



# HITS

Heidelberg Institute for  
Theoretical Studies

# Data management intro

Wolfgang Müller, HITS

Think beyond the limits!





# FAIRDOM Initiative



<http://fair-dom.org>

Findable  
Accessible  
Interoperable  
Reusable

Data  
Operations  
Models

- develop a community
- establish an internationally sustained Data and Model Management service
- Partners from U Manchester, HITS, U Zürich, ETH Zürich
- Funded by BMBF, ISBE, BBSRC, SystemsX (CH)
- Part of ELIXIR as “Node Resource” in UK and DE

# Topics

- Why data management?
- Some general thoughts leading up to the SEEK day and the openBIS day

# The reason: Reproducibility Crisis

- In a variety of subject matters
    - Many positive findings are false
  - Many papers to be trusted
  - But many not to be trusted
  - High profile misconduct incidents
- 
- Need for fast way to access data
    - Proving reproducibility
    - Making control easier
    - Making QC easier within labs

**Bharat Aggarwal** (US), a former Ransom Horne, Jr. Distinguished Professor of Cancer Research at the **University of Texas MD Anderson Cancer Center**,<sup>[8]</sup> resigned his position after fraud was discovered in 65 papers published by him in the area of **curcumin** as a treatment for cancer.<sup>[9]</sup> As of 2020 Aggarwal has had 29 of his research papers retracted, ten others have received an expression of concern, and 17 others have been corrected.<sup>[10][11]</sup>

**Why Most Published Research Findings Are False**  
John A. Ioannidis

Factors that influence the probability of a finding being true are:

- 1. The size of the study
- 2. The quality of the study
- 3. The consistency of the results
- 4. The plausibility of the results
- 5. The precision of the results
- 6. The statistical significance
- 7. The confidence interval
- 8. The p-value
- 9. The effect size
- 10. The heterogeneity of the results
- 11. The publication bias
- 12. The funding source
- 13. The author's reputation
- 14. The journal's reputation
- 15. The field's reputation
- 16. The time of the study
- 17. The location of the study
- 18. The population of the study
- 19. The intervention of the study
- 20. The outcome of the study

Wikipedia: The Free Encyclopedia

Wiki Loves Earth 2021 photo competition: take photos in nature and support Wikipedia.

## List of scientific misconduct incidents

From Wikipedia, the free encyclopedia

Scientific misconduct is the violation of the standard codes of **scholarly conduct** and **ethical behavior** in the publication of professional **scientific research**. A *Lancet* review on *Handling of Scientific Misconduct in Scandinavian countries* gave examples of policy definitions. In Denmark, scientific misconduct is defined as "intention[al] or gross negligence leading to fabrication of the scientific message or a false credit or emphasis given to a scientist", and in Sweden as "intention[al] distortion of the research process by fabrication of data, text, hypothesis, or methods from another researcher's manuscript form or publication; or distortion of the research process in other ways."<sup>[1][2]</sup>

A 2009 **systematic review** and **meta-analysis** of survey data found that about 2% of scientists admitted to falsifying, fabricating, or modifying data at least once.<sup>[3]</sup>

This is a **dynamic list** and may never be able to satisfy particular standards for completeness. You can help by **adding missing items** with **reliable sources**.

### Contents [hide]

- 1 Biomedical sciences
- 2 Chemistry
- 3 Computer science and mathematics
- 4 Philosophy
- 5 Physics and engineering
- 6 Plant biology
- 7 Social sciences
- 8 Other

# Consequence: drive for FAIR data

- Next slides:
- What is FAIR
- And what do funders ask for, at the example of DFG



# What is FAIR?

## Data should be like money



## To be useful, money must be...



findable



accessible



interoperable



reusable



# But data is more

Principle	Meaning
<b>Findable</b>	Identified, metadata enriched, searchable
<b>Accessible</b>	Locatable, standardized protocols
<b>Interoperable</b>	Metadata rich, sustained, metadata survives data
<b>Reusable</b>	Licensed data&metadata, versatile metadata

The image features a composite of Earth from space, divided into vertical strips. The strips show different parts of the planet, including blue oceans, white clouds, and brown landmasses. The text "The stick" is overlaid in the center in a white, sans-serif font.

The stick

# Funders want this



Wissenschaftliche Integrität

Deutsche Forschungsgemeinschaft

START
KODEX
WI DURCH DIE DFG
AKTUELLES

ENGLISH

## Kodex - Leitlinien zur Sicherung guter wissenschaftlicher Praxis

In diesem Portal finden Sie den Kodex "Leitlinien zur Sicherung guter wissenschaftlicher Praxis" der Deutschen Forschungsgemeinschaft (DFG). Ziel des Kodex ist es, eine Kultur der wissenschaftlichen Integrität in der deutschen Wissenschaftslandschaft zu verankern. In 19 Leitlinien werden angemessene Standards für wissenschaftliches Arbeiten beschrieben und praxisbezogen kommentiert. Hochschulen und außerhochschulische Forschungseinrichtungen setzen die Leitlinien als Voraussetzung für den Erhalt von Fördermitteln durch die DFG um.

