



Medical Delta Technological solutions for tomorrow's healthcare

Medical Delta – Life Sciences, Health & Technology



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mEdical dElta we add

Colophon

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Medical Delta

Technological solutions for sustainable healthcare

The COVID-19 pandemic has underlined the urgency of achieving a healthcare transition in the coming years. It has accelerated this transition by boosting the digitalization of healthcare and created new opportunities to get the right care in the right place.

In the meantime, the average life expectancy in Europe is increasing. Due to the aging of our population and other factors, such as the increase in chronic diseases, the demand for specific care will increase in the coming years, as will the costs. At the same time, we want to live longer, in good health and condition and in a sustainable world, but the number of caregivers is declining. To keep our healthcare system sustainable, a transition is recommended. In our opinion, technology plays an indispensable role in this transition.

To push this transition forward, the regional MedTech ecosystems in The Netherlands, in collaboration with universities, companies and governmental organizations, set up the MedTechNL Agenda. This agenda highlights the ambition that the MedTech sector wants to deliver: more focus on prevention and early diagnosis, more operations with minimal trauma and more care at home. Demand-driven innovation is crucial in realizing these ambitions and thus in realizing Medical Delta's contribution to the healthcare transition. Technological innovations are key, and it is of great importance that the technology itself fits into daily healthcare practice. This is only possible if scientists from different disciplines work together. Engineers and healthcare professionals collaborate, and patients' perspectives should also be taken into account from the beginning of each innovation process.

Over the past 15 years, Medical Delta has brought together scientists, engineers, medical doctors, small and medium enterprises, large companies, municipal and provincial governments, science funders and many others to collaborate. Medical Delta has captured the best scientists in their fields in 15 different consortia that are working together on 15 interdisciplinary scientific programs. Their research takes place at locations including the BioScience Park in Leiden, the TU Delft campus and the Convergence HealthTech campus in Rotterdam. Acceleration of innovation takes place in 10 Medical Delta Living Labs, where scientists, companies, healthcare professionals and patients collaborate.

This magazine introduces the 15 Medical Delta scientific programs. Each program aims to achieve ambitious goals in the coming years. You will learn how these Medical Delta programs are helping to accelerate technological solutions for sustainable healthcare.



Prof. dr. Frank Willem Jansen Chairman Medical Delta



Medical Delta overview

Medical Delta brings together knowledge and experience from three universities, two academic medical centres and four universities of applied sciences in the province of Zuid-Holland. Top researchers from these institutions work together on technological solutions for sustainable healthcare, from molecule to reimbursement. A strong, outstanding scientific and business community, Medical Delta aims at creating and exchanging knowledge, stimulating and facilitating talent, and accelerating the valorisation and implementation of research results and innovations into healthcare practice. To ensure this, collaborations include several living labs, real-life settings in which healthcare innovators develop and test new ideas together with end-users.

Imaging and Big Data for Life

Medical Delta Diagnostics 3.0: Dementia and Stroke Medical Delta Cancer Diagnostics 3.0: Big Data Science of in & ex vivo Imaging Medical Delta UltraHB: Ultrafast and Ultrasound for the Heart and Brain

Medical Instruments for Diagnosis and Treatment

Medical Delta NIMIT: Novel Instruments for Minimally Invasive Techniques Medical Delta Cardiac Arrhytmia Lab Medical NeuroDelta

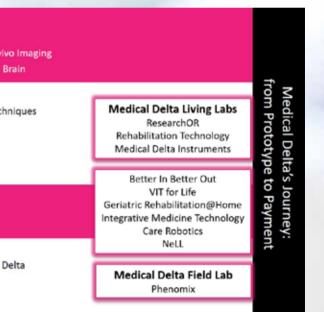
Medical Delta IMT: Improving Mobility with Technology Medical Delta Institute of Fetal and Neonatal Care

eHealth & self-management

Medical Delta eHealth & selfmanagement for a healthy society Vital Delta: Medical Delta's journey towards vitality and health

Personalised, precision and regenerative medicine

METABODELTA: Metabolomics for clinical advances in the Medical Delta HollandPTC Medical Delta program on HTA value proposition Medical Delta Regenerative Medicine 4D Medical Delta AI for Computational Life Sciences



Medical Delta Diagnostics 3.0: Dementia & Stroke

Dementia and stroke: how deep learning algorithms make faster diagnoses





Prof. dr. Wiro Niessen Biomedical Image Processing (EMC/TUD) Imaging Physics/Radiology (TUD)

Dr. Frans Vos

Prof. dr. Mark van Buchem Radiology (LUMC)

And the second second second

Dementia and stroke impose an enormous burden on individuals and society. To address this challenge, tools are required to identify individuals at risk - ideally very early in the disease process, to support the trend towards prevention.

> Image-based techniques are among the most promising of the technologies available to improve diagnosis, prognosis and treatment selection. In particular, the application of imaging technology combined with the use of advanced data analytics, such as deep learning, will be helpful. This way, Stroke imaging biomarkers will be combined with other biomarkers and clinical data. Initially, dementia and stroke will be addressed in the Medical Delta

Dementia

Diagnostics 3.0 program.

Dementia is a brain disorder characterized by loss of cognitive function such as thinking and remembering, which affects people's daily functioning. Using MRI imaging techniques and advanced analysis techniques, researchers want to understand better how dementia, such as in Alzheimer's disease, develops. Using images and data from large population studies and artificial intelligence

techniques such as deep learning, they can estimate if a person is at high risk of developing a disease. More accurate and faster diagnosis and prognosis can improve disease treatment and support steps towards disease prevention.

A stroke occurs when the blood flow to parts of the brain is cut off. This can happen due to a blood clot that blocks the blood flow or due to bleeding in the brain. These two different types of stroke require different treatments, and the right treatment must be selected.

Accurate and objective imaging systems can support treatment selection in the acute stage of stroke by determining which type of stroke the patient has dealt with, thus optimizing the treatment selection process.

Future perspective

In the future, a scan of a patient will not only be

inspected and interpreted by the doctor quantitatively; clinical decision making will be supported qualitatively based on deep learning algorithms that have been trained on previous cases.

In this way, doctors can detect a disease in an early phase, make a diagnosis faster, determine the best possible treatment more precisely and monitor the disease over a longer time.

"In this program we aim to learn optimally from imaging and healthcare data from previous patients, to treat the next generation of patients more effectively", says Wiro Niessen, professor in Biomedical Image Analysis at Erasmus MC and TU Delft.

Collaboration

This program brings together experts in different fields, such as imaging physicists, medical image analysis experts and clinicians, with the joint aim of developing, implementing and validating imaging

concludes Niessen.



techniques to improve patient care. The research relies on data and clinical expertise available through collaborations with population and clinical studies. "Through the collaboration across the different disciplines of the Medical Delta institutes, we create a whole which is greater than the sum of its parts,"

Using images and data researchers can estimate if a person is at high risk of developing dementia

Medical Delta Cancer Diagnostics 3.0: Big Data Science of in & ex vivo Imaging

Improving cancer diagnostics with new imaging techniques and machine learning

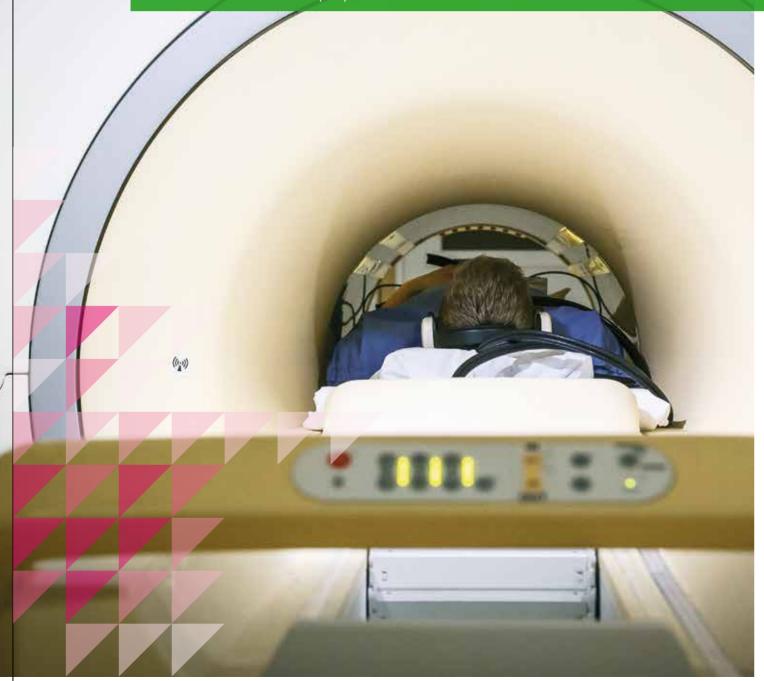






Prof. dr. ir. Matthias van Osch Radiology (LUMC) Prof. dr. Marion Smits Radiology and Nuclear Medicine (EMC)

Prof. dr. Sjoerd Stallinga Imaging Physics (TUD)



Different patients with the same type of cancer can respond very differently to a specific treatment, and as a result, outcomes vary greatly between patients. Many clinical, pathological, and genetic factors make the diagnosis and choosing the right treatment for an individual patient increasingly complex.

