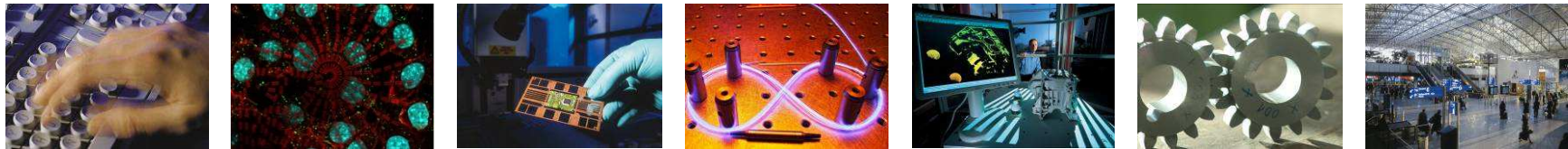

Fraunhofer Group for Life Sciences

Applied Science Dedicated to Life



Profile of Fraunhofer-Gesellschaft



- 72 institutes
- 25 000 employees
- Budget of 2,3 billion euros

Seven Fraunhofer Groups

- Information and Communication Technology ICT
- Life Sciences
- Microelectronics
- Light & Surfaces
- Production
- Materials and Components – MATERIALS
- Defense and Security VVS

Fraunhofer Group for Life Sciences

■ Facts and figures

Established in the year 2000

6 Member institutes, 1 research institution

Chairman since April 1, 2017: Prof. Horst-Christian Langowski

Co-Chairman: Prof. Norbert Krug

Office: Hannover

Head of Central Office: Dr. Claus-Dieter Kroggel

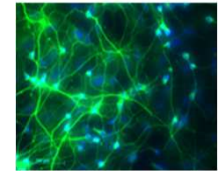
■ Business units under development

Diagnostics and therapeutics – from target screening to clinical studies

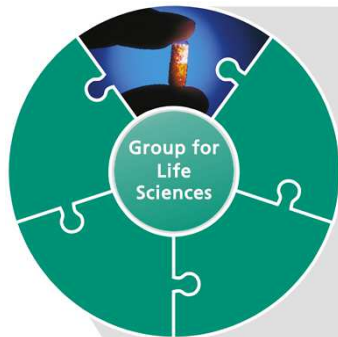
Medical technology – with biology and engineering to the approved product

Food – healthy and safe on the plate

Bio-based economy – sustainable value-adding from resources



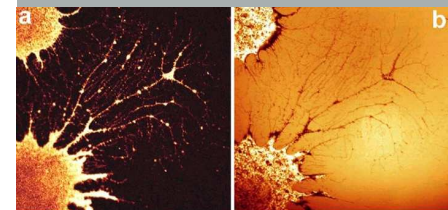
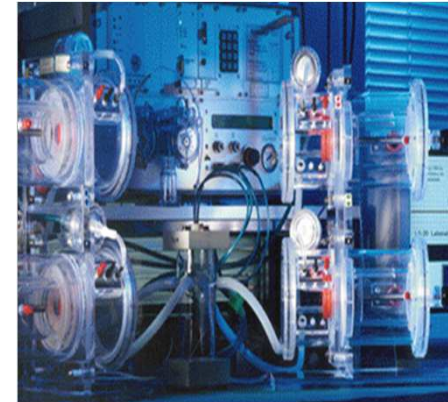
Medical Translational Research and Biomedical Technology: The Challenge of Innovative Diagnostics and Personalized Therapy



Our responses

- innovative biomarkers for prognosis and therapy control, e.g. RNA biomarker
- isolation and analysis of CTCs and DTCs for individualized tumor diagnostics
- insects and their microbiota as new source for therapeutics, e.g. highly effective antibiotic substances
- GMP platform with a wide range of expression systems, i.a. plant cell culture and plants (vertical farm)
- cell-free protein synthesis

Successful treatment through innovative diagnostics and optimally tailored, personalized therapy



