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CONSORTIUM



fundación para el
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MIT Massachusetts
Institute of
Technology

From **bench** to **bedside** and back again

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From **bedside** to **bench** and back again: The Catalyst and Converter programs

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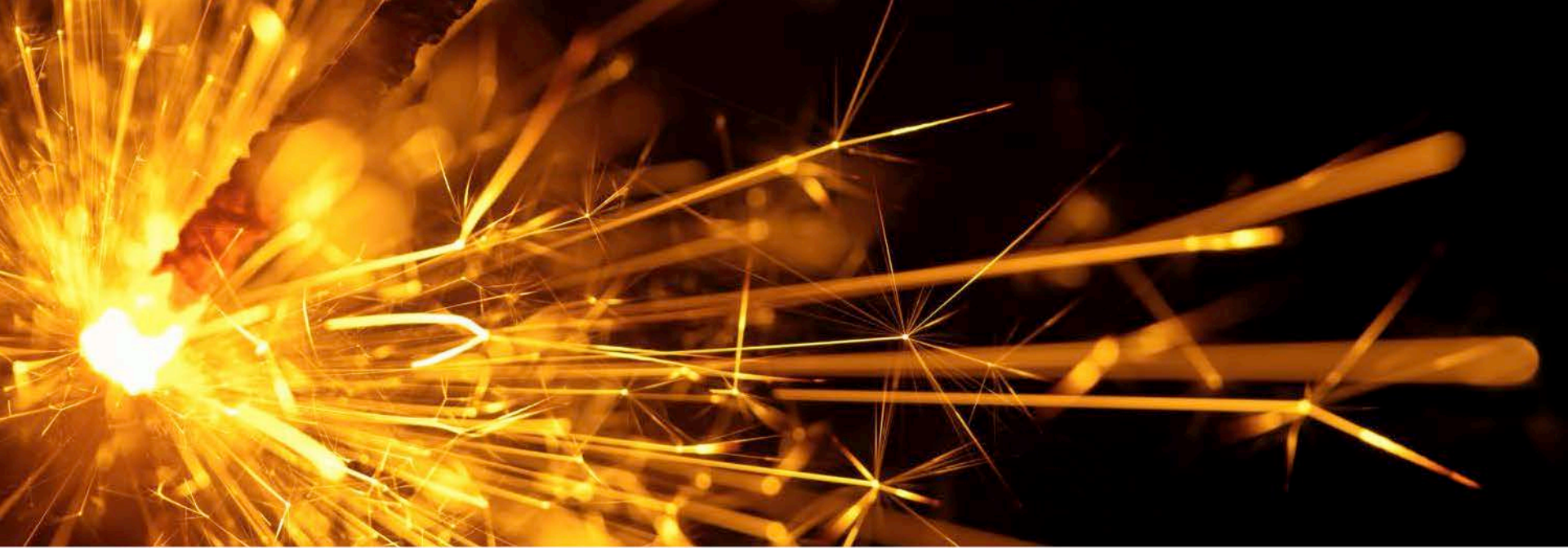
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Technology

Catalyzing change in the health technology innovation ecosystem

By changing how people think and work

For long-term societal and economic benefit



What is M+Visión?

An international collaboration for biomedical technology innovation led by Comunidad de Madrid and MIT

A new way of working that integrates the clinic, the research lab, and the office

A proven method to deploy R&D+i investment to reach patient impact



M+Visión: From concept to proof

Established a seasoned management team and **a community of 350 faculty, collaborators, and program contributors**

Developed and tested a **new innovation method, IDEA³**

Demonstrated results in impact-oriented biomedical technology R&D



Our ethos: Focus on people

Open, multi-professional community, sharing experiences through **meaningful work**

Developing people and ideas that will drive healthcare innovation

Leveraging existing institutions and infrastructures



Our community: A model ecosystem

Research universities

The ingenuity to invent new solutions

Healthcare institutions

The inspiration for change and the field of opportunity

Business sector

The expertise to translate ideas into successful innovations





Start up phase: Bets

1. Young talent would be capable of playing a central role
2. Community would embrace the opportunity
3. Need-driven approach would drive innovations and ecosystem development
4. Effort would yield follow-on benefits to MIT and Madrid