In Medical Delta Cancer Diagnostics 3.0, the newest imaging techniques will be used together with machine learning to provide diagnosis faster and to better predict the course of the disease.

Creating non-invasive methods

At the moment, biopsies are taken to obtain information about the genetic and molecular characteristics of the tumor. Taking a biopsy is an invasive method that can sometimes be risky. Researchers from the Medical Delta Cancer Diagnostics 3.0 program believe analyzing tumors could be improved by extracting more information from MRI images.

Advanced MRI techniques

Within this program, which initially focusses on brain tumors, advanced MRI techniques will be used to gather relevant information about the tumor. The researchers will do this using standard MRI scans, as well as the newest MRI techniques and hardware. "For example, we will use a 7 Tesla MRI, which has a higher magnetic field and provides images with a better resolution and more information," says Thijs van Osch, professor in the radiology department at LUMC.

The aim of comparing information gathered through biopsies and information visible on the MRI scans is to deduce the genetic and molecular characteristics of the tumor from the MRI-data: a 'virtual biopsy'.

Machine learning

In addition to obtaining better MRI images, this Medical Delta research program links them to machine learning to unlock all the information hidden in the different MRI-contrasts. Ultimately, this is expected to contribute to an even better treatment choice tailored to the individual patient, and to enable better monitoring of the tumor during treatment.

"It would be of enormous added value if we were able to better predict on the basis of all those MRI characteristics how the tumor will evolve and what that means for the patient in the Erasmus MC Zafm Leiden Univer





future," says Johan Koekkoek, neuro-oncologist in LUMC and MC Haaglanden.

For doctors it would very helpful to get guidance in making the right diagnosis and selecting the best treatment. "Currently, diagnosis is based on a visual assessment of the MRI scan by a radiologist," says Marion Smits, professor of Neuroradiology at Erasmus MC. "Machine learning techniques will help us to include a lot more information from the MRI scan in our assessment, and to make such diagnoses more objective."

Machine learning solutions are also in development in the field of digital pathology of biopsy material. Sjoerd Stallinga, professor of Computational Imaging at TU Delft: "Analysis and classification of tissue morphology is a very suitable task for modern deep learning methods."

Machine learning techniques will help us to include more information from the MRI scan and to make diagnoses more objective

Collaboration

The close collaboration that has been established between Erasmus MC, TU Delft and LUMC with MRI experts, computational scientists, engineers and clinicians is a key success factor for realizing the ambition to enable biopsy-free image-based diagnosis of cancers. The support to provide a personalized treatment selection and treatment monitoring is envisioned as a template for similar Medical Delta innovations in the care of other cancer types.

Medical Delta ultraHB: Ultrafast and Ultrasound for the Heart and Brain Ultrafast ultrasound for the heart and brain







Dr. David Maresca **Imaging Physics (TUD)**

Dr. Annemien van den Bosch Cardiology (EMC)

Prof. dr. Aad van der Lugt Radiology & Nuclear Medicine (EMC)

Renewing ultrasound techniques will open doors for a whole range of diagnostic and therapeutic possibilities in the cardiovascular and neurological field. The Medical Delta program Ultrafast Ultrasound for the Heart and Brain offers excellent opportunities to tackle the major existing and evolving healthcare threats.

> In the Netherlands, it is expected that 195,000 people will be diagnosed with heart failure in 2025, leading to a high number of hospitalizations and deaths. Atherosclerosis is a disease in which plaque builds up inside arteries. If plaque ruptures in an artery linked to the brain, this can lead to stroke. In neurology, brain tumors are also a large threat. In the Netherlands alone, more than 1,000 patients were diagnosed with a brain tumor in 2018.

The program focuses on three healthcare threats: heart failure, atherosclerosis and neurological disorders.

Ultrasound

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Ultrasound is the most commonly used medical imaging technique. It is harmless, relatively cheap and, because it uses real-life images, it gives immediate diagnostic feedback. The technology is developing rapidly: new ultrafast imaging brings the functioning of the heart and brain directly into view,

and special contrast agents make new diagnoses and treatments possible.

Technological developments

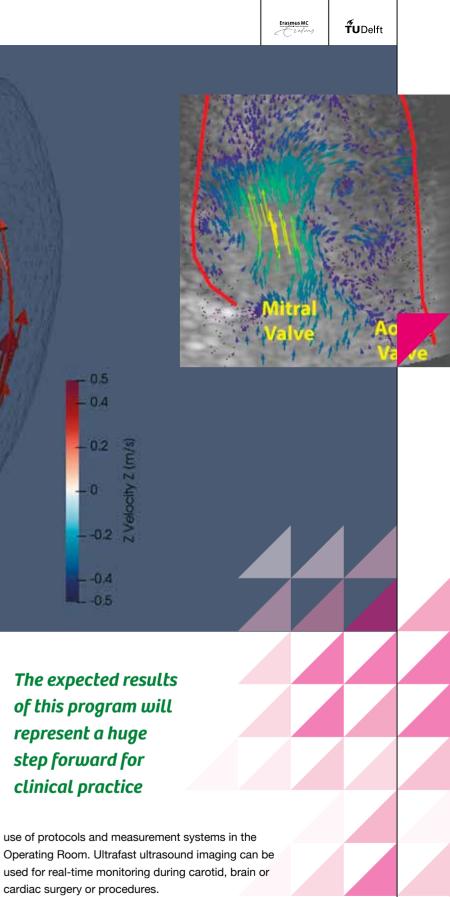
Three technological developments are expected to have an impact in this field. The first is ultrafast ultrasound imaging, involving thousands of images per second. This makes the blood flow in the heart and through organs and even the activation of brain parts directly visible ("functional ultrasound"). The second focuses on the realization of new sensors, making ultrasound into a truly 3D modality. At the moment, almost all applications show a 2D visualization: this creates a limitation for treatment. In this program, 3D image reconstruction will be used, to improve diagnosis and treatment options in healthcare.

The third is the development of ultrasound contrast agents that target disease-specific markers and can transport drugs with them.

This program will offer diagnostic methods with excellent opportunities to tackle the major existing and evolving healthcare threats, such as heart failure, atherosclerosis and neurological disorders. The technological breakthroughs will also be directly clinically tested in the cardiology and neuroradiology departments at Erasmus MC. The expected results will represent a huge step forward for clinical practice, where doctors eagerly await the new possibilities these techniques offer.