Translational Medicine: From Molecule to Patient

Bio-Hybrid and Cell-Based Tests – Early Go/No-Go Decisions



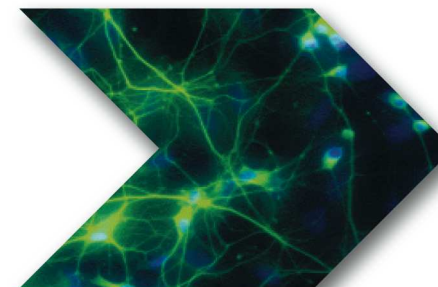
Target and drug discovery

- Databases: gene expression profiles
- High-throughput screening
- Array technologies
- Biochips: DNA chips, protein chips
- Cell-based assays and chips



Target validation

- Animal transgenic and in-vitro 3-D human disease models
- Large-animal models
- Behavioral animal models
- Cardiovascular and neuronal organotypic models and disease models

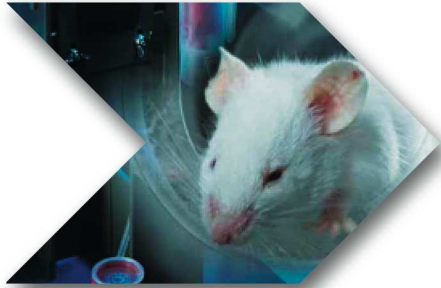


Lead optimization

- Structure optimization and humanization
- Aerosol generation
- High-content screening
- Human cell models
- Organotypic cell culture models

GXP Platform: GLP – GMP – GCP

APIs/Vaccines/Cell Therapeutics – System-independent Solutions – Clinical Studies Airways



Safety pharmacology and toxicology

- GLP
- In-silico prediction ADME
- Molecular and functional imaging
- Nano- and microparticle systems
- Biomarkers of disease



Manufacture of clinical investigational products

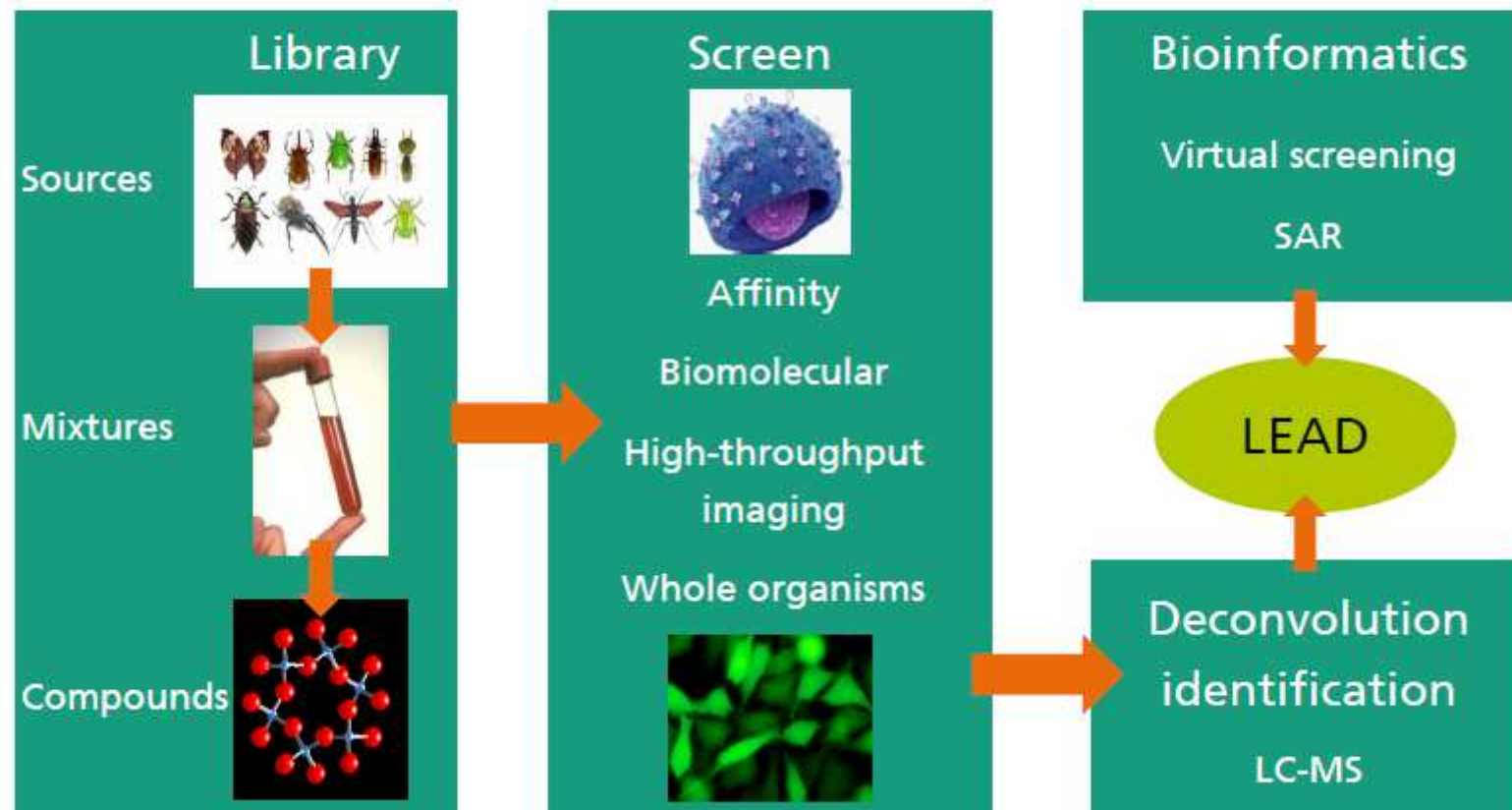
- Consulting on regulatory and economic aspects
- Evaluation of customer processes
- GMP facilities
- Compound libraries and high-throughput systems
- Expression systems