ZUM KODEX

### Über dieses Portal

Die praxisnahe und fachspezifische Kommentierung des Kodex steht im Zentrum des Portals; die Deutsche Forschungsgemeinschaft (DFG) sichert die Qualität der Beiträge. Ferner finden Sie Informationen dazu, wie sich die DFG im Themenfeld „Wissenschaftliche Integrität“ engagiert. Aktuelle Themen sowie Veranstaltungshinweise runden das Portal ab.

### Häufig gesuchte Schlagworte

AUTORSCHAFT
PUBLIKATION
ARCHIVIERUNG

DOKUMENTATION
BERUFSETHOS
FAIR-PRINZIPIEN

FORSCHUNGSDATEN
QUALITÄTSSICHERUNG

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Sign up

September 15, 2019
Journal article
Open Access

## Guidelines for Safeguarding Good Research Practice. Code of Conduct

Deutsche Forschungsgemeinschaft

The DFG's Code of Conduct "Safeguarding Good Research Practice" represents the consensus among the member organisations of the DFG on the fundamental principles and standards of good practice and are upheld by these organisations. These guidelines underline the importance of integrity in the everyday practice of research and provide researchers with a reliable reference with which to embed good research practice as an established and binding aspect of their work.

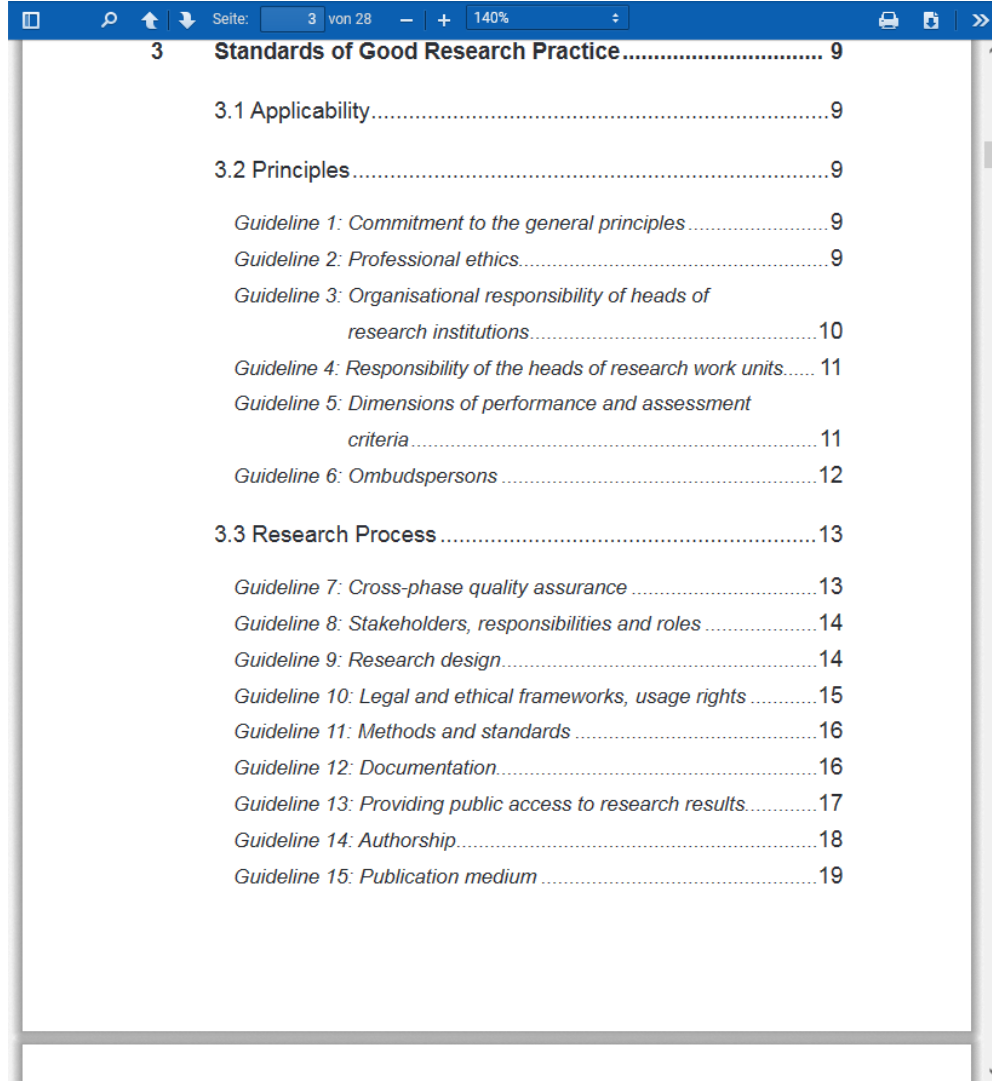
Available in German and in English.