Collaboration

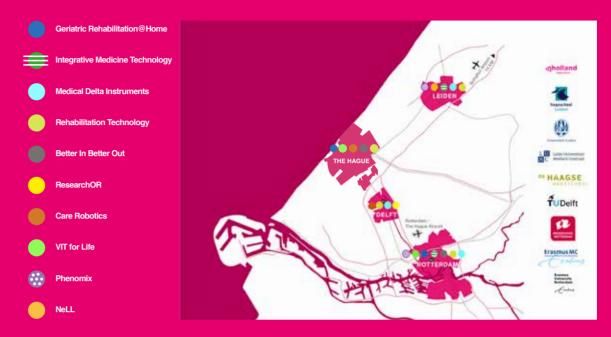
Erasmus MC, TU Delft and LUMC are involved in this program, which is also very well linked to two Medical Delta living labs. The first is Medical Delta Instruments, focuses on the use of minimally invasive instruments. Ultrafast ultrasound imaging can be used during minimally invasive procedures for realtime tissue characterization - this is important, for example, in surgical procedures. The second is Medical Delta ResearchOR, which focusses on the



Medical Delta Field and Living Labs: between prototype and practice

The Medical Delta Living Labs form a crucial link in the healthcare innovation chain. They test companies' and healthcare institutions' promising technological healthcare solutions with healthcare professionals and patients in real-life environments. Practical questions are the foundation for the public-private projects the Living Labs take on. The interdisciplinary nature of the work is characteristic of the Living Labs' approach: health lecturers collaborate with technology lecturers from other universities of applied sciences. They conduct their research together with end users, healthcare institutions and companies, and through this collaboration, the Living Labs have a social and economic impact on the region. The Living Labs also provide a bridge to research carried out at Medical Delta's academic knowledge institutions.

Locations Field- and Living Labs





Medical Delta Living Lab VIT for Life Paying more attention to health and prevention will reduce the pressure on our healthcare system. Health apps and innovations that stimulate a healthy lifestyle and exercise do not reach all people with an increased risk of lifestyle-related complaints. Medical Delta Living Lab VIT for Life focuses on these groups and aims to make inaccessible apps or complicated technology user-friendly. In addition, it integrates these technologies into the broader range of interventions provided by doctors and physiotherapists.

At Medical Delta Living Lab VIT for Life, Rotterdam University of Applied Sciences, The Hague University of Applied Sciences, Stichting WMO Radar and Kinderfysiotherapie Regio Westland work together with patients and companies. The Living Lab also collaborates with universities affiliated with Medical Delta. Medical De Better Out The fitter some program, the hi Medical Delta L on researching and technologie patients. At Medical Delt Hague Universi University of Ap and the Universi Hague work tog professionals a for healthcare. basis with resea institutions, incl



Medical Delta Living Lab Better In Better Out

The fitter someone enters a cancer treatment program, the higher their chance of a faster recovery. Medical Delta Living Lab Better In Better Out focuses on researching and developing e-health applications and technologies that improve the fitness of cancer

At Medical Delta Living Lab Better In Better Out, The Hague University of Applied Sciences, Rotterdam University of Applied Sciences, HMC Anthoniushove and the University Cancer Center Leiden-The Hague work together with companies, healthcare professionals and patients on technological solutions for healthcare. The Living Lab also works on a project basis with researchers from academic knowledge institutions, including TU Delft and Maastricht



Medical Delta Living Lab Geriatric Rehabilitation@Home

E-health applications can significantly ease the pressure on geriatric rehabilitation and improve care. There is also a greater need for medical specialist rehabilitation. Medical Delta Living Lab Geriatric Rehabilitation@Home focuses on the development of e-health applications that promote home rehabilitation of the elderly. At Medical Delta Living Lab Geriatric Rehabilitation@Home, Inholland University of Applied Sciences, The Hague University of Applied Sciences, Rotterdam University of Applied Sciences and Stichting Omring work together with companies, care institutions, informal caregivers and elderly people. The Living Lab also works on a project basis with researchers from academic knowledge institutions.



Medical Delta Living Lab ResearchOR

In the Medical Delta Living Lab ResearchOR. processes and systems are developed and validated to increase and guarantee efficiency and patient safety in the operative process. An operating room with measuring equipment and sensors is used for this. Reinier de Graaf Hospital and TU Delft are involved in this Living Lab.



Medical Delta Living Lab Integrative Medicine Technology

For some health issues, such as antibiotic resistance and pain complaints due to chronic diseases, regular medicine offers insufficient (good) solutions. Medical Delta Living Lab Integrative Medicine Technology is exploring non-pharmacological, integrated prevention and treatment approaches, such as the use of natural products, to make Integrative Medicine more available, accessible and acceptable in a responsible

way. The University of Applied Sciences Leiden, University of Applied Sciences Rotterdam, NVAA and Cure + work in this Living Lab together with patients, healthcare professionals and companies. The Living Lab also works on a project basis with researchers from academic knowledge institutions.



Medical Delta Living Lab Care Robotics

Robotics allow people to live independently at home for a longer period of time. Medical Delta Living Lab Care Robotics offers companies and other healthcare developers the opportunity to test, develop and validate their innovations in a practical manner together with end users. In addition, the Living Lab gives companies the opportunity to take up questions from healthcare practice and translate them into promising innovative technological solutions. The University of Applied Science the Hague, TU Delft, LUMC, Pieter van Foreest and the Reinier de Graaf Hospital collaborate in this Living Lab.



Medical Delta Living Lab Rehabilitation Medical Delta Fieldlab Phenomix Technology

Smart technology such as robotics, sensors, artificial intelligence and e-Health solutions offer opportunities for intensive rehabilitation. This can be done at home, in a rehabilitation center or at the physiotherapist. The Medical Delta Living Lab Rehabilitation Technology gives companies and other healthcare developers the opportunity to test their innovations together with end users and to develop them in practice. They can also take up questions from healthcare practice. The University of Applied Sciences the Hague, TU Delft, LUMC, Erasmus MC, Sophia Rehabilitation Center, Rijndam Rehabilitation Center and the Rijnlands Rehabilitation Center collaborate in this Living Lab.



Living Lab Medical Delta Instruments Minimally invasive medical instruments ensure smaller scars, less trauma, less blood loss, faster recovery and a lower risk of infection. The Living Lab Medical Delta Instruments connects technical and clinical Medical Delta researchers with companies and supports them in the development of high-guality prototypes of medical instruments. TU Delft, Erasmus MC, LUMC, AMC, Reinier de Graaf

Hospital and LIS (Leiden Instrument Makers School) collaborate in this Living Lab.



expanding the Fieldlab.



Medical Delta National eHealth Living Lab (NeLL)

E-health plays an important role in making healthcare future-proof. To make digital care part of regular care, Medical Delta Living Lab NeLL facilitates scientific research into e-Health solutions and their validation and testing. In addition, the Lab shares news, insights and knowledge. Within this Living Lab, patients, caregivers, consumers, students, scientists, entrepreneurs, organizations and institutions work together on tomorrow's healthcare.

A good picture of one's current health can be created by measuring metabolic products such as amino acids, hormones, glucose and adrenaline. With the resulting 'metabolic profiles', doctors can make earlier diagnoses and personal treatment plans. In the Medical Delta Fieldlab Phenomix, companies, healthcare institutions and scientists work together on practical applications of metabolomics research. Leiden University and Erasmus MC participate in the Fieldlab together with various companies from the biotech and pharmaceutical industries. It is located at the Leiden Bio Science Park and is being established with an EFRO subsidy. InnovationQuarter, Medical Delta and others are involved in setting up and

Medical Delta NIMIT: Novel Intruments for Minimally Invasive Techniques

Creating standardized methods to develop minimally invasive instruments





Prof. dr. Frank Willem Jansen Gynaecology (LUMC/TUD)

Prof. dr. Gijs van Soest Cardiology (EMC)

Prof. dr. Jenny Dankelman **BioMechanical Engineering (TUD/** LUMC)

In minimally invasive surgery, the surgeon operates through the small incisions. Compared to open surgery, minimally invasive surgery results in smaller scars, less trauma, less blood loss, faster recovery and a lower risk of infection.

