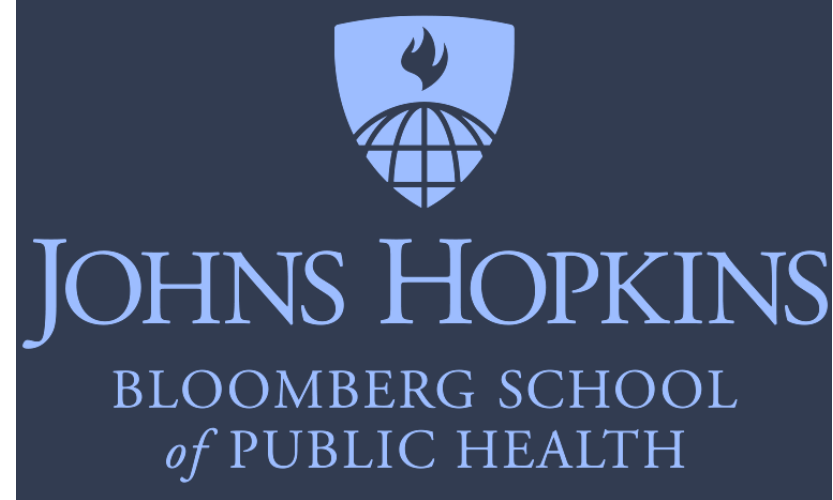
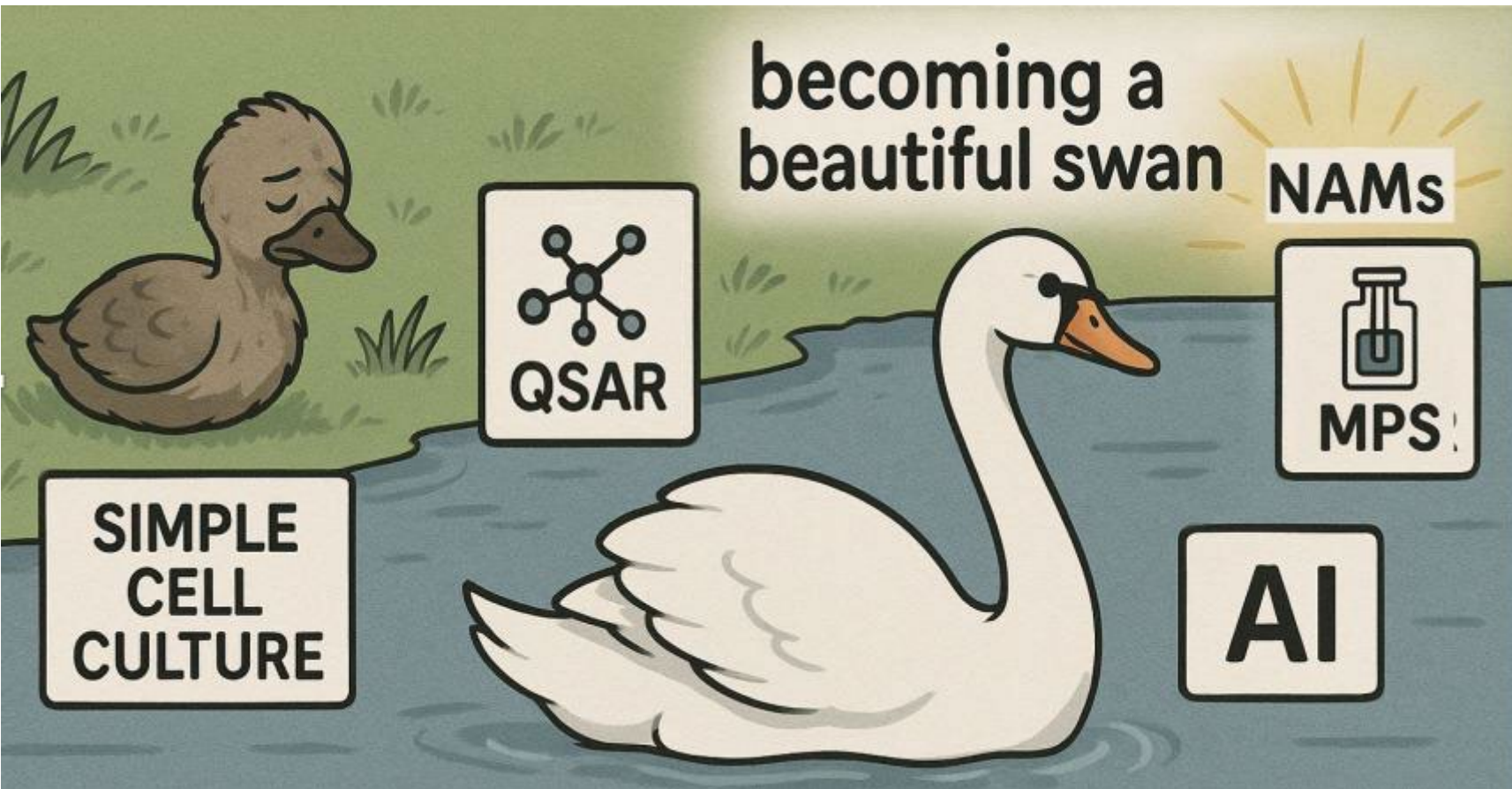


# Disruptive Technologies: MPS & AI

Thomas Hartung & team



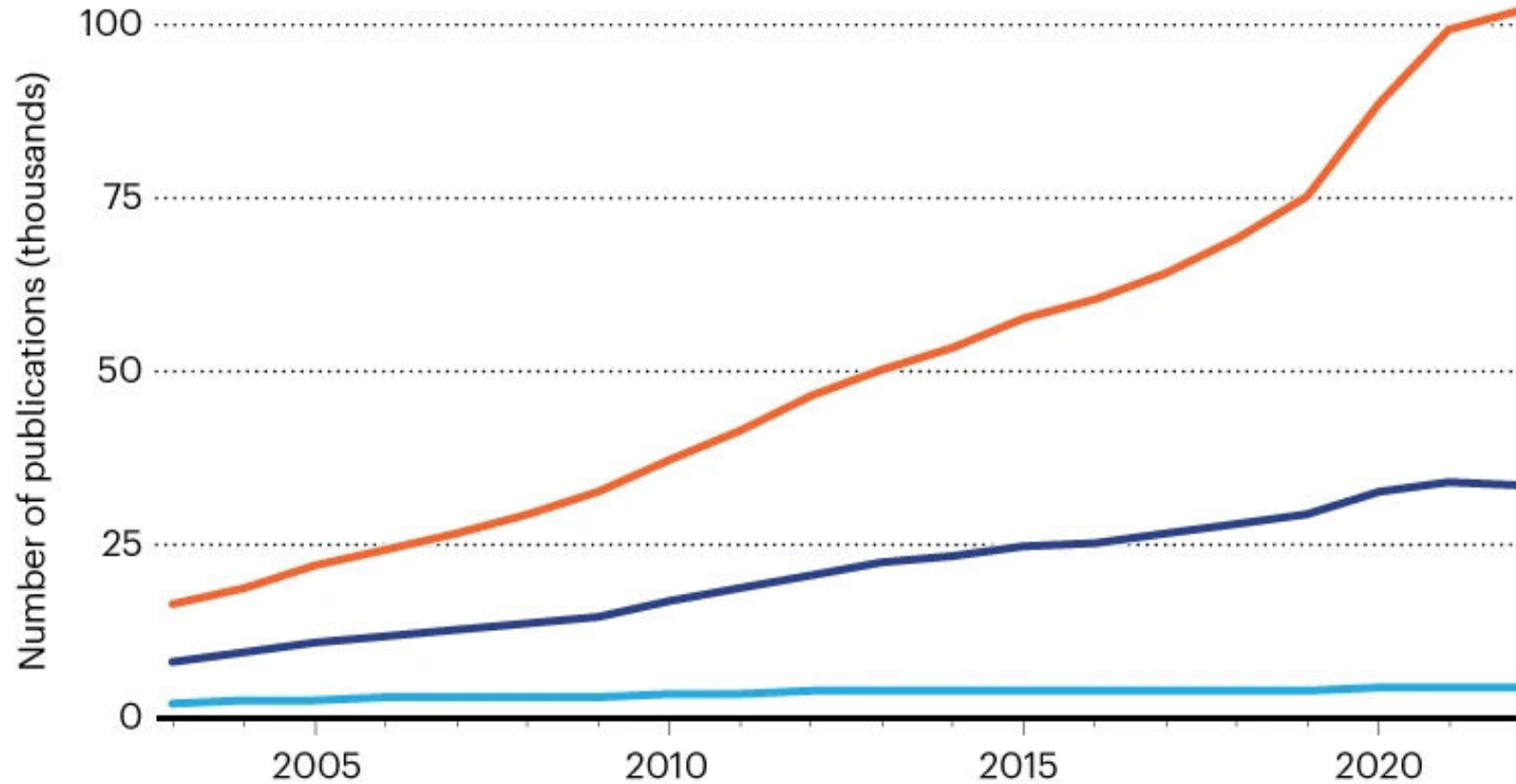
Slides available:  
<https://share.zight.com/KouXyZpl>



