

# smith

*Smart Medical Information  
Technology for Healthcare*

Markus Löffler  
Leipzig



UNIVERSITÄT  
LEIPZIG

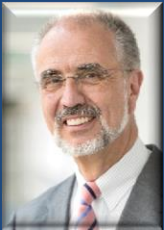


Friedrich-Schiller-Universität Jena

RWTHAACHEN  
UNIVERSITY

# smith

*Smart Medical Information  
Technology for Healthcare*



Prof. Dr. Wolfgang E. Fleig



Prof. Dr. Uli Hahn



Andreas Henkel



Prof. Dr. Gernot Marx



Volker Lowitsch



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Friedrich-Schiller-Universität Jena



- Long-standing cooperation in clinical research
  - Infectious diseases, intensive care medicine
- Complementary methodological expertise in
  - clinical trial research
  - epidemiology
  - medical informatics
  - systems medicine and disease modelling
  - natural language processing
- Management experience of large research clusters
- State-of-the-art intersectoral IT services for healthcare

# Use cases

**smith**

*Smart Medical Information  
Technology for Healthcare*



## **HELP**

Hospital-wide EMR-based computerized decision support system to improve outcomes of patients with bloodstream infections



## **ASIC**

Algorithmic Surveillance of ICU patients to improve personalized management of care



## **PheP**

Phenotype Pipeline, algorithms for phenotyping and NLP on EMR data

Source: UKJ (A. Schroll, M. Szabó), NVHR

# Use Case HELP

## EMR-based decision support for bloodstream Infections



PIs:  
Pletz, clinician  
Scherag, biometry  
Jena

- Setting: Normal wards and ICUs
- Application of EMR-based computerized decision support system to improve outcomes of patients with bloodstream Infections
- Development of **digitalized Antibiotic Stewardship** based on harmonized structured and unstructured data from diverse information systems
- **SMITH App** as user interface
- Outcomes: personalized management of bloodstream infections, reduced hospital readmission, reduced mortality

# Use Case ASIC

## Algorithmic Surveillance of ICU patients



PI:

Gernot Marx, Clinician

Andreas Schuppert, Modelling  
Aachen

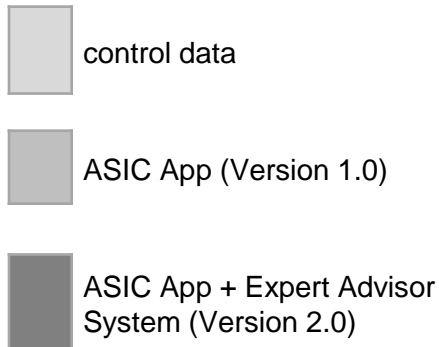
- Setting: Intensive Care Units (ICUs) (Respiration, ARDS, Sepsis)
- Application of „**High-Performance Computing**“ for model-based, clinical decision support
- Development of **virtual patient models** for clinical research (partners: Research Centre Jülich, Bayer AG)
- **SMITH App** as user interface
- Outcomes: personalized management of ARDS, reduced organ dysfunctions, reduced mortality