In addition to the direct benefits for the patient, there are economic the Medical Delta program, phantom models will be developed. benefits too: shorter hospital stavs and quicker recovery mean Phantoms are artificial structures representing human tissues. the patient can go back to work faster. Minimally invasive surgery and they can be used to test medical instruments for function and relies on the minimally invasive instruments that are designed usability. Such phantoms could also be used in place of some of the and developed in academia. Currently, many instruments being animal and clinical trials of the instruments. "It would be preferable if developed have unique designs and advanced functionalities. phantoms themselves were standardized and certified by regulatory However, these instruments lack standardization, they are complex. bodies. This would contribute to the certification of medical expensive and difficult to manufacture. This creates hurdles for instruments," says John van den Dobbelsteen, Associate Professor regulatory assessment and certification and delays innovation at the department of Biomechanical Engineering at TU Delft. benefits for the patient. The Medical Delta NIMIT program aims When creating a new instrument, it is important to keep documenting the practical workflows of existing instruments to design minimally invasive instruments that enable a rapid innovation cycle based on wel characterized, documented process and learn lessons from those devices. This should result in a steps and standardized tests. This will eventually lead to quicker standardized method for development. implementation in clinical practice. In order to achieve this, new It would be ideal to have a prototype lab where instruments can be design, manufacturing and assembly methods should be developed made in small, reproducible series. "We don't want to create and for minimally invasive instruments. The instruments produced test instruments to have the perfect picture for a scientific paper, should be simpler ad should support sustainable surgery. we want to test the instruments to be used in clinical practice," says Van Soest

Modular instruments

"As academic developers, we realized that we do not get far enough in the translational trajectory to transfer our technologies to clinical practice," says Gijs van Soest, Associate Professor at the Department of Cardiology at Erasmus MC.

"It is a shame to throw away every part of a good technical device that still functions well, after having only been in the body for half a minute," says Tim Horeman, Assistant Professor at the department of Biomechanical Engineering at TU Delft.

Circular design can help reduce this waste. The circular economy can be applied on three levels: reuse the instrument as a total assembly; reuse certain components or reuse the materials. The goal of this project is to develop a modular design approach for a variety of instruments for several purposes, for example the development of simple and smart hardware components with wellcontrolled properties that can also be reused for other applications.

Phantom models

The instruments must be tested following a set of regulations. In

Instruments should be produced simpler and support sustainable surgery





Collaboration

In this consortium, TU Delft and Erasmus MC provide the technical knowhow and labs with required equipment. LUMC and Erasmus MC provide the knowledge for safe application. Medical Delta Living Labs ResearchOR and Medical Delta Instruments are also connected to this consortium.

Because this program leads to standardized methods to develop instruments faster and better following regulations, the results can also be of value for other programs at Medical Delta.

Medical Delta Cardiac Arrhythmia Lab

Unravelling disease mechanisms of cardiac arrhythmia for better treatment











Prof. dr. Natasja de Groot Cardiologist-Electrophysiologist Veen - Signal Processing (TUD) (EMC)

Prof. dr. ir. Alle-Jan van der

Prof. dr. ir. Wouter Serdijn **Bioelectronics (TUD)**

Prof. dr. Bianca Brundel Molecular Biologist (VUmc)

The number of patients diagnosed with cardiac arrhythmia is rapidly increasing due to ageing, obesity, diabetes and elevated blood pressure. Cardiac arrhythmia is therefore the cardiovascular epidemic of the 21st century. Treatments are often not successful, but a new patient-specific therapy can improve existing treatments.

> Cardiac arrhythmia is the cardiovascular epidemic of the 21st century. The number of patients diagnosed with cardiac arrhythmia is rapidly increasing due to ageing, obesity, diabetes and elevated blood pressure.

Providing a patient-specific therapy can improve treatment

Atrium fibrillation is the most common arrhythmia. It is a progressive disease, which means that episodes of arrhythmia progress from short-lasting episodes to episodes which are continuously present and no longer terminate spontaneously. Atrium fibrillation may cause stroke and heart failure and is even associated with death.

Treatments are often not successful, but a new patient-specific therapy can improve existing treatments. This can be achieved by measuring the



degree of electropathology ('staging the arrhythmia'). However, this is still not yet possible because there are no diagnostic tools to measure electropathology. It is therefore also not possible to recognize patients at risk of arrhythmia early. In addition, there are no therapies specifically targeting electropathology.

Treatment of Arrhythmia

Electrical signals recorded from young hearts usually have a simple morphology, as electrical waves propagate smoothly through cardiac tissue. Over the years, these electrical signals may become more complex (electropathology) due to damaged cardiac tissue. When the degree of electropathology exceeds a certain threshold, cardiac arrhythmia may occur. Treatment of cardiac arrhythmia is still often not successful, says Natasja de Groot, professor and cardiologist-electrophysiologist at Erasmus MC. Current treatment consists of either an 'electrical shock' to restore the normal heart rhythm, drugs or ablative therapy (eliminating the cardiac tissue that is

causing arrhythmia). Unfortunately, these therapies have side-effects and are only moderately effective.

Unravel electropathology

The aim of the Medical Delta Cardiac Arrhythmia Lab is to reduce the cardiac arrhythmia burden by unravelling arrhythmia-related electropathology and designing and testing novel bio-electrical diagnostic tools and therapies targeting electropathology. This enables staging of the cardiac arrhythmia and selection of the appropriate treatment in the individual patient, thereby improving therapy outcome. A first step is to unravel electropathology by quantifying electrical parameters. For this purpose, a 192 electrode-array has been designed to record electrical signals directly from the surface of the heart during open heart surgery. Advance signal processing techniques are then used to comprehend electrical activation patterns during arrhythmia. Linking electrical signals with the structure of cardiac tissue is essential to unravel the mechanisms of

techniques.

Collaboration

arrhythmia. The future goal is to assess the degree of electropathology using non-invasive mapping

Arrhythmia-on-a-chip

To further unravel the mechanisms of arrhythmia, this consortium aims to design an arrhythmia-on-achip platform enabling the investigation of electrical conduction in relation to e.g. genetic defects.

This program is a collaboration between biologists, engineers and medical doctors from Erasmus MC, LUMC and TU Delft. They combine their unique expertise on advanced signal recording and processing techniques, cardiac mapping tools and arrhythmia related molecular mechanics.

ŤUDelft

Medical NeuroDelta: Ambulant Neuromonitoring for Prevention and Treatment of Brain Disease

Neuromonitoring for prevention and treatment of brain disease







Prof. dr. Chris de Zeeuw **Neuroscience (EMC)**

Prof. dr. ir. Wouter Serdijn **Bioelectronics (TUD)**

Brain disorders have a disastrous influence on people's lives, but they are notoriously hard to study and difficult to treat. For example, migraine, which is characterized by repeated attacks of severe headache, is difficult to investigate in patients, because it is impossible to predict when an attack will

strike.

When patients are asked to go to the hospital to investigate their brain disorder, they will often notice they do not get an attack. This happens because the hospital setting is so different from the patient's natural environment. "In order to better understand the disease, it is important that the patient's brain activity can be measured over a longer period of time in their home environment, preferably using a non-invasive method," savs Arn van den Maagdenberg, professor of molecular and functional neurogenetics at LUMC.

The same holds for autism, a developmental disorder that starts in young children. "Autism is caused by a disturbed interaction between the cerebellum and cerebrum. People with autism have problems with social interaction and they show repetitive behavior. We need better means to monitor how the brain deteriorates in autism," says Chris de Zeeuw, professor and head of the Department of Neuroscience at Erasmus MC.

Observing patients in their home environment

In this Medical Neurodelta program, researchers focus primarily on autism and migraine - two disorders with widespread brain dysfunction. For both disorders, it is important to observe patients in their home environments to understand why disease worsens over time or how attacks begin. A non-invasive method would provide a much wider time window in which brain activity can be studied, giving doctors and scientists extremely valuable insights.

Wireless long-term neuromonitoring systems

The Medical Neurodelta program focuses on the development of wireless long-term neuromonitoring systems to register brain activity. Bio-electronics, acoustrodes and optrodes in combination with artificial intelligence and machine-learning will be applied for diagnosis and treatment.

The new system should not only monitor brain activity, but it should also modulate brain activity. Advances in light probes and ultra- sound, made at TUDelft, make this feasible. In Rotterdam,

High-frequency ultrasound would allow monitoring throughout the day in the home environment, which would be unique in the world







within CUBE. Medical Neurodelta researchers have already started to use high-frequency ultrasound to measure the activity in large parts of the brain; within the project, they will use low frequencies to stimulate specific brain areas to correct brain activity. According to professor De Zeeuw, ultrasound would also allow monitoring throughout the day in the home environment, which would be unique in the world. Compared with MRI, for example, this would be much easier - it would not be possible to place an MRI scanner in a patient's house.