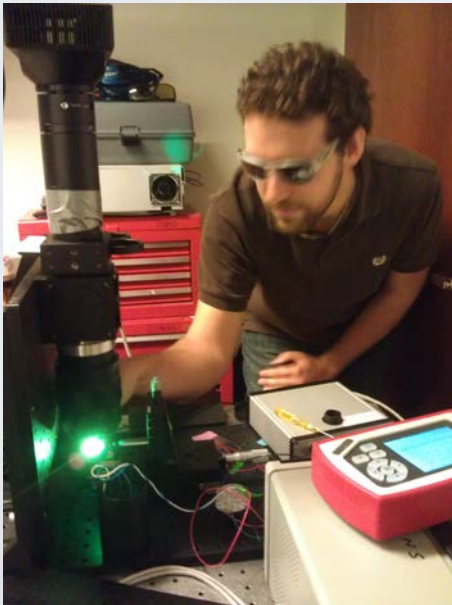
The Catalyst program

<u>Block 1:</u> <u>Project Definition</u> 4-5 months All fellows in cohort Throughout, they regularly test assumptions and impact proposition directly with diverse stakeholders	<i>Needs</i>	<i>Solution ideas</i>	<i>Validate</i>	<i>Proposal</i>
	Identify medical needs and specify the “impact proposition” for addressing them.	Propose and assess solutions that would address the medical need.	Define the use case and outline definitive evidence or study that would demonstrate the impact	For selected opportunities (= need + idea), propose an action plan for a project to realize the opportunity. Each Fellow is a member of from 1-3 project teams.



The Catalyst program

<i>Block 2: Proof-of-Project 6-9 months By each project team</i>	<i>Project launch</i>	<i>Early experiments</i>	<i>Proof of project</i>	<i>Updated project plan</i>
<i>Throughout this block they aggressively seek to produce preliminary evidence for their idea</i>	Recruit collaborators who can provide key expertise and infrastructure. Finalize initial resource requirements.	Initial exploration into project execution. Training and approvals as needed.	Studies and other actions that provide evidence that project is viable.	Define a set of milestones, objectives, and associated resources needed to achieve proof-of-principle



The Catalyst program

<i>Block 3: Case for continuity 12-24 months By each project team Overall aim is to have clear continuity plan for the project and each Fellow</i>	<i>Project execution</i>	<i>Documenting evidence</i>	<i>Continuity opportunities</i>	<i>Graduation</i>
	Project priority is to provide evidence that idea is viable for the envisioned use case	Document progress through IP disclosure and prosecution and publication in conferences and journals	Identify needs and opportunities to continue project development after the program ends. Identify and secure next stage career opportunities for Fellows	Fellows: proceed to academic, medical, or business positions. In some cases they will continue to be involved in project development.



Our results: Flagship *Catalyst* program

20	>131	88	25	7	75%
projects with healthcare impact potential	collaborators	research groups at 31 institutions	invention disclosures and 15 patent application s	new grants awarded	collaborators opening new lines of research