# Generating Evidence

## ASIC

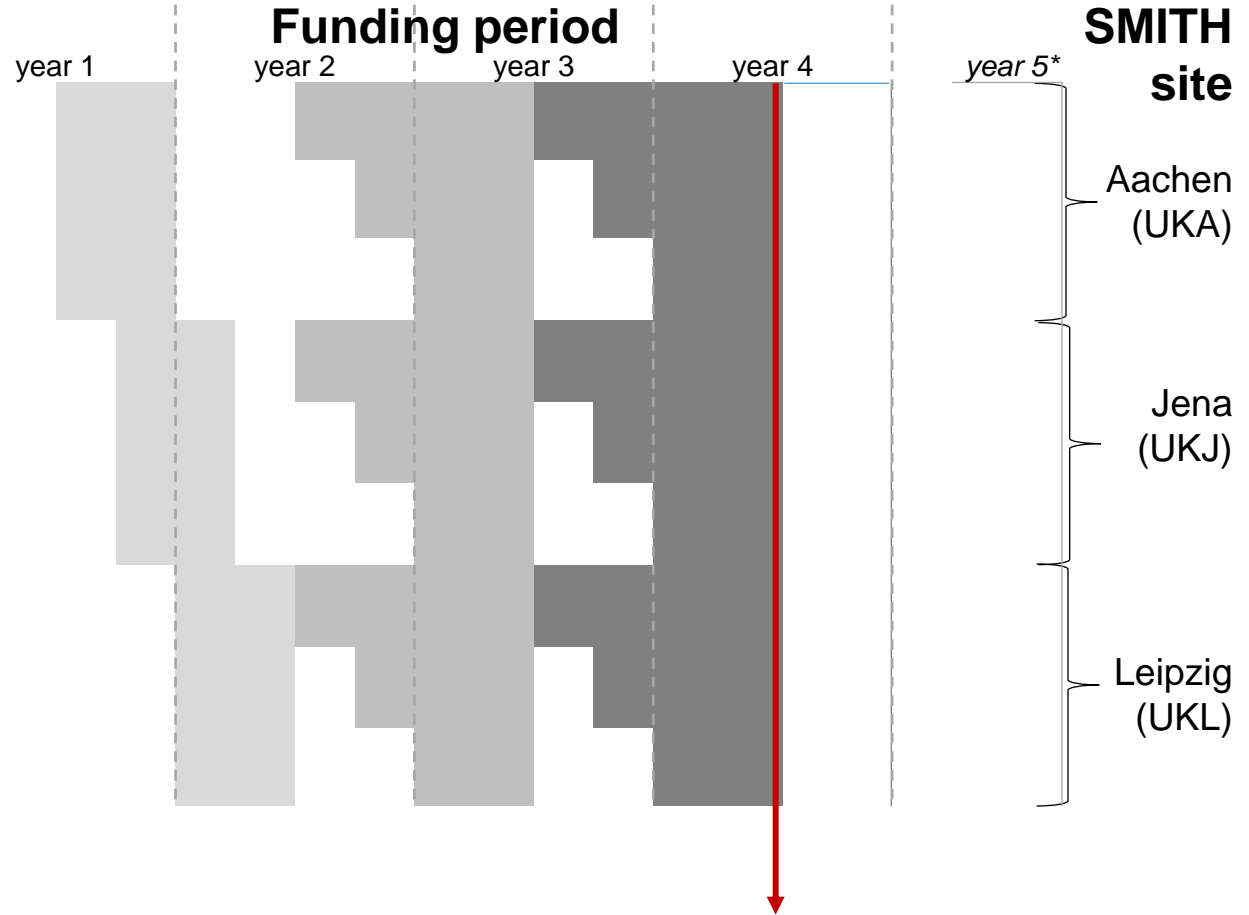
### Study design

(sequential stepped wedge designs)



Biometry

Andre Scherag



Outcome: hard clinical endpoints



# Use Case PheP

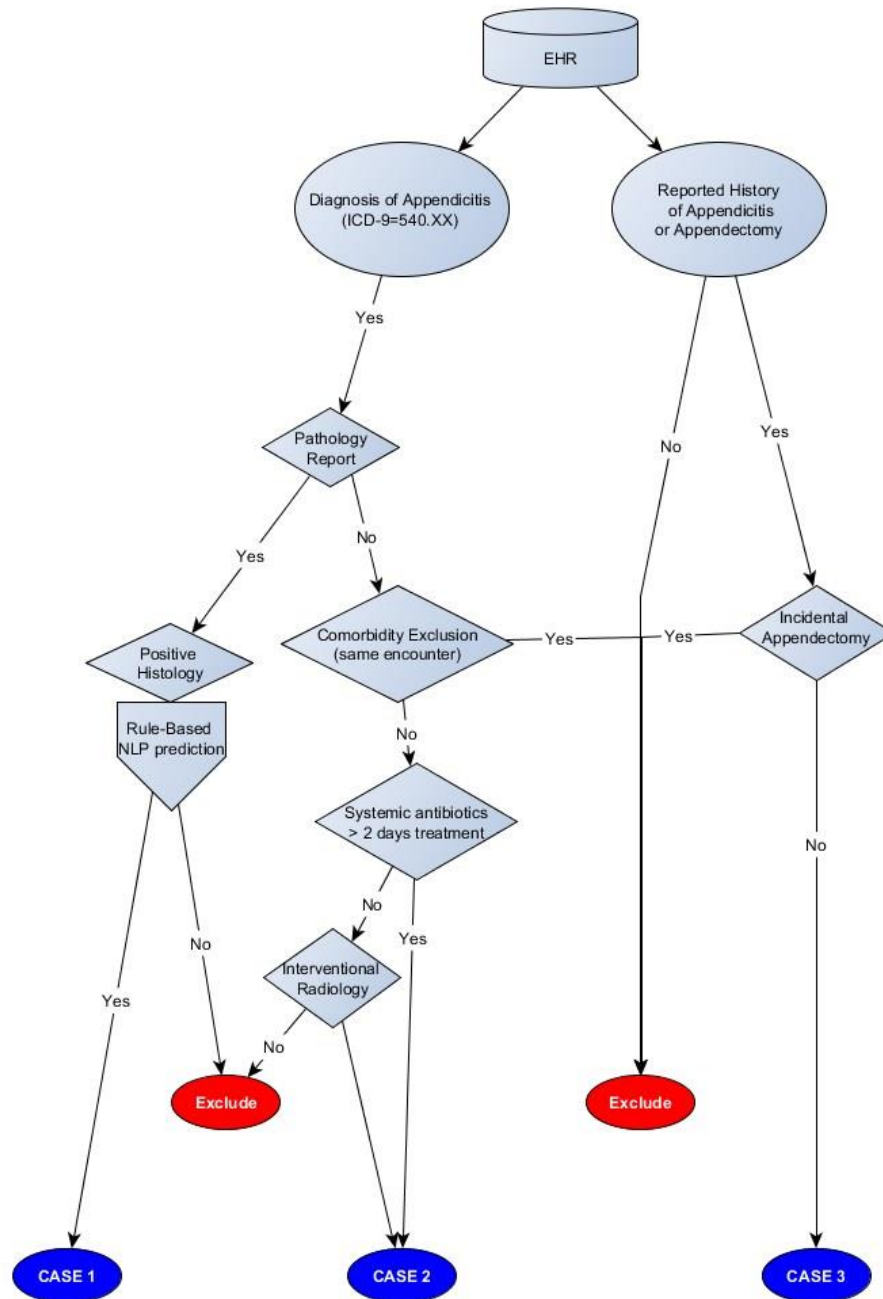
## Phenotype pipeline and NLP



PIs:  
M Löffler, classifiers & models  
U Hahn, NLP

- **Algorithms for phenotyping** by using structured and unstructured data from electronic medical records (EMR) → classification, annotation
- Development of
  - a rules engine and factory and a meta data repository
  - a natural language processing engine and factory and text corpus
- create a technology to automatically mine EMR and generate phenotype classifications and annotations (the uses cases are two examples)





