



# The Digital Twin in Clinical Endocrinology

Iraj Nabipour, MD

Bushehr University of Medical Sciences

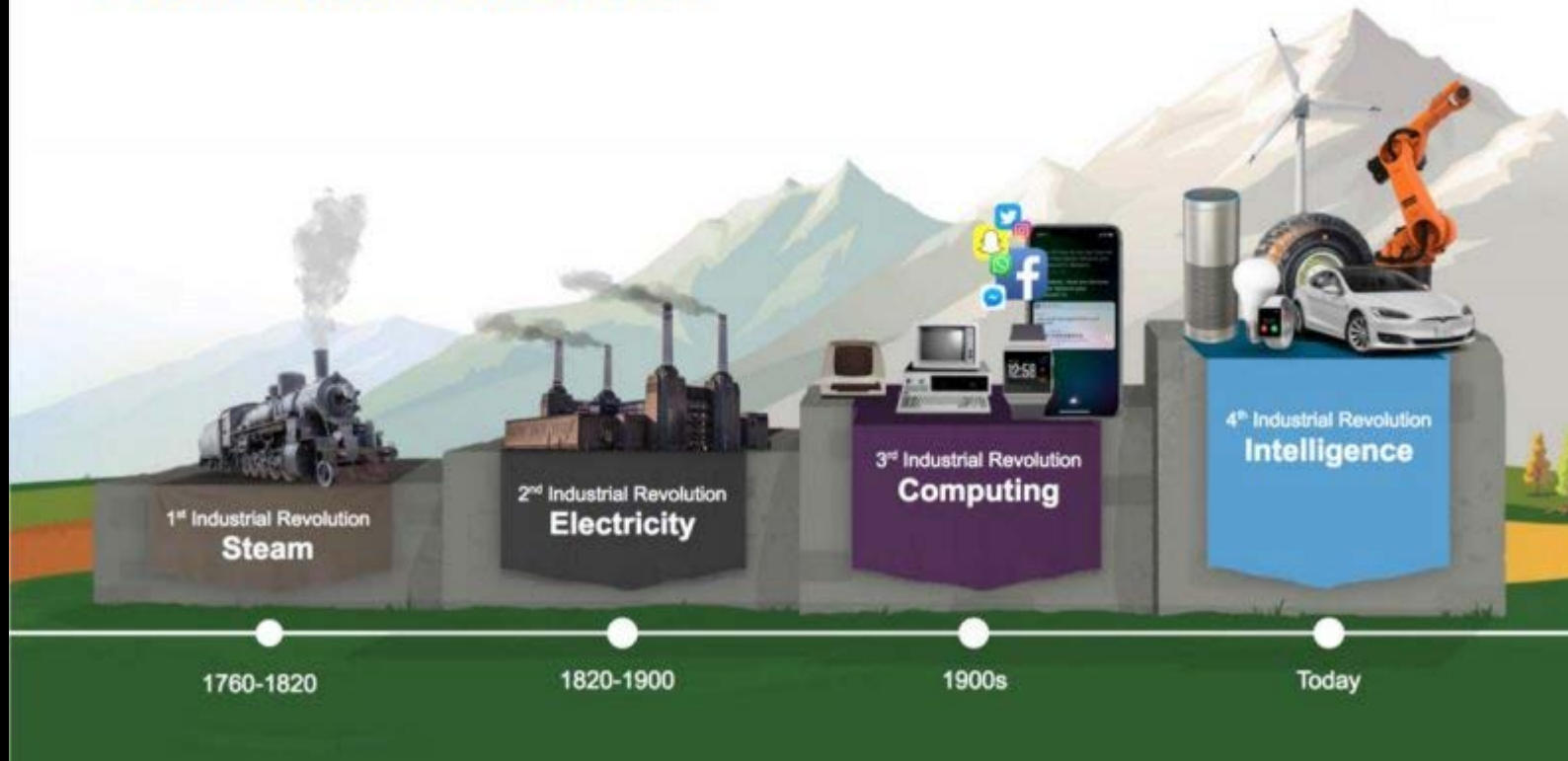
# **DIGITAL TWIN**

## **A REVOLUTION IN MEDICINE**

**Dr. Iraj Nabipour**



## Fourth Industrial Revolution



These advances are merging the physical, digital and biological worlds in ways that create both huge promise and potential peril.

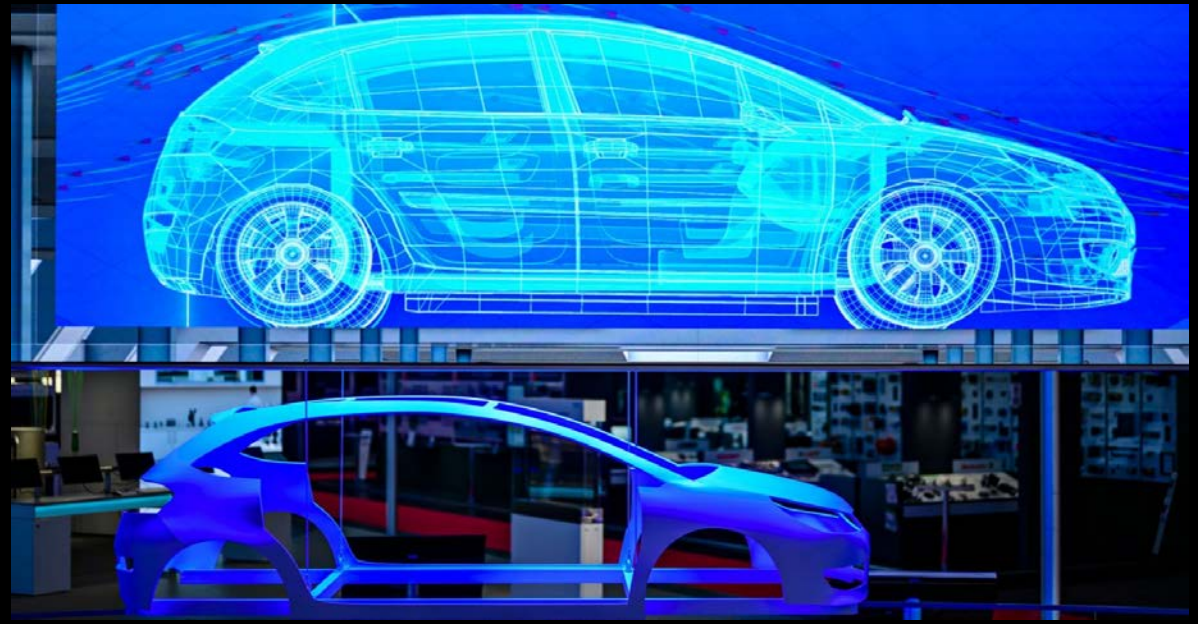


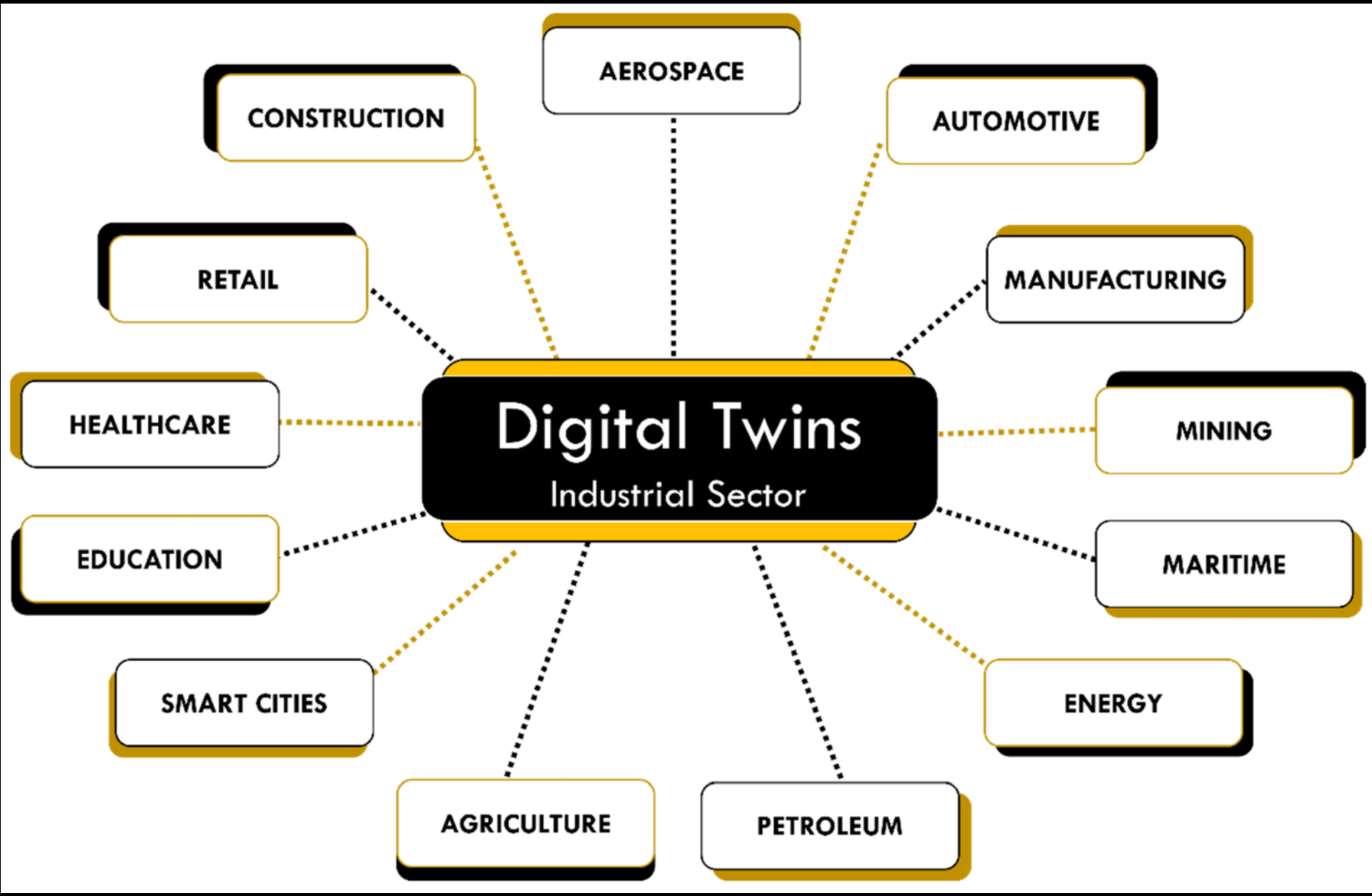


The concept of a physical twin, a precursor to Digital twin, is rather old and dates to NASA's Apollo program.

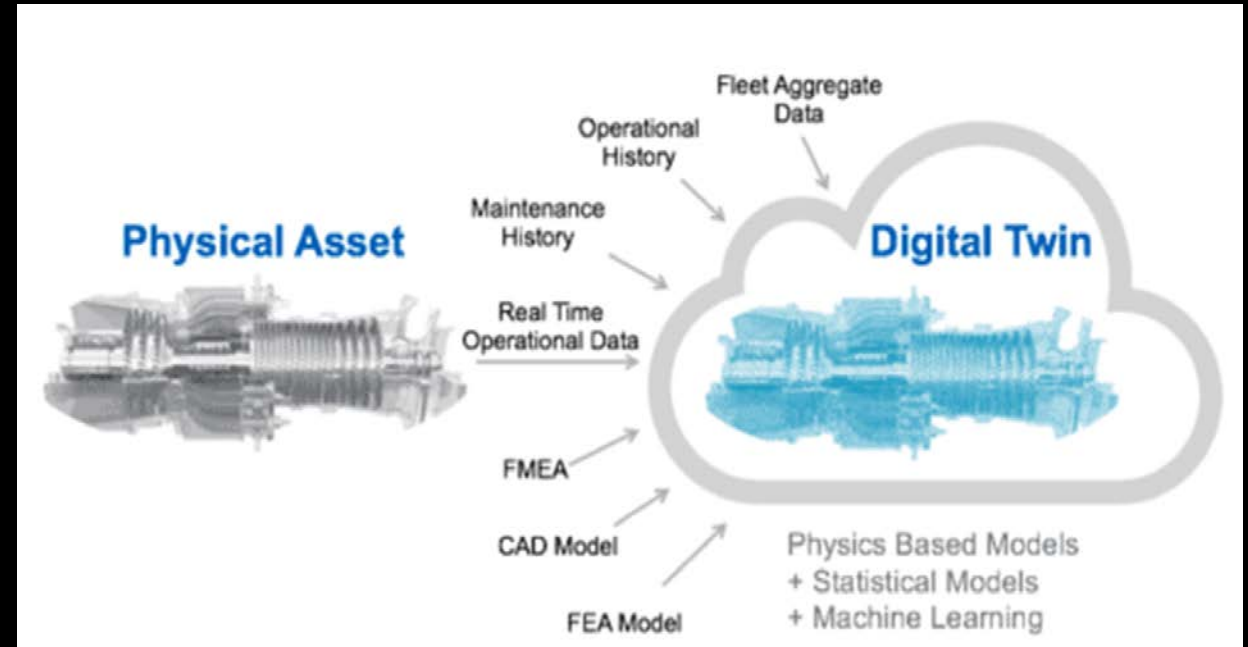
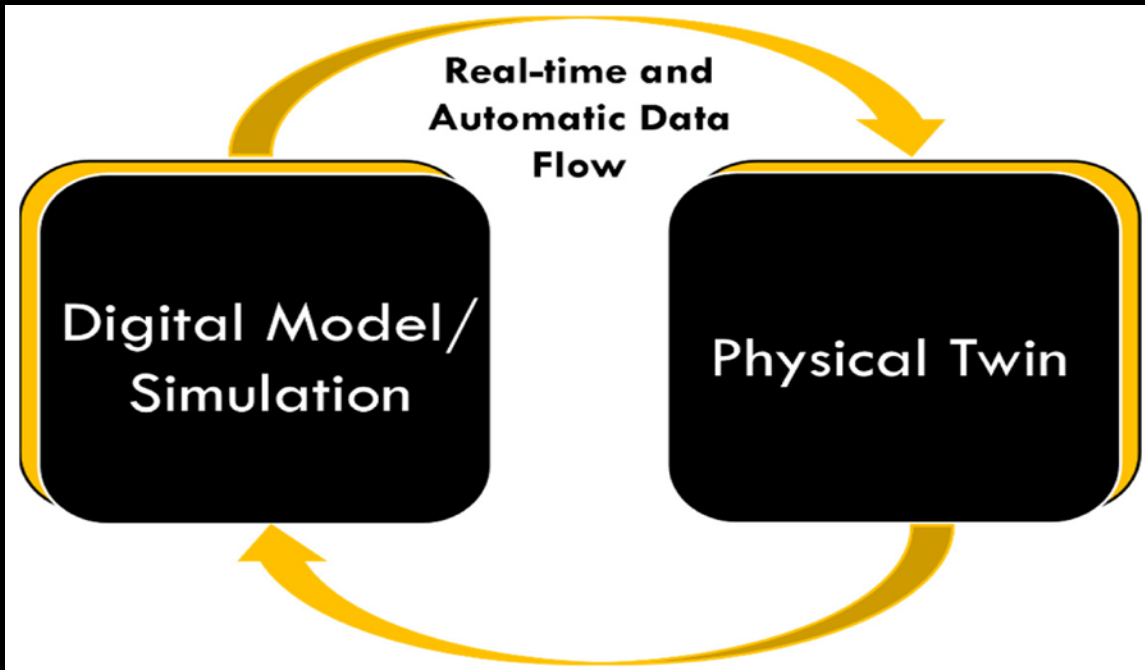