6,483 views
7,757 downloads
See more details...

Indexed in

Publication date:  
September 15, 2019  
DOI:  
10.5281/zenodo.3923602  
Keyword(s):  
Wissenschaftliche Integrität, gute wissenschaftliche Praxis, wissenschaftliches Publikationsverhalten, Leitlinien zur Sicherung guter wissenschaftlicher Praxis, Deutsche Forschungsgemeinschaft, DFG, Kodex, good scientific practice, research integrity, scientific misconduct, German research foundation, code of conduct  
Imprint:  
Deutsche Forschungsgemeinschaft  
Communities:  
Path2Integrity  
License (for files):  
CC Creative Commons Attribution Share Alike 4.0

# Rough overview of GSP guidelines of DFG



The screenshot shows a PDF viewer window with a blue header bar. The header bar contains navigation icons and text: 'Seite: 3 von 28', a zoom level of '140%', and a right arrow icon. The main content area displays a table of contents for a document titled 'Standards of Good Research Practice'. The table lists sections and their corresponding page numbers. The sections are: 3 Standards of Good Research Practice (9), 3.1 Applicability (9), 3.2 Principles (9), Guideline 1: Commitment to the general principles (9), Guideline 2: Professional ethics (9), Guideline 3: Organisational responsibility of heads of research institutions (10), Guideline 4: Responsibility of the heads of research work units (11), Guideline 5: Dimensions of performance and assessment criteria (11), Guideline 6: Ombudspersons (12), 3.3 Research Process (13), Guideline 7: Cross-phase quality assurance (13), Guideline 8: Stakeholders, responsibilities and roles (14), Guideline 9: Research design (14), Guideline 10: Legal and ethical frameworks, usage rights (15), Guideline 11: Methods and standards (16), Guideline 12: Documentation (16), Guideline 13: Providing public access to research results (17), Guideline 14: Authorship (18), and Guideline 15: Publication medium (19).

<b>3</b>	<b>Standards of Good Research Practice</b>	<b>9</b>
3.1	Applicability	9
3.2	Principles	9
	<i>Guideline 1: Commitment to the general principles</i>	9
	<i>Guideline 2: Professional ethics</i>	9
	<i>Guideline 3: Organisational responsibility of heads of research institutions</i>	10
	<i>Guideline 4: Responsibility of the heads of research work units</i>	11
	<i>Guideline 5: Dimensions of performance and assessment criteria</i>	11
	<i>Guideline 6: Ombudspersons</i>	12
3.3	Research Process	13
	<i>Guideline 7: Cross-phase quality assurance</i>	13
	<i>Guideline 8: Stakeholders, responsibilities and roles</i>	14
	<i>Guideline 9: Research design</i>	14
	<i>Guideline 10: Legal and ethical frameworks, usage rights</i>	15
	<i>Guideline 11: Methods and standards</i>	16
	<i>Guideline 12: Documentation</i>	16
	<i>Guideline 13: Providing public access to research results</i>	17
	<i>Guideline 14: Authorship</i>	18
	<i>Guideline 15: Publication medium</i>	19

- Comprehensive standards of good research practice
- Many suggestions with an impact on data management:
  - Quality control
  - Long-term preservation
  - Publishing of research



The background of the image is a composite of several vertical strips showing different views of Earth from space. The strips are arranged side-by-side, creating a mosaic effect. The colors range from deep blues of the oceans to light blues and whites of the clouds, and some strips show brownish-yellow landmasses. The text 'The carrot' is written in a large, white, serif font, centered horizontally across the middle of the image. The text is slightly transparent, allowing the underlying image to be seen through it.

The carrot



The background is a composite image of Earth from space, divided into vertical strips of different colors and textures. The strips show various views of the planet, including blue oceans, white clouds, and brown landmasses. The text is overlaid in the center in a white, sans-serif font.