## ALTERNATIVES ON THE UP

In research on seven diseases, the number of publications reporting the use of new approach methodologies (NAMs), which do not use animals, has grown since 2003.

— NAMs alone — NAMs and animals — Animals alone



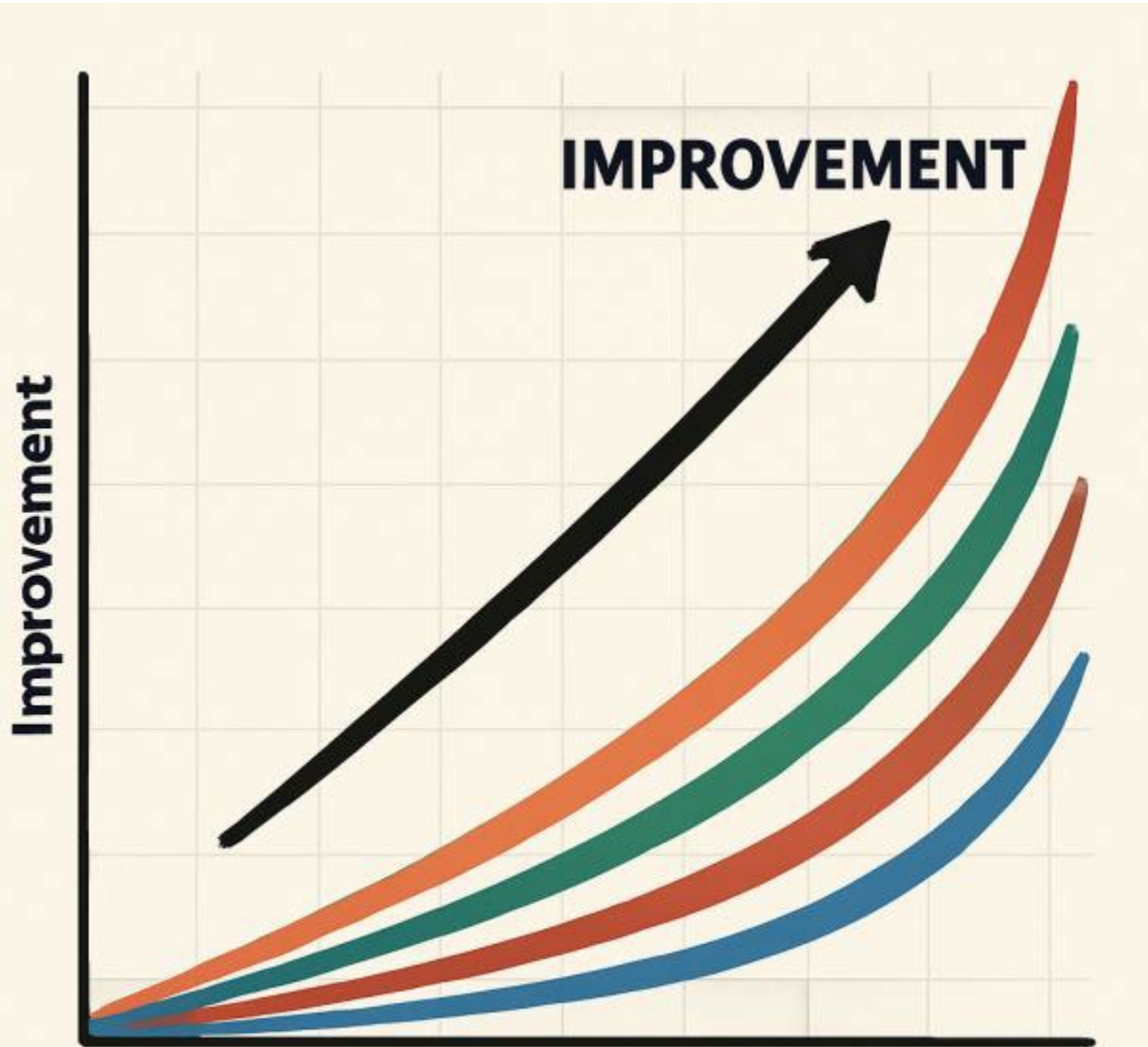
Governments have announced plans to reduce the number of animals, such as mice, used in experimental procedures.

## CAN WE REALLY GET RID OF ANIMAL TESTING IN RESEARCH?

Advances in organ and computer models are raising the prospect that some animal experiments could be eliminated. But there are still huge hurdles to overcome. **By Diana Kwon**