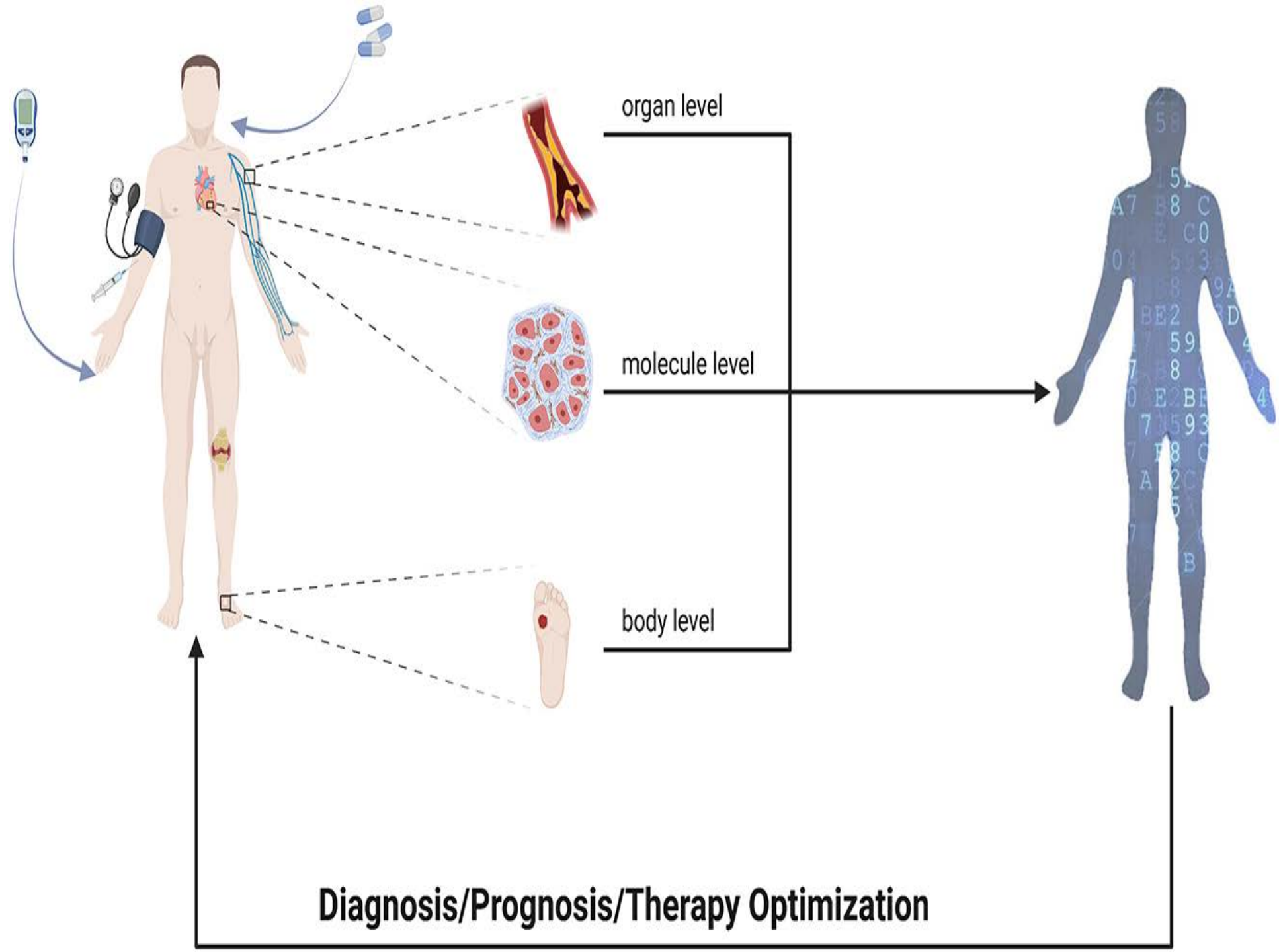




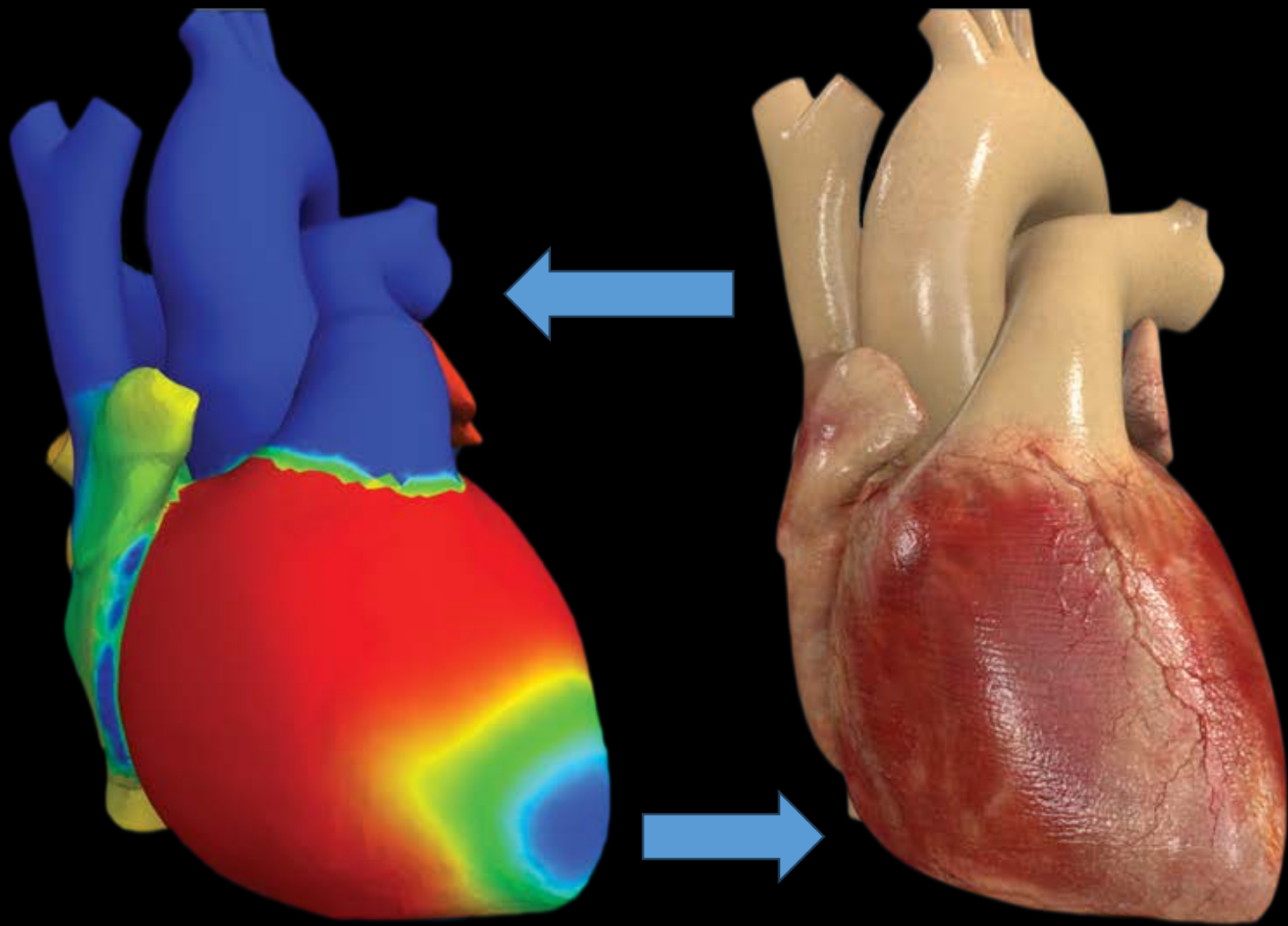




1. Physical twin: A real-world entity (living/non-living) such as part/product, machine, process, organization, or human, etc.
- (ii)
2. Digital twin: The digital representation of the physical twin with the capability to mimic/mirror its physical counterpart in real time.
- (iii)
3. Linking mechanism: The bidirectional flow of data between the two which operates automatically in real-time.

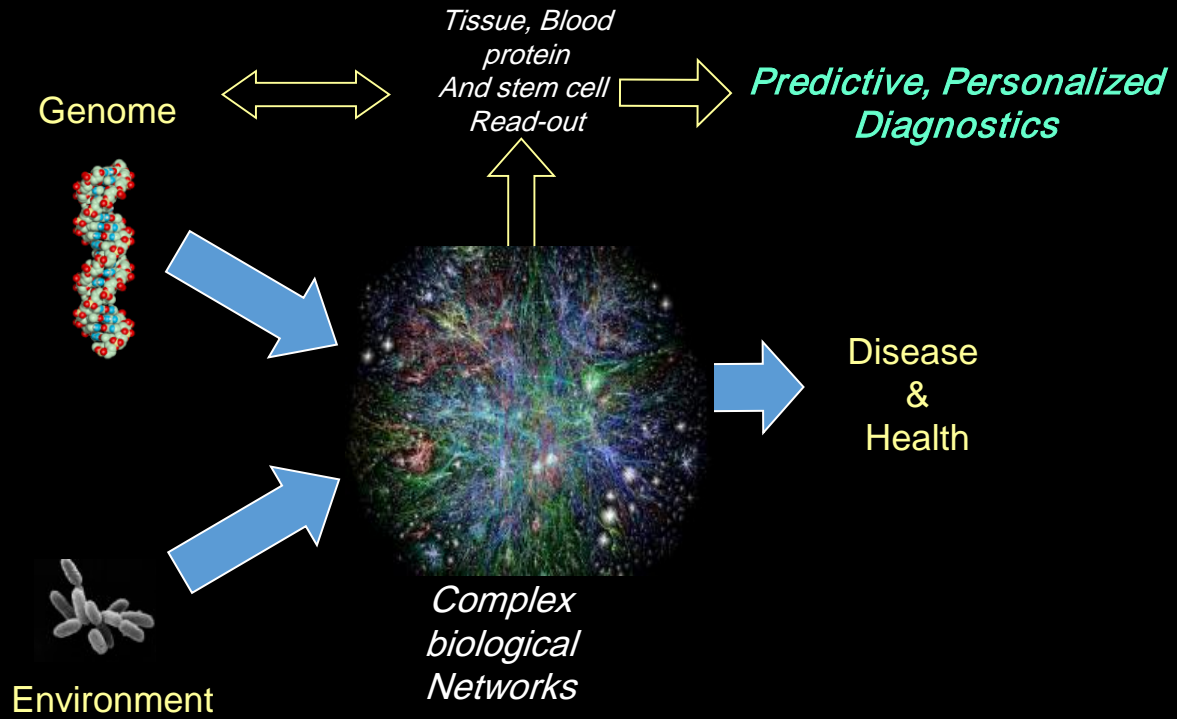








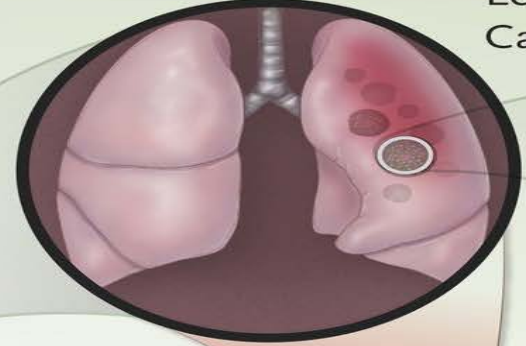
# There are two types of Biological Information



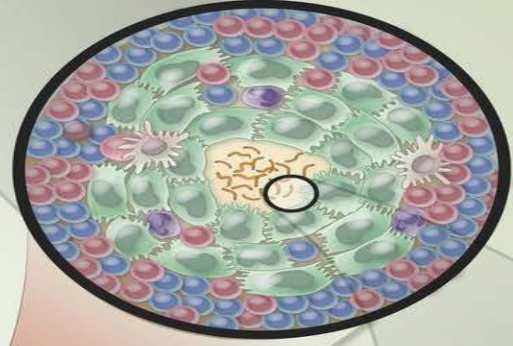
**18th century**  
Sick person  
Phthisis



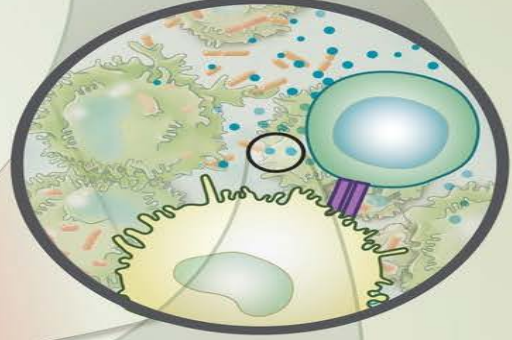
**Early 19th century**  
Lesions of organs and tissues  
Caseating granulomata



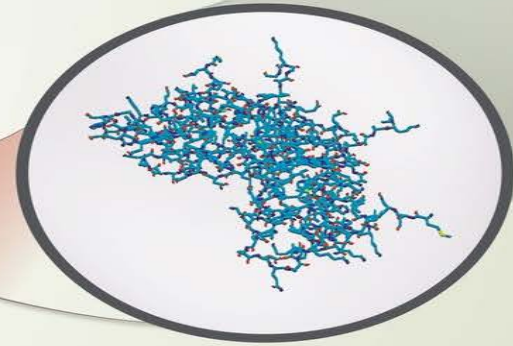
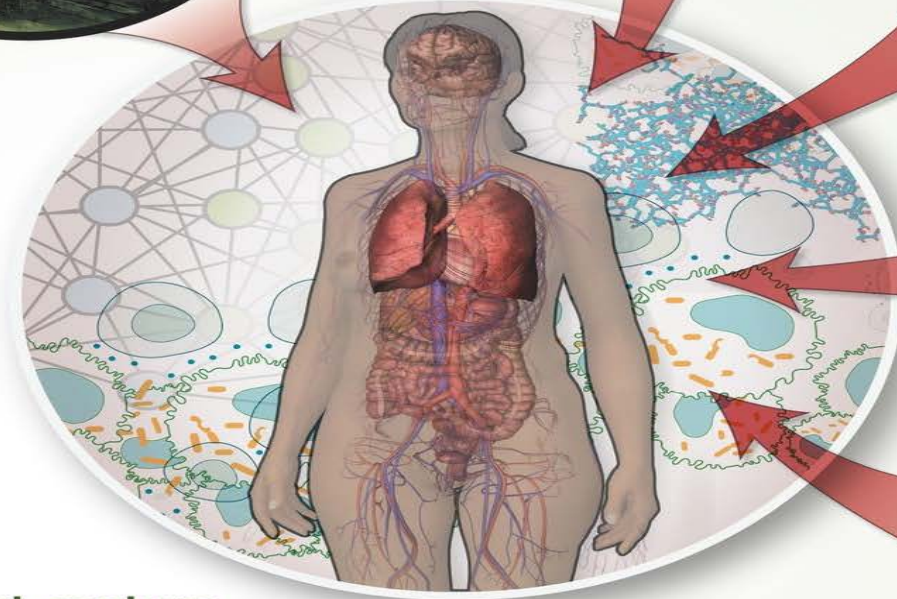
**Late 19th century**  
Lesions of cells  
and microbes  
*M. tuberculosis*



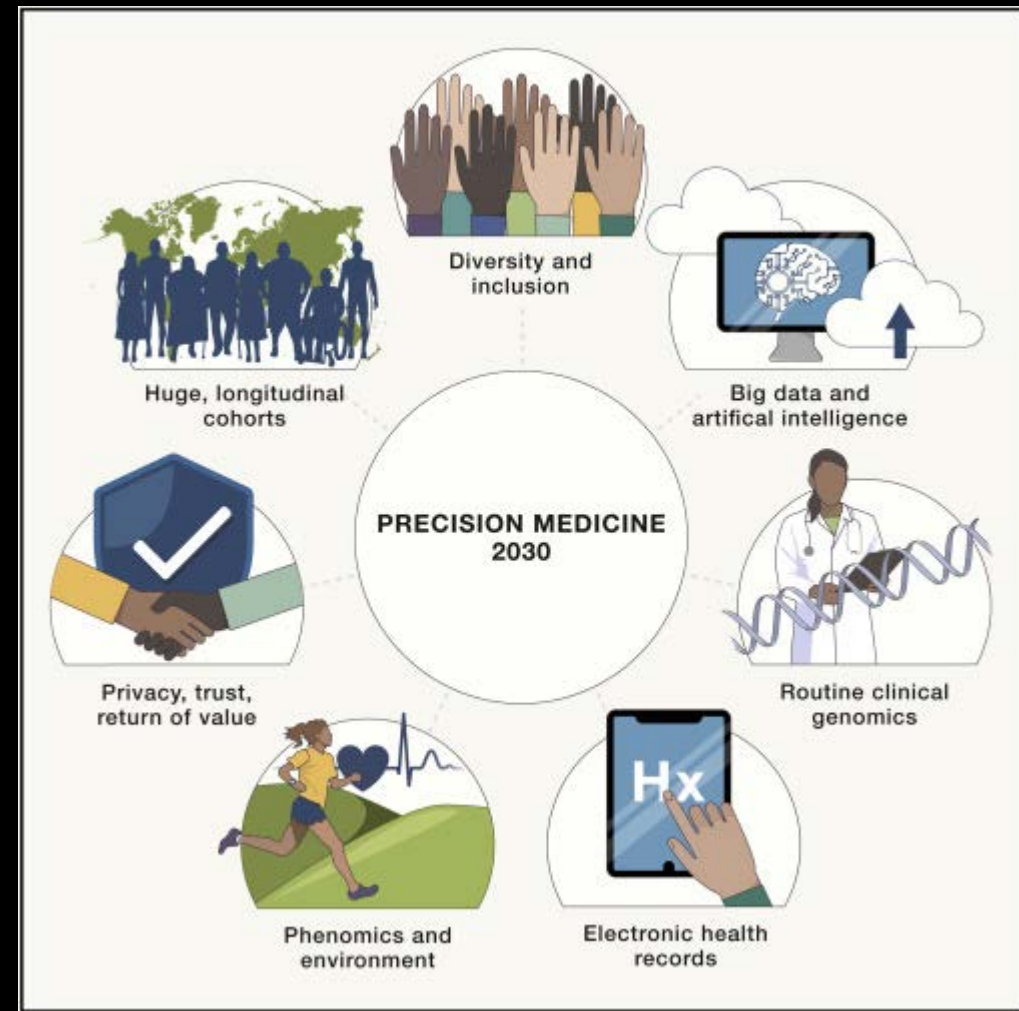
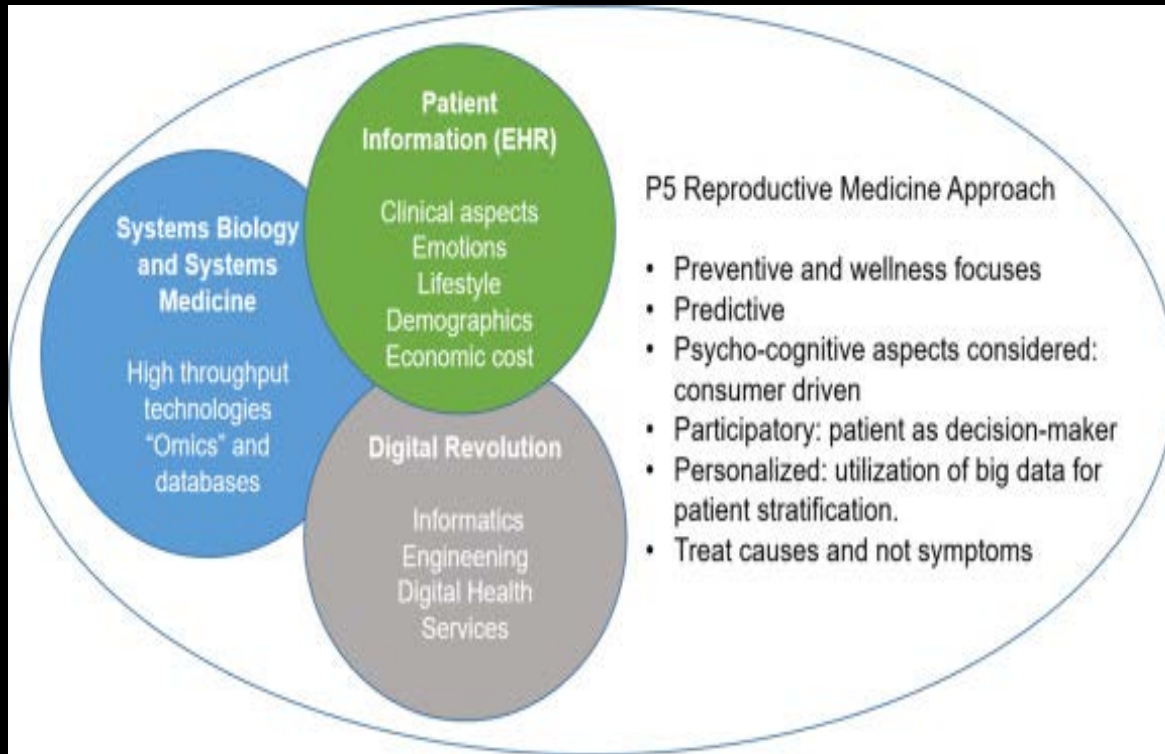
**Late 20th century**  
Lesions detected  
at molecular level  
Interferon testing