Imagine: You want to  
discover a  
project/group/institute

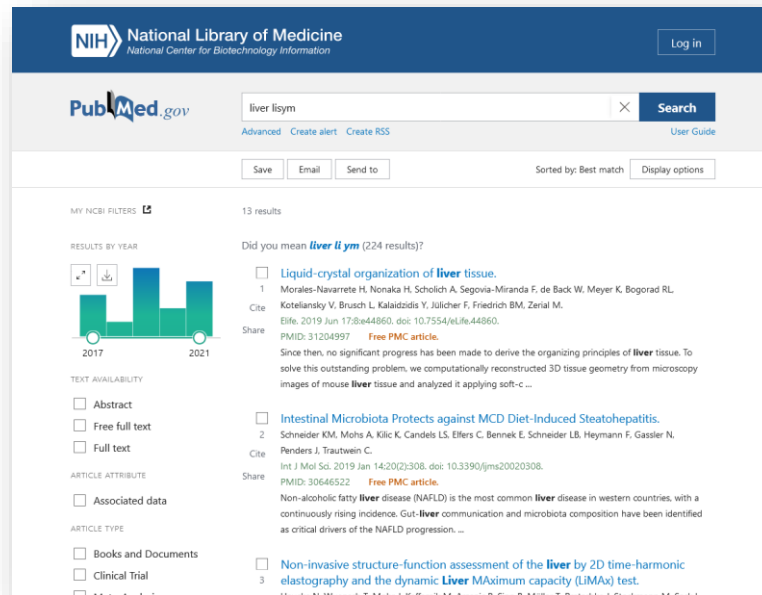


The background is a composite image of Earth from space, divided into vertical strips of different colors and textures. The strips show various parts of the planet, including oceans, continents, and clouds. The text "For example: LiSym" is overlaid in white, sans-serif font, centered horizontally and vertically.

For example:  
LiSym

# Publications

- PubMed: „Liver Lism“ → 13 results







# Now, let's look at

- Models
- Data

# Models: Model DBs & GitHub/GitLab



JWS Online Model Database Simulation Database Manuscript Database Help Servers

New user Reset password Log in

**JWS ONLINE**

**Biological systems modelling**

JWS Online is a Systems Biology tool for the construction, modification and simulation of kinetic models and for the storage of curated models.

**Construct your own model**

With the new **JWS Online model builder** you can build a model from scratch using a simple interface. Models can be simulated directly in the **JWS Online simulator**. The builder adheres very closely to the **SBML** model specification and supports the **MIRIAM** and **SBO** standards for annotation, and provides a useful online annotation tool.

**Simulate models**

**PEtab -- a data format for specifying parameter estimation problems in systems biology**

PEtab is a data format for specifying parameter estimation problems in systems biology. This repository contains the PEPab specifications and additional documentation.

**About PEPab**

PEtab is built around **SBML** and based on tab-separated values (TSV) files. It is meant as a standardized

EMBL-EBI Services Research Training About us EMBL-EBI

**BioModels**

Individual Models Search

What is this box used for? Examples: MAPK cascade, homo sapiens lung cancer Search box

Home Browse Submit Curation Help About us Contact us Feedback Login Register

BioModels is a repository of mathematical models of biological and biomedical systems. It hosts a vast selection of existing literature-based physiologically and pharmacologically relevant mechanistic models in standard formats. Our mission is to provide the systems modelling community with reproducible, high-quality, freely-accessible models published in the scientific literature. More information about using BioModels such as [model submission](#), [updates](#), [publications](#) can be found in the [FAQ](#).

Submission / Update Manually Curated 1,002 models Non-curated 1,264 models Auto generated 833 models GO Chart 1,132 classes BioModels Parameters 228,842 records

**Model of The Month** March, 2021

**Ito2019 - Gefitinib resistance of lung adenocarcinoma caused by MET amplification**

**Browse by Journal** This shows models distribution based on journals. Move the mouse over a bubble to see the information.

**Find us on Twitter** Tweets by @biomodels

With support from @git4training team, we are organizing "Mathematics of Life", a beginners course to learn systems biology modelling. All Open to participants from

ag of your personal data in order to function. By using the Notice and Terms of Use. I agree, dismiss this banner

**PLOS COMPUTATIONAL BIOLOGY**

RESEARCH ARTICLE

**PEtab—Interoperable specification of parameter estimation problems in systems biology**

Published January 19, 2021 • <https://doi.org/10.1371/journal.pcbi.1008048>

**Abstract**

Reproducibility and transparency of the results of data-based modeling studies are essential. This has been a long-term goal in the field of systems biology. In this paper, we present the PEPab (Parameter Estimation Problems in Systems Biology) format, a standardized format for specifying parameter estimation problems in systems biology. The format is designed to be interoperable with existing formats and to be easy to use. It is based on the SBML (Systems Biology Markup Language) format and is designed to be compatible with existing tools. The format is designed to be interoperable with existing formats and to be easy to use. It is based on the SBML (Systems Biology Markup Language) format and is designed to be compatible with existing tools.

**Author summary**

Parameter estimation is a common and essential task in modeling. In many studies, data are sparse and noisy, and the models are complex. This makes the task of estimating the parameters of a model difficult. In this paper, we present the PEPab (Parameter Estimation Problems in Systems Biology) format, a standardized format for specifying parameter estimation problems in systems biology. The format is designed to be interoperable with existing formats and to be easy to use. It is based on the SBML (Systems Biology Markup Language) format and is designed to be compatible with existing tools.