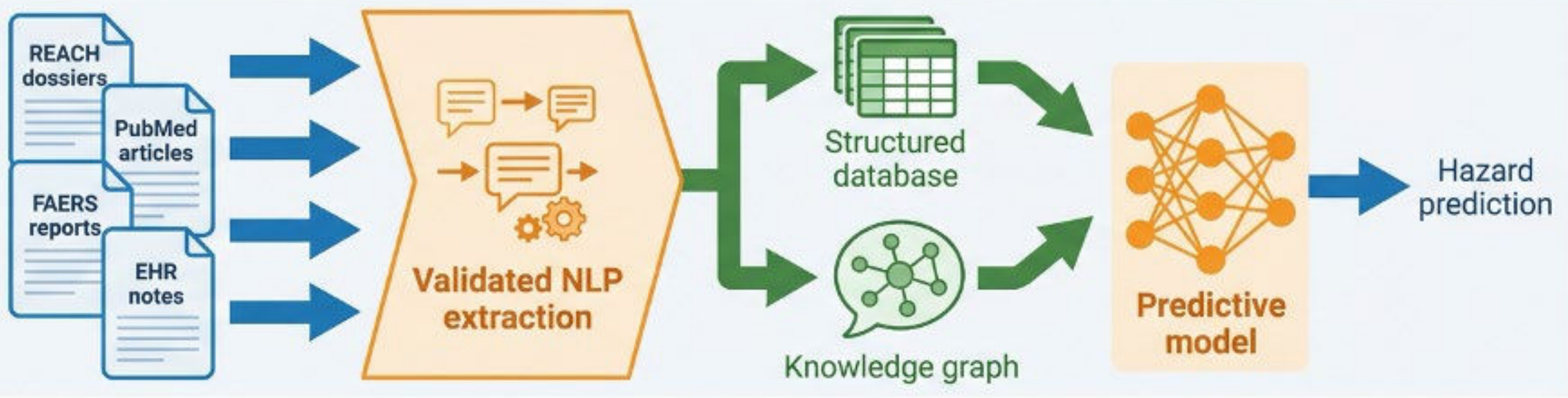
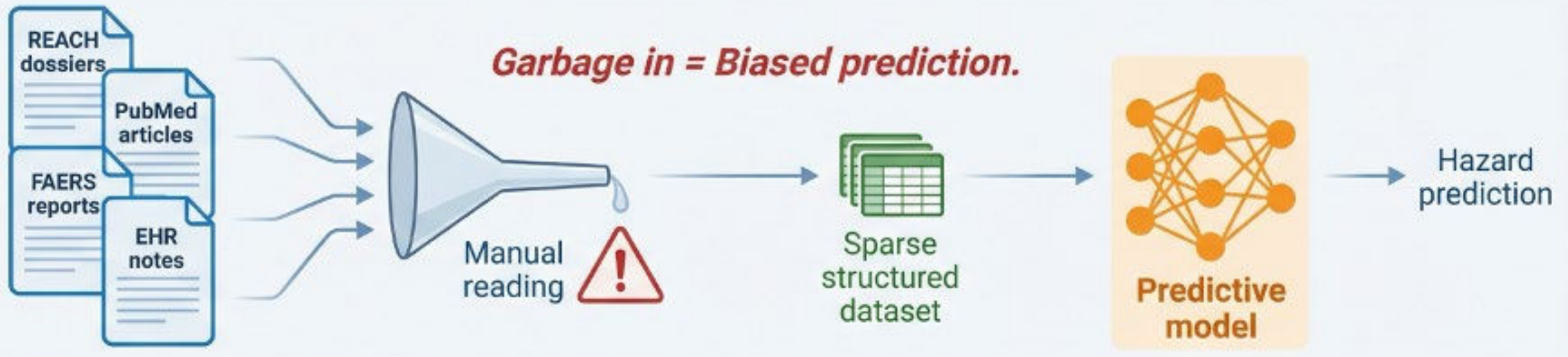
# NAM use rising

# Transformative (Disruptive) Technologies



- **Mass-spectroscopy** sensitivity doubling every 3-4 years
- **Microphysiological Systems** publications doubling every 4 years
- **Genome Sequencing** halving every 1.4 - 2 years
- **Computer** capacity doubling every two years (Moore's law)
- **Data** in the world are doubling every 18 months
- **AI** is doubling in capacity every 3 months since Deep Learning was introduced in 2010
- **AI price** falling 280x in last 18 months

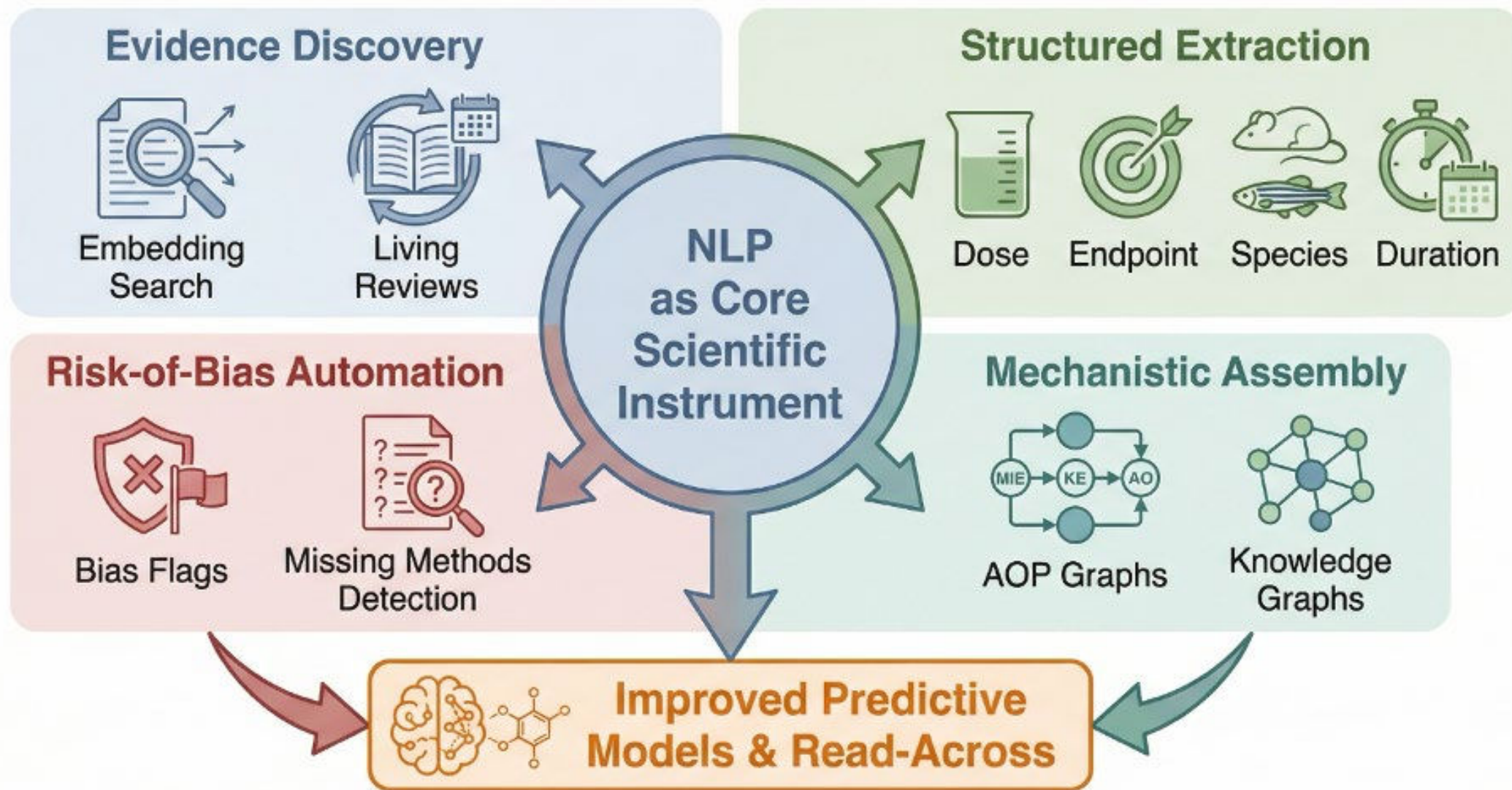
# The Reading Bottleneck in Predictive Toxicology



*Natural Language Processing in Predictive Toxicology: From Convenience Tool to Core Scientific Instrument*

Weida Tong and Thomas Hartung, under review

# Core NLP Applications in Predictive Toxicology



*Natural Language Processing in Predictive Toxicology: From Convenience Tool to Core Scientific Instrument*

Weida Tong and Thomas Hartung, under review

## Big Data

- High-content (~omics & imaging)
- High-throughput (Robotized testing, e.g., Tox21 & ToxCast)
- Sensors
- Literature, Internet
- Legacy studies

## ToxAicology



## Big Computer

### AI & Machine Learning

- Natural Language Processing (Large Language Models)

## Big Sense

- Data retrieval
- Evidence integration (systematic reviews, risk assessments)
- Predictive toxicology
- Digital pathology
- Reporting

6,000 properties  
70 bill. data from  
2,000 databases

Any chemical  
t.b.d. % correct?