**21st century**  
The challenge of reassembly

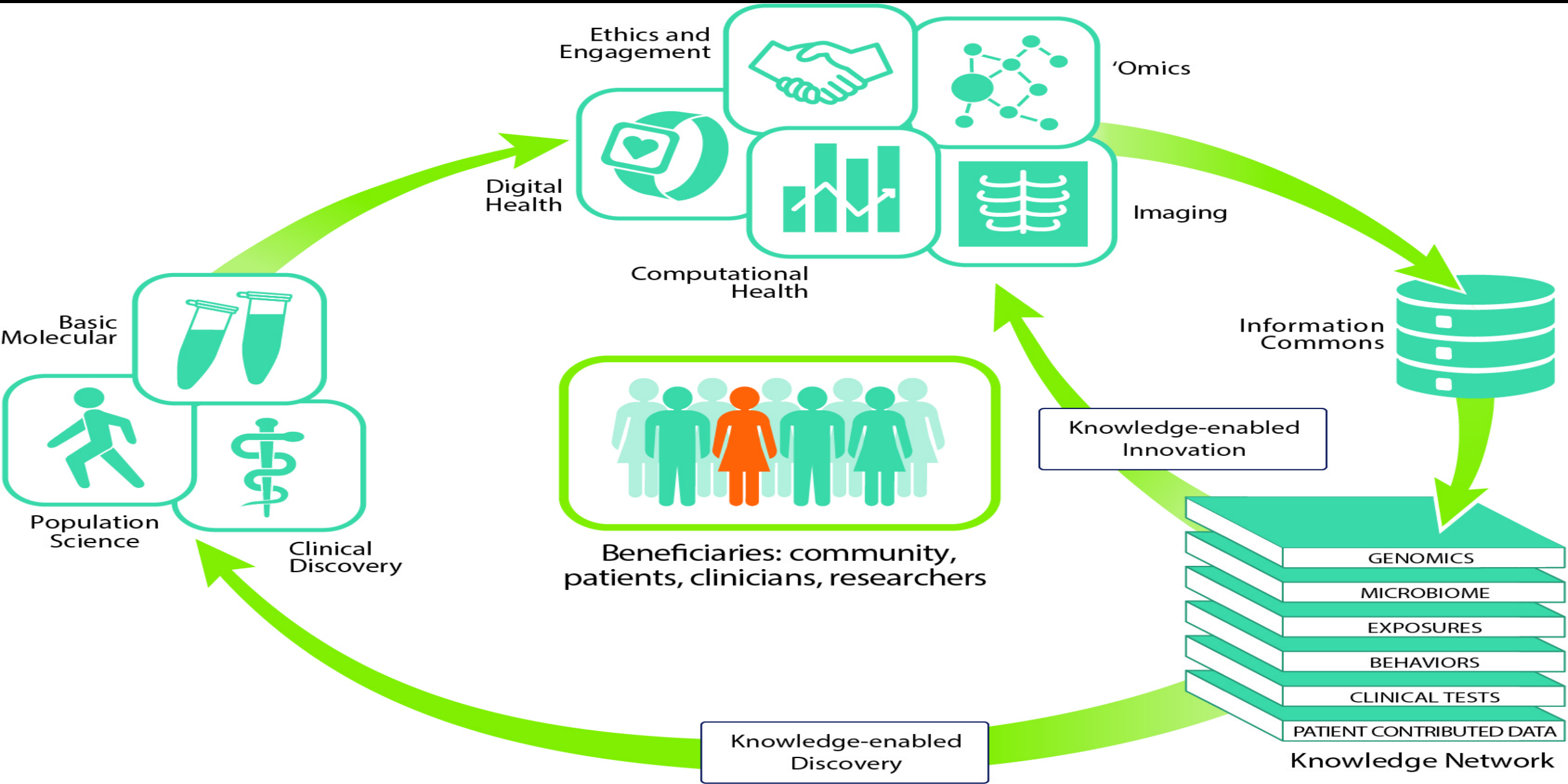






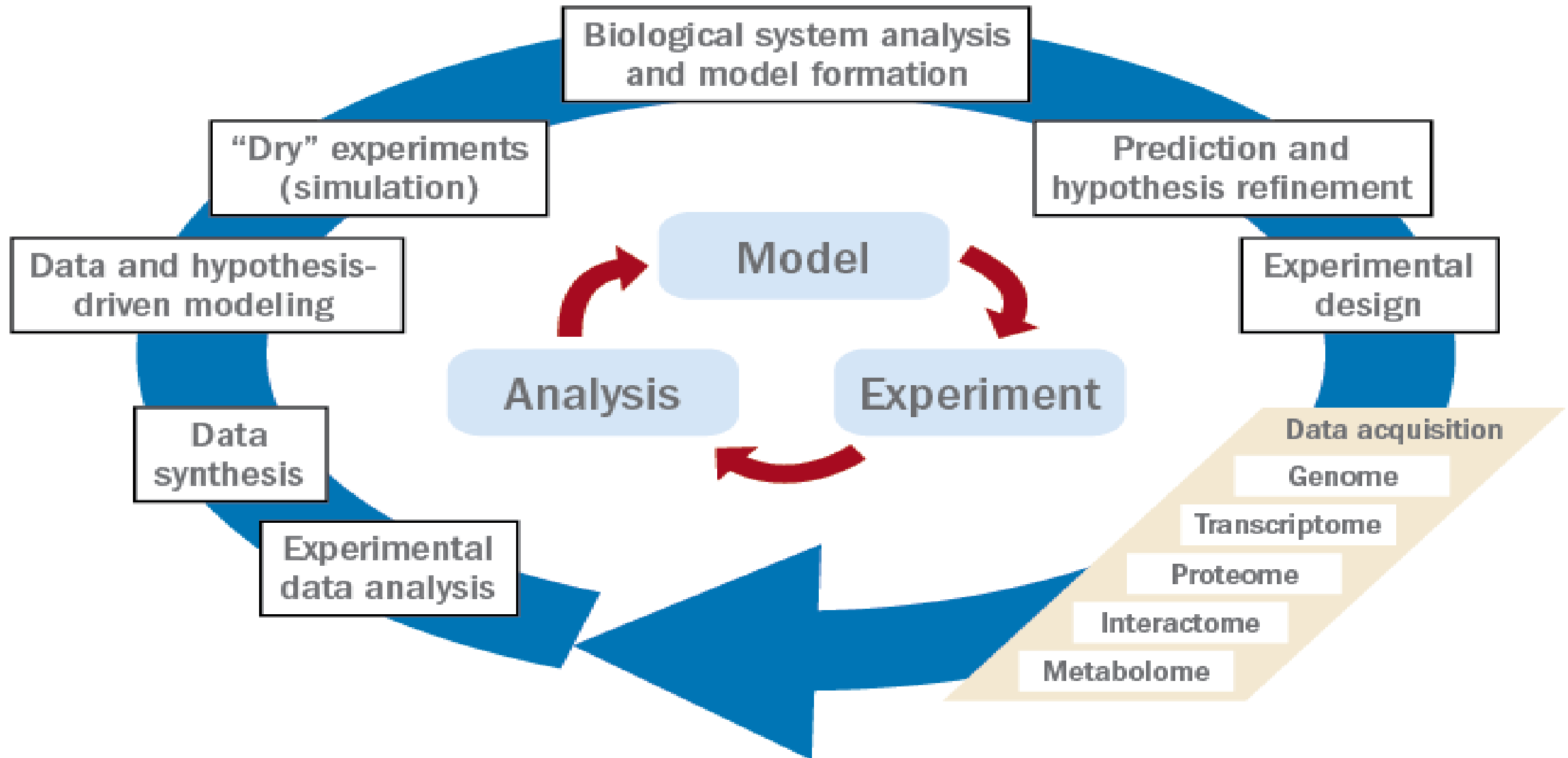
Systems medicine is defined as the application of the systems approach to the prevention of, understanding and modulation of, and recovery from developmental disorders and pathological processes in human health (Clermont et al. 2009).

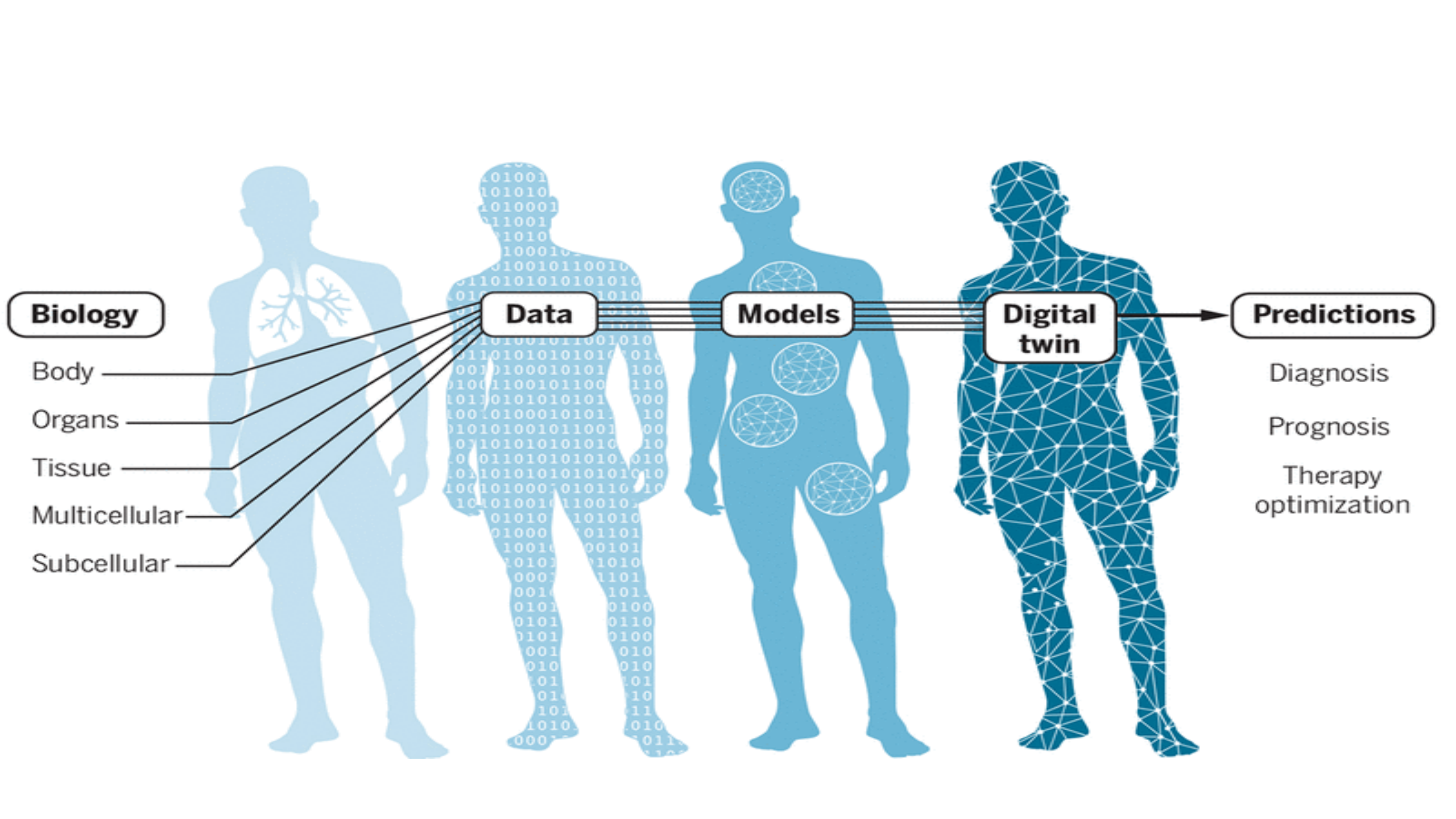


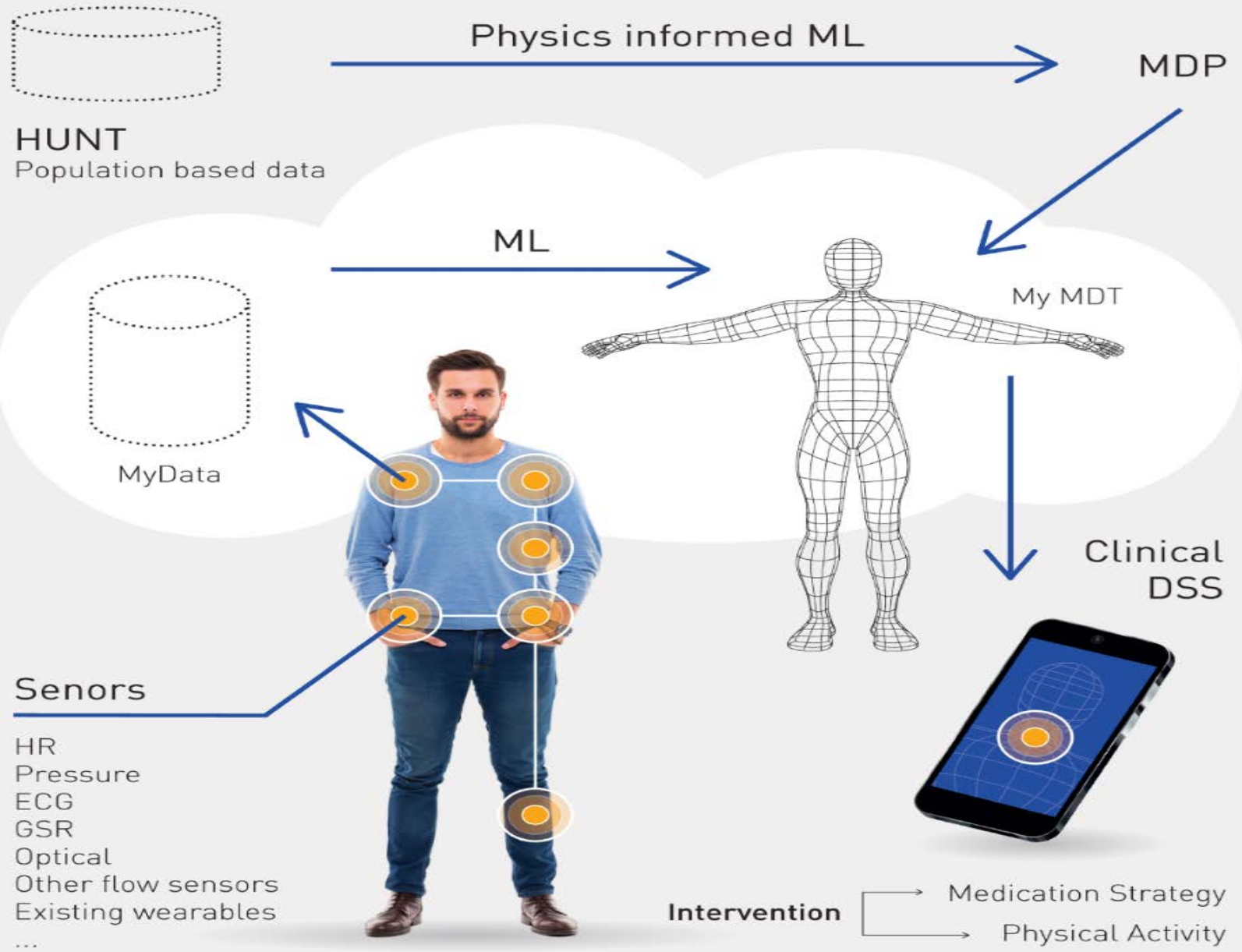


“Precision medicine is an emerging approach for disease treatment and prevention that takes into account individual variability in genes, environment, and lifestyle for each person.”

# The process of systems biology research







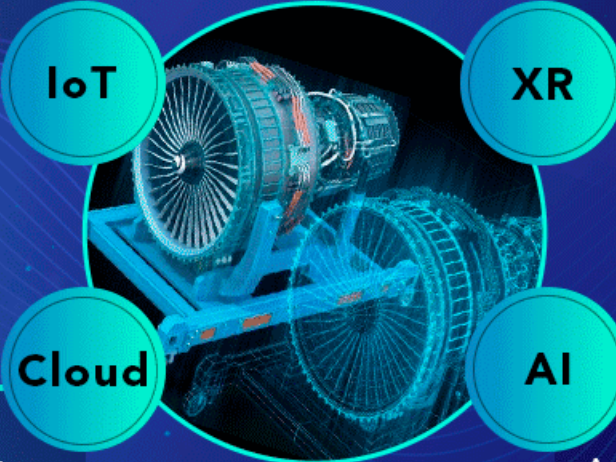






# TECHNOLOGIES USED IN DIGITAL TWINS

IoT sensors enable constant data transmission, which is used to create a digital duplicate of the physical object



Due to its visualization capabilities, XR allows to digitally model physical objects

Cloud computing allows to store gained data in the virtual cloud and easily access them from any location



As an advanced analytical tool, AI automatically analyze obtained data, provide valuable insights and make predictions