Testing in mice

In the program, ways to monitor and modulate brain activity will be first tested in mice, paving the way for clinical applications in humans. Testing in mice is already being done, but these experiments are conducted in non-natural settings. Professor De Zeeuw asks: "If you investigate autism with mice housed in isolation, how can you see the effects of an intervention?" In this new study, researchers will make use of their recently developed FlashTrack system, in which individual mice can move freely and can be easily identified within a group. Using this system, several mice can be put together and followed over time.

Collaboration

In this consortium, Erasmus MC, TU Delft and LUMC are collaborating. According to professor Van den Maagdenberg, the Medical Delta area is small enough that there is a regular crossover and large enough to incorporate different expertise and make real progressing. It is a unique opportunity to create added value.

In other projects within the overall Medical Neurodelta program, other brain diseases, such as epilepsy and brain dysfunction in infants, are also being investigated. The same technology can be applied to investigate these disorders.

Medical Delta: Improving Mobility with Technology Improving mobility with technology



(TUD)

Prof. dr. ir. Jaap Harlaar



BioMechanical Engineering Neurorehabilitation (EMC)





Prof. dr. Frans van der

Helm - BioMechanical

Engineering (TUD/LUMC)



(TUD/LUMC)

Prof. dr. Sita Bierma

General Practice (EMC)



Prof. dr. Rob Nelissen Orthopaedics (LUMC/TUD)

Our society is aging. Motoric disabilities increase with age, which means there will be more and more people with restricted mobility. The Medical Delta Improving Mobility with Technology program focuses on the development of accurate diagnostics and treatments to improve mobility.

Prof. dr. Gerard Ribbers

This program consists of two tracks: the rehabilitation track and the orthopaedics track, each with three themes.

Rehabilitation: practice in a safe manner

The rehabilitation track focuses on outcome prediction, precision diagnostics and unsupervised training. The latter aims at maintaining balance in people who learn to walk again after a stroke. Rehabilitation techniques in the field of robotics are used for treatment

"Therapy intensity is a major determinant for a positive outcome, but there is simply not enough individualized supervision by professionals to guide the patient", says Jaap Harlaar, professor of Clinical Biomechatronics at TU Delft. "The aim of our project is to let people practice more intensely in the rehab center and at home in a safe manner, by developing equipment they can use to practice unsupervised."

Recently Heike Vallery, professor at TU Delft, demonstrated the feasibility of a new wearable balance assistance backpack to help people walk with more stability and to keep balance. Together with professor Ribbers from the rehabilitation center Rijndam, professor Vallery will develop this backpack and other wearables to incorporate safe gait training into clinical practice. Implementation in clinical practice will involve the Medical Delta Living Lab Rehabilitation Technology.

Orthopaedics: multiscale modelling

The second track focuses on orthopedics: the safety of arthroplasties, sports injuries and, in particular, osteoarthritis. Osteoarthritis is the leading cause of physical disability worldwide and involves the degeneration of joint cartilage. Working together, researchers from the Medical Delta combine their expertise in epidemiology, the biomechanics of gait, tissue and cell research, and dynamic imaging

techniques. In this project, the researchers focus on multiscale modeling, in which biomechanical computational models are made at all levels and combined to gain better insights into the effects of load on cartilage.

Precision diagnostics

Choosing the right therapy to improve patients' mobility requires knowledge of the etiology. This 'precision diagnostics' is a focus for both tracks. "We want to maximize the treatment effect by selecting the optimal option; we don't want to over-treat or under-treat the patient," says Harlaar.

Collaboration

In this consortium, Erasmus MC, TU Delft and LUMC will collaborate closely. The Living Labs will also be involved in early prototype testing.



We want to maximize the treatment effect by selecting the optimal option

Medical Delta Institute of Fetal & Neonatal Care

Preventing morbidity and mortality for both mother and child





Dr. Alex Equink **Obstetrics and Gynaecology** (EMC)

Dr. Monique Haak Obstetrical Ultrasound and the BioMechanical Engineering Fetal Heart Program (LUMC)



Prof. dr. Thomas Hankemeier Analytical Biosciences (UL/ EMC)



In the Netherlands, about 175.000 women give birth every year. In most cases, pregnancy and childbirth are uncomplicated and end with a healthy child and happy parents. Unfortunately, about 1300-1400 children die in the period around birth. The 100 days before conception and 1000 days thereafter are fundamental for the development of a child and also largely determine their future health.

The major causes of perinatal mortality are premature birth, concenital anomalies and fetal growth restriction. The aim of the Medical Delta Institute of Fetal and Neonatal Care is to prevent morbidity and mortality caused by complications during pregnancy, birth or the neonatal period for both mother and child. The program will focus on substantially decreasing the incidence of the three major complications that can occur during pregnancy and childbirth. The MOMETA program (monitoring and metabolomics of complicated pregnancies and neonates), one of the projects within this Medical Delta institute, will develop innovative techniques in maternal-fetal monitoring to provide a strong and healthy start for every child.

Wireless heart rate monitoring

The program is divided into two parts. The first part will focus on developing new technological and medical solutions to monitor maternal and fetal heart rate.

To date, a pregnant woman is only admitted to a hospital when clear signals of illness are already present, which is often too late. Monitoring is done by a wired system, which is intrusive and limits patient mobility. In this program, a smart and wireless monitoring system will be developed to measure the heart rate of mother, fetus or neonate continuously. The system will ideally be combined with a big data analysis system and an automated warning system to detect early signs of clinical deterioration.

Metabolic monitoring for sepsis

The second part focuses on monitoring neonates and early diagnosis and prediction of neonatal sepsis (blood poisoning) using metabolomics.

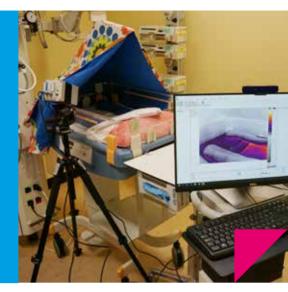
At the moment, sepsis in the neonatal period is often diagnosed when the newborn is already critically ill. The early diagnosis of sepsis and early prediction of treatment outcome of sepsis would be very desirable. A monitoring device for sepsis will be developed focusing on discovering and validating metabolic biomarkers for the diagnosis of sepsis and prediction of treatment outcome.











The data from these two parts of the program will be combined in a big data analysis to support early prediction, decision making and intervention to prevent maternal, fetal and neonatal morbidity and mortality.

Collaboration

In the Medical Delta Institute of Fetal and Neonatal Care, a unique close collaboration is realized between the maternal fetal medicine specialist and neonatologist, supported by technological experts and engineers from TU Delft, Erasmus MC, LUMC and Leiden University. This unique collaboration of clinicians is a strength in this program. This institute is linked to the Medical Delta Living Lab Care Robotics.

A smart and wireless monitoring system will be developed to measure the heart rate of mother or neonate continuously

Medical Delta eHealth & selfmanagement for a healthy society

How eHealth is changing healthcare







Prof. dr. Andrea Evers Health Psychology (UL)

Prof. dr. Niels Chavannes **General Practice and** eHealth in disease managemen (LUMC)

Prof. dr. Mark Neerincx Human-Centered Computing (TUD)



Dr. Valentijn Visch Industrial Design, Design Aesthetics (TUD)

Rita van den Berg Health-related physical fitness and lifestyle

interventions (EMC)

Existing healthy lifestyle programs can be effective in the short term, but many people eventually relapse into their unhealthy behavioral patterns. A paradigm shift is needed in health behavior - one that aspires to make healthy living attractive, immediately gratifying and convenient in the short term thereby consolidating healthy living in the long term.

> It plays a key role in preventing chronic diseases and in maintaining a healthy lifestyle. The goal of this program is to create instructions on how to reach vulnerable groups and to create a customized integrated eHealth solution. An integrated program, such as the benefit for all program, rewards people for their healthy lifestyles. They get points for everything they do, including logging in to the app. Short-term success is not the main measure, it is the long term that counts. People can fall back, as long as they pick it up again. This integrated solution should also help people with multiple diseases by giving them one approach to use, instead of multiple apps.