2026

1 toxicity  
1,000 data  
150 chemicals  
65% correct

2015

9 toxicities  
600,000 data  
190,000 chemicals  
87% correct

2018

2025

4,000 properties  
260 mill. data from  
12 databases  
26 mill. chemicals  
91% correct

# The Next

- Building trust
- Deploy ToxAcology



npj | digital medicine

Perspective

Published in partnership with Seoul National University Bundang Hospital



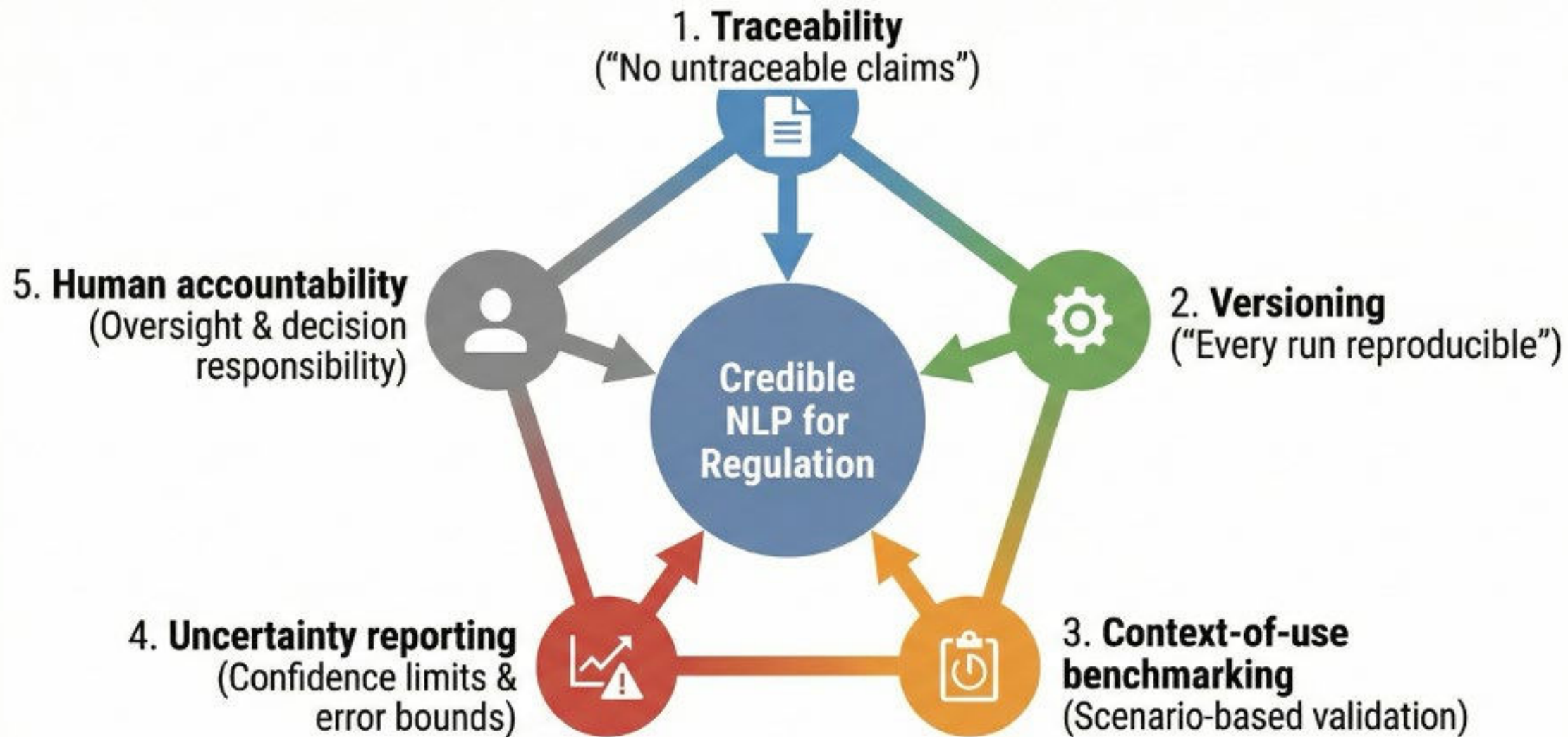
<https://doi.org/10.1038/s41746-025-01596-0>

## Is regulatory science ready for artificial intelligence?



Check for updates

Thomas Hartung<sup>1,2</sup>, Maurice Whelan<sup>3</sup>, Weida Tong<sup>4</sup>✉ & Robert M. Califf<sup>6</sup>



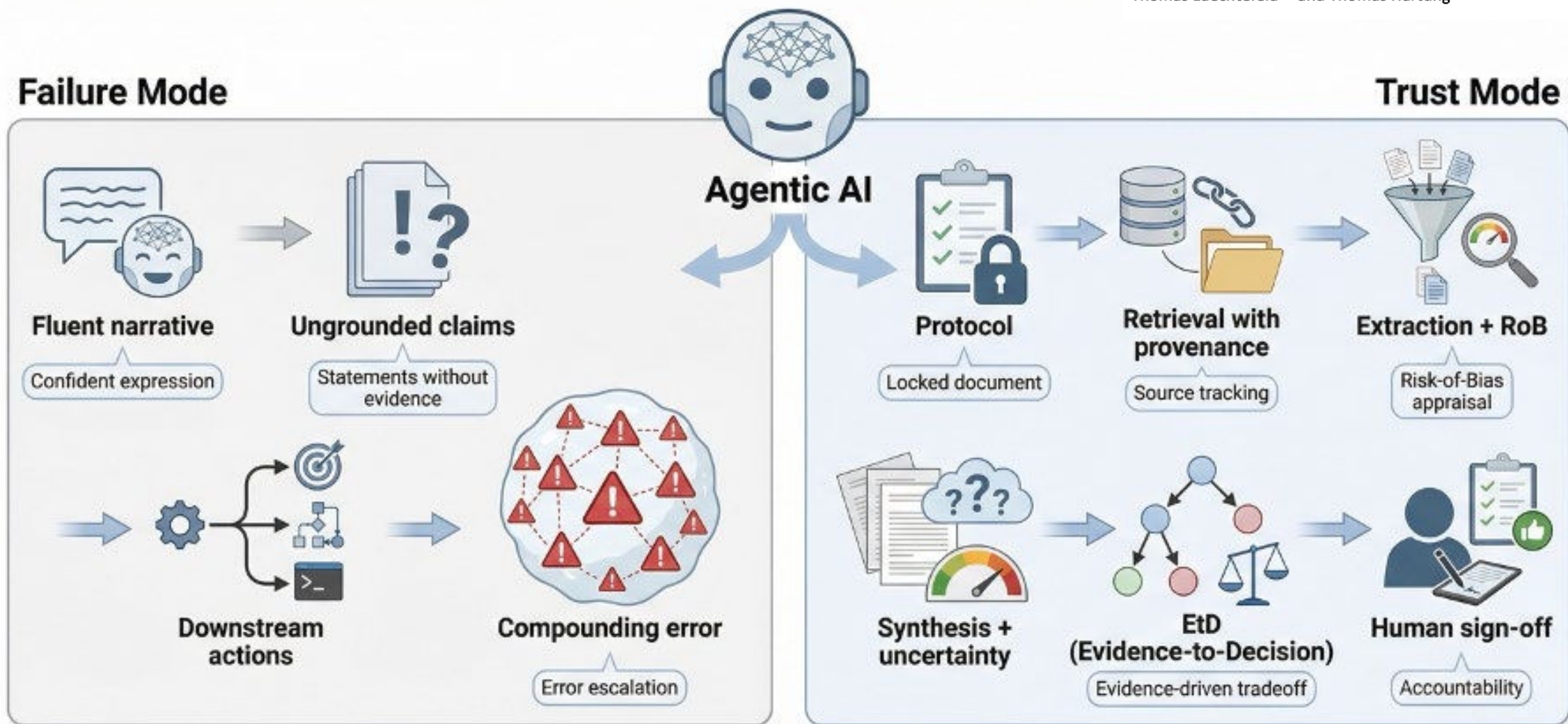
*Natural Language Processing in Predictive Toxicology: From Convenience Tool to Core Scientific Instrument*