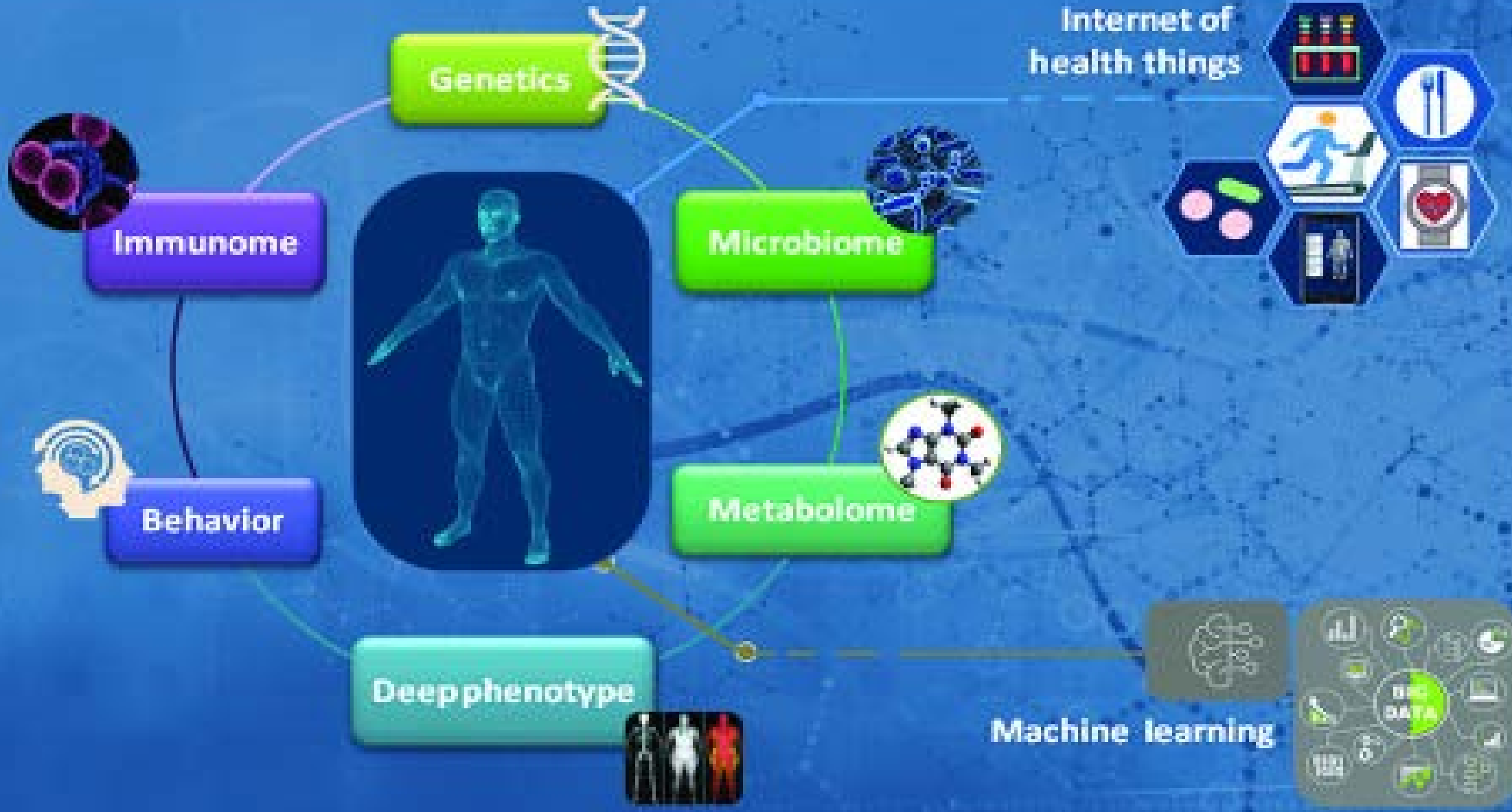
# DEEP MEDICINE

HOW ARTIFICIAL  
INTELLIGENCE  
CAN MAKE  
HEALTHCARE  
HUMAN AGAIN

**ERIC TOPOL, MD**

With a foreword by  
**ABRAHAM VERGHESE,**  
*author of Cutting for Stone*





Genetics

Immunome

Behavior

Microbiome

Metabolome

Deep phenotype

Internet of health things

Machine learning

BIG DATA

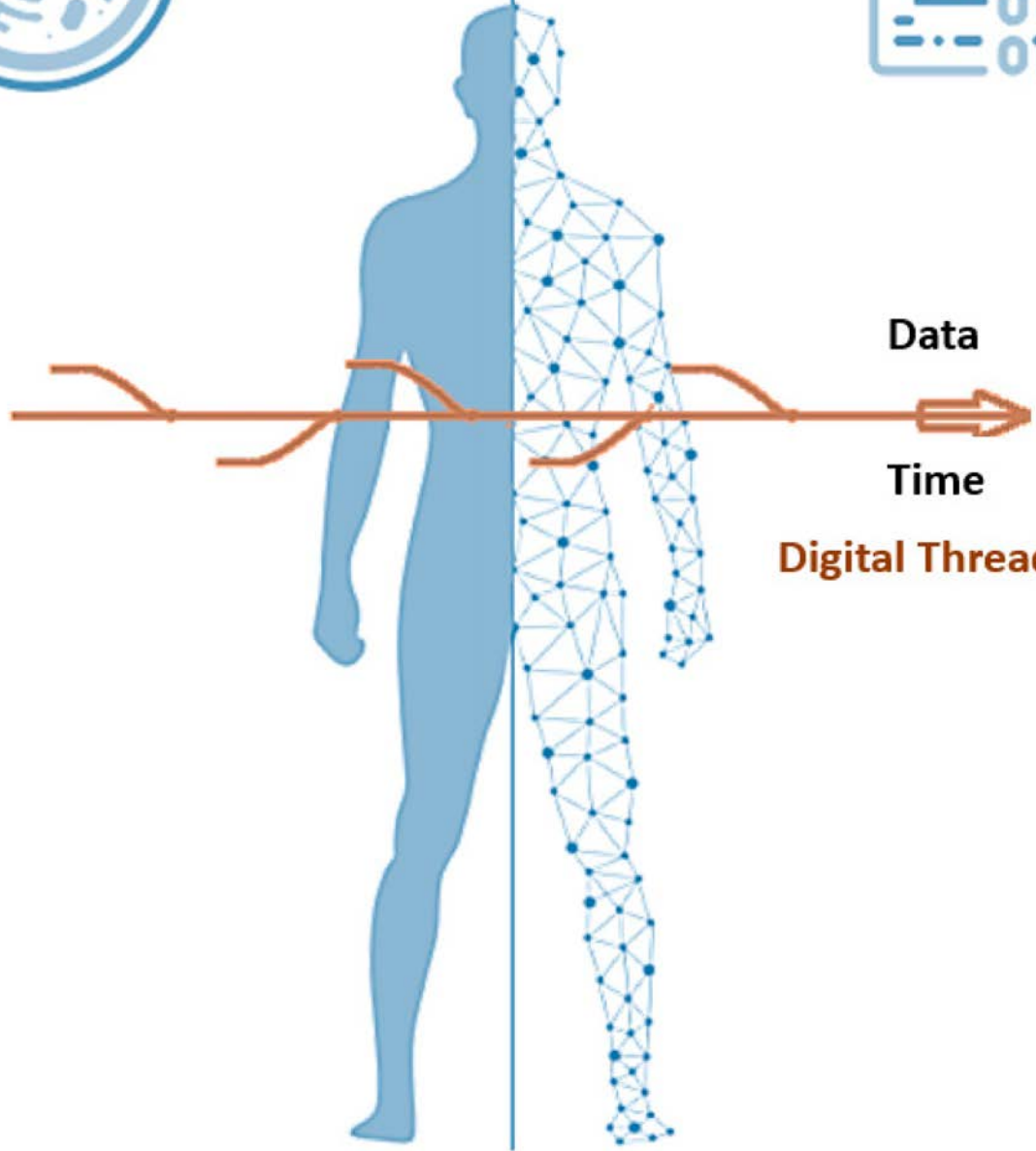




Real World



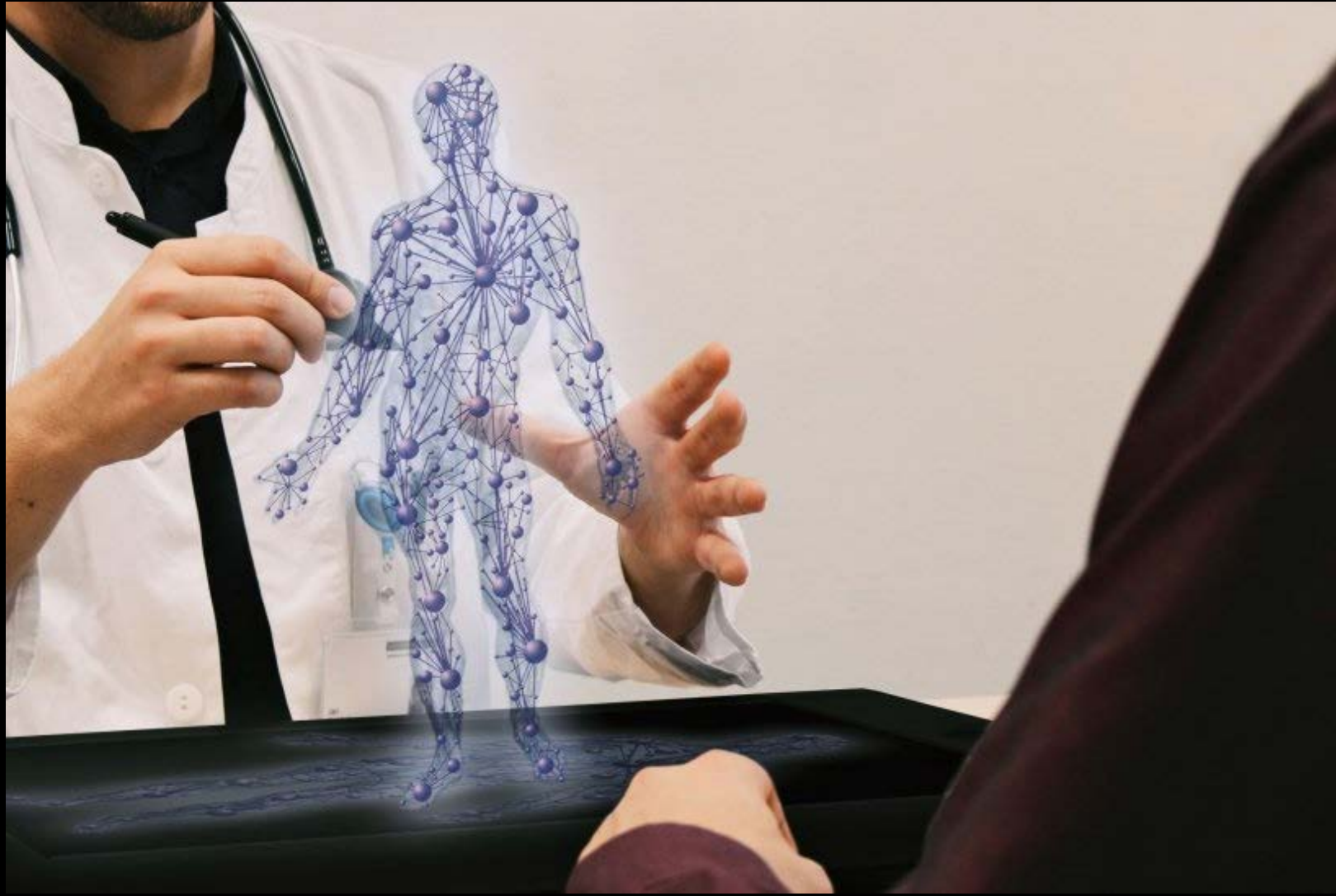
Digital Twin



Data

Time

Digital Thread

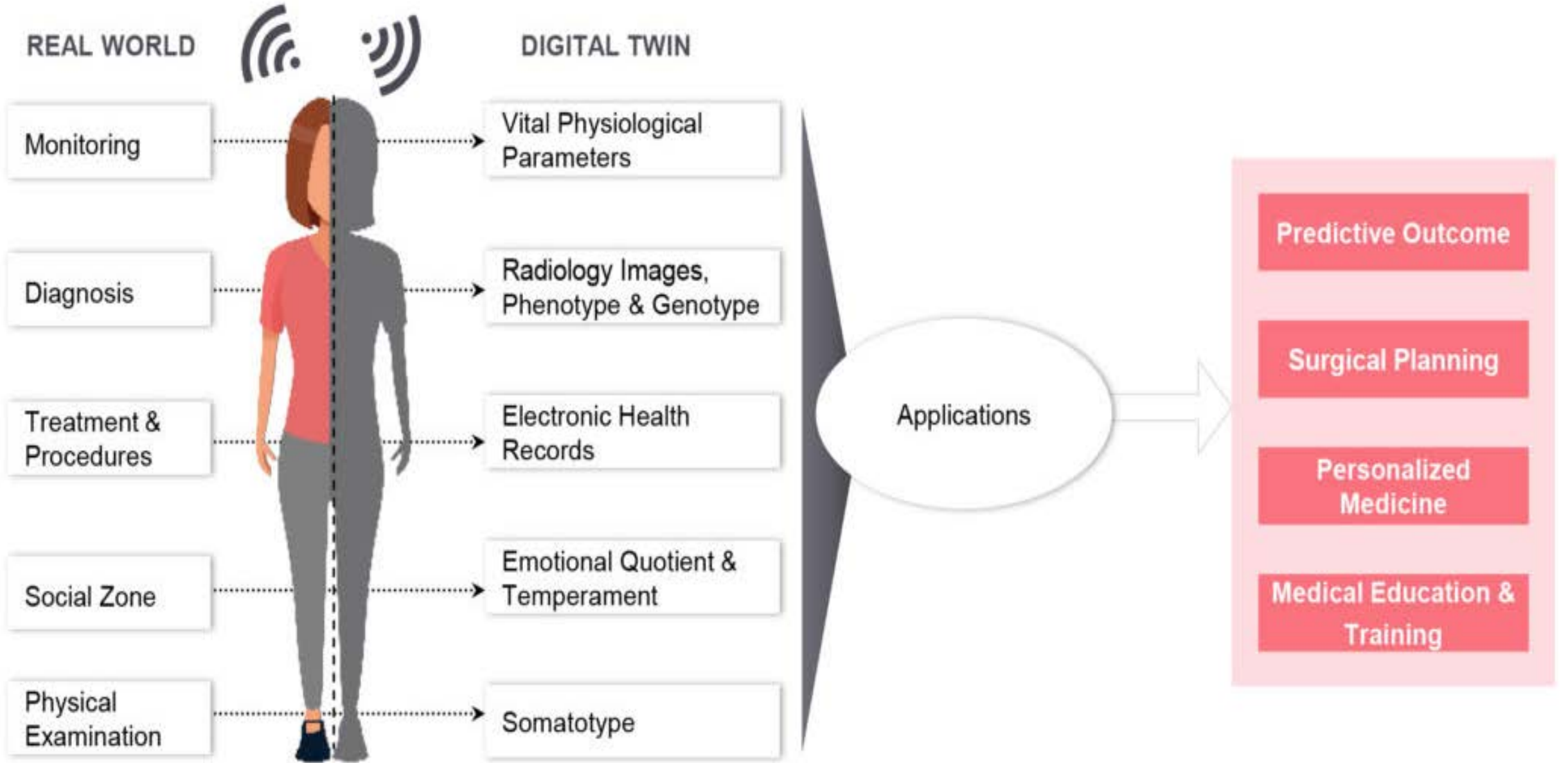


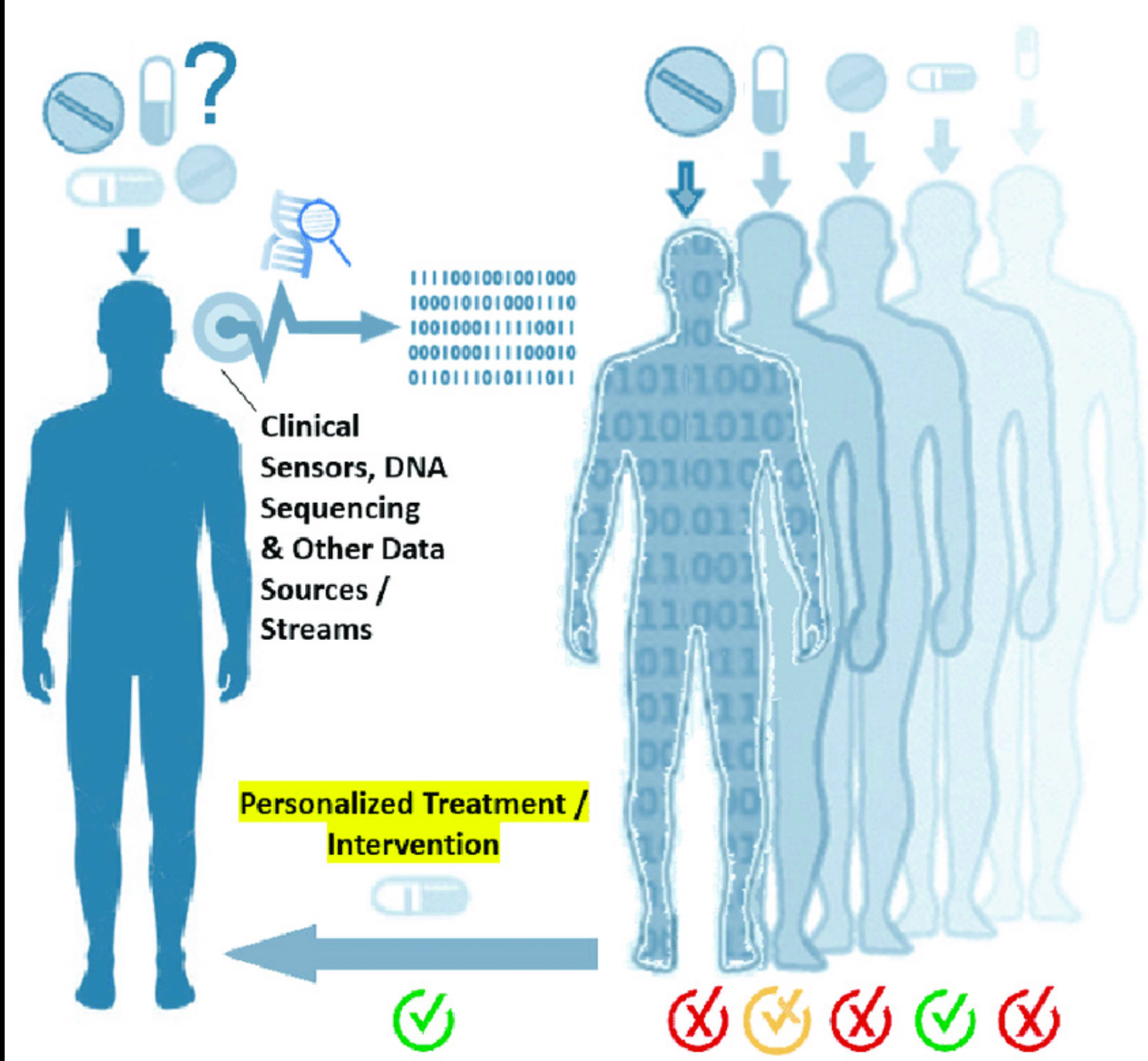


# Swedish Digital Twin Consortium

Contact Us

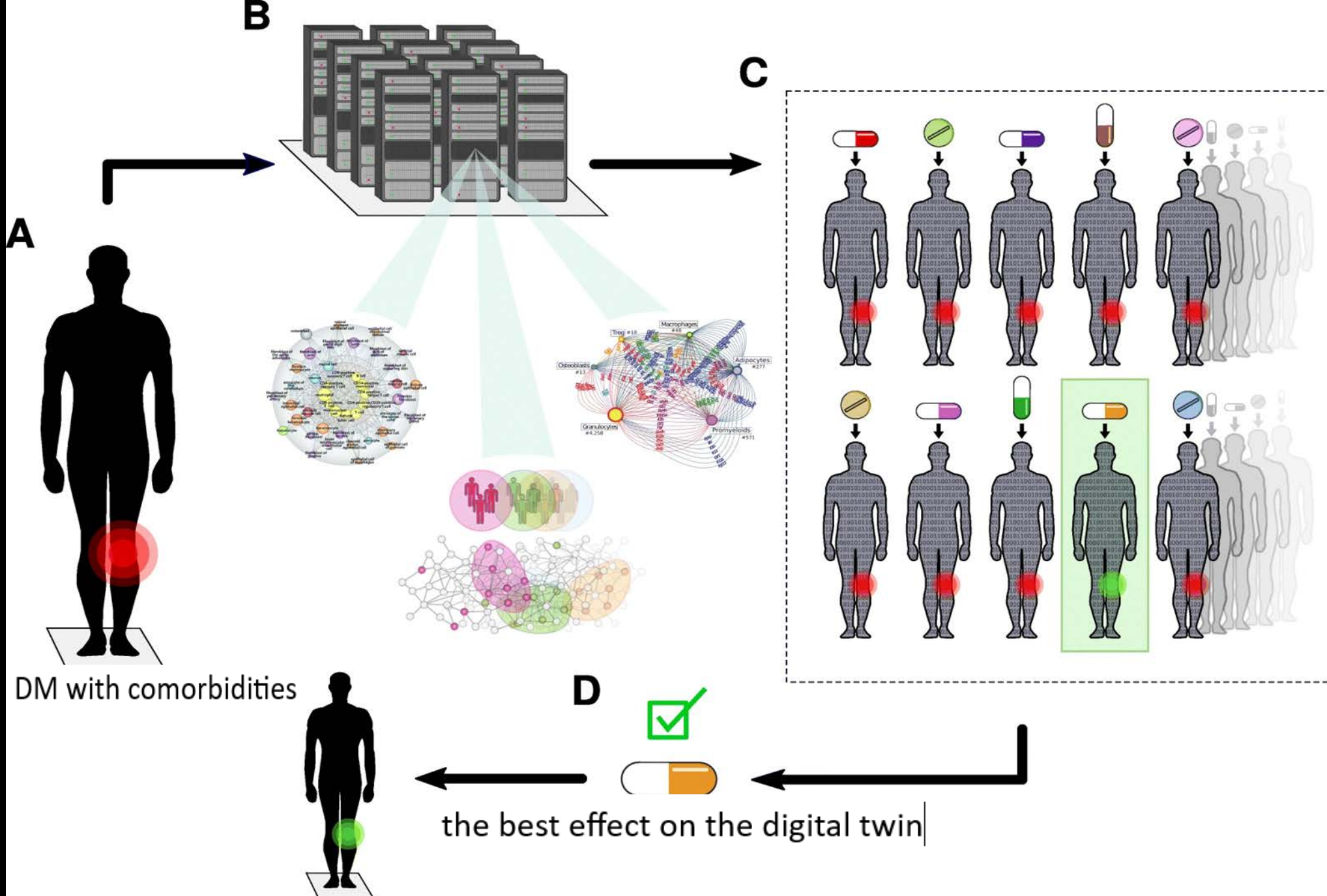
## Acquiring Data





The digital twin concept for personalized medicine.



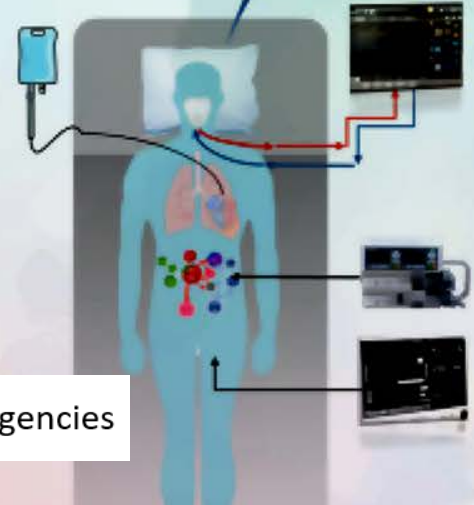


The digital twin concept for personalized medicine.