Web Table 2. Characteristics of Heart Failure Status Definitions			
HF Status	Presence of ICD9 AND positive mention of HF	Heart Failure Date	Ejection Fraction
Definite	Yes	365 day window	No EF EF <50% (HF Type = 1) EF ≥50% (HF Type = 2)
Probable	Yes (or ≥5 unique dates of either)	365- 1825 day window	No EF EF <50% (HF Type = 1) EF ≥50% (HF Type = 2)
Possible	Either or none if EF <50	Unable to assign date	any
Control	None	N/A	No EF EF ≥50

**Heart Failure Date Assignment Rules:** Taking the cross product of all the unique

# NLP-Ejection fraction

**Web Table 4. Regular Expressions\* for Reporting of Ejection Fraction in Unstructured Reports**

<ul style="list-style-type: none"> <li>• Calculated EF ###</li> <li>• Calculated LVEF ###</li> <li>• Calculated LV ejection fraction ###</li> <li>• Calculated Left Ventricular ejection fraction ###</li> <li>• Calculated Ejection Fraction ###</li> <li>• Calculated Ejection Fraction ###. Visual estimate ###-###</li> <li>• Estimated EF ###</li> <li>• Estimated EF = ###</li> <li>• Estimated EF ###-###</li> <li>• Estimated Ejection Fraction ###</li> <li>• Estimated Ejection Fraction ###-###</li> </ul>	<ul style="list-style-type: none"> <li>• Estimated Left Ventricular Ejection Fraction ###</li> <li>• Estimated Left Ventricular Ejection Fraction ###-###</li> <li>• Estimated Left Ventricular Ejection Fraction range ###-###</li> <li>• EF ###</li> <li>• Ejection Fraction ###</li> <li>• LVEF ###</li> <li>• LVEF ~ ## - ###Left Ventricular Ejection Fraction ###</li> <li>• Visual Estimate of LVEF ###</li> <li>• Visual estimate of Left Ventricular Ejection Fraction ###</li> <li>• Visual Estimate of EF ###</li> <li>• Visual Estimate of Ejection Fraction ###</li> </ul>
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*\*The regular expression list above includes the variations identified during the algorithm validation at Mayo Clinic and Group Health and thus is not an exhaustive list of every possible combination of the use of characters such as "=" or "~".*

# Natural Language Processing

**Objective: Building a nucleus for text analysis of German medical records**

- **Automating** content-focused analyses
- Build up of accessible well annotated **clinical corpora** (gold standard for system evaluation, training data for machine learning of classification models)
- Development of novel, **semantics-focused software** (primarily dealing with clinical named entities, their relations, and temporal dependencies among events)
- Formation of a dedicated **NLP-pipeline** useable by the **DICes**
- **Collaboration** with leading national players in biomedical Natural Language Processing (NLP), from industry (Averbis, Freiburg and ID, Berlin) and academia (JULIE Lab, Prof Hahn, FSU Jena)

# Outcome of a first corpus project

(Uli Hahn, M Loeffler)

- **Collection** of 2,360 German-language discharge summaries from roughly 3,000 patient records (EHR) taken from the three sites involved (J: 960, L: 850, AC:550)
- **Annotation with the BRAT tool** carried out by 8 annotators across the three sites (J: 5, L: 2, AC: 1)
- **Annotation of all medication information** contained in the discharge summaries on drug name, dosage, mode, frequency, duration, reasons for administration
- **Evaluation of the quality of annotations** by measuring the inter-annotator agreement (IAA) for the 5 annotators from Jena, using a centroid F-score: drug name, dosage and frequency (.95), mode (.75), duration (.70), reasons (<.50)
- **Evaluation of a-prototype medication recognition system** using F-score: dosage and frequency (.82), drug name (.50), mode (.21), reasons (–)