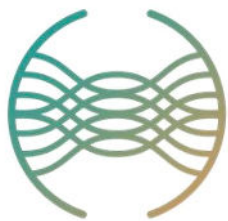
Weida Tong and Thomas Hartung, under review

# From trailblazer to “*trustblazer*”: two pathways for agentic AI

Evidence-based AI: from trailblazer to trustblazer?

Thomas Luechtefeld<sup>1,2</sup> and Thomas Hartung<sup>2,3,1</sup>





**MPS WORLD SUMMIT**

CONNECT, EXCHANGE, EDUCATE

**2022, 2023, 2024, 2025**



**INTERNATIONAL MPS SOCIETY**

CONNECT, EXCHANGE, EDUCATE

**2023**



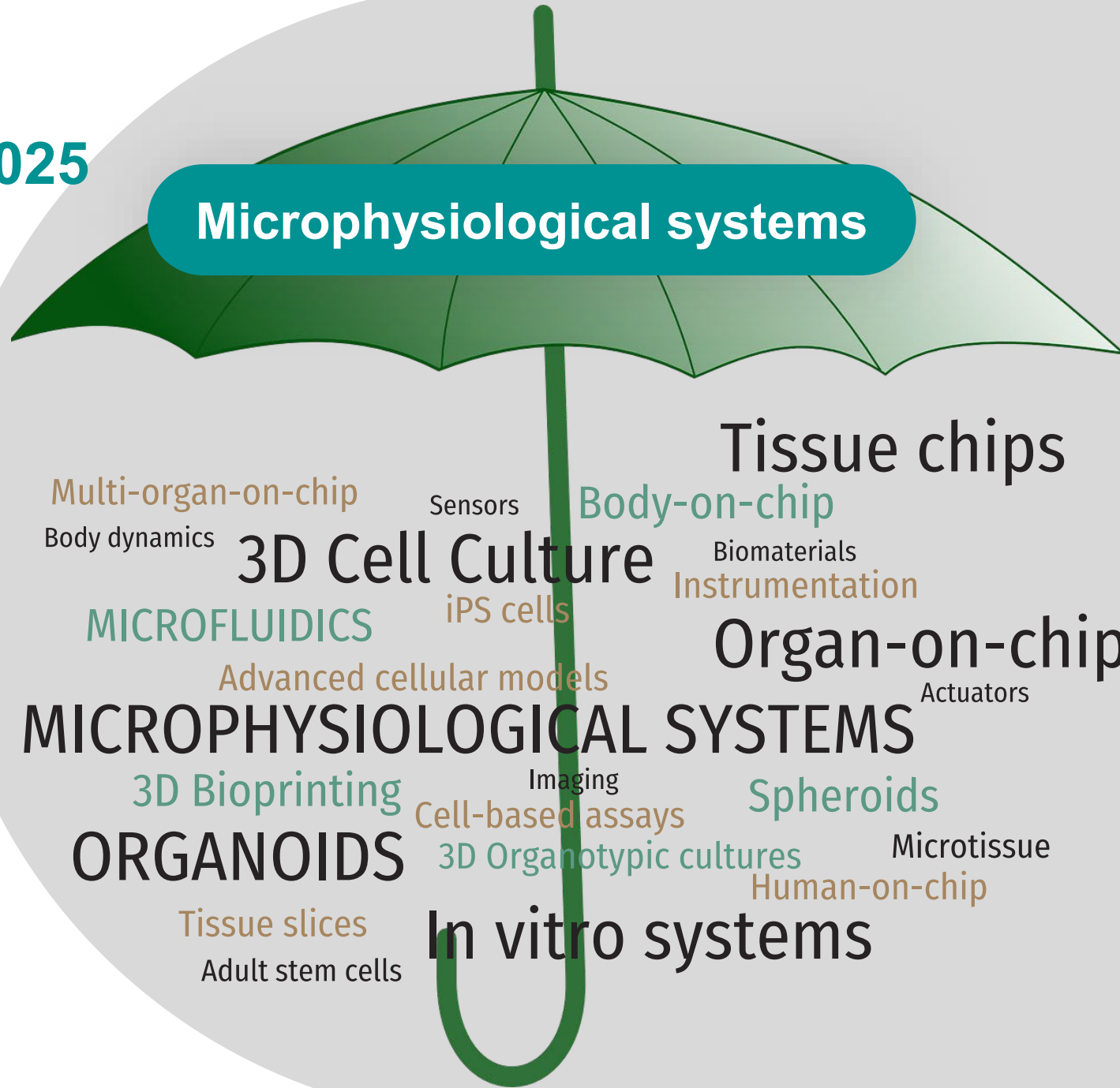
**MPS workshops**

**2016, 2020,  
2025**

**GCCP**

**2.0**

**2022**



**Tissue chips**

**Body-on-chip**

**3D Cell Culture**

**Organ-on-chip**

**MICROPHYSIOLOGICAL SYSTEMS**

**ORGANOIDS**

**In vitro systems**

**3D workshop 2014**

# Microphysiological Systems World Summit

WASHINGTON, DC  
MAY 25–29, 2026

ABSTRACT SUBMISSION OPEN UNTIL January 11<sup>th</sup> 2026

CONNECT

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**MPS WORLD SUMMIT**  
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# Guidance Document on Good Cell and Tissue Culture

## Practice 2.0 (GCCP 2.0)

**ALTEX 2022, 39:30-70**



*David Pamies<sup>1</sup>, Marcel Leist<sup>2,3</sup>, Sandra Coecke<sup>4</sup>, Gerard Bowe<sup>4</sup>, Dave Allen<sup>5</sup>, Gerhard Gstraunthaler<sup>6</sup>, Anna Bal-Price<sup>4</sup>, Francesca Pistollato<sup>4</sup>, Rob deVries<sup>7,8</sup>, Helena T. Hogberg<sup>9</sup>, Thomas Hartung<sup>2,9</sup> and Glyn Stacey<sup>10,11,12</sup>*



- **Quality of cell model**  
**(GCCP)**
- **Quality of reporting**  
**(GIVReSt)**
- **Quality of results**  
**(validation)**



Food for Thought ...

## Guidance for Good *In Vitro* Reporting Standards (GIVReSt) – A Draft for Stakeholder Discussion and Background Documentation

Ronit Mohapatra<sup>1</sup>, Marcel Leist<sup>2,4</sup>, Sonja von Aulock<sup>5</sup> and Thomas Hartung<sup>1,2,3</sup>



Food for Thought ...