Many people come into contact with eHealth while making an appointment with their dentist or doctor online, using a smartphone activity tracking app or taking heart rate measurements. eHealth is booming. eHealth is about patient empowerment and is considered the future of health and well-being in our digital society.



"eHealth offers many solutions. However, there is a huge discrepancy between what is being developed and what is actually being used. Numbers indicate that the health gap between different socioeconomic groups is getting bigger. This is a large problem", says Andrea Evers, professor in health psychology at Leiden University. The eHealth technologies being developed mainly benefit those who know how to handle them.

In this Medical Delta program, the development of eHealth self-management solutions that are accessible for vulnerable groups is key. Vulnerable groups are difficult to reach and consist of people with a lower socioeconomic status (SES) or people with multiple diseases or comorbid diagnoses. "Vulnerable groups do not have access to suitable solutions. It is important to learn more about how people can be motivated to adopt a healthy lifestyle. They need to be rewarded and encouraged in a different way; for example, a financial incentive is

more important when targeting low SES groups", says Evers.

Data-driven prediction models

Data-driven prediction models using large data sets and machine learning will be used in this program to investigate what works best for whom, and why. Analyzing and interpreting the data will enable personalized solutions to show which people benefit most from which type of intervention.

Collaboration

The consortium includes researchers at Leiden University, LUMC, TU Delft and Erasmus MC working in collaboration. Researchers at these institutes excel in eHealth research, behavior change and smart technology solutions, and they are able to translate science into clinical applications. All consortium partners actively participate in the Netherlands eHealth Living Lab (NeLL).

ŤUDelft Univ Leide LU MC

Short-term success is not the main measure, it is the long term that counts

Vital Delta: Medical Delta's journey towards vitality and health Improving the vitality and health in the Delta region



Dr. AnneLoes van Staa Lector Healthcare Transitions (Rotterdam University of Applied Sciences)

> A population that is physically, mentally and socially more vital and healthier; from young to old. That is what the program Vital Delta: Medical Delta's journey towards vitality and health strives for. The practice-oriented research program was set up by a consortium of the four universities of applied sciences within Medical Delta (Rotterdam University of Applied Sciences, The Hague University of Applied Sciences and Inholland University of Applied Sciences).

> > In the research program, citizens, students, clients, healthcare professionals and researchers work together with colleges, universities, healthcare institutions and companies. The interdisciplinary program improves the health and well-being of the population in the delta region of Leiden, The Hague and Rotterdam by promoting the development of innovative (technological) health solutions through practice-oriented research.

> > The eight-year program started in 2018 and focuses on four work packages:



and safe implementation of healthcare and welfare technology

Fysically Vital develops, evaluates and implements innovations to promote a healthy lifestyle among vulnerable target groups

Socially Vital stimulates and improves cooperation between professionals from the social and health domain in particular

Self Vital aims to let people take control of their own lives and health

Collaboration

The cooperation between healthcare, the social domain and technology is the starting point for each of these work packages. To ensure this, each work package is led by two lecturers from different universities of applied sciences and coordinated by a work package leader.

Vitale Delta receives a grant from the SIA Steering Body. In addition, Medical Delta and the four universities of applied sciences also contribute to the realization of the program.

Supported Vital focuses on the design, development



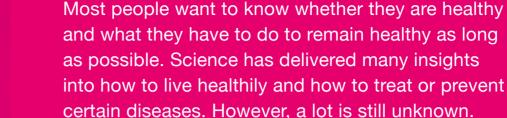
METABODELTA: Metabolomics for clinical advances in the Medical Delta Growing old healthy by predicting and preventing diseases













Prof. dr. Simon Mooiiaart

Epidemiology (LUMC)



Prof. dr. Aad van der Lugt Radiology (EMC)

Genes predict the risk of developing certain diseases in an individual. However, genes do not indicate if, and when, such a disease will develop. This is also determined by environmental factors such as nutrition and lifestyle.

Measuring the metabolites in the blood can tell you what is going on in the body. Metabolites provide information about someone's current health state. Metabolites are the result of the interplay between genes and these environmental factors. Metabolites are small chemical substances, like glucose or fats, that are products of the processes and reactions taking place within and between cells. Measuring metabolites therefore reveals information about such disturbances. Many of these metabolites are present in blood.

Metabolic profile

If many diseases can be early predicted with metabolomics, obtaining metabolic profiles for every person as a health monitoring approach becomes promising. The metabolic profile would give a warning before any symptoms appeared. "This kind of metabolic profile might be acquired in just a few years in a single drop of blood. In such way metabolomics can contribute to change healthcare in a fundamental manner," says Thomas Hankemeier, professor of Analytical BioSciences at Leiden University.

Measuring all the relevant metabolites is a challenge. Researchers are already able to measure thousands of metabolites in blood and urine samples using advanced analytical technologies, also called metabolomics. Metabolomics can be used to find a combination of metabolic biomarkers that predict complex diseases, such as dementia. It is even more interesting to investigate the underlying chemical processes causing the disruption in the metabolic equilibrium, to resolve or compensate the disruptions before the disease develops; early diagnosis is helpful in that case.

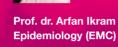
The METABODELTA: three themes

In the METABODELTA program, three themes will be addressed.

Prof. dr. Thomas Hankemeier Analytical biosciences (UL/ EMC)

Prof. dr Eline Slagboom Molecular epidemiology LUMC)

Prof. dr. Cock van Duiin Epidemiology (EMC/UL)













The first - "From metabolomics for discovery to clinical practice" - is focused on the translation of metabolomics discoveries to clinical and point-of-care applications. For that, novel technologies for large-scale metabolomics will be developed and implemented. In parallel, a robust and clinical metabolomics analyzer will be developed. The aim of the project is to integrate the metabolomics analysis steps into a small analyzer that can be used as a point-of-care metabolomics analyzer. These technologies will allow monitoring of the metabolic health state for diagnosis and choice of the proper intervention.

The second theme is focused on using metabolomics to understand the main metabolic processes determining heterogenous response of elderly people to interventions. Using the metabolome as monitoring a tool to improve the health of our ageing society would help to stimulate vitality. And finally, the third theme focuses on the development of novel preventive intervention strategies for dementia by understanding the interplay between the microbiome, blood metabolome and vascular and neuropsychiatric pathology.

Collaboration

In METABODELTA, leaders in the development of metabolomics technology and clinical diagnosis join with leaders who have access to unique cohorts and other clinical studies, to apply metabolomics in population and clinical studies to study healthy ageing. METABODELTA will make metabolomics more accessible within the Medical Delta. "Together we are bringing the ideal of growing old healthily into closer reach," concludes Hankemeier. This program also aims to make the data obtained in its research comparable to metabolomics data acquired by other labs worldwide, to validate findings, implement others' findings into research being done in the Medical Delta, and ultimately, apply the data for the benefit of healthcare.

HollandPTC Medical Delta program on HTA value proposition

Determining the cost effectiveness of proton therapy





Prof. dr. Marco van Vulpen Radiotherapy (EMC/LUMC/TUD) **Dr. Hedwig Blommestein** Health Policy & Management Health Technology (EUR)

Since healthcare costs are rising and resources are scarce, it is important to consider the societal impact in addition to the clinical impact of new technologies, such as proton therapy. It is essential for patients and society that scarce resources are spent as efficiently as possible. As proton therapy carries hight investment and operational costs, but may also provide high clinical value, a so-called Health Technology Assesment (HTA) is essential.

> A HTA includes an economic evaluation to determine the cost- effectiveness of proton therapy compared to photon therapy. Furthermore, HTA also covers the logistical factors, the implication of technological advances and its impact on overall healthcare delivery costs.