# Datacollections, Zenodo, web apps

**Gene Expression Omnibus**

GEO is a public functional genomics data repository supporting MIAME-compliant data submissions. Array- and sequence-based data are accepted. Tools for gene expression profiles.

**Transcriptomic cross-species analysis of chronic liver disease reveals a consistent regulation pattern between humans and mice**

The results and the analysis scripts presented on this website ensures the reproducibility of all bioinformatics related findings presented in *"Transcriptomic cross-species analysis of chronic liver disease reveals a consistent regulation pattern between humans and mice"*. Please see below to get more information about the individual analysis and some tips on how to reproduce the results.

**Abstract**

**Background and aims**

Mouse models are frequently used to study chronic liver diseases (CLD). To assess their translational relevance, we quantified the similarity of commonly used mouse models to human CLD based on transcriptome data.

**Methods**

Gene expression data from 372 patients were compared to data from acute and chronic mouse models comprising 227 mice, and additionally to nine published gene sets of chronic mouse models. Genes consistently altered in humans and mice were mapped to liver cell types based on single-cell RNA-sequencing data and validated by immunostaining.

**Results**

Considering the top differentially expressed genes, the similarity between humans and mice varied among the mouse models and depended on the period of damage induction. The highest recall (0.4) and precision (0.33) were observed for the model with 12 months damage induction by CCl<sub>4</sub> and by a Western type damage, respectively. Genes consistently upregulated between the chronic CCl<sub>4</sub> model and human CLD were enriched in inflammatory and developmental processes, and mapped to cholangiocytes, macrophages, endothelial and mesenchymal cells. Downregulated genes were enriched in metabolic processes and mapped to hepatocytes. Immunostaining confirmed the regulation of selected genes and their cell type specificity. Genes that were upregulated in both acute and chronic models showed higher recall and precision with respect to human CLD than exclusively acute or chronic genes.

**Conclusion**

**zenodo**

February 27, 2021

**Transcriptomics directory of liver disease in human and mouse**

108 views, 6 downloads

DOI: 10.5281/zenodo.4148008

Publication date: February 27, 2021

License (for file): CC BY-NC-ND 4.0 International

**PK-DB: pharmacokinetics database for individualized and stratified computational modeling**

Grzegorzewski J, Brandhorst J, Green K, Elthneradur D, Dupont Y, Barbach F, Köller A, Kie D, De Angelis S, König M. Nucleic Acids Res. 2021 Jan 8;49(D1):D1358-D1364. doi: 10.1093/nar/gkab990. PMID: 33151297

**Data**

Any pharmacokinetics study contains subjects under investigation. These subjects are characterized by properties like their sex, age, body weight, health status, and further accessible pharmacokinetics influencing characteristics. In PK-DB this data is saved as groups and individuals. Next, some kind of interventions are performed on the subjects, which is mostly a dosing of a substance to the body of the subject. Finally, pharmacokinetics measurements are performed on the subject. These are often some kind of concentration profiles in some tissue of the subject. Additionally, derived pharmacokinetics parameters e.g. AUC, clearance, or half-lives are commonly reported. Correlations between these outputs are often shown in form of scatter plots.

Studies	Groups	Individuals	Interventions	Outputs	Timecourses
614	1897	10074	1546	181824	127386



# Summarising

- Many specialized data sources
- Many useful data locations
- Quite some data merged – curated
- Looking at all of them: too cumbersome
- Data management enables to have
  - one place to find them all
  - one place to standardize
- Needs to be adapted to
  - Remote data
  - Local data



FAIRDOM



SEEK

Be the hub for  
heterogeneous, big  
projects, store data &  
models, too



FAIRDOM

Be the data management  
for groups (and possibly  
bigger)



openBIS

# Now

- Talk by Piotr Zadora, DKFZ  
Covering the use of openBIS  
in the Klingmüller Department at DKFZ
- Then some more work about challenges  
to address in DM or DMM
- Then „the SEEK day“
- Followed by „the openBIS day“