# Custom GPTs to Aid in Compliance Checking for Reporting Standards in Academic Publishing

Ronit Mohapatra<sup>1</sup> and Thomas Hartung<sup>1,2,3</sup>

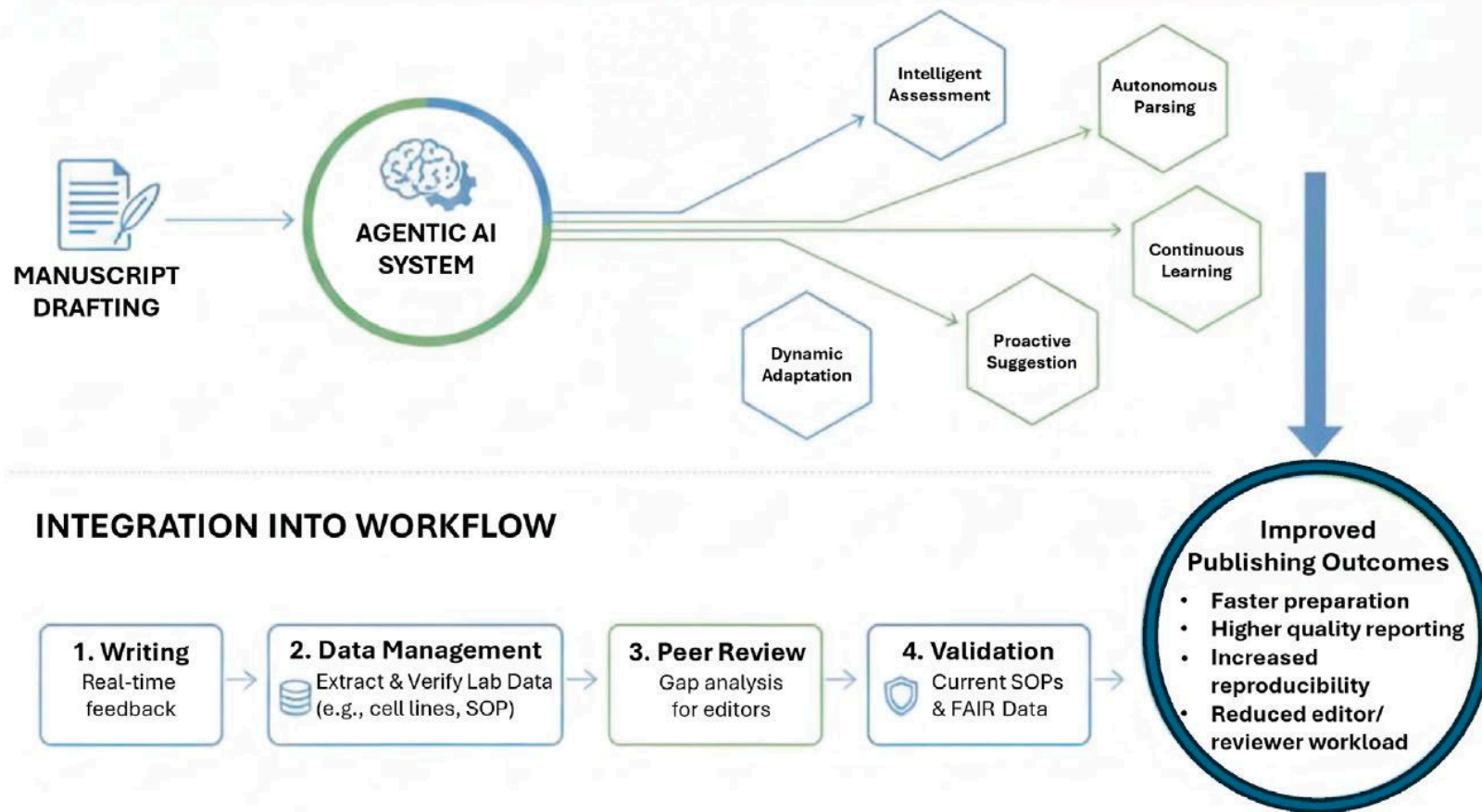


## Agentic AI to check compliance with:

- GIVReSt
- Tox-R-Tool
- ARRIVE guidance

### AGENTIC AI: THE COMPLIANCE COPILOT

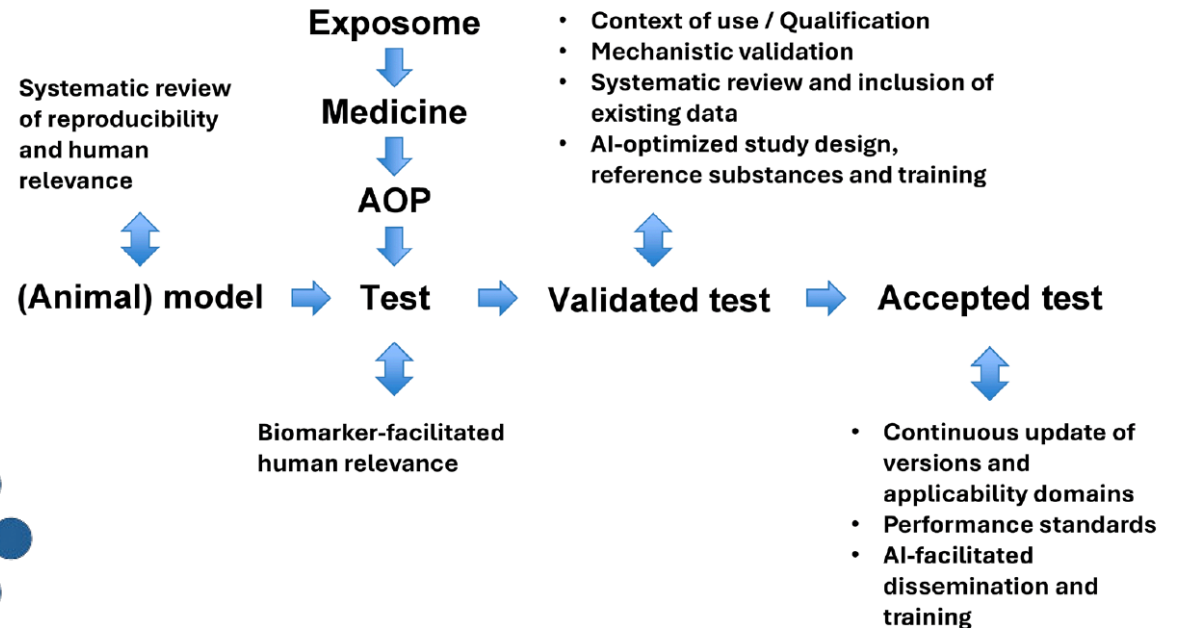
Automating Scientific Reporting & Standards Compliance



# Emerging Ontologies for MPS

Item	Form	Covers organoids?	Covers MPS/OoC?	What it contributes to ontology-building
NAMO (2025–)	Ontology + LinkML schema (generates OWL/JSON schema, etc.)	Yes	Yes	A unifying semantic framework spanning organoids + OoC + other NAMs
MPS-Db (2016; 2020)	Structured database + metadata framework	Indirect	Yes	Published, operational metadata structure for describing/analyzing MPS experiments
GIVReSt (2025)	Reporting standard	Yes	Yes	Defines minimum reporting elements that can be formalized into ontology properties
MIOR (2025)	Organoid reporting standard	Yes	Indirect	Organoid-focused minimum information standard aligned with FAIR interoperability goals
MISpheroid (2021)	Knowledgebase + “minimum information string”	Spheroids (related)	Indirect	Concrete parameter set for 3D model identity; useful pattern for organoid/MPS identity metadata
MIACARM (2016)	Minimum information standard (cell assays)	Mentions organoids	Indirect	Explicitly promotes use of controlled terms/ontologies for exchangeable assay metadata
ISO/CD 25448 (2026 stage)	Vocabulary standard (in development)	Often under “MPS” umbrella	Yes	Canonical definitions and terminology scope that an ontology can align to
Piergiovanni et al. (2021)	Standards/roadmap analysis paper	Yes (as part of “MPS” scope)	Yes	Establishes the need for harmonized terminology and classification

# Next Generation Validation for Next Generation Risk Assessment



Food for Thought ...