Proton therapy

Radiation therapy is used in the treatment of cancer to kill cancer cells. The usual radiotherapy with photons works well enough for many patients, but sometimes the tumor is too close to vulnerable organs or is relatively insensitive to the usual radiation. In those cases, proton therapy can be an option. Proton therapy is a new way to treat cancer. The program focuses on low-grade brain tumor and head and neck cancer. Proton therapy is different from traditionally used photon therapy, as it makes use of very local and precise dose deposition on tumors. This causes less damage to the surrounding normal tissue, resulting in fewer side effects. Proton therapy is a form

of radiation that uses protons instead of photons. Protons are small, charged particles that are in the core of an atom. To irradiate a tumor, the protons are accelerated to more than half the speed of light. The proton beam is directed at the tumor. The speed of the protons can be adjusted in the accelerator in a way the protons only destroy the tumor tissue, and surrounding healthy tissue is damaged as little as possible. However, it is more expensive than photon therapy and has high investment costs.

Two types of cancer

In this consortium, an HTA will be performed on proton therapy for the first time. The program initially focuses on two types of cancer. One is the low-grade brain tumor, which grows relatively slowly and often occurs in young people (30-45 years old). The other is head and neck cancer, which grows relatively fast and is more common among elderly people (60-75 years old). Currently, the patients involved in the assessment are being treated at HollandPTC and data collection has

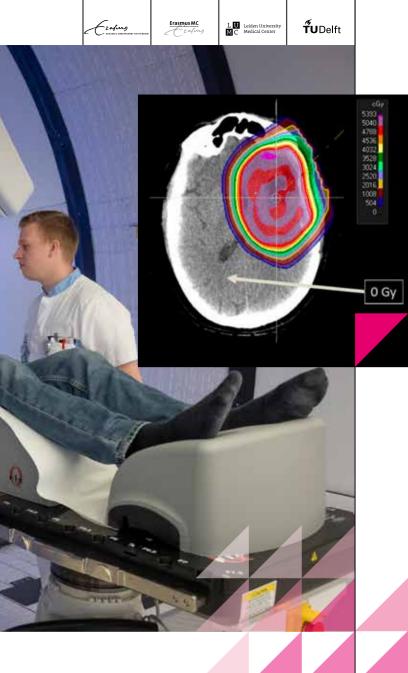
started. HollandPTC, founded by Erasmus MC, LUMC and TU Delft, is a treatment center for proton therapy Including these two different kinds of cancer will give a broad overview of proton therapy for the HTA. This research will lead to a generic model that can be easily adjusted to map the cost-effectiveness and value of proton therapy for all other types of cancer, which will be very useful in the future.

Collaboration

BEAM

Collaboration between the Erasmus University in Rotterdam, Leiden University and HollandPTC is realized in this program. One researcher will evaluate the cost structure of the proton center as well as the total cost of proton therapy from a societal perspective. Another researcher will be working on determining the value proposition of proton therapy for the two types of cancer, to eventually create a uniform model for decision making and cost- effectiveness analysis of proton therapy.

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The program focuses on low-grade brain tumor and head and neck cancer

Medical Delta Regenerative Medicine 4D: Generating complex tissues with stem cells and printing technology

Generating complex tissues with stem cells and printing technology





(TUD)



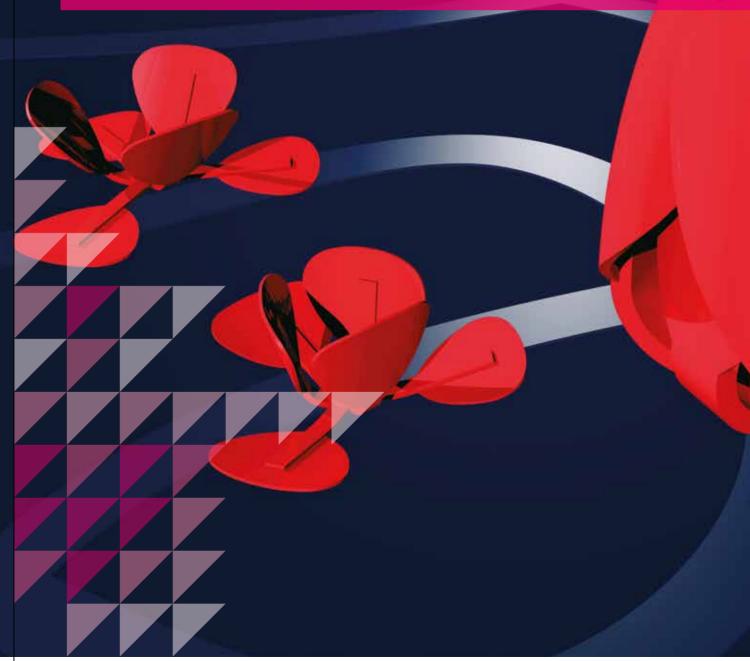
Prof. dr. Amir Zadpoor Biomechanical Engineering



Prof. dr. Rob Nelissen Orthopaedics (LUMC/TUD)



Prof. dr. Luc van der Laan Experimental Transplantation and Intestinal Surgery (EMC)



Prof. dr. Ingrid Meulenbelt Molecular Biology of Osteoarthritis (LUMC)

Regenerative medicine focuses on the development of new treatments to repair or regenerate diseased tissues and organs, such as cartilage, bone or liver, to restore function and improve patients' quality of life.

Patients with osteoarthritis, a joint disease affecting cartilage and bone, that leads to reduced mobility and pain, could benefit from regenerative medicine. It could also help patients with liver diseases, as the shortage of donor livers for transplantation has driven the field to look for alternative solutions. Regenerative medicine focuses on the development of new

treatments to repair or regenerate diseased tissues and organs, such as cartilage, bone or liver, to restore function and improve patients' quality of life.

The Regenerative Medicine Medical Delta program follows two tracks. The first focuses on reconstructing cartilage and bone defects, the second focuses on disease models.

Reconstruction: creating complex tissue constructs

4D printing is a new technology developed at TU Delft. Flat objects can fold into 3D objects over time when stimulated by pH, light, temperature or cells – think of origami structures. This technique will be used to create complex tissue constructs to direct cell behavior. Using this advanced technology together with stem cell technology, the researchers are building a proofof- principle cartilage-bone unit to repair defects with a biological implant.

Disease models: mimicking diseases

Disease models are used to mimic a disease as precisely as possible in a laboratory environment by putting together cells to construct complex tissues. In this program, induced pluripotent stem cells will be used. These are stem cells that can differentiate into all different cell types – for example, a cartilage cell or liver cell.

The disease models will be used to better understand diseases. Using the models, medication can be screened, thereby speeding up the quest for new pharmacological treatments. Erasmus MC Cafm **ŤU**Delft



Future perspective

According to Gerjo van Osch, professor of Connective Tissue Regeneration in the Departments of Orthopaedics and Otorhinolaryngology at Erasmus MC, repairing defects or having biological implants instead of metal ones would be an improvement.

However, this program aims to develop disease models in such a way that diseases can be treated at an early stage or even be prevented, so patients wouldn't need an implant at all.

Disease models are used to mimic a disease as precisely as possible

Collaboration

Regenerative medicine is a multidisciplinary field in which medicine, biology and engineering provide solutions jointly. The Medical Delta region is an ideal habitat in which to stimulate these developments. This program combines unique 4D-printing technology developed in Delft with strong knowledge on stem cells, hydrogels and disease models in Rotterdam and Leiden. This collaboration brings the developments in the individually strong research groups to a higher level.

Medical Delta AI for Computational Life Sciences

Creating new opportunities for biomedical research











Prof. dr. ir. Boudewijn Lelieveldt - Biomedical Imaging (LUMC / TUD)

Prof. dr. ir. Marcel Reinders **Bioinformatics (TUD /** LUMC)

Prof. dr. Mario van der Stelt Dr. Michel van de Velden Molecular Physiology (UL) Econometrics (EUR)

Prof. dr. Gerard van Westen Artificial Intelligence & Medicinal Chemistry (UL)

The possibilities to access the molecular data of cells and tissues are increasing, but this also creates new challenges. How can one find the information needed for research when there is such an enormous amount of data, for example?

The scientific program Medical Delta AI for Computational Life Sciences uses AI techniques to unlock biomedical data, discover new candidate drugs, detect special cell abnormalities or connections, and find useful information for research. This opens new doors for scientific research into the human body and in particular the development and effect of new medicines.