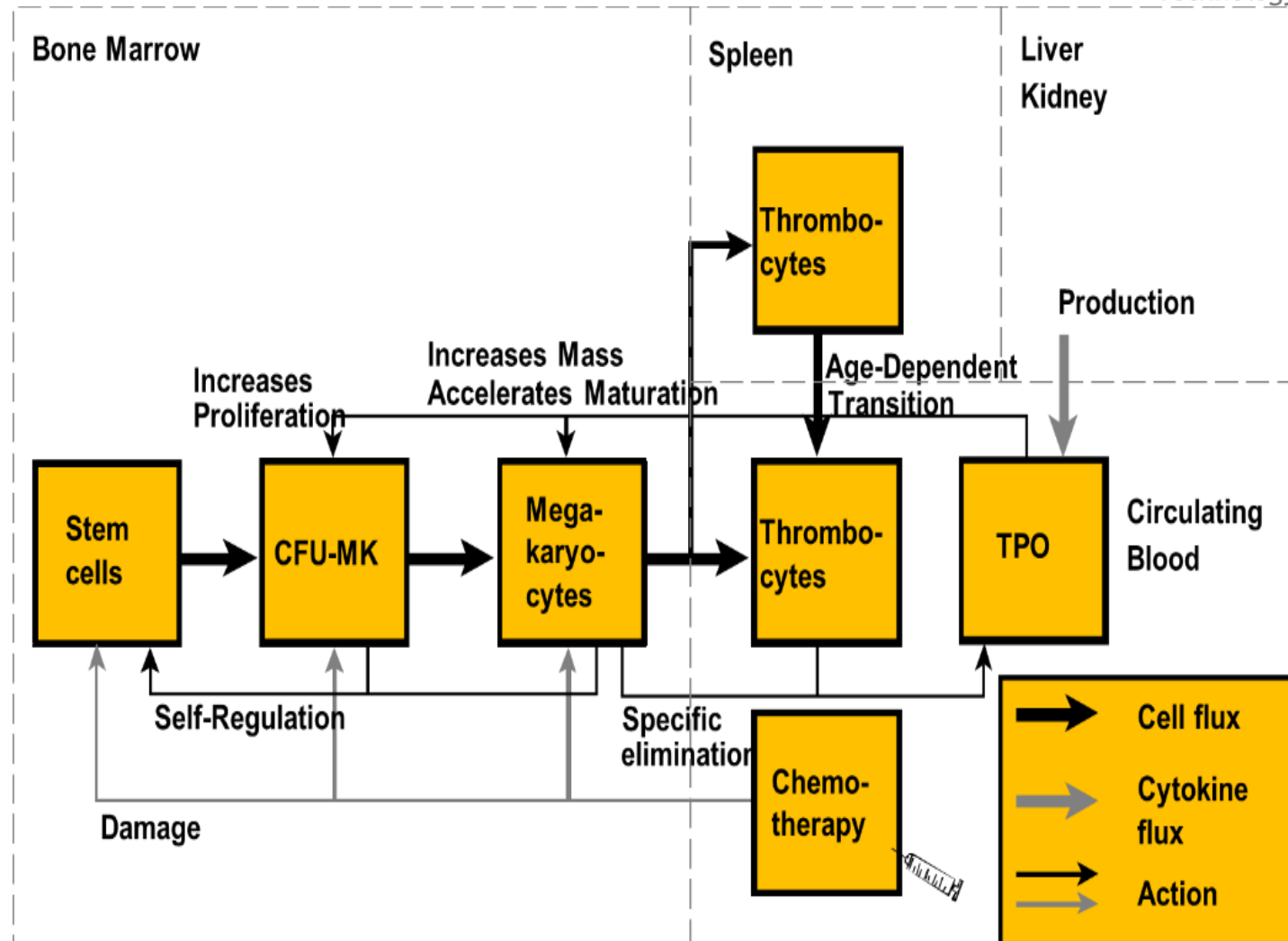
# PheP can be used to

- analyse real world patient care
- search for particular phenotypes (eg CAP, genetic)
- investigate and improve EMR data quality
- Integrate systems medicine disease models into the physicians desk top (perspective emerging)
  - eg chemotherapy dosing and timing
  - based on hematotoxicity models (e:Med projects)
  - eg biomarker based treatment decisions

# Model of thrombopoiesis

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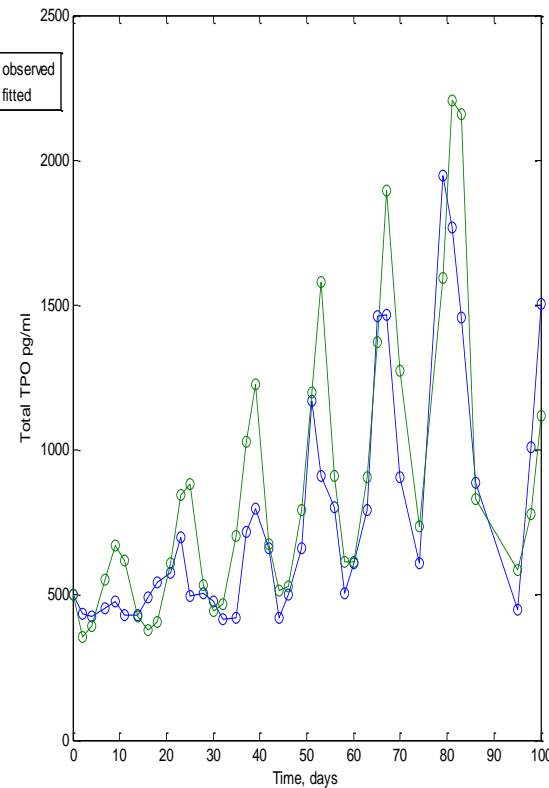
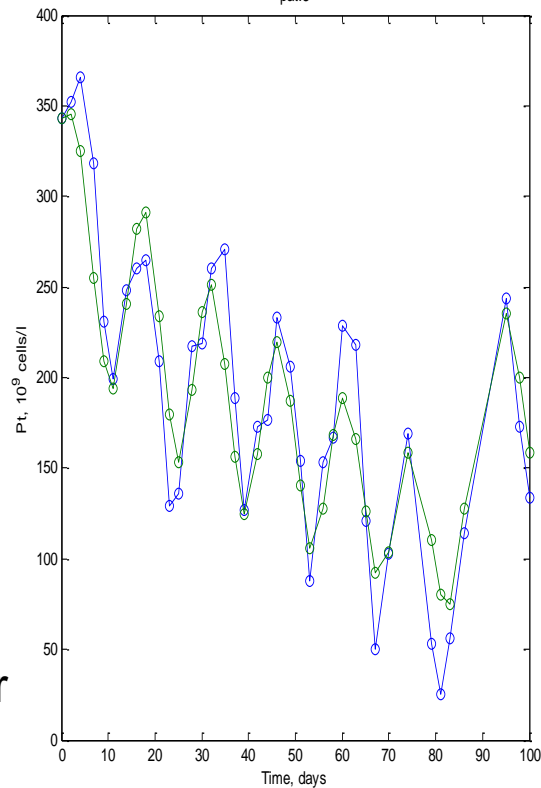