## The Validation of Regulatory Test Methods – Conceptual, Ethical, and Philosophical Foundations

Thomas Hartung



Food for Thought ...

## E-validation – Unleashing AI for Validation

Thomas Hartung<sup>1,2</sup>, Alexandra Maertens<sup>1</sup> and Thomas Luechtefeld<sup>1,3</sup>



Food for Thought ...

## Leveraging Biomarkers and Translational Medicine for Preclinical Safety – Lessons for Advancing the Validation of Alternatives to Animal Testing

Thomas Hartung<sup>1,2</sup>, Nicholas M. P. King<sup>3</sup>, Nicole Kleinstreuer<sup>4</sup>, Marcel Leist<sup>2,5</sup> and Danilo A. Tagle<sup>6</sup>



Food for Thought ...

## Challenges and Opportunities for Validation of AI-Based New Approach Methods

Thomas Hartung<sup>1,2</sup> and Nicole Kleinstreuer<sup>3</sup>



# The Promise and Potential of Brain Organoids

*Lena Smirnova\* and Thomas Hartung\**

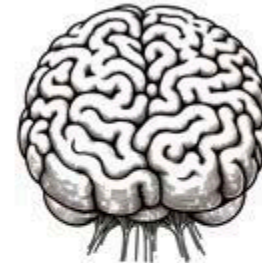


**Lena Smirnova**

## BRAIN ORGANOID OPPORTUNITIES



## BRAIN ORGANOID AND BRAINSPHERE VARIANTS

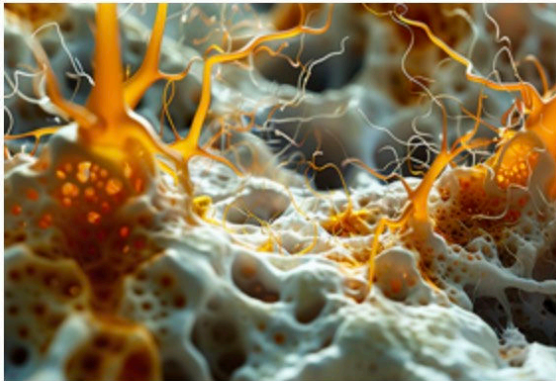


- Different species; patient-derived or healthy; genetically modified or not
- Reaggregation of tissue-derived cell suspension or stem cell-derived (embryonic stem cells or iPSC)
- Use of extracellular matrix, hydrogels or none
- Scaffold or none; possible electrode integration
- Use of growth factors to induce differentiation or not; addition of cell types (micro-glia)
- Gravity-based aggregation (e.g., hanging drop or microtiter plates) or bioreactor (e.g., spinner or shaker)
- Brain region-specific organoids; addition of blood-brain-barrier
- Fusion and assembloids of different brain regions or organs (cell lineages)
- Use of microfluidics and vascularization for perfusion or not



For best mobile view,  
please rotate your phone

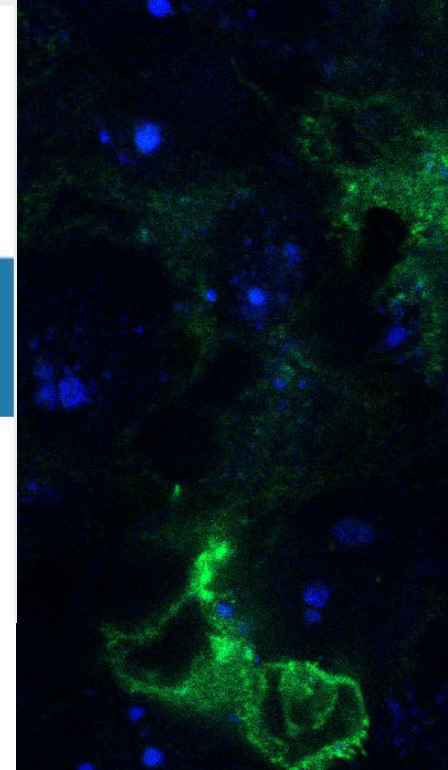
## Four Pilot Projects Selected to Advance Human-Based Research Models



### Developmental Neurotoxicity

This lab-based system is designed to assess how the brain develops and reacts to harmful chemicals. It combines advanced genetic tools with fluorescent markers to track six essential processes, including how neurons form, how brain cells connect with each other, and how brain networks take shape. Once validated, this system could provide a reliable, cost-effective alternative for assessing developmental brain toxicity.