'Computational Life Sciences' combines computer science and life science. Using AI techniques, Computational Life Sciences helps researchers better understand biological mechanisms and make predictions about interactions between molecules and cells and with drugs. In this way, it helps scientists search more specifically for the molecules and genes that are of interest for a particular biological study. It helps them clarify hypotheses and better predict what medicines must contain in order to work properly. This creates new opportunities for, among other things, drug development that can be carried out faster, more specifically and better.

Medical Delta Professor and Scientific Leader Prof. Dr. Boudewijn Lelieveldt compares this to putting a new space telescope into operation. "If it suddenly becomes possible to see much further than one ever thought was possible, an astronomer has to think carefully about where to aim the telescope. Otherwise, he or she would get lost in the infinite possibilities. With the help of AI techniques, Computational Life Sciences helps us search for the molecules and genes that are interesting for specific biological research and drug development."

Leading scientists in the fields of bioinformatics, drug development, AI and computer science collaborate with pharmaceutical companies and with clinicians on the research projects within the program. In addition to the development and application of Computational Life Sciences with AI techniques in general, the program focuses on two lines of research: antibiotic resistance for tuberculosis and targeted medication for brain disorders.

Antibiotic resistance to tuberculosis

Tuberculosis is one of the deadliest infectious diseases in the world. Ten million people contract the disease every year, and 1.5 million of those people die (source: RIVM). A major problem in the fight against this disease is antibiotic resistance. Medical Delta AI for Computational Life Sciences wants to support research into antibiotic resistance with computer modelling and determine the effect of bacterial mutations by the use of algorithms, among other things. By gaining a better understanding of the mechanism of antibiotic resistance, the researchers hope to be able to search more specifically for ways to disrupt resistance and thus develop better candidate drugs.

Targeted medication for brain disorders

In the development of new medicines for brain disorders, it has proved difficult to translate data from laboratory animal research directly into the effects

runs faster and better.

Collaboration



of medicines on humans. For example, the control of cells by genes differs between humans and mice. Medical Delta AI for Computational Life Sciences is looking for methods to convert research data correctly, so that the development phase of medicines

In this program, scientists from Leiden University, LUMC, Erasmus University and TU Delft in the field of bioinformatics, drug development, Al and Computational Life Sciences work together with clinicians and pharmaceutical companies. Scientists from Erasmus MC are also involved in the program.

Medical Delta's Journey from Prototype to Payment From prototype to payment

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Health Care Governance (EUR)

In the early 1960s, it appeared that thalidomide (Softenon), used as a sleeping pill and sedative for pregnant women, led to babies being born with malformed limbs. Because of this disaster, people realized that new drugs and technologies need to be regulated.

> The idea that society must play a role in the development of technologies grew stronger and became the standard.

New technologies are too important to leave to doctors and technicians; the development of new technologies should be guided by the assessment of its potential risks, costs, benefits and impact on society.

In the Medical Delta program From Prototype to Payment, this societal perspective is positioned in parallel to the other 14 scientific programs.

Health Technology Assessment

A Health Technology Assessment (HTA) is a policy framework used to decide on market access and the reimbursement of new technologies. HTA provides the methods to assess ethical issues, organizational consequences and cost-effectiveness. Ethical issues relate to doing good to all. Today, big

data is used for early detection and diagnosis of diseases. If we could better predict when someone would get sick, what would the consequences be? Would people want to know this? What would it mean for the insurance business? Would we continue to show solidarity with those who get sick? Organizational consequences relate to the effects of implementing a new technology. Do we need to train technicians? Will physicians delegate tasks to nurses? What will be the financing model of the new technology? A device that will be used once inside the hospital will have a completely different business model than a device that will be used multiple times outside the hospital. Another question is who would buy the device?

The hospital, insurance companies or the patients themselves? The development of new technologies should be guided by the assessment of its potential risks, costs, benefits and impact. Antoinette de Bont,

The development of new technologies should be guided by the assessment of its potential risks, costs, benefits and impact

professor Sociology of Innovation in Health Care at the Erasmus University Rotterdam, is considering these type of questions in this Medical Delta program.

Added value

Cost-effectiveness of a new technology is an essential part of the assessment. De Bont comments: "We look at how much a certain technology costs and what added value it offers. New technologies will be

compared to technologies that are currently used. An example would be MRI scanners. The scanners used today work well, but even better MRI scanners have been developed

Collaboration

technologies.

It is important to involve many stakeholders in this Medical Delta program - as mentioned, new technologies are too important to leave to doctors and technicians. Collaboration with the National Health Care Institute (Zorginstituut Nederland) is essential. Next to care institutions, other decision makers, such as the Health and Youth Care Inspectorate and the Ministry of Health, Welfare and Sport, will be involved to develop the methods for evaluating health

Ezofung



YOUNG Medical Delta (YMD) is a platform for students, young researchers and young professionals who are at the starting point of their career in life sciences, health and technology.

Young Medical Delta connects students and young professionals who share a passion for medical technology and opens up new opportunities for them to explore. Young Medical Delta has participating associations from different universities, as well as the universities of applied sciences and PhD students from the different institutes that are connected to the Medical Delta. Through its connection with Medical Delta, Young Medical Delta is in contact with a large network of researchers who are working on the cutting-edge in their field. Young Medical Delta organizes its own events as well as events in collaboration with its partners. These events enable discussion and the discovery of recent research in the field of medical technology. One of such events is a yearly hackathon, where participants are motivated to find a solution to a problem that has been submitted from within the healthcare system. With the organization of symposia, Young Medical Delta presents ongoing research and allows for participants to network and find new collaboration opportunities.

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Drs. J. (Jasper) Faber, Young Medical Delta

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Medical Delta Professors

Our Medical Delta professors are appointed to at least two of the five academic institutions represented within Medical Delta (TU Delft, Erasmus MC, Erasmus University Rotterdam, Leiden University and LUMC). The professors who receive this appointment are committed to interdisciplinary research in the field of health & technology.





Prof. dr. ir. Boudewijn Lelieveldt LUMC & TU Delft

Prof. dr. Rob Nelissen LUMC & TU Delft





Prof. dr. ir. Nico de Jong TU Delft & Erasmus MC

Prof. dr. ir. Lucas van Vliet TU Delft & Universiteit Leiden



Prof. dr. ir. Richard Goossens TU Delft & Erasmus MC



Prof. dr. ir. Ton van der Steen Erasmus MC & TU Delft



Prof. dr. Jenny Dankelman TU Delft & LUMC



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Prof. dr. Andrea Evers Universiteit Leiden, Erasmus Universiteit & Erasmus MC



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About Medical Delta

Medical Delta wants to realize sustainable care with technological solutions. As a key player in the health and technology ecosystem, it aims to create an impact on people, care, knowledge and economy through interdisciplinary scientific research and practiceoriented Field and Living Labs. In doing so, Medical Delta helps make healthcare better, while keeping it accessible and affordable.



Medical Delta brings together three renowned universities, two university medical centers and four universities of applied sciences in the province of Zuid-Holland. Medical Delta started a major scientific program in 2019. In the coming years, almost 300 scientists will work together in 15 different consortia on technological solutions for sustainable care. Companies, healthcare institutions, the province and major cities in the Delta are closely involved, including through Living Labs, where innovations with end users are tested in healthcare practice.

The objective for the period 2019-2023 is for the Medical Delta research programs to grow in size, achieve excellence and develop technological solutions for sustainable care.

Medical Delta was founded in 2006 by TU Delft, Leiden University, Erasmus University Rotterdam, LUMC and Erasmus MC. Since 2016, four universities of applied sciences have joined Medical Delta: The Hague University of Applied Sciences, Hogeschool InHolland, Hogeschool Rotterdam and Hogeschool Leiden. Medical Delta is managed by board members Frank Willem Janssen and Gertine van der Vliet. The board is advised by the Scientific Council and the Social Council. The Scientific Council is represented by Medical Delta professors, educational directors, lecturers and Young Medical Delta representatives. Medical Delta is facilitated by a small, effective agency.



Gertine van der Vliet Msc. Managing Director Medical Delta



Prof. dr. Frank Willem Jansen Chairman Medical Delta