# داده‌های اندازه‌گیری شده

بیولوژی /  
فیزیولوژی / فیزیک /  
پاسخ مکانیکی

مدل‌ها / ریاضیات



Endocrine emergencies

بیمار ICU

همزاد  
دیجیتالی

Digital Twin



بازتاب / پیش‌بینی

مراقبت بهینه



پزشک

# The Endocrine Digital Twin

Body level



Organ level



Tissue level



Cellular level

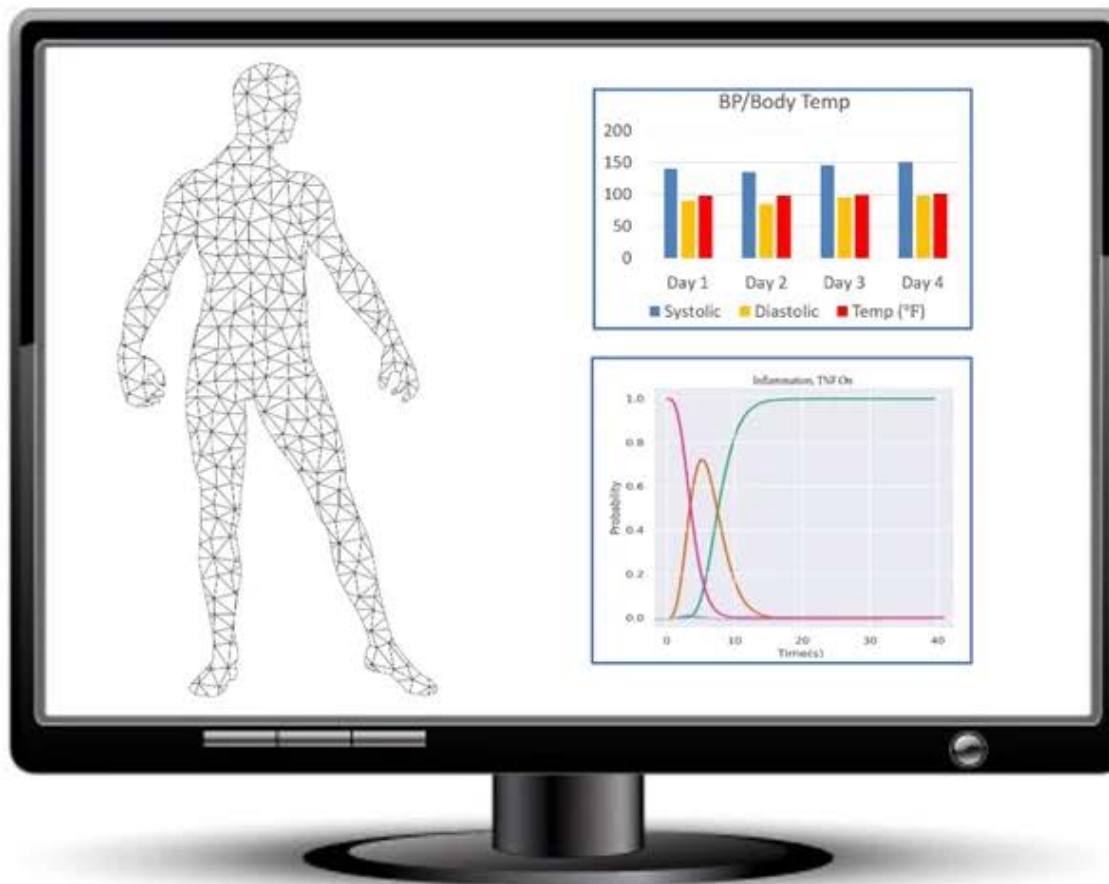


Molecular level



Data Integration & Analysis

Modelling & in silico simulations



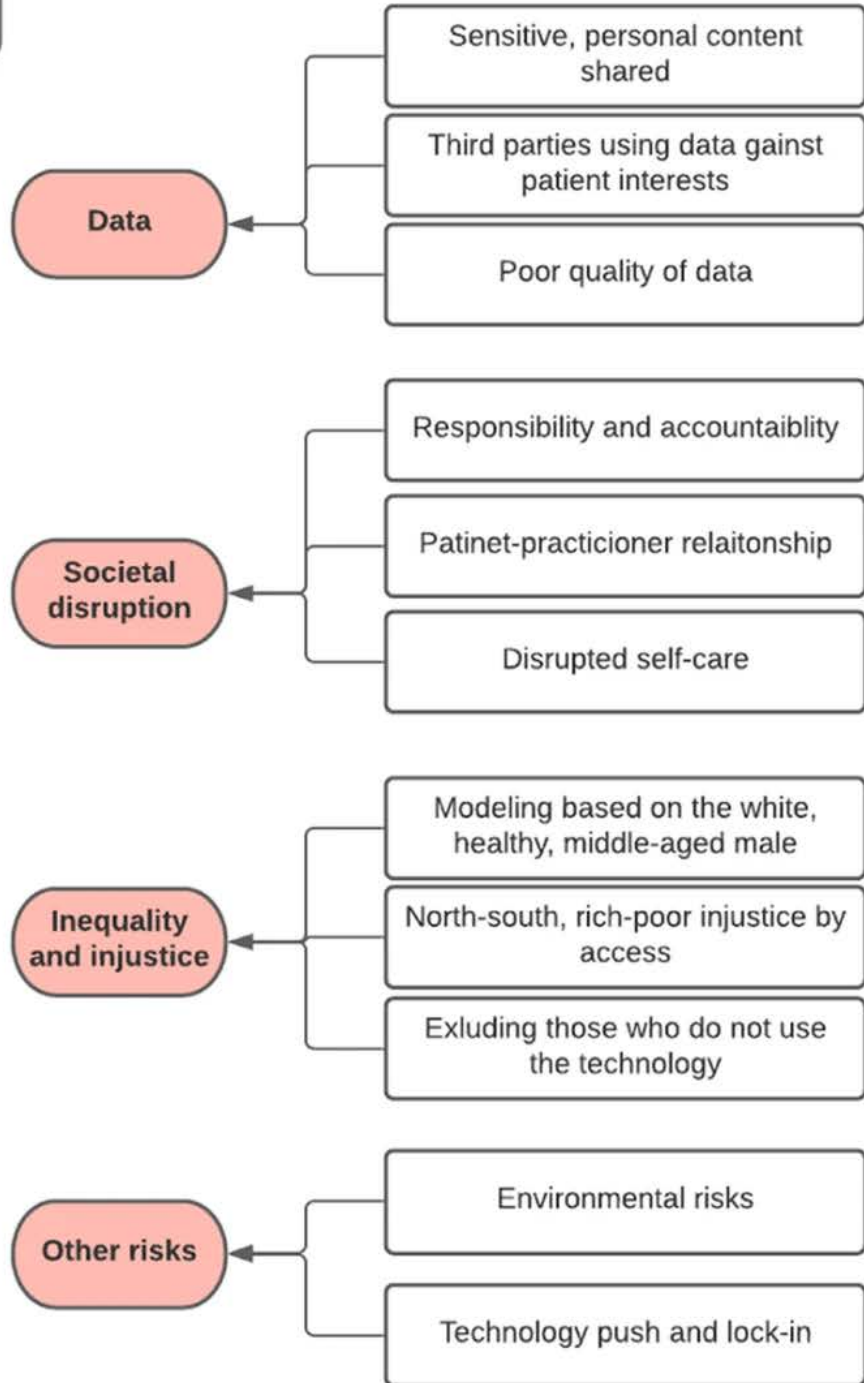
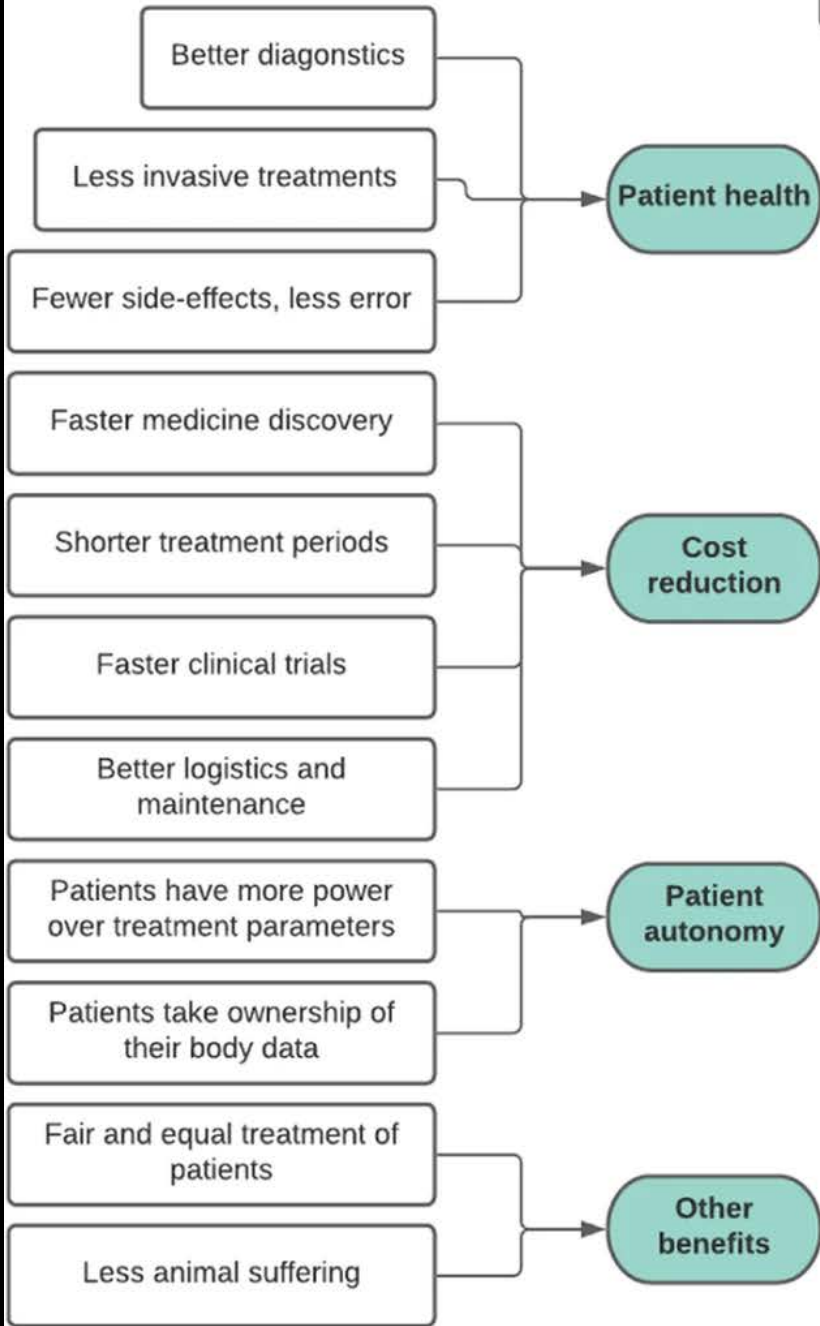
Diagnosis

Prognosis

Therapy Optimization



# Digital Twin



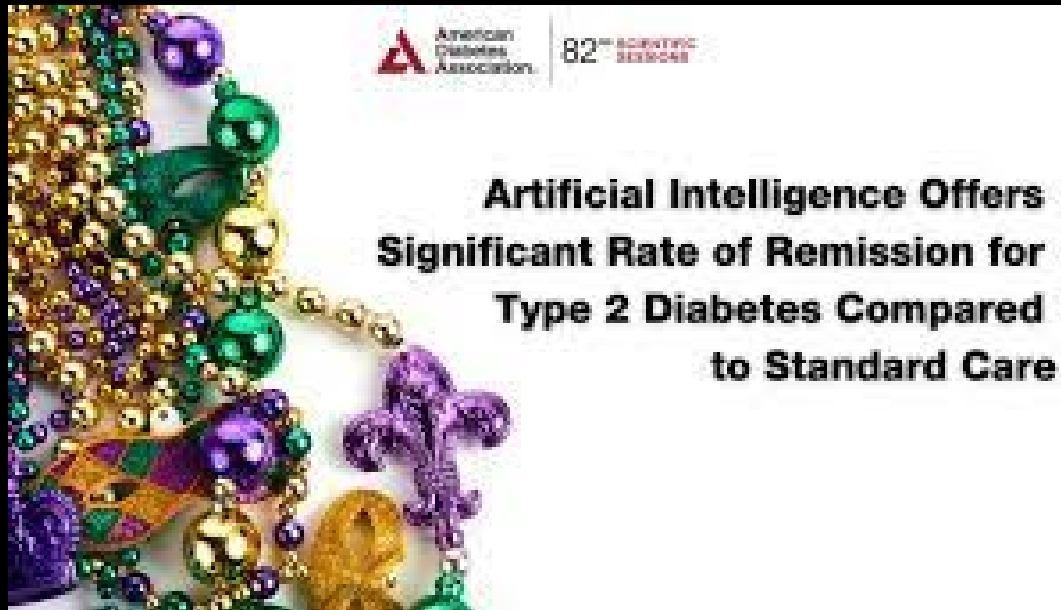
3000+ health signals / day are  
send through our body sensors



Whole Body  
Digital Twin

In return, you get 'precision treatment'  
for your body to heal its metabolism

# The Artificial Intelligence-based Whole Digital Twin for diabetes remission



- A more recent 2021 ADA consensus statement defined diabetes remission as HbA1c<6.5% at least 3 months after stopping glucose-lowering pharmacotherapy.




RESEARCH ARTICLE

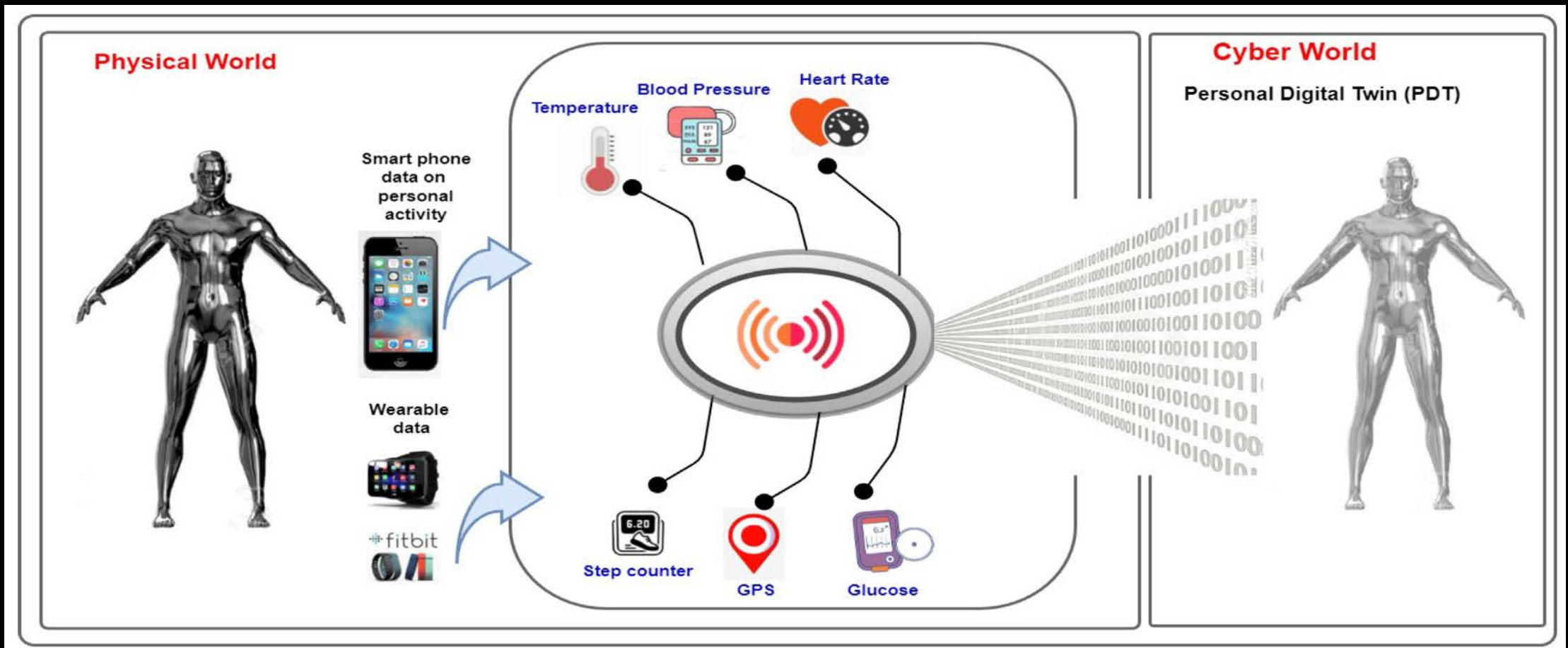
Open Access



# Type 2 diabetes reversal with digital twin technology-enabled precision nutrition and staging of reversal: a retrospective cohort study

Paramesh Shamanna<sup>1</sup>, Shashank Joshi<sup>2</sup>, Lisa Shah<sup>2</sup>, Mala Dharmalingam<sup>3</sup>, Banshi Saboo<sup>4</sup>, Jahangir Mohammed<sup>2</sup>, Maluk Mohamed<sup>2</sup>, Terrence Poon<sup>2</sup>, Nathan Kleinman<sup>5\*</sup> , Mohamed Thajudeen<sup>2</sup> and Ashok Keshavamurthy<sup>2</sup>

A randomized controlled trial of the tech was conducted on individuals with type-2 diabetes, to use the Whole Body Digital Twin platform (n = 199) or receive standard of care (n = 63) for 180 days.

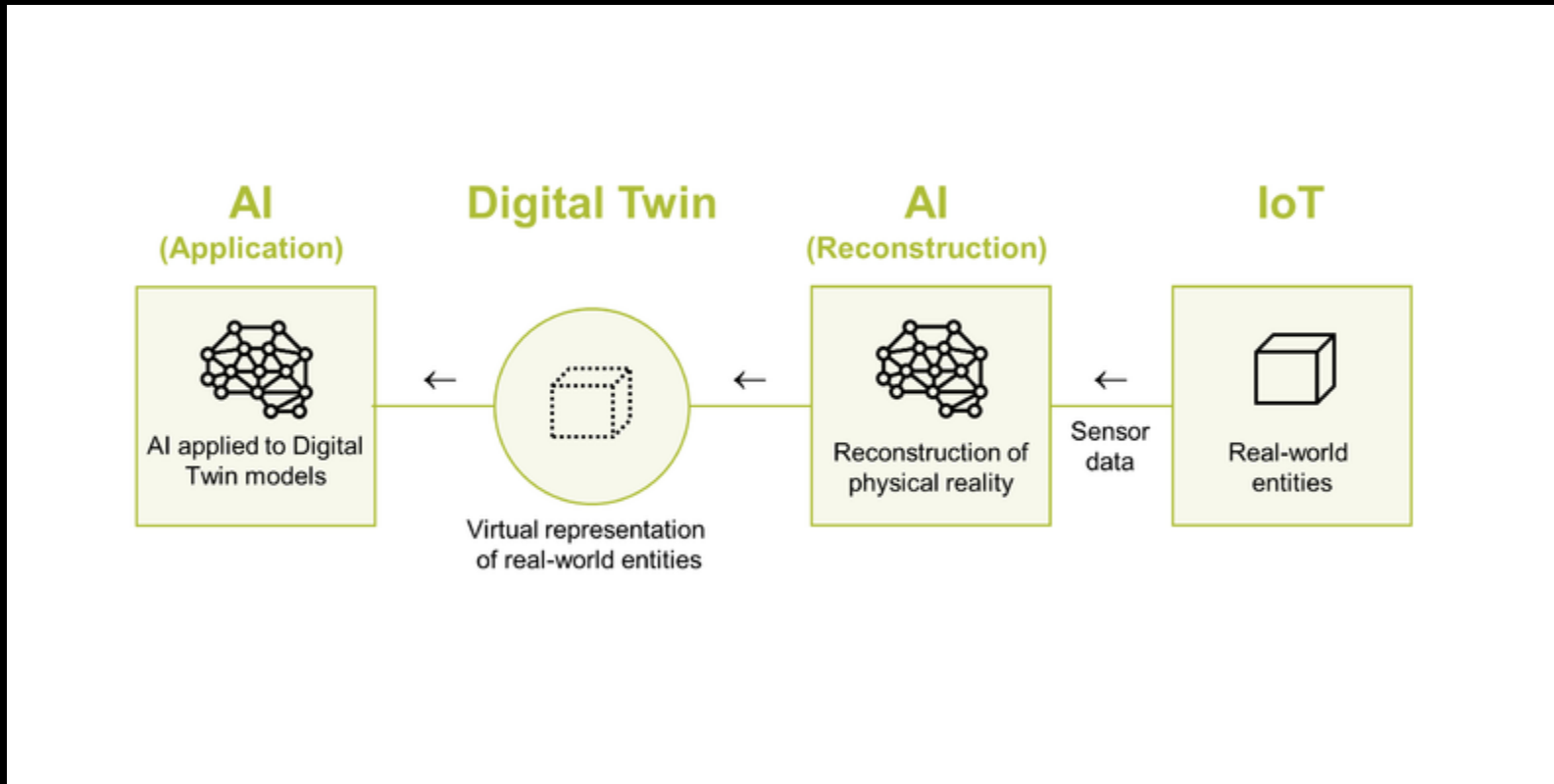


The platform collects daily data from continuous glucose monitors (CGM), sensor watches, blood pressure meters, smart scales, detailed patient food intake information, and a mobile app to track and analyze the body's health signals in order to personalize the patient's treatment and provide daily precision nutrition guidance to the patient.