Clinical trials of phases I and II

- GCP-compliant protocol development
- Toxicology and pharmacology
- Dose escalation studies in patients with malignant diseases
- Segmental lung challenge

The Fraunhofer IME Knowledge-based Screening Portal



Insects as Resources for New Therapeutics



Wound-healing
promoting
molecules



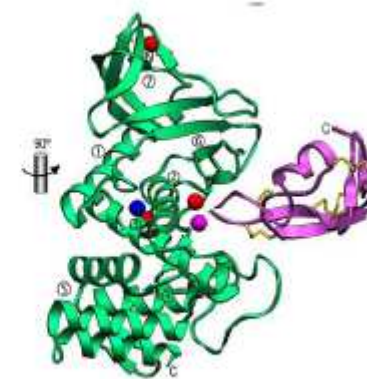
Metallo-proteinase-
inhibitor



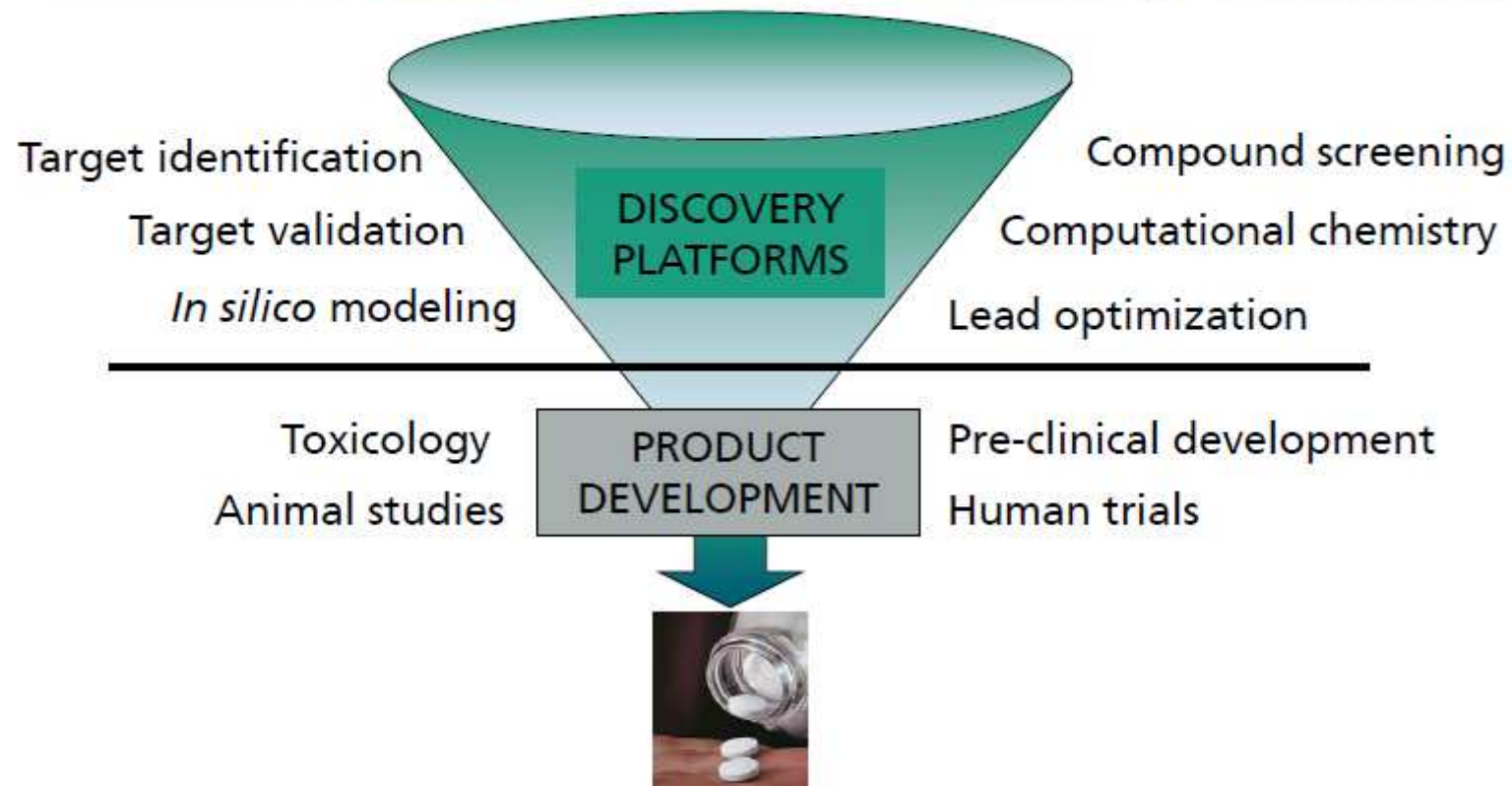
Biofilm-degrading
enzymes



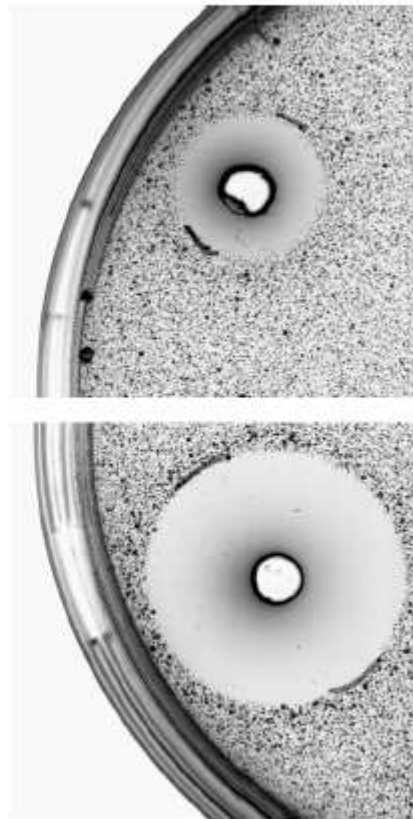
Antimicrobial
compounds



IME Insect Biotechnology Tool Box



Biofilm* Degrading Enzymes

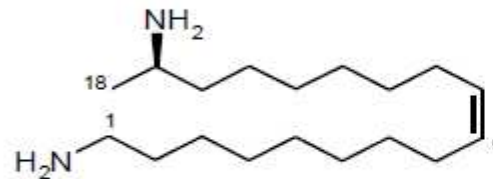


- Detection of high antimicrobial activity in hemolymph of untreated *Eristalis* larvae (= standing army costs)
- Antimicrobial activity increases remarkably upon injection of bacterial lipopolysaccharides (= war costs)

*IUPAC definition: Aggregate of microorganisms in which cells that are frequently embedded within a self-produced matrix of extracellular polymeric substance (EPS) adhere to each other and/or to a surface.