# Hematotoxicity

## BEACOPP 14

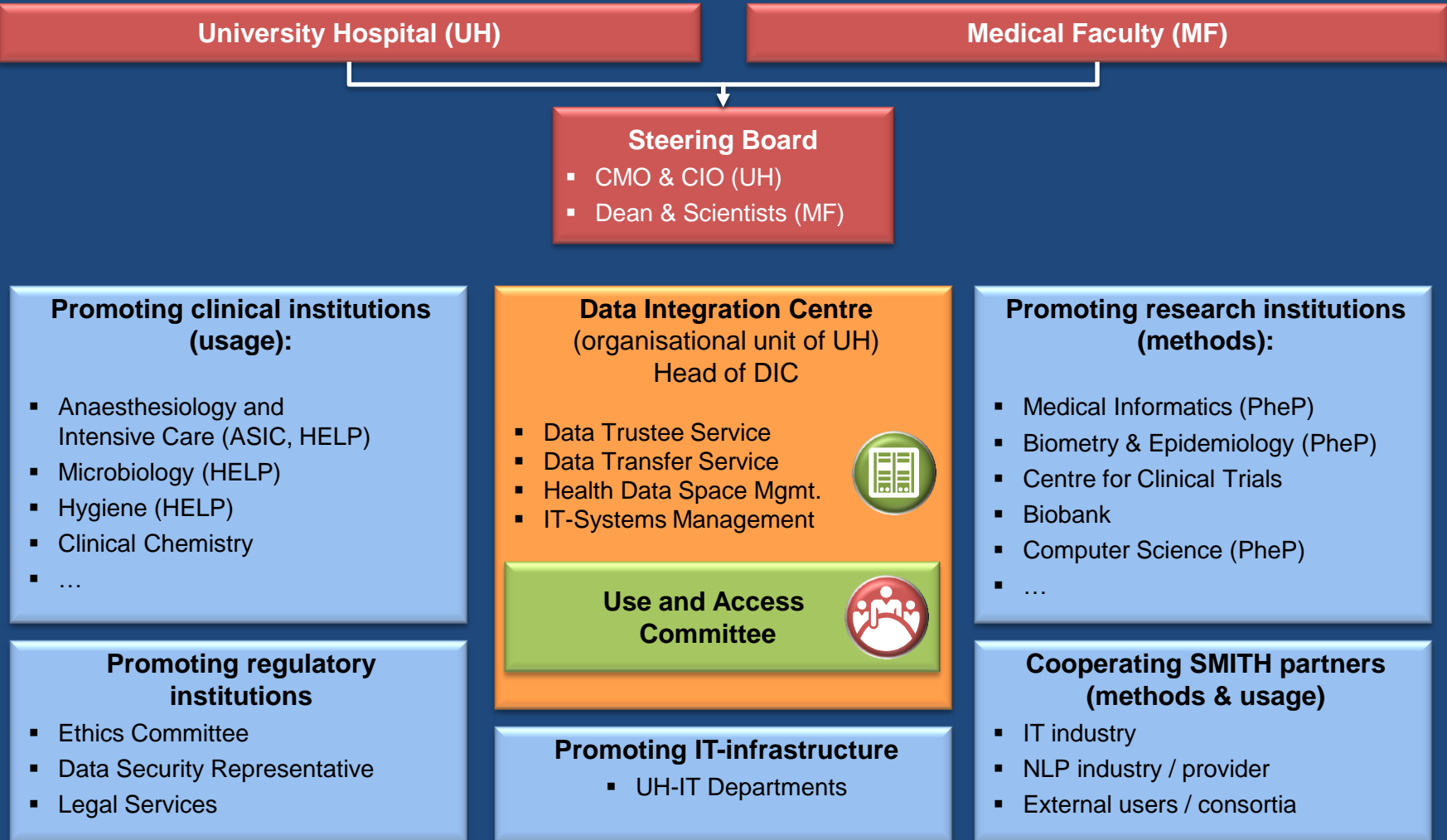
Engel et al,  
pat.3



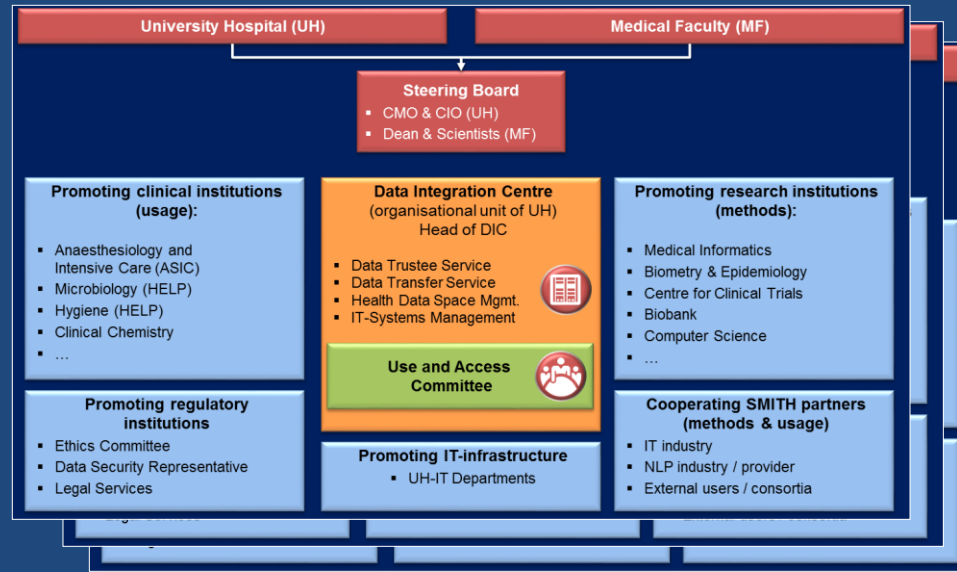
Use model for  
prediction

# Data Integration Centres

## - Generic Concept -



# Data Integration Centres



## Heads of DICs

UH Aachen



Dr. Silke Haferkamp

UH Jena



