### **German Center for Child and Youth Health Ulm Site**

# TRADITION





RESEARCH

preimplantation 🕸 Participation 💣 Immune System 🗁 Obesity and Metabolism @ Mental Diseases 👶 Ethics 🐵 Epigenetics & Coping

## **Population based cohorts**

Key questions/objectives

- Risk/protective factors for healthy weight and metabolism
- Somatic, psychosocial pathology

#### Assets at UCH

- Two representative longitudinal UCH birth cohorts (including mother/child biobank)
- Statewide kindergarten and school\* cohort
- Nationwide follow-up neonatal screening on endocrine disorders
- Nationwide health insurance data (WIdO)

\* 7.000 teachers | 160.000 children and families per year





Cardio-vascular-risk factors in elementary school children (state cohort)

FAM13A affects fat distribution and adipocyte function Nat Commun. 2020; 11: 1465

#### **Proposal for DZKJ**

- National Cohort (development, biobank)
- Risk factors and digital interventions
- Dissemination and implementation research

Glucose uptake (CPM/µg protein) 30 Basal Insulin-stimulated



[3H] Glucose uptake

(D12 SGBS adipocytes)

scRNA

siFAM13A

60

45

## **Obesity and metabolism**

#### Key questions/objectives

- Biologically defined obesity subtypes
- Prenatal metabolic programming
- Need for nationwide bio-psycho-social prevention

#### Assets at UCH

- Nationwide obesity/diabetes registries
- Molecular and functional diagnostic platforms
- Structured care for pregnant women
- Statewide kindergarten/school-based prevention





**Obesity:** 125.000 patients 3.257 T2DM







#### **Proposal for DZKJ**

- (EpiGen-)OMICs-based obesity subtypes
- UCH healthy metabolism pregnancy program
- Expanding UCH prevention program nationwide
- Stratified therapies in clinical studies
- Reframe, antistigmatisation, coping



Start of Metreleptin

## Immune System and Cellular Therapy

#### Key questions/objectives

- Priming and checkpoints of immune development
- Somatic (epi)genetic variation of immunity
- Modulation of immunity by metabolism
- Detection of disturbed immune function and transition to malignancy

#### Assets at UCH

- Diagnostic platforms, cohorts, registries
- >1.000 pediatric stem cell transplantations (>30% PID)
- Extensive human and animal models
- Multi-OMICs immune phenotyping





SARS-CoV2 anti-Spike IgG in preparation

Methylome B-cell differentiation Within BLUEPRINT, Nat Genet 2015; 47: 746

#### **Proposal for DZKJ**

- Understanding normal immune development
- Diagnostic criteria of immune dysfunction
- Improving cellular and gene therapy
- Coping with long-term immunedysfunction
- Screening for hemoglobinopathies



0

M6 (528

M7 (4,838) M8 (34.604

M9 (13,044)

M16 (1 319

M17 (1.604

M19 (6,581

0000



## Mental Diseases as Systemic Immuno-Metabolic Disorders





#### Key questions/objectives

- Mental diseases as disturbed immunometabolism
- Prenatal priming of mental disease
- (Epi)genomic burden and mental disease
- Resilience as immunometabolic strength

#### Assets at UCH

- Proof-of-concept evidence
- Comprehensive interdiciplinary methodology
- Preclinical model systems
- Patient cohorts and intervention studies





#### **Proposal for DZKJ**

- Characterization of mental-immunometabolic interface
- Multi-OMICs/-systems development and risk assessment
- Biological systems profiling as intervention measure
- Pre-clinical human readouts of immunometabolism

## **Digital Behavioral Health and Coping**

# ulm child health



#### Key questions/objectives

- Health and coping insights by smart sensing
- Al-enhanced innovations
- Nationally scalable implementation models

#### Assets at UCH

- Interoperabel IT solutions and AI-ready databases
- Nationally largest body of digital behavioral youth health solutions
- Six interdisciplinary UU IT and trauma centres of competence
- Nationwide digital behavioral health network



eHealth Program Ulm University



#### Time to Onset of Major Depressive Disorder in Back Pain JAMA Psychiatry 2020; 77: 1001



#### **Proposal for DZKJ**

- Digital Behavioral Health Hub as incubator within DZKJ
- Rapid innovation cycle from prototypes to implemented solutions
- Specific focus on diminishing health care gaps and disparities: Leave no one behind

## **UCH Infrastructures**

#### **Core Facilities**

- Pediatric Clinical Trials Center as part of ZKS
- Animal Research and Transgenic Mice
- Translational Imaging (electron, confocal and multiphoton, small animal, whole-body incl. PET/MRI, automatic high content)
- Omics (NGS, Affymetrix, Proteomics, Functional Peptidomics) and Bioinformatics
- Cytometry, including Mass Cytometry
- Metabolism (SeaHorse, Power-O2k, ESR)
- GMP-grade clean rooms
- Automatic Metabolic and Behavioral Phenotyping (mice)
- eSano Digital Health Intervention Platform

### **Cooperation Partners on Campus, e.g.**

- Institute for Diabetes Technology (IfDT)
- Boehringer Ingelheim Ulm University BioCenter (BIU)
- Institutes of Quantum Optics and Complex Quantum Systems

#### Nation-wide patient registries





**Department of Pediatrics and Adolescent Medicine** 

#### **Proposal for DZKJ**

- Sharing access to crucial technologies across sites
- Establishing cores within DZKJ when appropriate
- Improvement and development of new technologies,
  e.g. in the area of diabetes and quantum imaging
- Combining existing and establishing new cohorts

#### Cohorts





## Our vision for DZKJ:

Improving health and well-being of every child and adolescent from (pre-)birth to adult age.