# Announced Last month!



## Mini- Brainbow

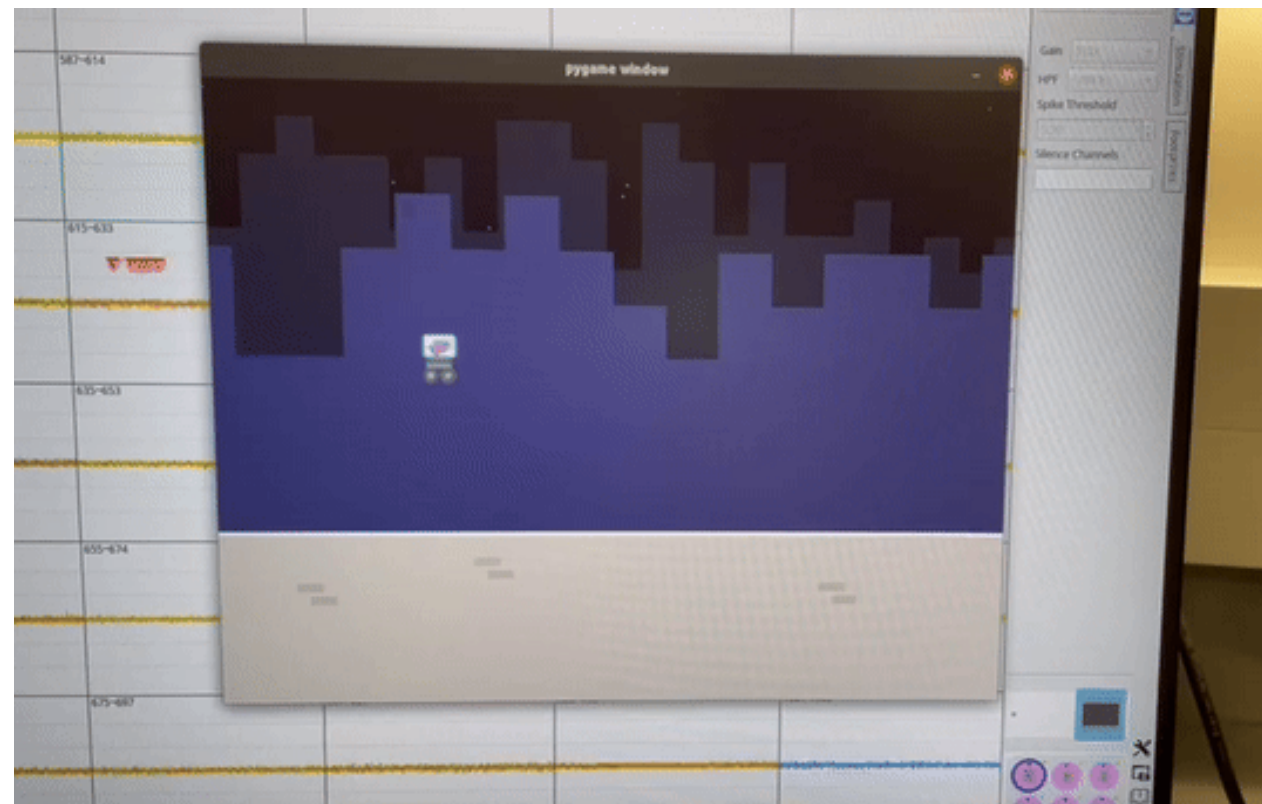
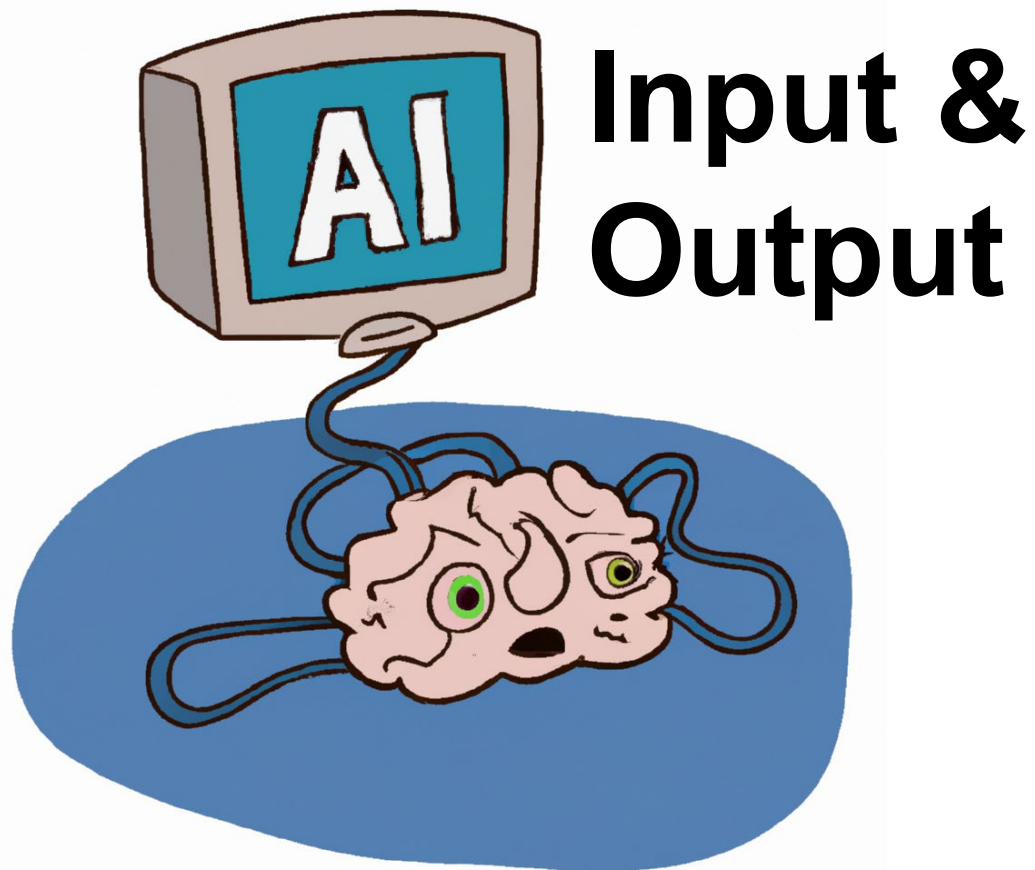


Neurons    Astrocytes  
Oligodendrocytes  
Synapses    3D electrophys

## Oligodendrogenesis and myelination tracing in a CRISPR/Cas9-engineered brain microphysiological system

July Carolina Romero<sup>1</sup>, Cynthia Berlinicke<sup>2</sup>, Sharon Chow<sup>1</sup>,  
Yukan Duan<sup>2</sup>, Yifei Wang<sup>1</sup>, Xitiz Chamling<sup>2</sup> and  
Lena Smirnova<sup>1\*</sup>

# Combining MPS, $\equiv$ Organoid Intelligence (OI) sensors and AI



# INTERNATIONAL CONFERENCE ON ORGANOID INTELLIGENCE

Mark your calendars for a first-of-its-kind, in-person event bringing together the brightest minds and boldest leaders working across OI, biological and wetware tech.

Featuring targeted talks, hands-on sessions, and interactive demonstrations, this will be an incredible opportunity to learn, connect and help lead this field forward.



SAVE THE DATE

10.26.26

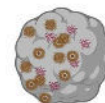
— 10.27.26

Johns  
Hopkins  
Bloomberg  
Center

Washington, D.C.  
washingtondc.jhu.edu/  
bloomberg-center/

## The next big step

### OI and Medicine



Food for Thought ...

**Organoid Intelligence (OI) - The Ultimate  
Functionality of a Brain Microphysiological System**

*Lana Snirnov<sup>1</sup>, Iry E. Moresco Pavesio<sup>2</sup> and Thomas Hartung<sup>1,2</sup>*



### Neurodevelopmental & cognitive disorders

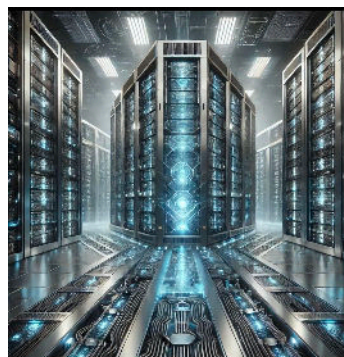
- Autism
- Intellectual disabilities
- Neurological rare diseases associated with cognitive impairment



### Neurodegeneration

- Alzheimer's disease
- Amyotrophic Lateral Sclerosis
- Multiple Sclerosis

### Public Health

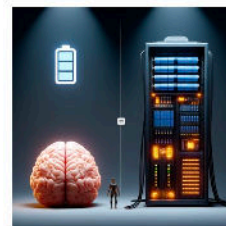


\$600 million; 680 m<sup>2</sup>

One million time  
more energy

**Fake Intelligence**

The work of a supercomputer  
with the energy of a laptop



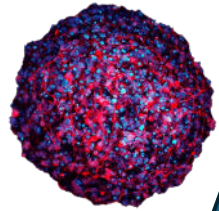
1.4 kg  
Storage ~ 2.5 PetaByte

**Actual Intelligence**

# Disruptive Technologies

Microphysiological systems (MPS)  
and other data-rich technologies

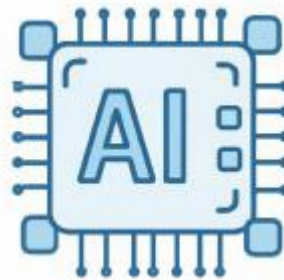
(open) Scientific literature  
& Data



Big Data

Exposome  
Intelligence (EI)

Early Testing  
with small  
amounts



Green Toxicology  
& Drug fail-early

Benign by design

Personalized Medicine

Human Exposome

*“safer products”*

*“better treatments”*

*“safer people”*

# REVOLUTION



- Integrate Disruptive Technologies with Existing Knowledge
- Accelerate Drug Development
- Optimize Prevention and democratize Healthcare Access

**AI, MPS, ~omics,  
sensors....**

**Green Toxicology  
Human Exposome**

Slides available:  
<https://share.zight.com/KouXyZpl>