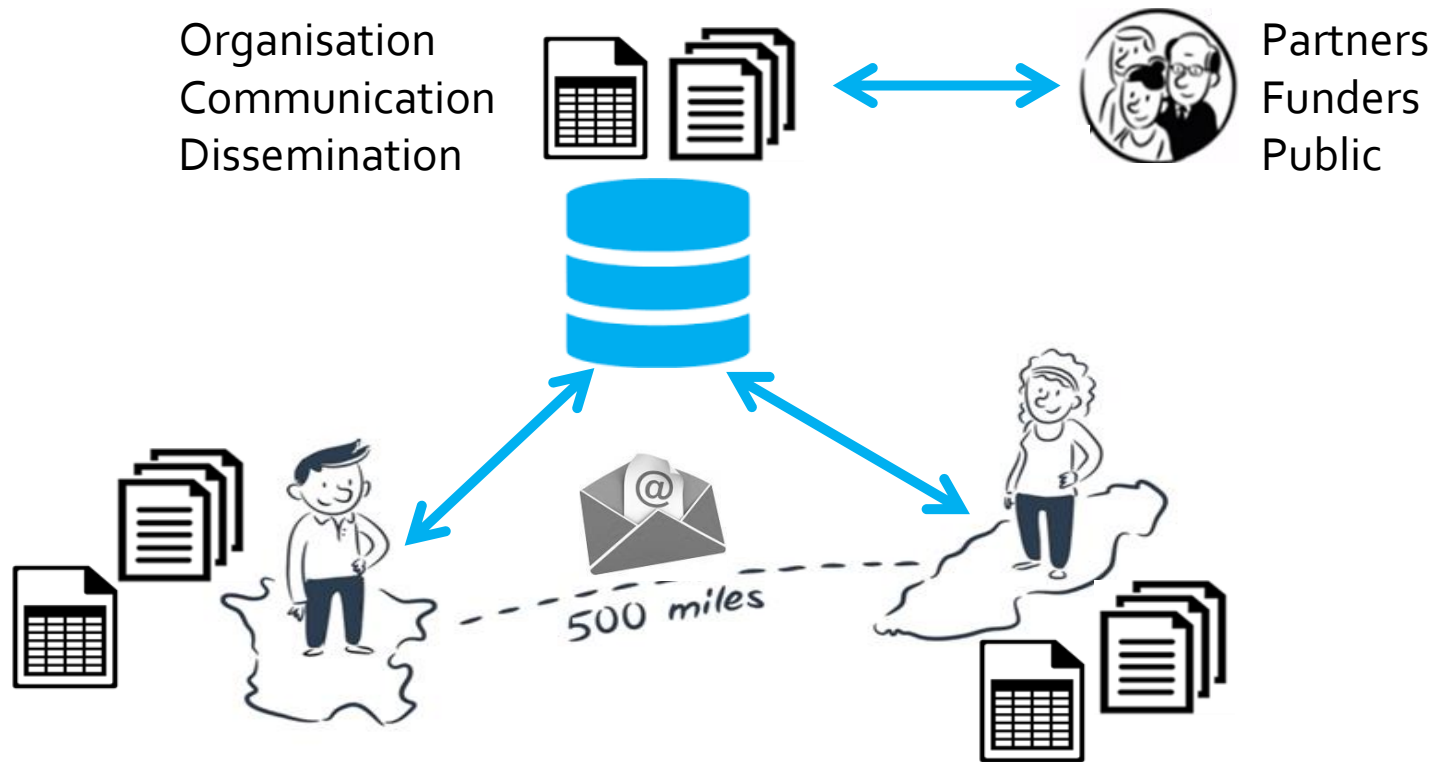
The background is a composite image of Earth from space, divided into vertical strips of different colors and textures. The strips show various parts of the planet, including oceans, continents, and clouds. The text "Some more general words about DM" is overlaid in white, centered on the image.

Some more  
general words  
about DM



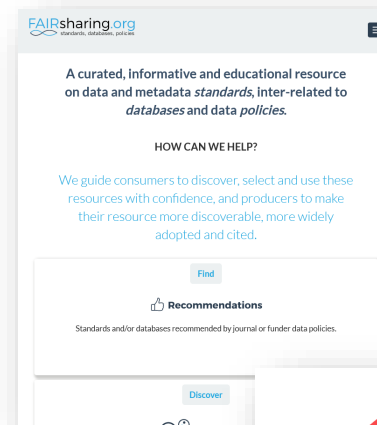
- Strong push by lead
- Onboarding by proficient users
- Jupyter Notebooks in openBIS
- Data models  
(FAIRDOM support but scientific tech  
organiser, internally, too)

# Goal: Project Data Management



# Challenges to address

- Social & organisational
- Conceptual
  - Data structures
  - Ontologies
  - Standards
- Tooling
  - Which tools to use
  - How to combine