Dr. Danny Ammon

UH Leipzig



Dr. Thomas Wendt

# Technical Standards for Data Integration



- PIX / PDQ
- ATNA
- BPPC / APPC
- XDS
- XCA
- XUA



- CDA
- FHIR
- CQL



- SNOMED-CT
- LOINC
- ICQ/OPS
- IHE-D Value Sets

# SMITH DIC – Architectural Aspects

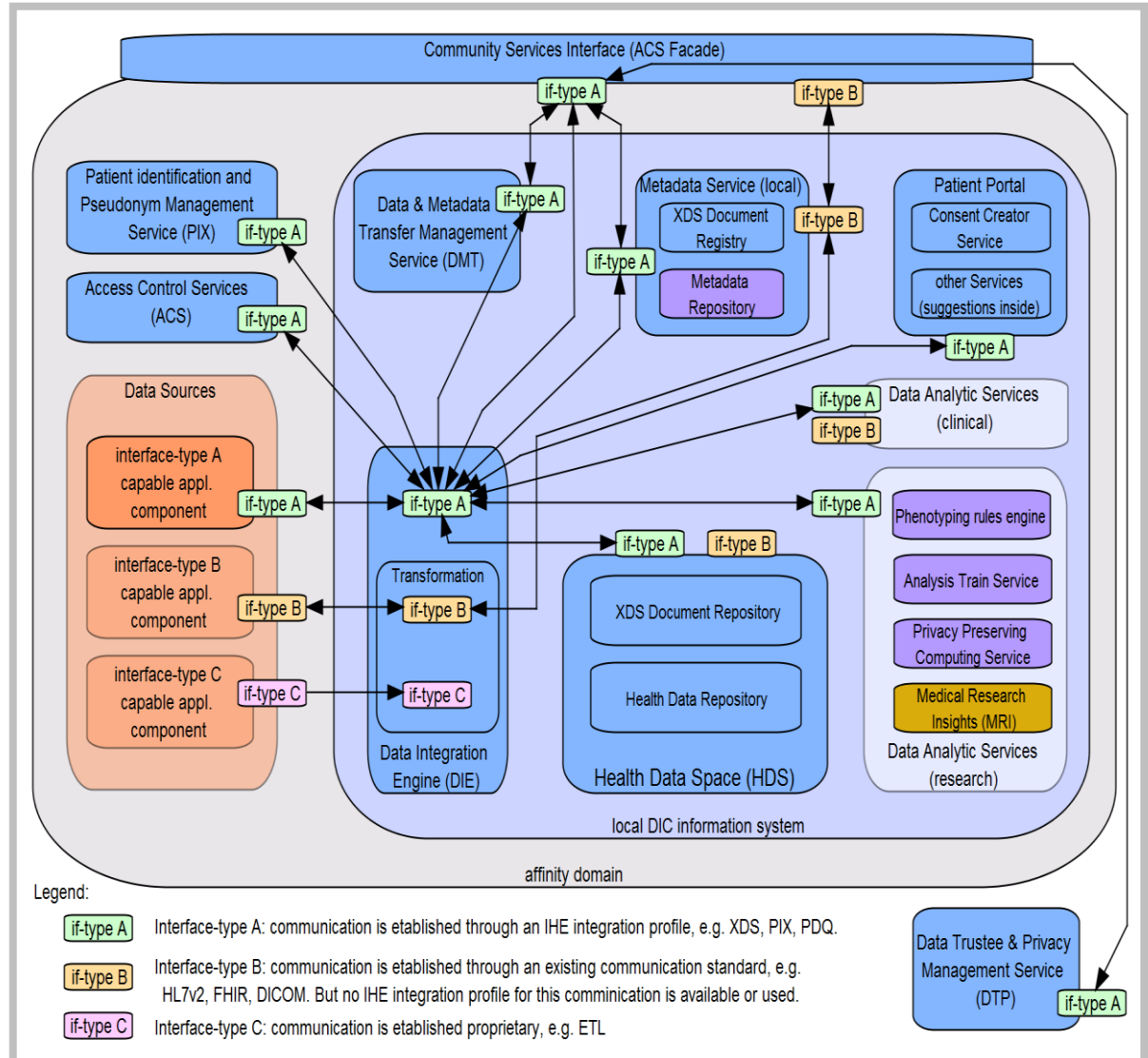