Catalyst projects, 2012–2015

Diverse in subjects and technical areas

Early intervention

Projects that can enable earlier, better responses to disease and injury

- + Cell
- + Colo
- + Fetal
- + NeuroQWERTY
- + Skin

Managing health

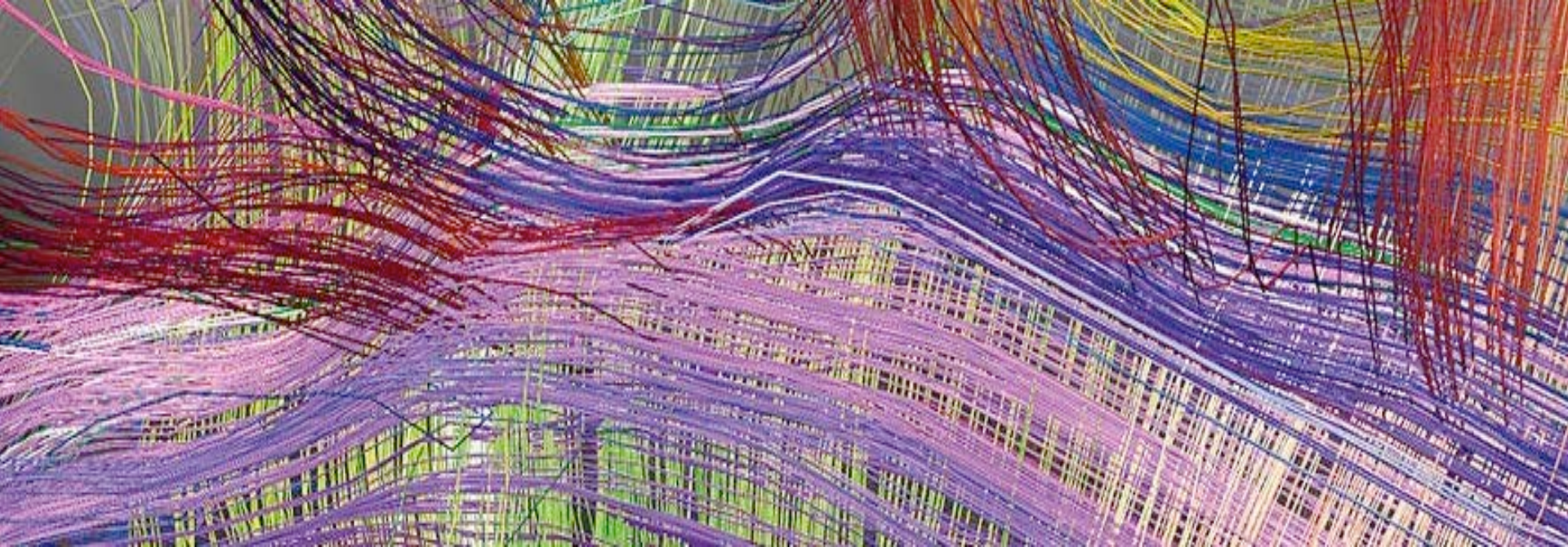
Projects that can help people achieve and maintain better health