Drugs from Bugs

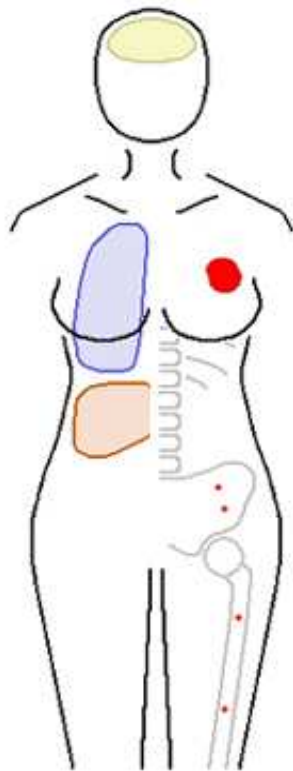
- Harlequin ladybird (Asian lady beetle) shows enduring resistance against diverse pathogens, which allows it to outperform and therefore dominate the most abundant native European ladybirds.
- Resistance due to chemical defense compounds in hemolymph
- Identification of **harmonine** with promising activity against *Mycobacterium tuberculosis* (tuberculosis) and *Plasmodium falciparum* (malaria)



Vilcinskas, A., Stoecker, K., Schmidtberg, H., Röhrich, C., Vogel, H. (2013). *Invasive harlequin ladybird carries biological weapons against native competitors*. SCIENCE.

Personalized Approaches in Translational Medicine

Isolation and Discovery of Circulating and Disseminating Tumor Cells



Circulating (CTC) and disseminating (DTC) tumor cells are isolated from:

- Bone marrow
- Blood
- Lymph nodes
- Cerebrospinal fluid



Detection of individual DTCs by e.g. cytokeratin staining

Clinical Research Center Hannover (CRC Hannover)

State-of-the-Art Research Facility for Early-Phase and Proof-of-Concept Studies



- The CRC Hannover is a nationally unique medical research center.
- It is on its way to becoming the leading center for patient-oriented and translational medicine in Europe.
- CRC Hannover's state-of-the-art infrastructure is jointly used by three leading research institutes: the Fraunhofer ITEM, the Hannover Medical School (MHH), and the Center for Infection Research (HZI).

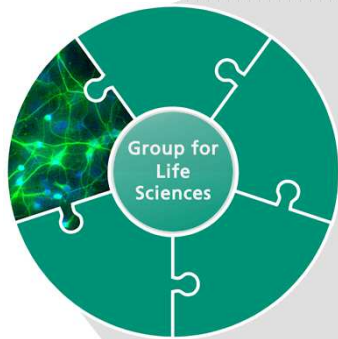
Clinical Research Center Hannover (CRC Hannover)

Numbers and Data

- 3 Partners: Fraunhofer ITEM, MHH, and HZI
- In the direct vicinity of Fraunhofer ITEM and MHH
- 6,000 square meter space
- 30 Intermediate care beds (for clinical trials phase I and IIa)
- 20 Beds for study participants with low monitoring needs
- 15 Rooms for specialized diagnostics
- Modern imaging methods (MRI)
- Biomarker laboratory
- Biobank



Regenerative Medicine: The Challenge of Qualified Biobanking and Controlled Self-Healing