## - Communication -

### Health Data Repository

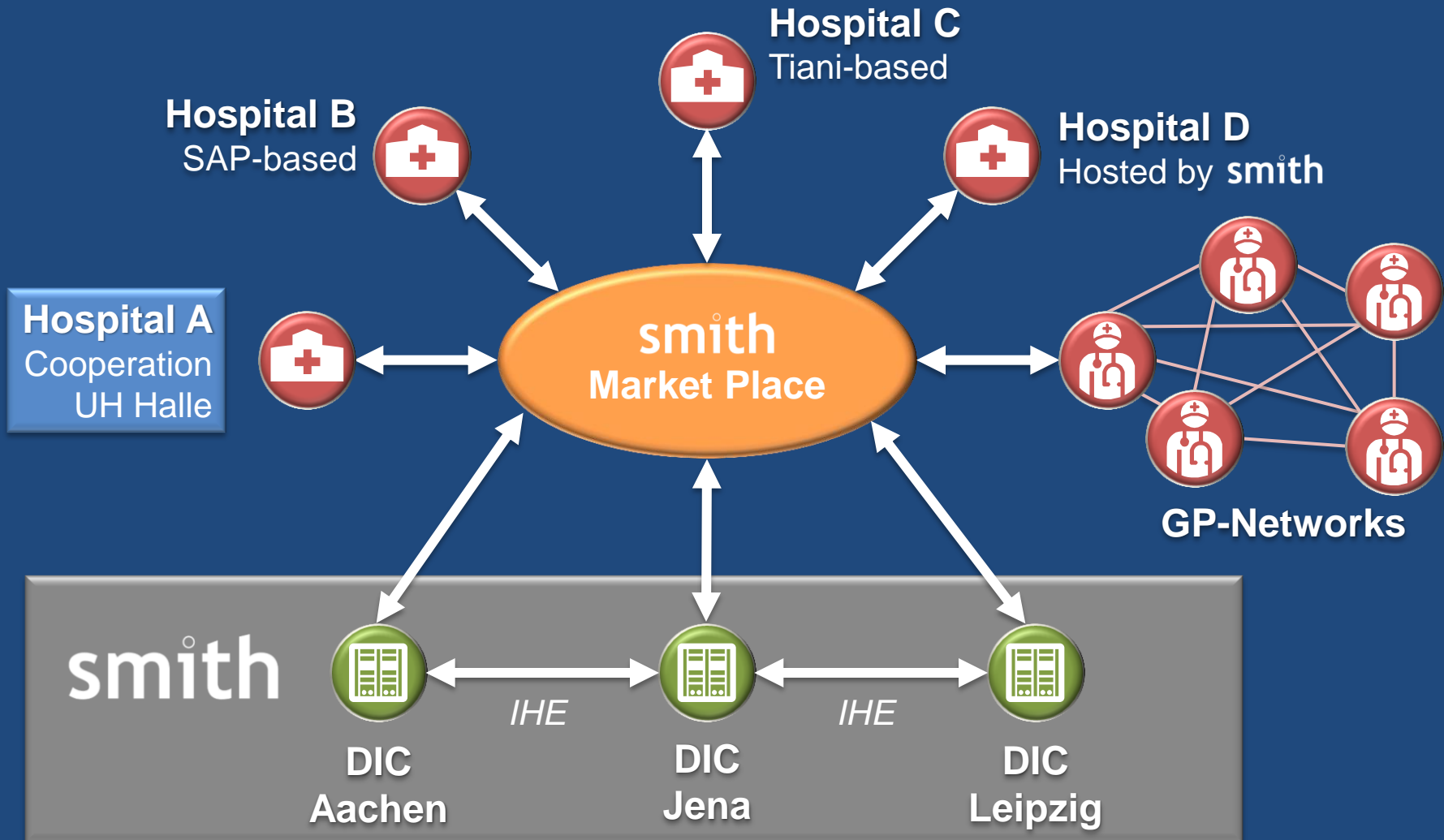
- Central persistency for structured clinical data
- FHIR interface

### Medical Research Insights

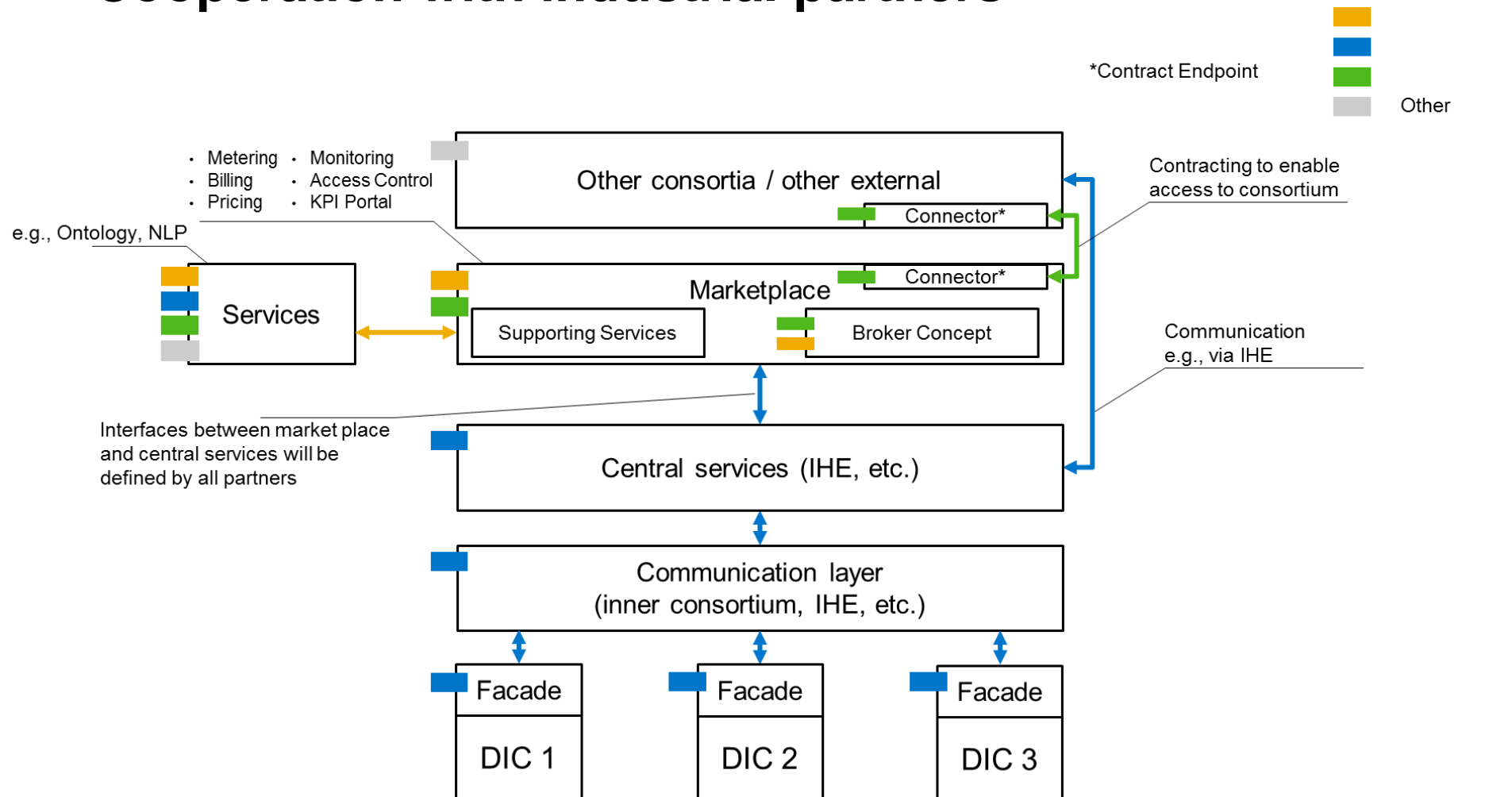
- Analyze patient data to verify or develop research hypotheses
- Identify candidates for studies