- + Brain
- + Eye
- + Hydration
- + Leuko
- + Scolio

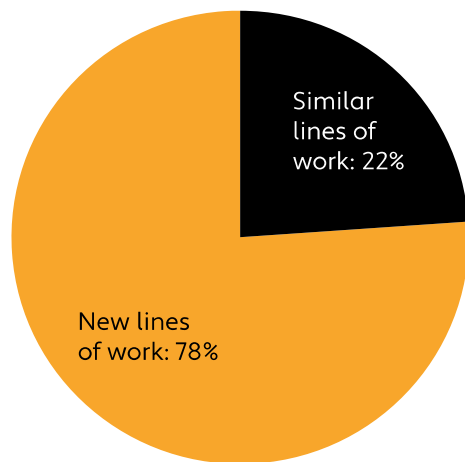
Better care delivery

Projects that can make clinical care safer and more effective

- + CT
- + Lumbar
- + mPET
- + MS-SAR
- + Pancreas



Our results: New lines of work, accelerated IP



New lines of research
expand opportunities

M+Visión invention disclosures: 25+

Patent app's: 15+

Licenses &
options: 3+

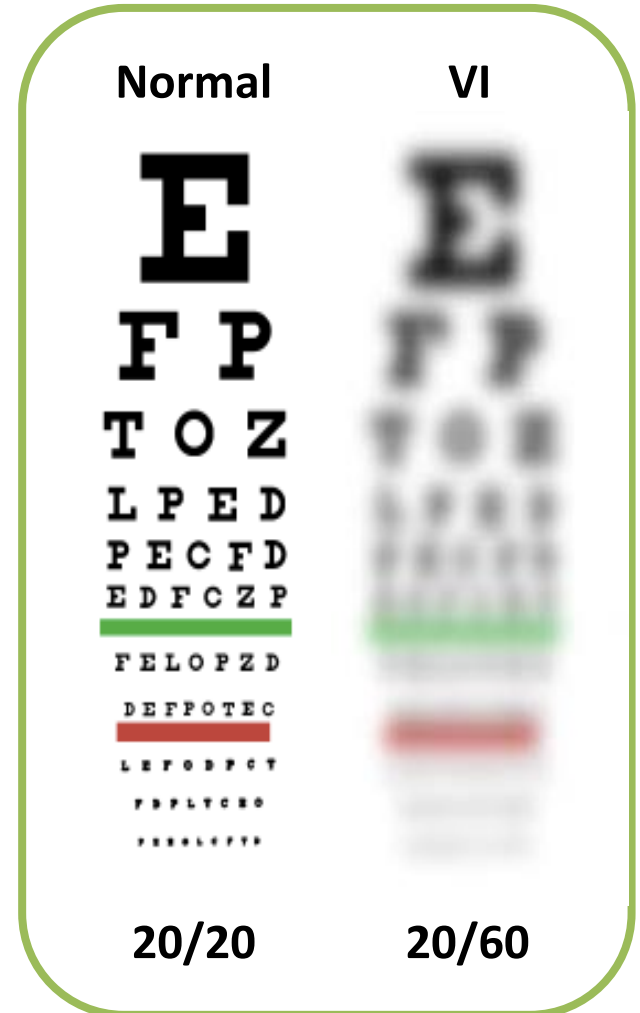
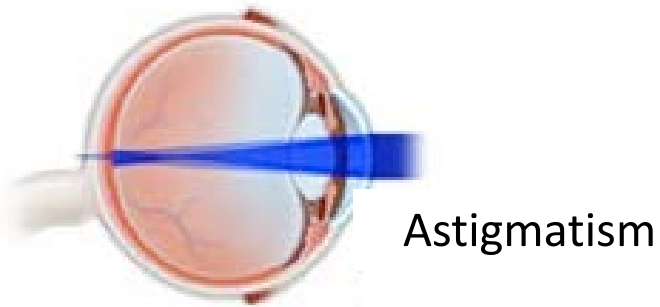
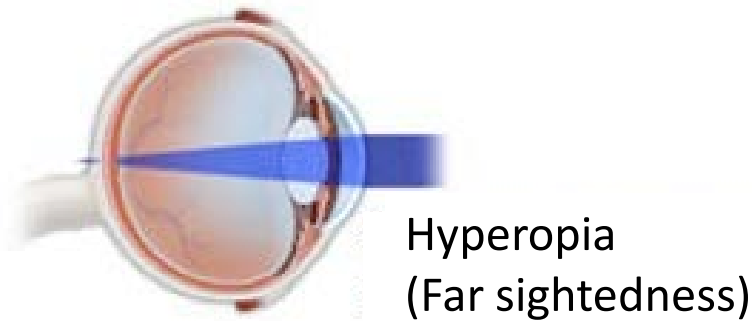
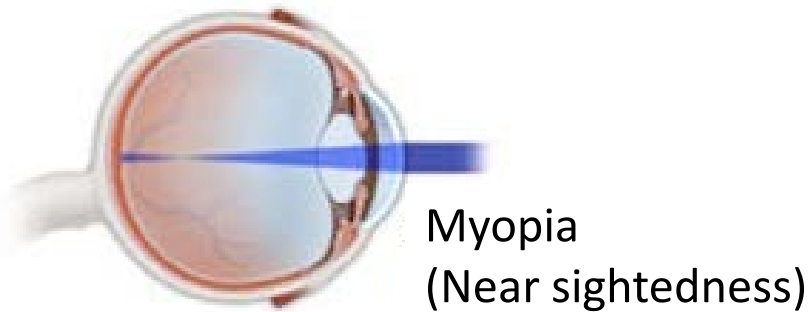
MIT invention
disclosures: 14