The program employs a machine learning predictive model using data from food logs and multiple sensors, including continuous glucose monitoring fitness watches and comprehensive blood tests. The technology provides individualized and timely recommendations through a mobile app to users and their care teams.

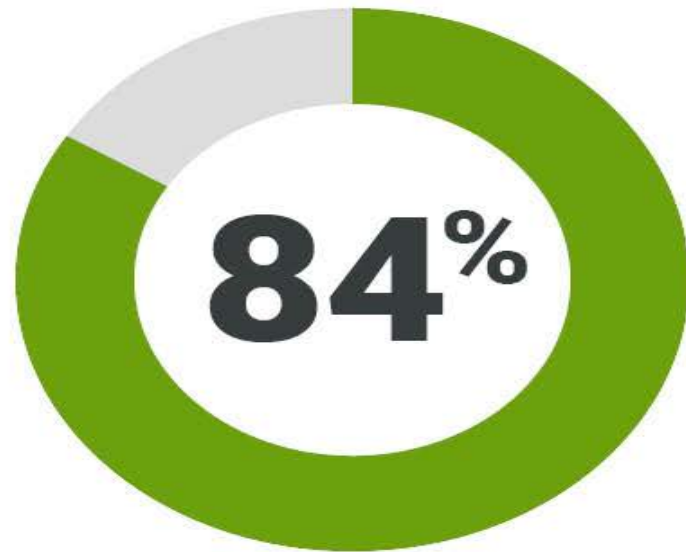




Machine learning algorithms analyzed the patient's macronutrients, micronutrients, and biota nutrients to measure and predict glucose response to specific foods. Participants were provided with specific daily food recommendations to avoid glucose spikes.

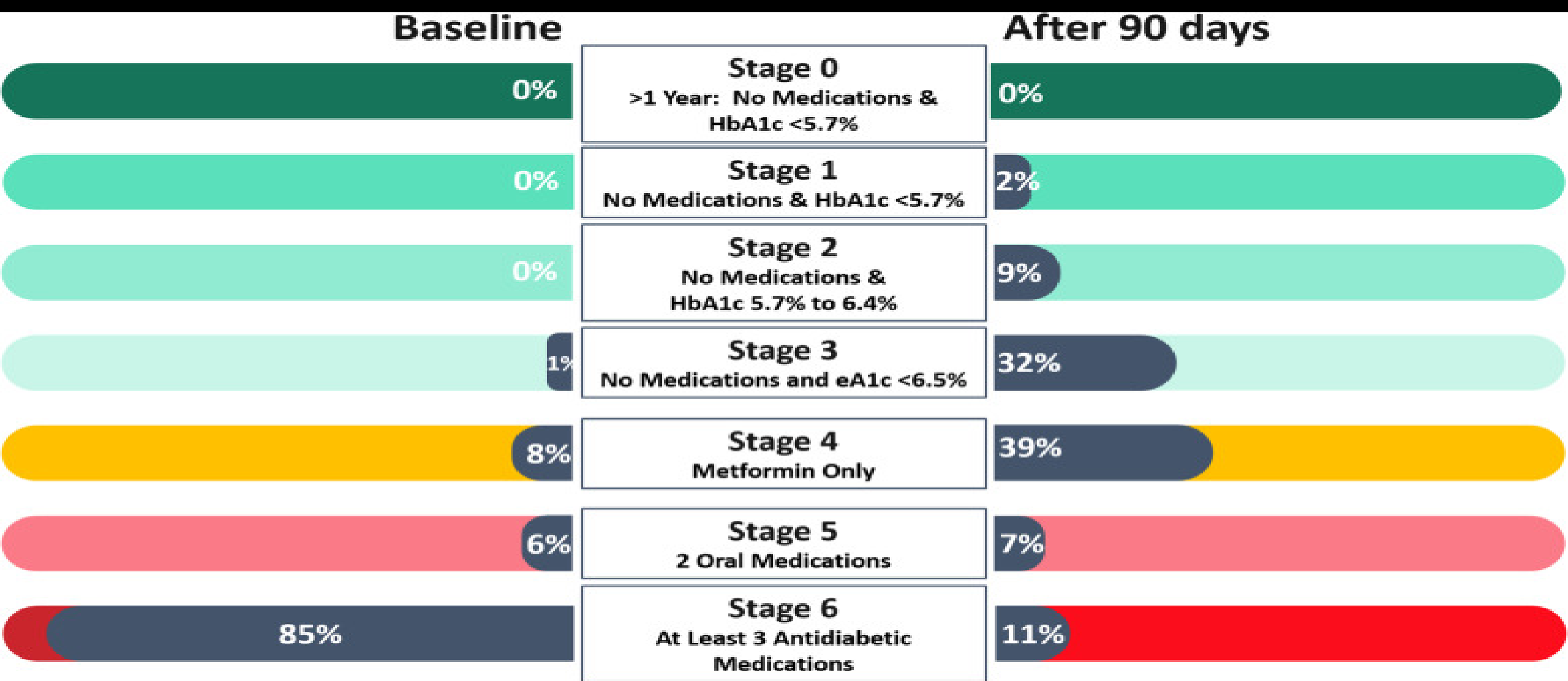
Change in HbA1c from baseline to 180 days and the percentage of participants achieving type 2 diabetes remission were the primary outcomes.

**Of adults with type 2 diabetes using Whole Body Digital Twin technology:**

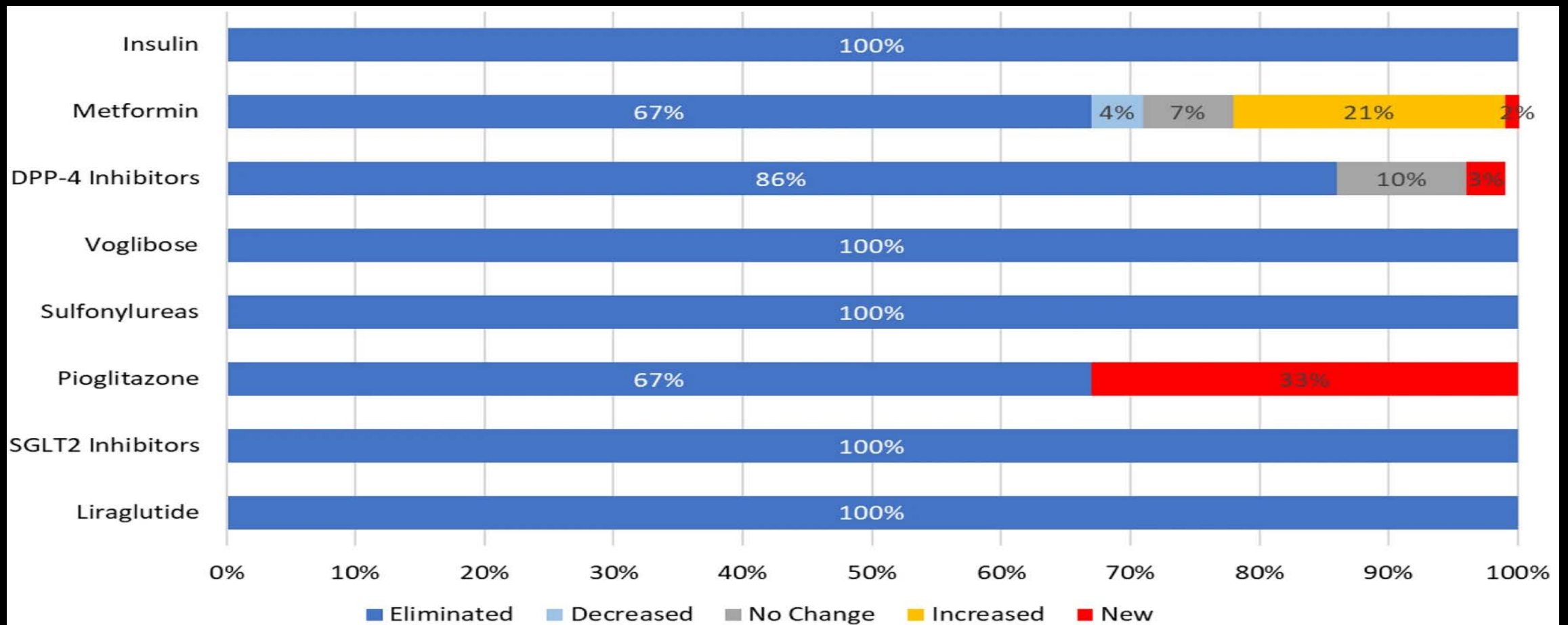


**Achieved type 2  
diabetes remission as  
defined by ADA criteria  
at 180 days**





In interim analysis, majority of participants in the intervention group reached an HbA1c <6.5% at 180 days using no medications or metformin only  
 The remission was independent of the level of weight loss.



All nine participants in the intervention group who used insulin at baseline stopped using insulin prior to 90 days.





## Reduce medication, lose weight, and recover from metabolic diseases

"In total, nearly 84% of the group who followed their digital twin's guidance for six months were determined to be in remission by the end of that period, per the ADA's standards—meaning they maintained normal blood glucose levels for at least three months without taking diabetes medication."

– **American Diabetes Association (ADA)\***

The Whole Body Digital Twin™ is a digital representation of your unique metabolism and delivers precise, personalized guidance about foods, sleep, activity, and breathing through the easy-to-use app.

Twin Health's program combines the Whole Body Digital Twin™ with a dedicated care team that monitors your sensor data, offers personalized recommendations, and supports you on your health journey.

[\\*Artificial Intelligence Offers Significant Rate of Remission for Type 2 Diabetes Compared to Standard Care. American Diabetes Association \(2022, June 4\).](#)

“We were pleased to see the magnitude of remission of diabetes achieved in our study, validating the power of understanding the distinctions of each individual’s unique metabolism to provide personalized recommendations,” Shamanna said.

# Beating metabolic diseases is now possible with your Whole Body Digital Twin™

Prediabetes | Diabetes | Obesity | PCOD | Metabolic Wellness for Preventive Care

The Twin Health Program uses advanced Whole Body Digital Twin™ technology to help heal your disrupted metabolism — the root cause of many chronic diseases.



*Shamanna :“Our results demonstrate the potential of Whole Body Digital Twin technology to change the conventional, medication-driven management of type 2 diabetes to achieving remission of type 2 diabetes with a life free of medication in a large proportion of people. Precision lifestyle changes implemented by continued inputs delivered digitally by artificial intelligence has the potential to deliver both glycemic and extra glycemic benefits.”*



YOUR BODY CAN  
TELL YOU HOW TO  
**REVERSE  
DIABETES**  
SAFELY

HERE'S THE  
TECHNOLOGY  
TO LISTEN



ASK YOUR DOCTOR ABOUT  
TWIN DIABETES REVERSAL  
TECHNOLOGY.

T&C Apply.



**REMISSION**  
of your chronic metabolism conditions  
**Like Prediabetes & Obesity**  
with **Whole Body Digital Twin™**





Heal your metabolism to help normalize your blood sugar



Reduce costly medications including insulin



Increase energy, improve mood, and decrease symptoms





[OVERVIEW](#) > [NEWS](#)

## OPTOMICS: Digital twin technology improving type-2 diabetes healthcare

The EU Horizon 2020 OPTOMICS project will develop and validate a digital-twin model, which aims at developing non-invasive and inexpensive prognostic and treatment planning methods for type-2 diabetes patients.

**date:** 18/02/2021



This so-called digital-twin model aims to improve prediction and early detection of individuals likely to develop the disease, which will improve the overall possibility for prevention. Simultaneously, the method will reveal potential risks for developing disease complications, all while personalizing patient treatment.

The project involves in-depth molecular phenotyping of individuals at the DNA, protein and metabolite level.

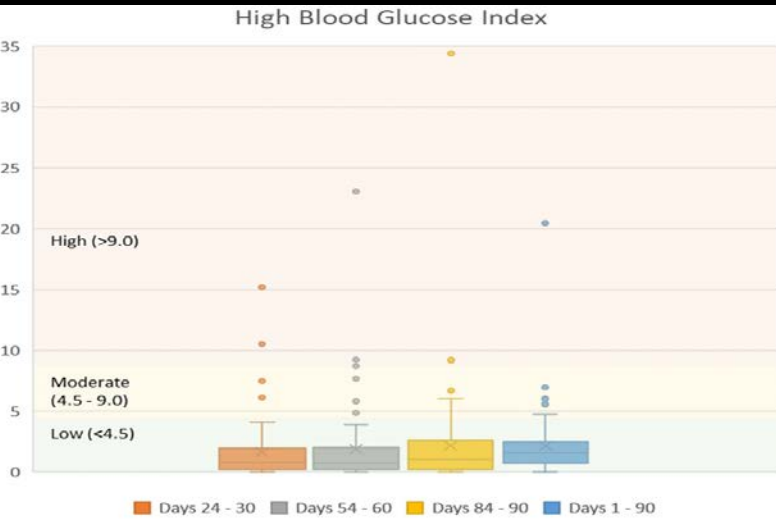


OPEN

## Retrospective study of glycemic variability, BMI, and blood pressure in diabetes patients in the Digital Twin Precision Treatment Program

Paramesh Shamanna<sup>1</sup>, Mala Dharmalingam<sup>2</sup>, Rakesh Sahay<sup>3</sup>, Jahangir Mohammed<sup>4</sup>, Maluk Mohamed<sup>4</sup>, Terrence Poon<sup>4</sup>, Nathan Kleinman<sup>5</sup>✉ & Mohamed Thajudeen<sup>4</sup>

The objective of this retrospective observational cohort study was to measure glycemic variability and reductions in body mass index (BMI), blood pressure (BP), and use of antihypertensive medications in type 2 diabetes (T2D) patients participating in the digital twin-enabled Twin Precision Treatment (TPT) Program. Study participants included 19 females and 45 males with T2D who chose to participate in the TPT Program and adhered to program protocols. Nine additional enrollees were excluded due to major program non-adherence. Enrollees were required to have adequate hepatic and renal function, no myocardial infarction, stroke, or angina  $\leq 90$  days before enrollment, and no history of ketoacidosis or major psychiatric disorders. The TPT program uses Digital Twin technology, machine learning algorithms, and precision nutrition to aid treatment of patients with T2D. Each study participant had  $\geq 3$  months of follow-up. Outcome measures included glucose percentage coefficient of variation (%CV), low blood glucose index (LBGI), high blood glucose index (HBGI), systolic and diastolic BP, number of antihypertensive medications, and BMI. Sixty-four patients participated in the program. Mean ( $\pm$  standard deviation) %CV, LBGI, and HBGI values were low ( $17.34 \pm 4.35$ ,  $1.37 \pm 1.37$ , and  $2.13 \pm 2.79$ , respectively) throughout the 90-day program. BMI decreased from  $29.23 \pm 5.83$  at baseline

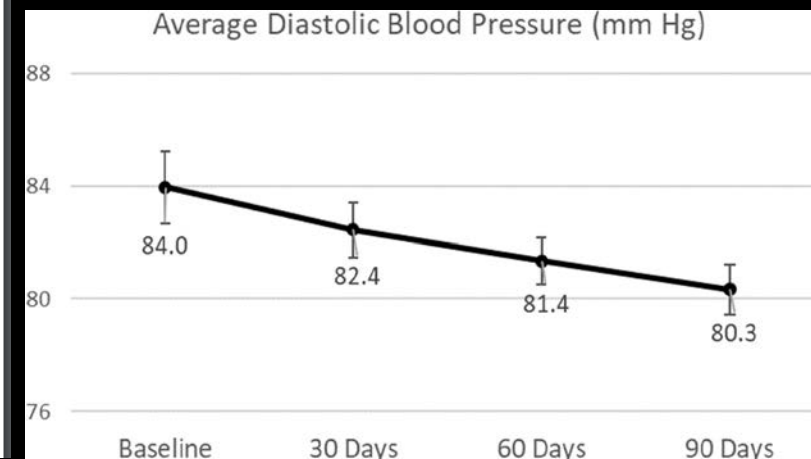
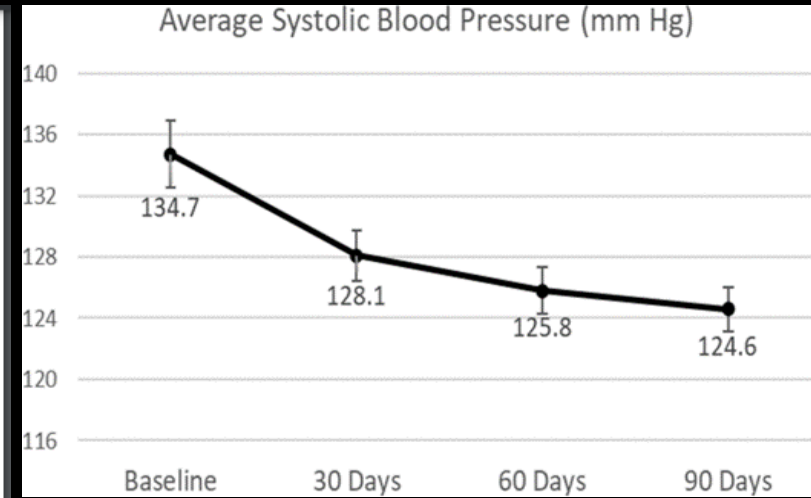


## scientific reports

### OPEN Retrospective study of glycemic variability, BMI, and blood pressure in diabetes patients in the Digital Twin Precision Treatment Program

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During 90 days of the TPT Program, patients achieved low glycemic variability and significant reductions in BMI and BP.

Antihypertensive medication use was eliminated in nearly all patients.