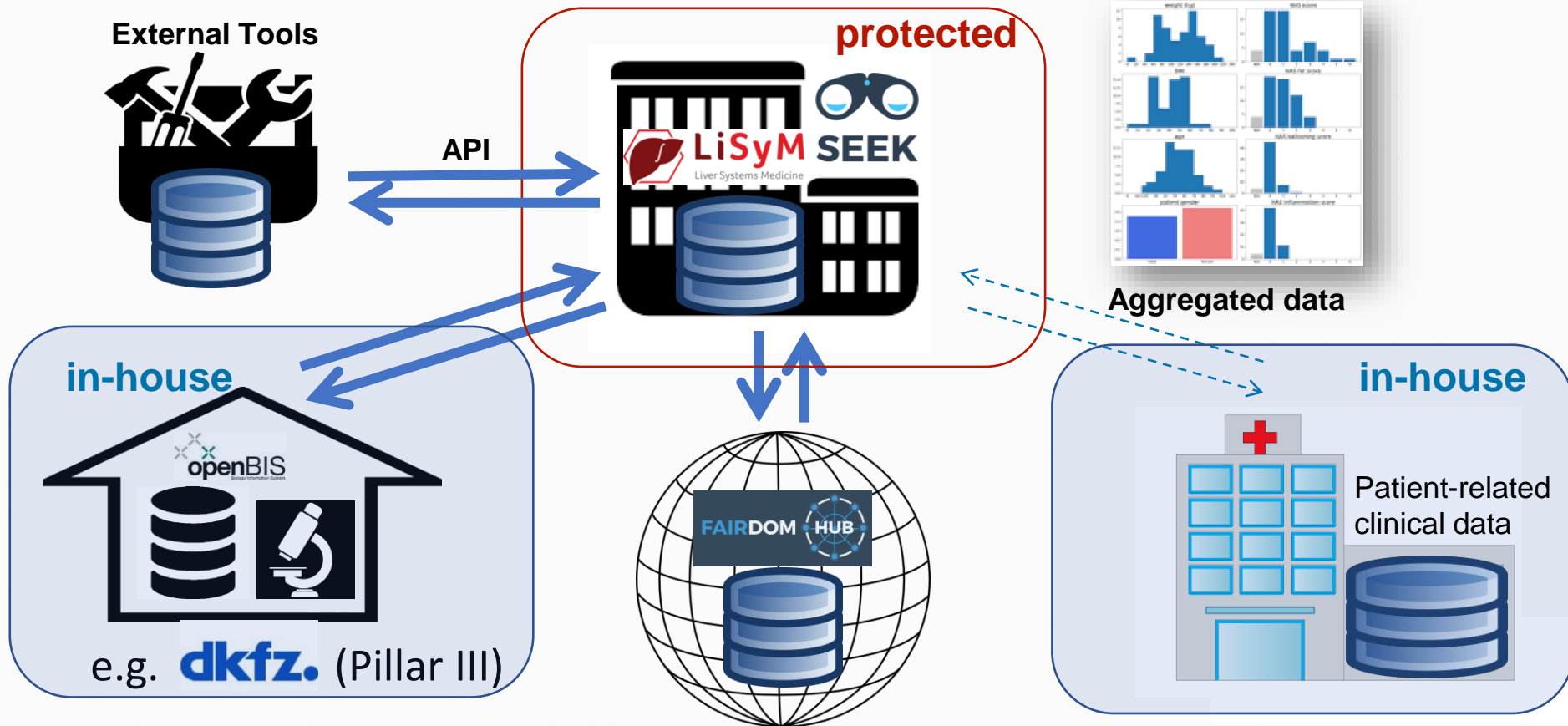
# Assembling data management processes

- Collect raw and processed (secondary) data, models & metadata together with experimental context
- Organise and link assets
- Prepare reproducible publications
- Use standardised metadata
- Share with colleagues and public
- Integrate with legacy, home grown, external systems
- Reuse tools and community archives



**RDMkit**  
<https://rdmkit.elixir-europe.org>

# Structuring deployment: LiSyM Example





# Assembling vs. Integrating: integrated JWS modelling tool in SEEK-based FAIRDOMHub

FAIRDOM HUB

Search here...

Search

Register

Log in

solventogenesis Version 1

No description specified

SEEK ID: <https://fairdomhub.org/models/43?version=1>

1 item is associated with this Model:

Haus1.xml (SBML and XML document - 22.1 KB)

Organism: Not specified

Model type: Not specified

Model format: SBML

Execution or visualisation environment: JWS Online

Model image: No image specified

Selected: solventogenesis (Model)

Description: No description

SEEK ID: <https://fairdomhub.org/models/43>

solventogenesis

By continuing to use this site you agree to the use of cookies

I agree

FAIRDOM HUB

Search here...