# Roll-out Concept



# SMITH Market Place – Structure & Cooperation with industrial partners





# Partnerships & Cooperations

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Smart Medical Information  
Technology for Healthcare



*DIC network and security  
technology*



*IHE-conform data  
integration (configuration  
and support)*



*implementation of  
marketplace*




*general cooperation*

Healthcare IT  Solutions

*EPR / roll-out*

**averbis**  
text analytics

*NLP procedures and  
terminology server  
(use case PheP)*

ID Information und  
Dokumentation im  
Gesundheitswesen 

*text analysis  
and metadata  
classification (DIC)*



*reference data  
(use case ASIC)*



*mutual advancement  
of communication  
standards*



*IHE-conform data  
integration (development  
and implementation)*



*high-performance  
computing*

evimed®



*Industrial Data Space  
Medical Data Space*



*general cooperation*

# DIC – Build up

## Consortial contributions of tools and skills

### Interoperability

Standard, tools and Processes  
(UKJ, UKA, SAP, März, Tiani, Cisco)

### Market Place

Internal SMITH Tools  
& Connecting  
external partners  
(UKA, SAP, HC-IT-Solution)

### Metadata Repository

Method and Items  
(UL, UA, März)

## Data Integration Center

### Use Case

Decision support  
systems  
(UKJ, UKA, UL,  
Diagnostic-Industry)

### Semantic Text Extraction Methods

NLP-Factory  
(UJ, Averbis, ID)

### Phenotype Classifiers

Rules-Factory  
(UL, USE Case teams)

# Overarching timelines

## Interoperability:

1 - 18 Initial set up of interoperability in DIC

19 - 30 Interoperability implemented in DIC

## Clinical use cases:

19 First patient in use case control cohorts

30 First patient in use case interventional cohorts

42 Recruiting closed and analysis performed

## Roll out:

19+ Starting add DIC build up

42 DICs active, 2 hospitals, 3 GP networks

# Education

## Curricula for students:

PI: Alfred Winter

- 3 new BSc/MSc programs
  - „Data Science“ (Aachen),
  - „Medical Data Sciences“ (Jena),
  - „Biomedical and Health Informatics“ (Leipzig)
- 1 international postgraduate MSC (Aachen, International Academy)

# MSC Curricula

	D1	D2	D3	D4	D5	D6	D7	D8
Ac	●	●	●	◐	×	●	●	×
J	●	●	●	×	×	●	●	●
L	●	◐	●	●	●	●	◐	◐

- Addressed by existing program
- ◐ Partly addressed by existing program
- × Needed

<b>D1</b>	Basics of medicine and principles of medical decision-making in diagnostics and therapy.
<b>D2</b>	Basics of molecular biology, bioinformatics and computational biology.
<b>D3</b>	Statistical foundations of medical research and evidence-based medicine.
<b>D4</b>	Architecture of complex information systems for medical research and care.
<b>D5</b>	Management of complex information systems for medical research and care.
<b>D6</b>	Representing and modeling medical information and knowledge (incl. ontologies).
<b>D7</b>	Managing and processing medical signal/image data.
<b>D8</b>	Accessing, managing and mining biomedical big data.

Table 9-1:

# Education

## 3 Additional Professorships

- Leipzig (W3 Medical Data Science; 2017)
- Jena (W2 2018)
- Aachen (W3 2018)

# Beitritt zum SMITH-Konsortium

Wir sind grundsätzlich offen für Beitritte von Standorten, die unsere Vorgehensweise zu DIC, Interoperabilität und Use Cases nachvollziehen wollen.

Wir bitten solche Standorte um eine schriftliche Bekundung

Wir werden dann

- Kontakt aufnehmen
- zu einer Informationsveranstaltung am 3.Nov einladen
- dann die Vorgehensweise zu einem gemeinsamen Antrag abstimmen



**Danke für die Aufmerksamkeit**