Article

# Human Digital Twin for Personalized Elderly Type 2 Diabetes Management

Padmapritha Thamocharan <sup>1</sup>, Seshadhri Srinivasan <sup>1,2,\*</sup>, Jothydev Kesavadev <sup>3</sup>, Gopika Krishnan <sup>3</sup>,  
Viswanathan Mohan <sup>4</sup>, Subathra Seshadhri <sup>1</sup>, Korkut Bekiroglu <sup>5</sup> and Chiara Toffanin <sup>6</sup>

<sup>1</sup> Kalasalingam Academy of Research and Education, Srivilliputhur 626126, Tamil Nadu, India

<sup>2</sup> TVS-Sensing Solutions Pvt Ltd., Madurai 625122, Tamil Nadu, India

<sup>3</sup> Jothydev's Diabetes Research Center, Trivandrum 695032, Kerala, India

<sup>4</sup> Madras Diabetes Research Foundation & Dr. Mohan's Diabetes Specialities Centre,  
Chennai 600086, Tami Nadu, India

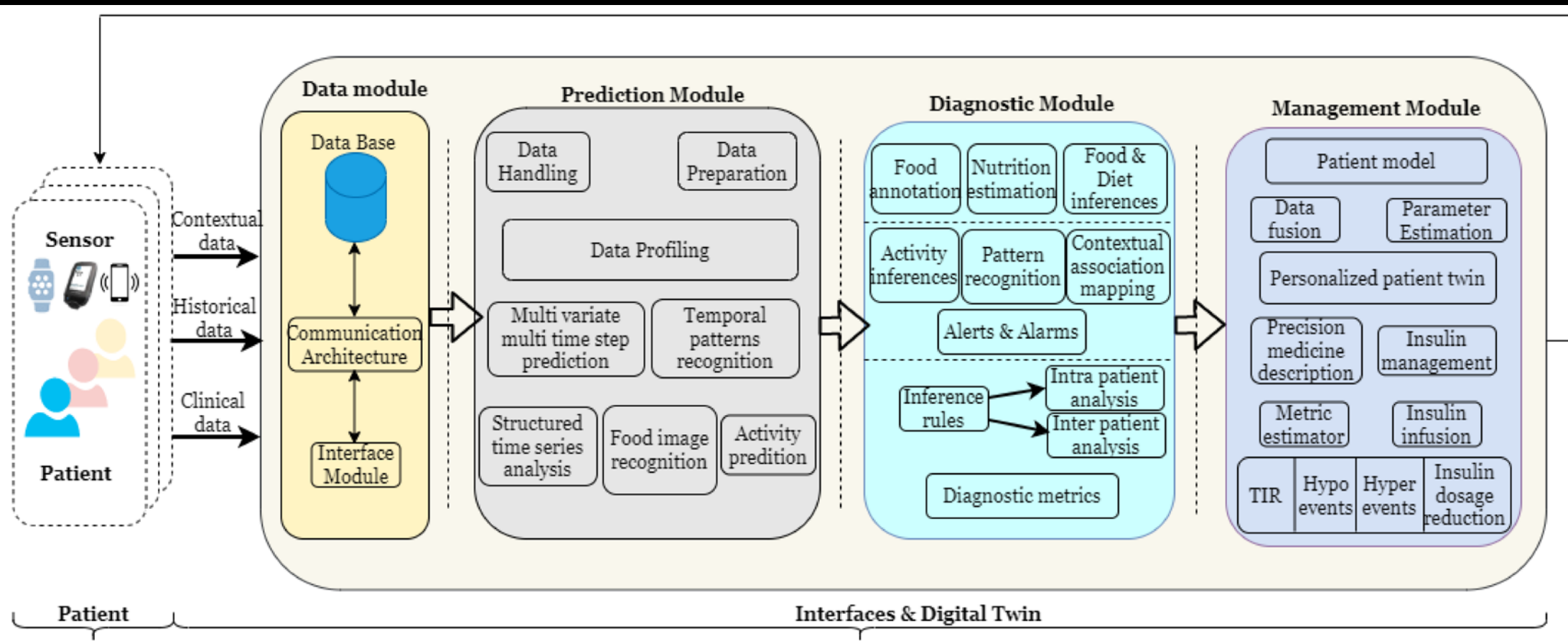
<sup>5</sup> SharkNinja, Needham, MA 02494, USA

<sup>6</sup> Department of Electrical, Computer and Biomedical Engineering, University of Pavia, 27100 Pavia, Italy

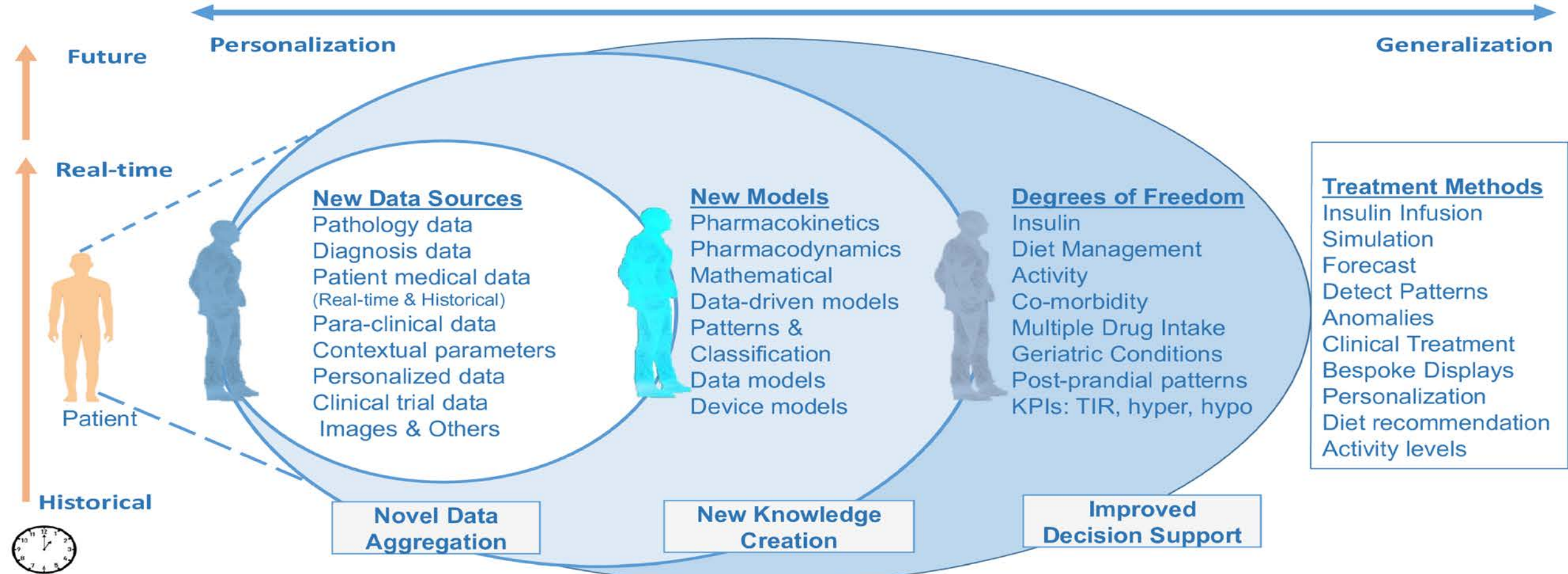
\* Correspondence: seshadhri.s@tvsss.co.in

**Abstract:** Managing Elderly type 2 diabetes (E-T2D) is challenging due to geriatric conditions (e.g., comorbidity, multiple drug intake, etc.), and personalization becomes paramount for precision medicine. This paper presents a human digital twin (HDT) framework to manage E-T2D that exploits various patient-specific data and builds a suite of models exploiting the data for prediction and management to personalize diabetes treatment in E-T2D patients. These models include mathematical and deep-learning ones that capture different patient aspects. Consequently, the HDT virtualizes the patient from different viewpoints using an HDT that mimics the patient and has interfaces to update the virtual models simultaneously from measurements. Using these models the HDT obtains deeper

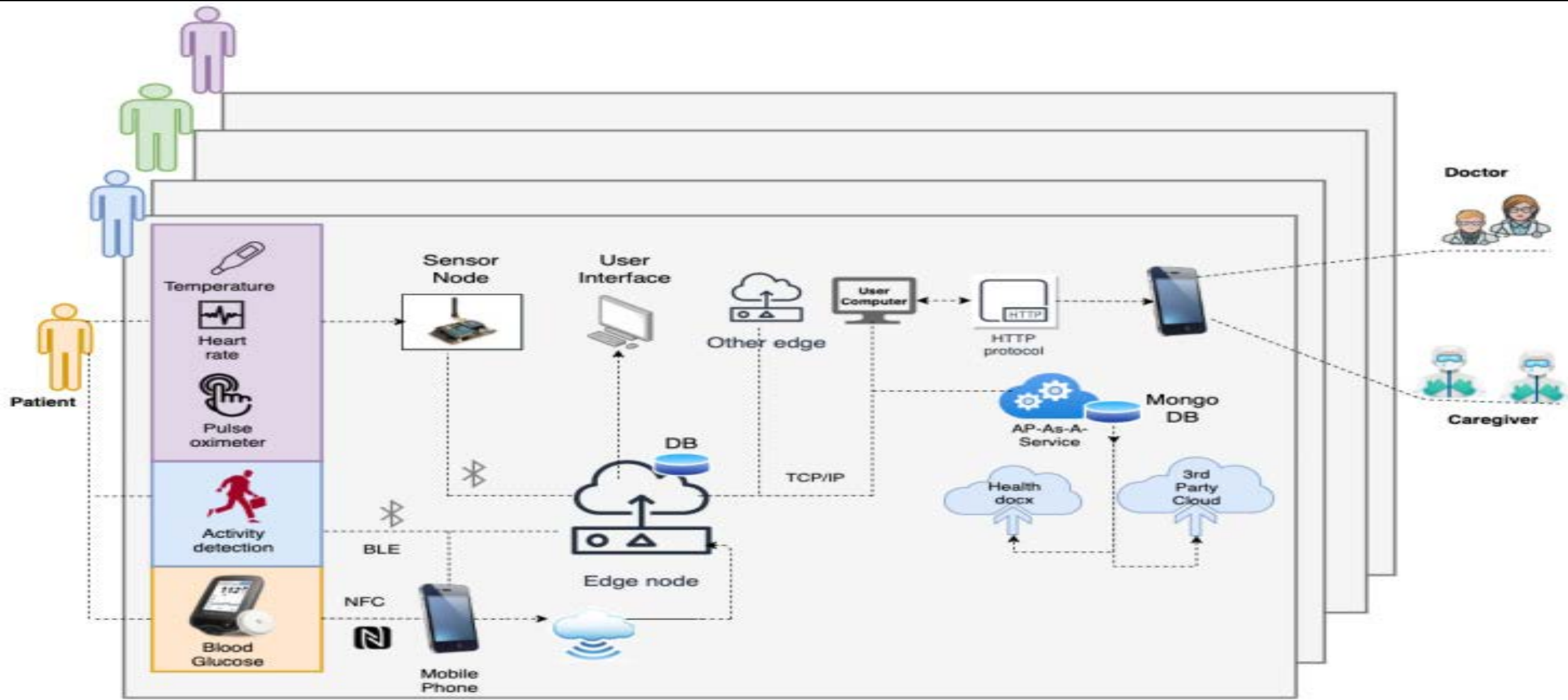




The human digital twin framework architecture



Precision Treatment Methods Through framework leads to new data, novel models that create additional knowledge, and improved decision support through new degrees of freedom.



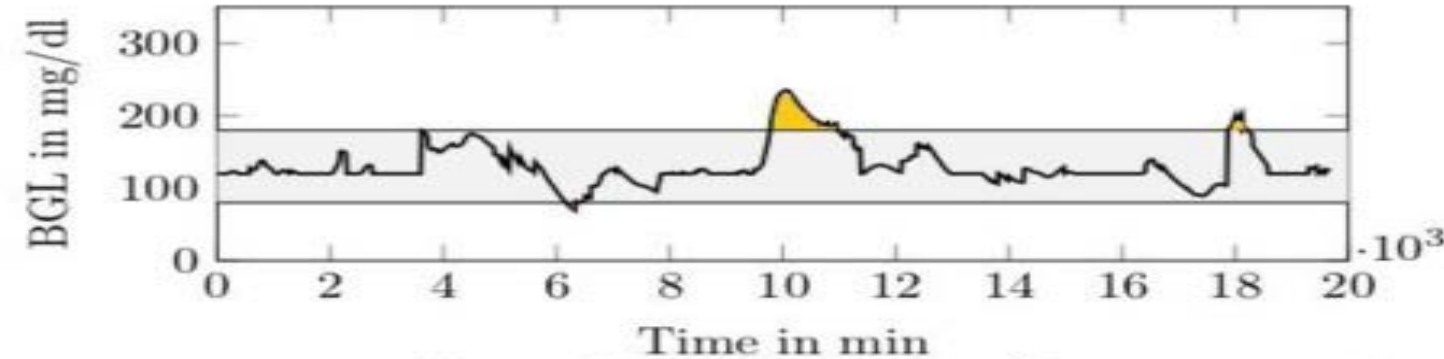
IoMT Architecture—Data transfer using IoT sensor devices from patients' (left), mounted devices to edge nodes (center) to the HDT architecture and finally to medical personnel and caretakers using a mobile app.

# Personalized Insulin Recommender



Patient Id: JDC 30569 – P1  
Status: on Insulin pump  
Diabetes: T2D

## Controlled BGL



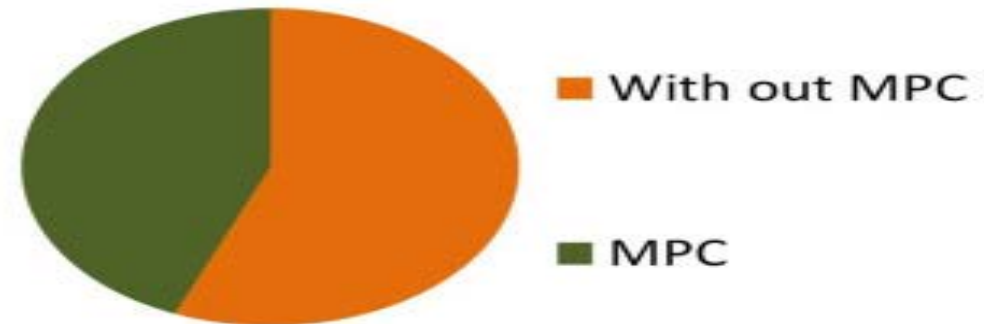
## Metrics

Avg BGL	Target Range	Time in Hyper	Time in Hypo
130 mg/dl	83 %	15 %	2 %

## Avg. CHO intake /day

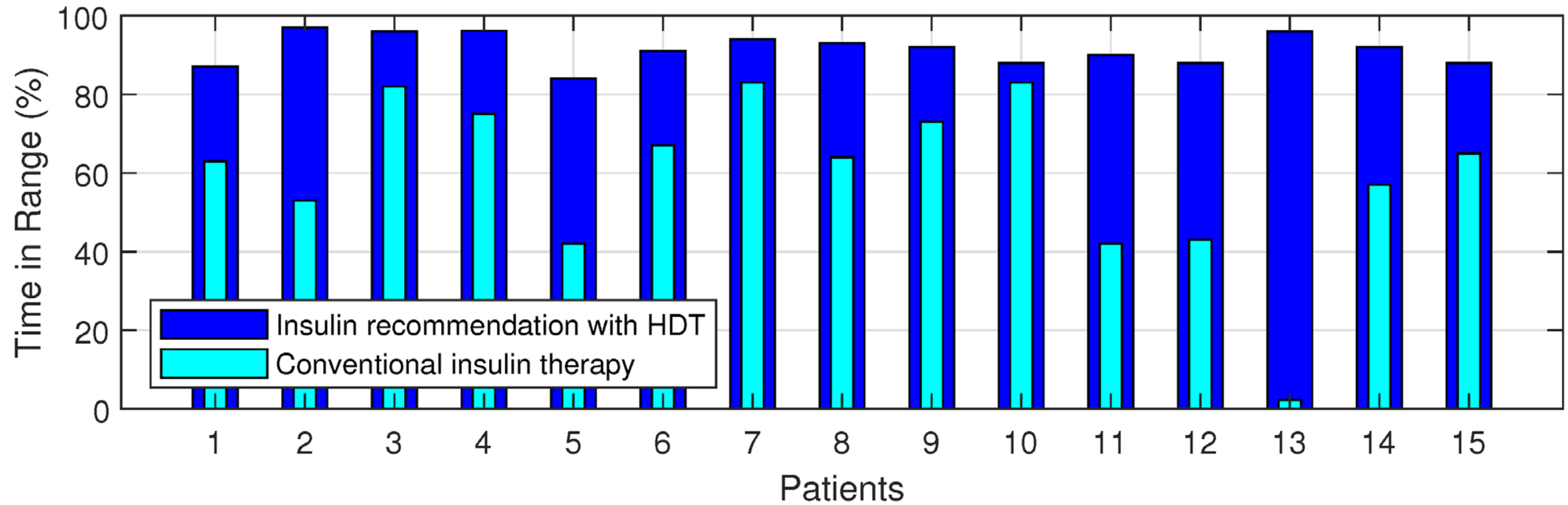


## Insulin infusion (U) 15 days

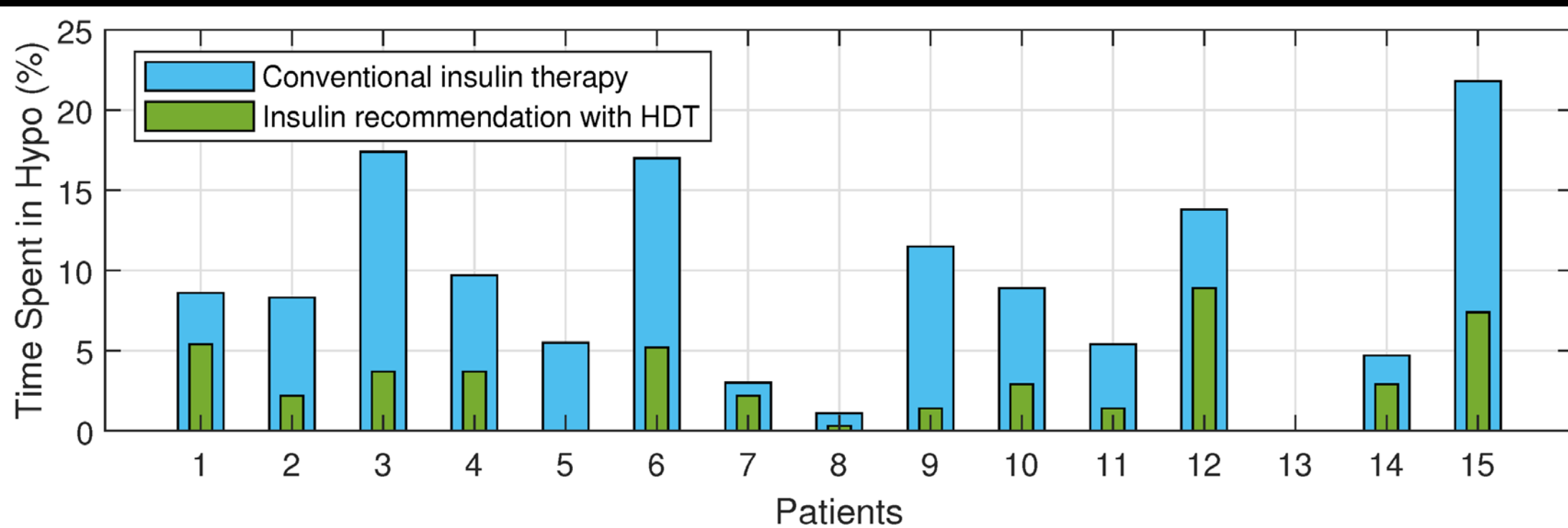


Simulated web page view of the patient's data.

