	 establishing an industry-leading project-data-analysis platform enabling comprehensive data access, tracking, comparison and visualization
support collaboration and faster decision-making	 streamlining and simplifying the compilation of project reports reducing elapse time to complete a particular stage of research
promote improved quality decisions	 ensuring provisioning of all project-relevant information providing multifaceted analysis capabilities tailored to project needs
enable a learning organization	 supporting rapid hypothesis testing (across projects) for knowledge generation facilitating cross-project learning using a sustainable platform to ensure data quality and

Acknowledgements



Business Team

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Project Organization

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Christophe Chabbert

D360 Support

Shijun Yu



Doing now what patients need next