Patent
app's:
6

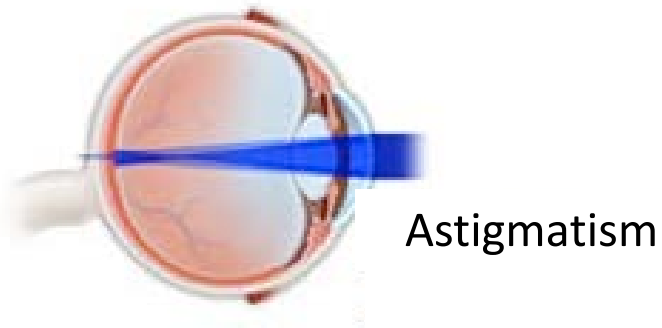
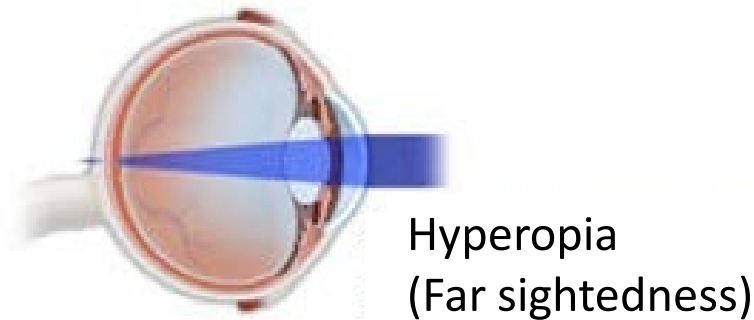
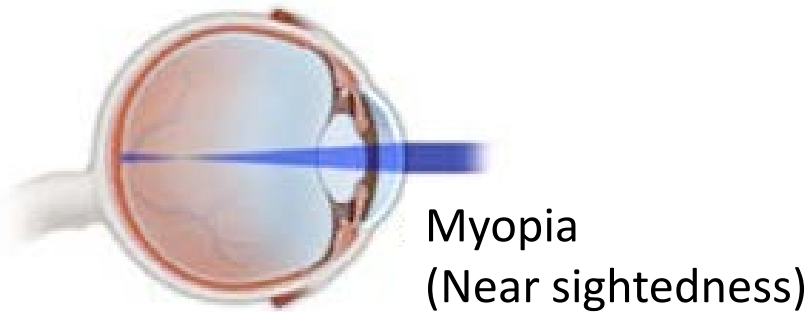
Licenses &
options: 3

Volume and pace of IP
development exceeds
institutional norms

Refractive errors can lead to visual impairment

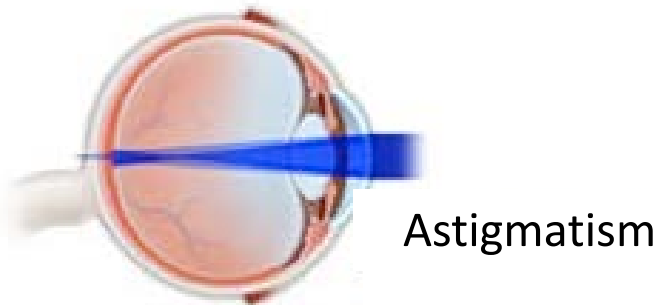
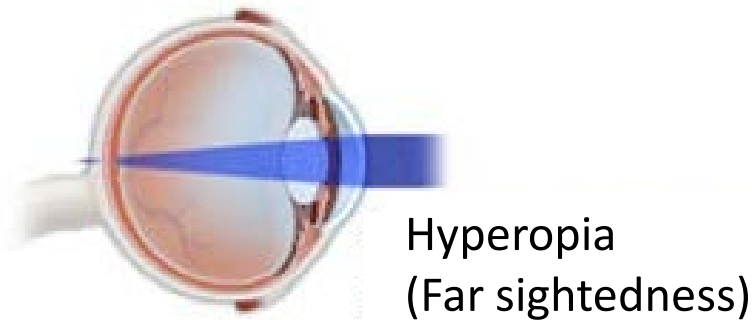
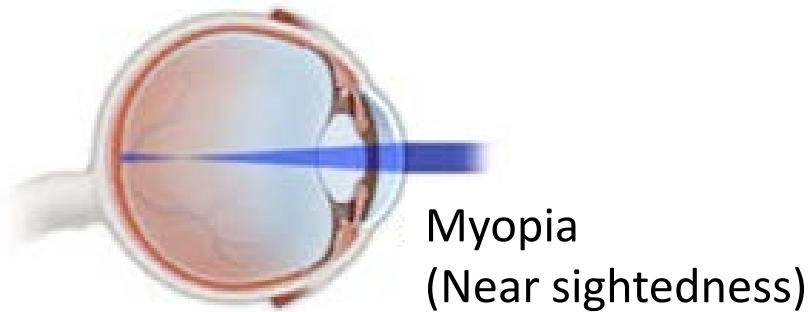