Search

Register

Log in

PGK 70C SBML - JWS

Online Model Simulation Version 1

Back to Model

kouril6

Detail

Download

Schema

Time evolution

Steady-state

Parameter scan

Reaction plots

Manuscript

Notes

Documentation

Reactions

Parameters

Fixed species

Initial values

Functions and Rules

Events

Constraints

pep

ADP

BPG

P3G

ATP

pyr

v1

v2

v3

pi

☒ Snap to grid

☐ Show wireframe

☐ Show modifiers

☐ Show compartments

Gravity\*  
0.3

Repulsion\*  
1000

Off On

Pool threshold\*

By continuing to use this site you agree to the use of cookies

I agree

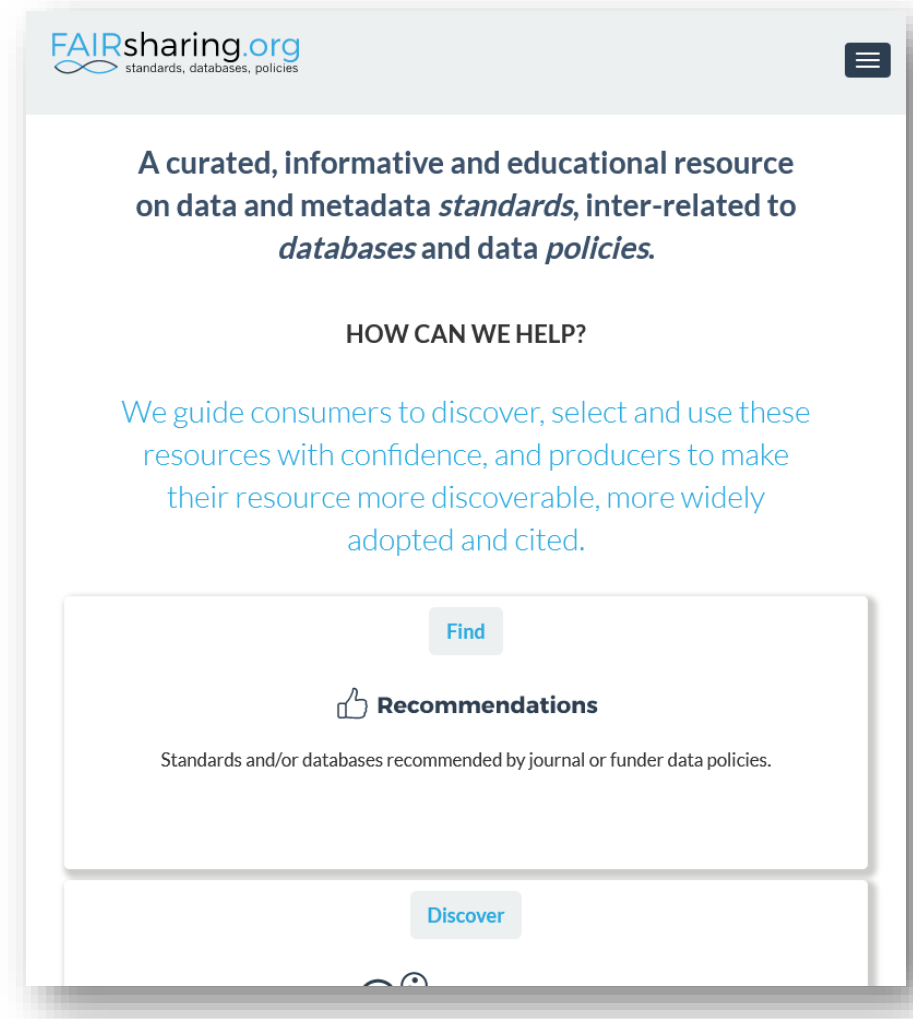
[https://fairdomhub.org/models/284/simulate?version=1&constraint\\_based=0](https://fairdomhub.org/models/284/simulate?version=1&constraint_based=0)

# Ontologies and standards: FAIRsharing.org

## How to structure your data?

Reuse other people's work

- Community standards
  - 1532 Standards  
(e.g. the MIBBI standards)
  - 1773 Databases
  - 140 Policies



# Social

- **Insight:** Data management means **change**
- **Challenge: How to include people?**
  - How to get information
  - Useful suggestions how to do better
  - Refinement of workflows
  - **Strive for painless ingestion and useful outcome**
- **Proposal:** Have responsables on per-project basis
- Regular meetings with data management responsables
- Who should be responsible?  
Depends on **your** organisation

The FAIRDOM approach:  
Have „PALs“, a focus group in  
the project



PALs kickoff of FAIRDOM  
2013

# Organisational: Data sharing policy

Example: LiSyM DSP, derived from ERASysAPP DSP

- Expression of intent
- Who is responsible?
- Who shares with whom?
- When?
- What?
- What in case of disagreement?





# Spend the right amount of effort on this

Try to hit  
the sweet  
spot here



Too  
little

Too  
much

# Now: Front End

Find, Access and Organise assets



- Upload data
- Link to data regardless of physical store
- Sharing
- ISA structure
- Yellow pages and collaboration
- Supplementary data for publications
- Standards-compliant

# Thanks, Questions?

Wolfgang.mueller@H-ITS.org