Our responses

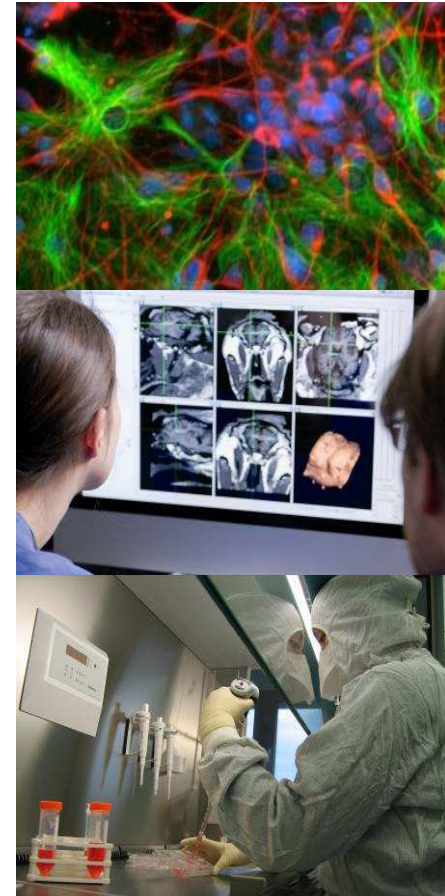
- Fraunhofer Bioarchive as an unrivaled source of live cell samples
- iPSCs
innovative approaches to adult stem cell differentiation
advanced technologies for genetic characterization and quality control procedures
- Development of patient-specific cell therapeutics
- small- and large-animal models as well as cell culture techniques, e.g. for stroke research
- 3D in vitro and ex vivo test systems for long-term and multiple exposition

Healthy aging



Cell-therapeutic Approaches – All Development Steps for Cell-therapeutic Agents from One Source

- **Development** and testing of cell-based therapeutic approaches
- **Focus:** Ischemic diseases (stroke, heart attack), neurodegenerative diseases (Alzheimer's), joint diseases, oncology
- Feasibility studies – study design and proof of concept in in-vitro and in-vivo (small-animal) models
- **Pre-clinical development** – safety, efficacy and, efficiency testing in different in-vivo models (small and large animals) with unrivaled predictivity for use in man
- GLP-compliant safety and efficacy testing
- **Clinical trials** – GMP-compliant manufacture of investigational medicinal products for clinical trials



Healthy Foods: The Challenge of High Consumer Acceptance and Disease Prevention



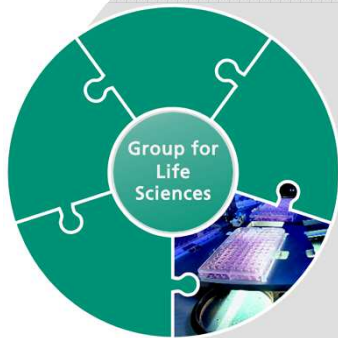
Our responses

- innovative screening and rapid tests to analyze single components and substance patterns
- gentle disinfection and sterilization methods
- safe and high-grade foods as a result of intelligent packaging
- Aroma and texture research for maximum eating pleasure
- tasty recipes for calorie-reduced and functional foods
- new ingredients for bioactive foods
- In-vitro and in-vivo proof of efficacy of bioactive food components

Health through a specific diet



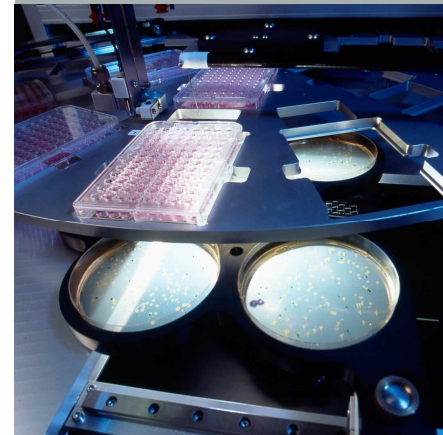
The New Potential of Biotechnology: The Challenge to Learn from Nature for Industrial Exploitation



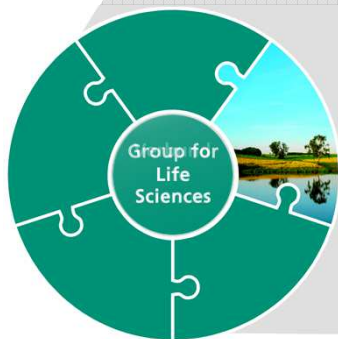
Our responses

- innovative methods to use renewable raw materials and also waste
- Supply of alternative, inexpensive fermentation raw materials, for example from lignocellulose
- Isolation of new and optimized biocatalyzers by metagenomics
- Exploitation of the wide range of insect enzymes for industrial use
- Transfer of process steps to plant production systems
Fraunhofer Center for Chemical-Biotechnological Processes, Leuna