Refractive errors can lead to visual impairment



**There is still
no effective
prevention
method**

Refractive errors can lead to visual impairment



Cost-effective
treatment



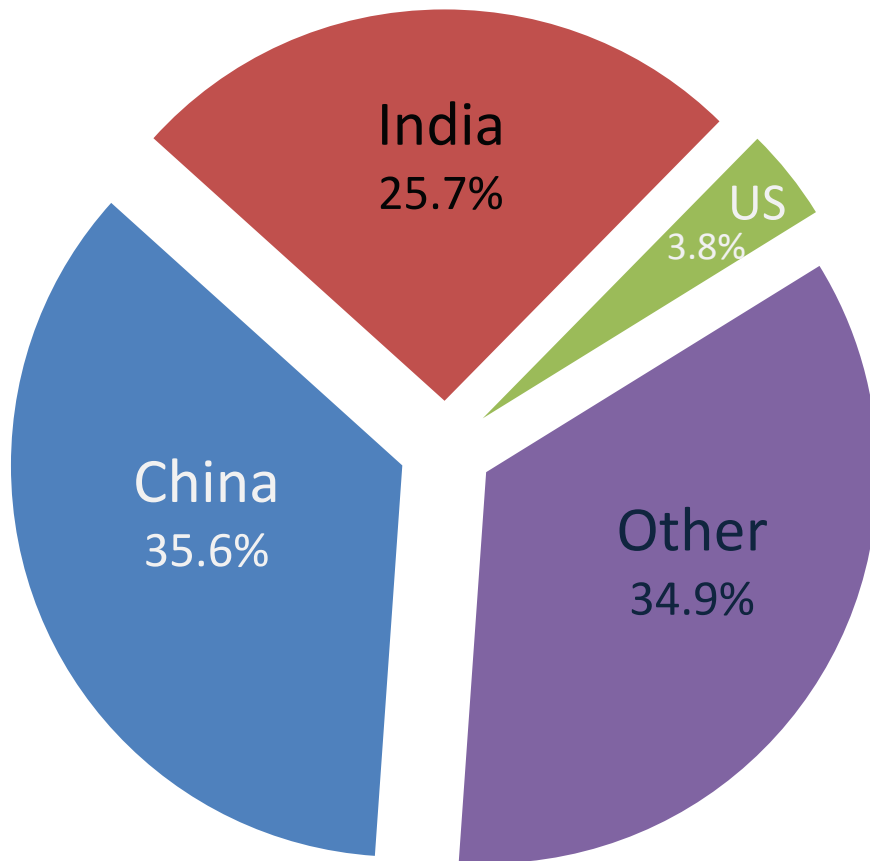
Visual impairment
due to
Uncorrected
Refractive Errors
(URE):

153 million

>10% children

Prevalence of URE is high everywhere

Visual impairment due
to URE



Country	Prevalence
China	4.4%
India	4.0%
USA	1.9%
Global	2.7%

Barriers for correction

Cost of lenses



2 €

Access to prescription

- Trained optometrists
 - United Kingdom
 - 1 optometrist: 8000 people
 - Ghana
 - 1 optometrist: 500,000 people
- Technological limits
 - Expensive
 - Difficult to use

Barriers for correction

Cost of lenses



2 €

Access to prescription

- Trained optometrists
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 - 1 optometrist: 8000 people
 - Ghana
 - 1 optometrist: 500,000 people
- Technological limits
 - Expensive
 - Difficult to use