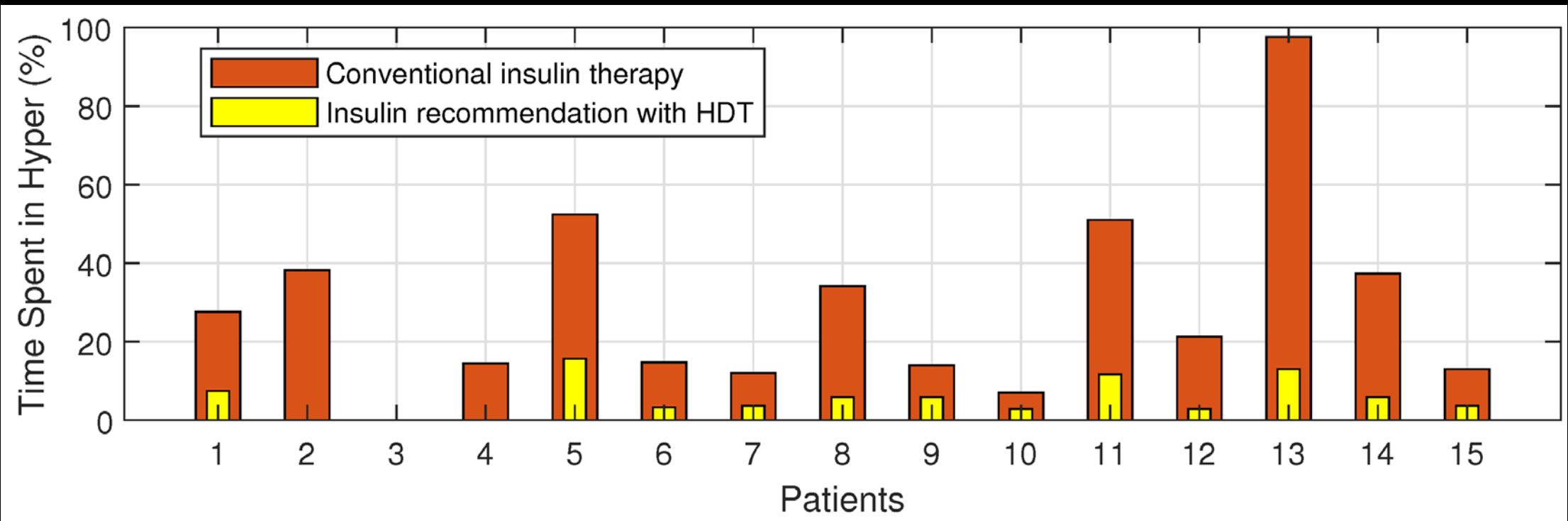




This shows that the TIR is increased from 3–75% to 86–97%.



This shows that HDT-based personalized BGL management can reduce the time spent in hypo from 0–22% to 0–9%,



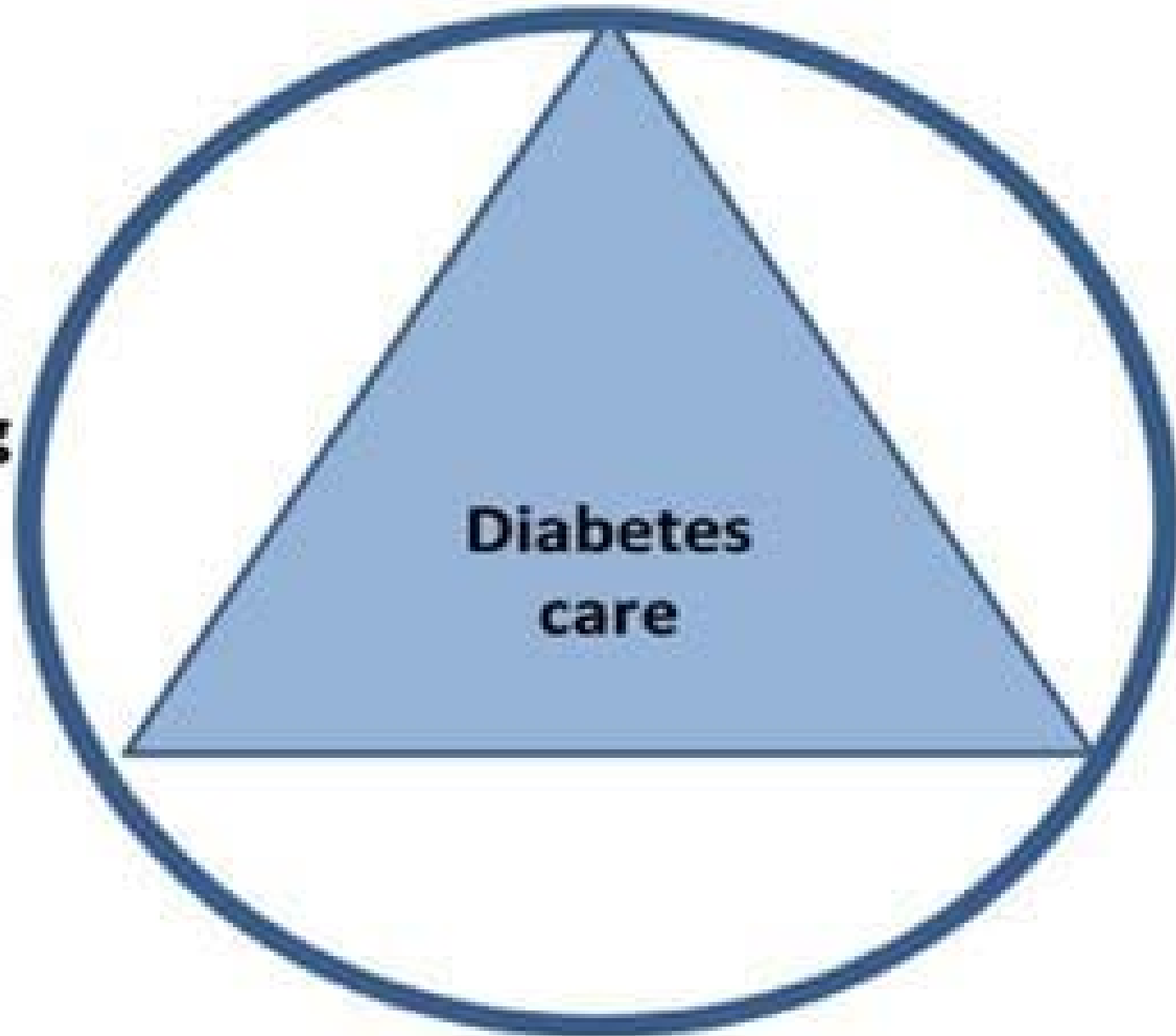
This shows that HDT-based personalized BGL management can reduce percentage of time spent in hyper from 0–98 % to 0–12%.





**Patients with  
Diabetes**

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care**

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Professional**

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**BASIC RESEARCH**

# Toward an artificial intelligence-assisted framework for reconstructing the digital twin of vertebra and predicting its fracture response

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**Abstract**

This article presents an effort toward building an artificial intelligence (AI) assisted framework, coined ReconGAN, for creating a realistic digital twin of the human vertebra and predicting the risk of vertebral fracture (VF). ReconGAN consists of a deep convolutional generative adversarial network (DCGAN), image-processing steps, and finite element (FE) based shape optimization to reconstruct the vertebra model. This DCGAN model is trained using a set of quantitative micro-computed tomography (micro-QCT) images of the trabecular bone obtained from cadaveric samples. The quality of synthetic trabecular models generated using DCGAN are verified by comparing a set of its statistical microstructural descriptors with those of the imaging data. The synthesized trabecular microstructure is then infused into the vertebra cortical shell extracted from the patient's diagnostic CT scans using an FE-based shape optimization approach to achieve a smooth transition between trabecular to cortical regions. The final geometrical model of the vertebra is converted into a high-fidelity FE model to simulate the VF response using a continuum damage model under compression and flexion loading conditions. A feasibility study is presented to demonstrate the applicability of digital twins

**Endless Electric Motion****Medical & healthcare**

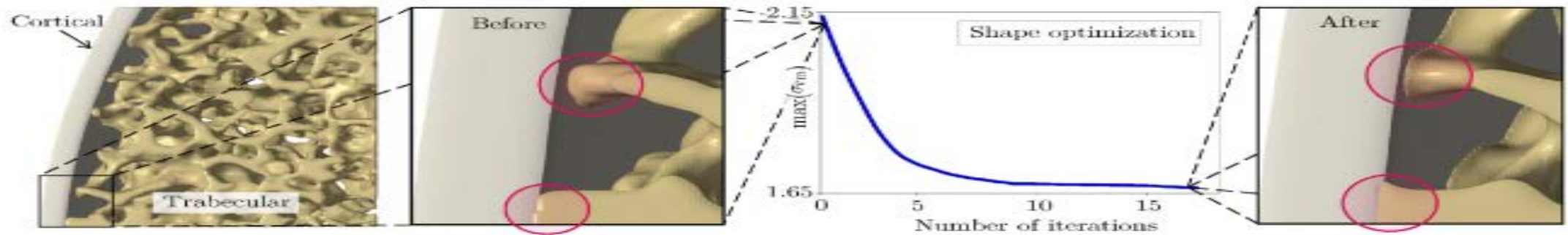
## AI-assisted digital twin could predict spinal column fractures in cancer patients

News | ⌚ 2 min read

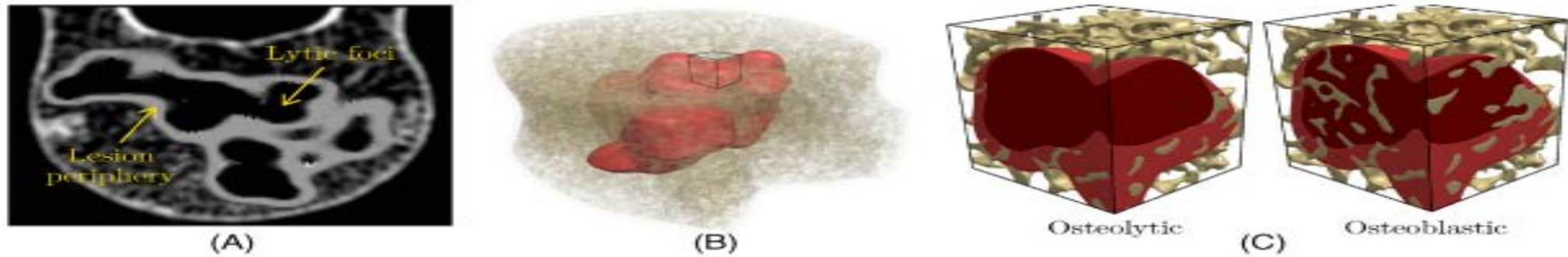
06 May 2022

An AI-assisted digital twin of human vertebrae could predict how cancer affects the probability of spinal column fractures, a study from Ohio State University has found.





**FIGURE 8** Performing finite element-based shape optimization to obtain a tapered transition from cortical to trabecular bone in the virtual vertebra model, which avoid unrealistic stress concentrations in these regions during Vertebral Compression Fracture (VCF) simulations



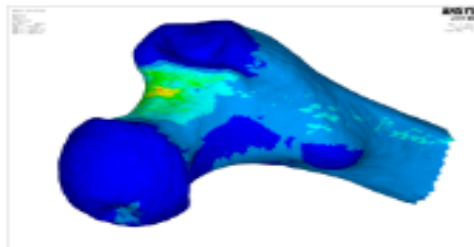
**FIGURE 9** (A) Co-registered CT-MRI scan of a vertebra with a metastatic tumor; (B) 3D reconstruction of tumor morphology added to the virtual vertebral model generated using ReconGAN; (C) comparing lytic and blastic tumors in a small portion of the trabecular bone

An AI-assisted framework, coined ReconGAN, was developed for synthesizing realistic digital twins of the human vertebra and predicting its fracture response.

ReconGAN enables integrating diagnostic imaging data (e.g., CT scans) with virtual microstructural models of the trabecular bone, to create realistic geometrical models of the whole vertebra.



- To develop an efficient digital twin solution for the estimation of long bone fracture risk in elderly patients (CT2S solution)
- To not only be able to evaluate the bone strength at present, but also predict the changes in the next 10 years (ARF10 solution)
- To extend the digital twin solution to predict other types of fracture, such as the vertebral
- To enable the running of large scale in silico clinical trials (BoneStrength solution, with a current target for bone treatments)
- To port these solutions on multiple HPC platforms across Europe and solve potential scalability issues





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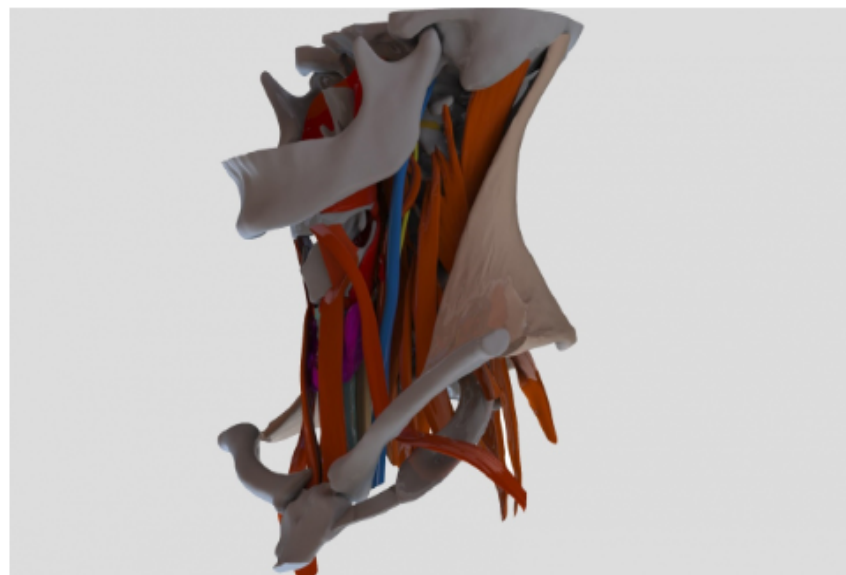
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Academic Competition "Gazprom" for school students

## LETI Scientists Developed a Digital Twin of a Neck for Effective Thyroid Surgery Planning



The model will also allow studying the effect of surgical intervention on the biomechanics of the patient's neck.

10.10.2022 290

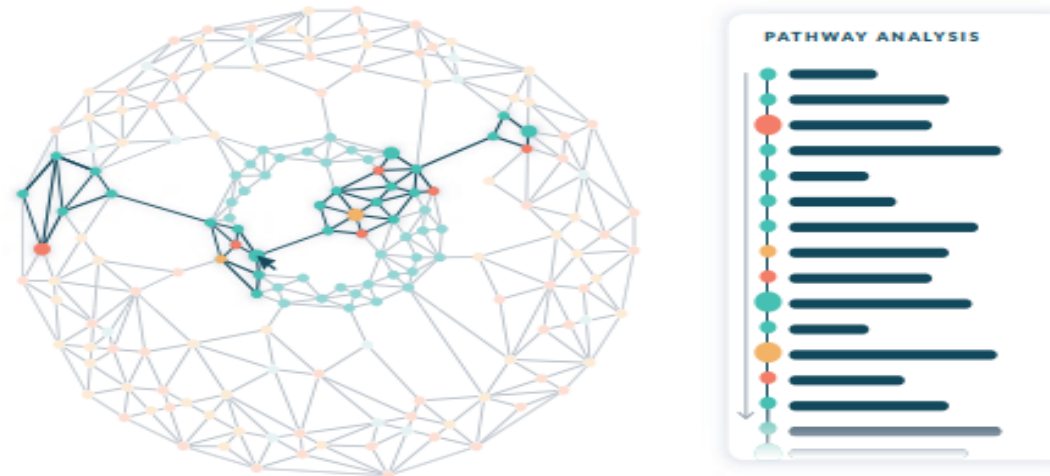
## How digital twins of human cells are accelerating drug discovery

Designed to reshape drug discovery and drug repositioning, systems biology company DeepLife has created a data-driven technology platform that creates digital twins of human cells, to evaluate response to new drug candidates.

The rapid proliferation of omics data, which provides essential information regarding biomolecular activity within cells, is transforming drug discovery. Equipped with this data, DeepLife, a next generation systems biology company, has established a platform for creating digital twins of human cells, enabling scientists to rapidly evaluate how unhealthy cells respond to drug candidates in silico. DeepLife has deployed and established proof-of-concept for its platform, and is now actively seeking partners for target identification and drug repositioning projects enabled by its digital twin technology.

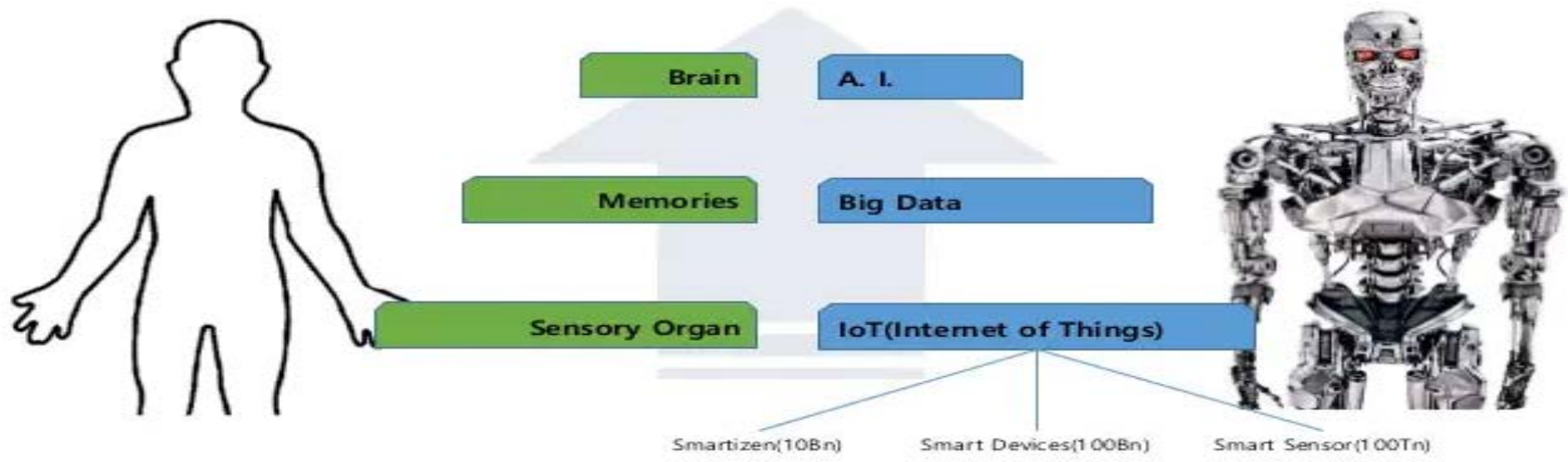
All diseases, and efforts to treat them, start at the cellular level. Small changes in the trillions of chemical interactions that make up human cells, which can be triggered by mutations or external forces, can cause cells to enter pathological states that ultimately manifest in diseases. The massive scale and complexity of the inner workings of cells has traditionally impeded efforts to identify the drivers of diseases through the iterative reconstruction of cell mechanisms, but science is now taming the challenge.

*"The convergence of recent technological*



**Fig. 1 | The DeepLife Digital Cell technology platform.** With the digital cell technology developed by DeepLife, biologists can access interpretable representation of cell mechanisms highlighting pathways and most probable druggable targets.

DeepLife has mapped more than 30 atlases with more than 20 million single. The atlases span brain, blood, liver, lung, intestines, and other tissues and organs, and DeepLife updates them monthly using more than 20 qualified data repositories.





# همزاد دیجیتالی انقلابی در پزشکی

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یکشنبه، ۰۹ مهر ۱۴۰۲

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## کتابخانه

در این صفحه شما می‌توانید به کتاب‌هایی با موضوع آینده‌پژوهشی و یا آینده‌نگاری پزشکی دسترسی داشته باشید.

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