New raw materials for the future



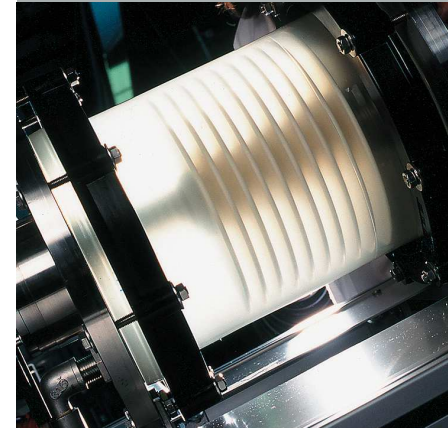
Process, Chemical, and Herbicide Safety: The Challenge of Environmental and Consumer Protection



Our responses

- substance-specific strategies for chemical evaluation
- Inclusion of data pools and in-silico methods
- broad range of alternative in-vitro and in-vivo test methods
- comprehensive ecological evaluation of herbicides including complex environmental simulation
- semi-decentralized supply and disposal systems for safe and effective water management
- Broadband sensor for continuous monitoring of drinking water

New technologies create safety for man and the environment



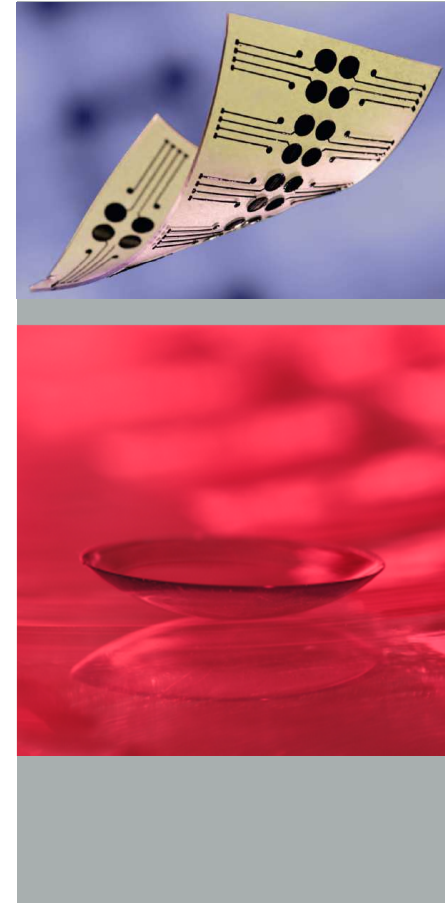
New BU within the Group – Result of Strategic Decisions

Medical Technology – From Idea to Approved Products

New
BU

Our responses

- Neuroprosthetics, e.g. elastic nano-functionalized polysiloxane structures
- wireless energy and data transmission for highly sensitive actuation of prostheses
- combining therapeutic and diagnostic functions in a single medical device
- Biotelemetry
- biomedical ultrasound
- Plasma technologies for cleaning and disinfection as well as functionalization of surfaces
- broad range of 3D scaffold creation technologies



6 Institutes, 1 Research Institution, 21 Locations

● Main locations:

Hannover: Fraunhofer Institute for Toxicology and Experimental Medicine ITEM

Aachen: Fraunhofer Institute for Molecular Biology and Applied Ecology IME

St. Ingbert: Fraunhofer Institute for Biomedical Engineering IBMT

Stuttgart: Fraunhofer Institute for Interfacial Engineering and Biotechnology IGB

Freising: Fraunhofer Institute for Process Engineering and Packaging IVV

Leipzig: Fraunhofer Institute for Cell Therapy and Immunology IZI

Lübeck: Fraunhofer Research Institution for Marine Biotechnology EMB

○ Branches:

Schmallenberg: IME; **Hamburg:** IME; **Sulzbach:** IBMT; **Braunschweig:** ITEM; **Dresden:** IVV; **Golm:** IZI

◎ New project groups:

Leuna: IGB, Fraunhofer Center for Chemical-Biotechnological Processes CBP

Straubing: IGB, Project Group BioCat "Catalytic Processes for a Sustainable Supply of Raw Materials and Energy on the Basis of Renewable Resources"

Giessen: IME, Project Group on Bio-Resources

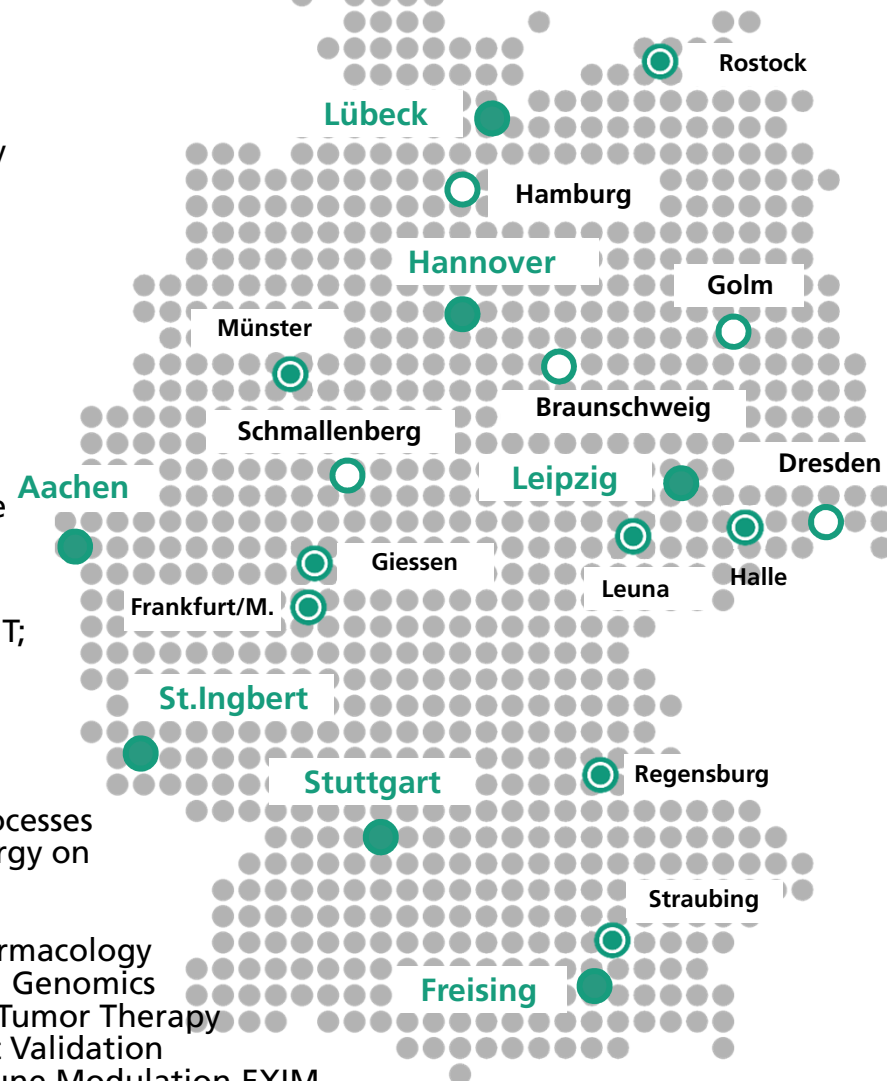
Frankfurt/Main: IME, Translational Medicine + Pharmacology

Münster: IME, Branch Lab for Functional + Applied Genomics

Regensburg: ITEM, Project Group on Personalized Tumor Therapy

Halle: IZI Project Group on Drug Design and Target Validation

Rostock: IZI, Project Group on Extracorporeal Immune Modulation EXIM



Do not hesitate to contact us

We are pleased to help you find answers to any questions you might have or solutions you are looking for.

Please contact our Central Office:

Dr. Claus-Dieter Kroggel
Head of Central Office

Fraunhofer Group for Life Sciences
Nikolai-Fuchs-Str. 1
30625 Hannover
Germany

Phone +49 511 5350 - 449
claus.kroggel@vls.fraunhofer.de



Dr. Claus-Dieter Kroggel
Head of Central Office