Solution

A simple and
autonomous device
that prescribes
aberrations
automatically



plenoptika.com



Affects more than 5 M people worldwide

The prevalence is expected to double by 2040

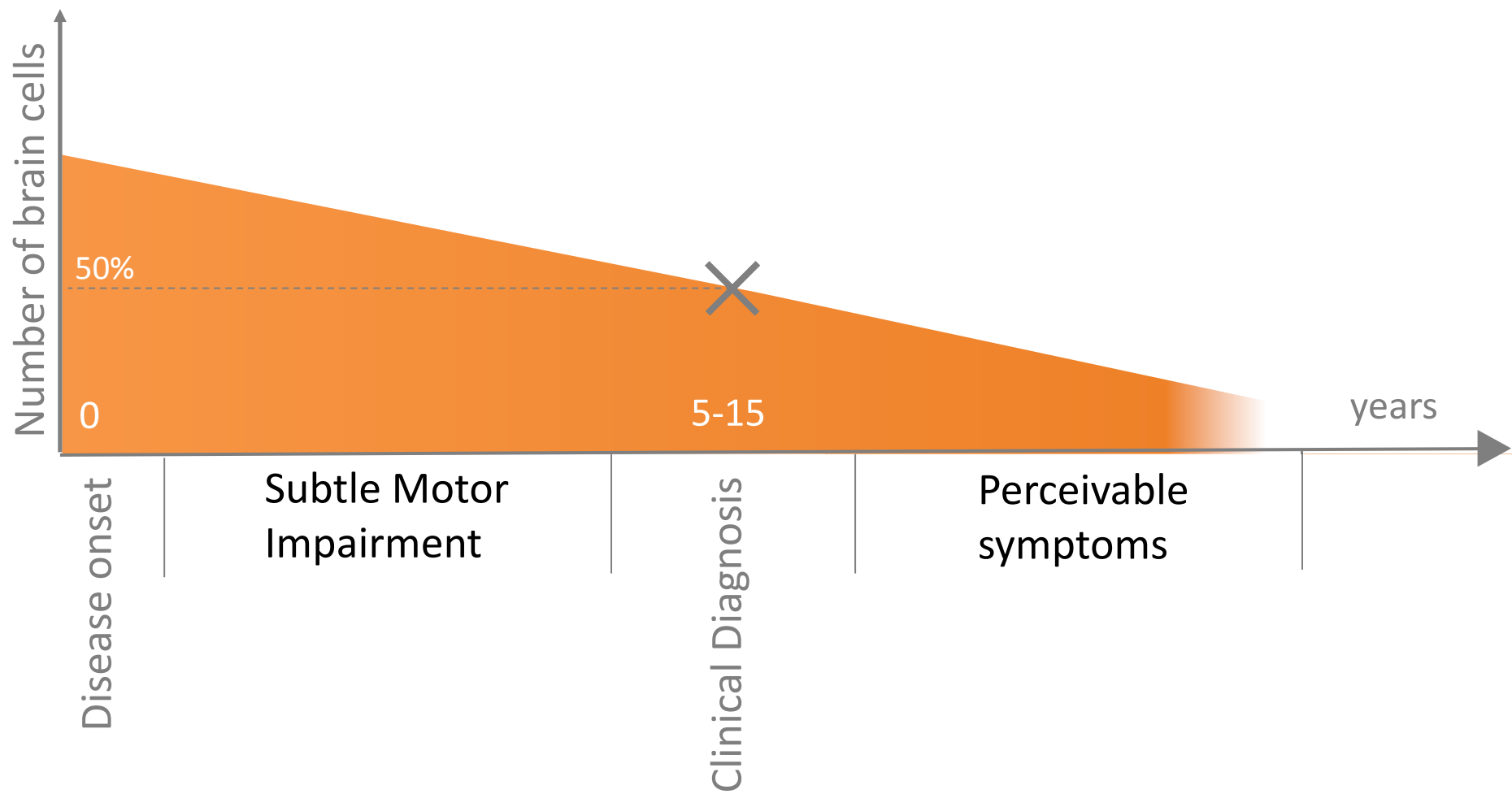
Health providers costs: \$ 29.3 Bn (US and Europe)

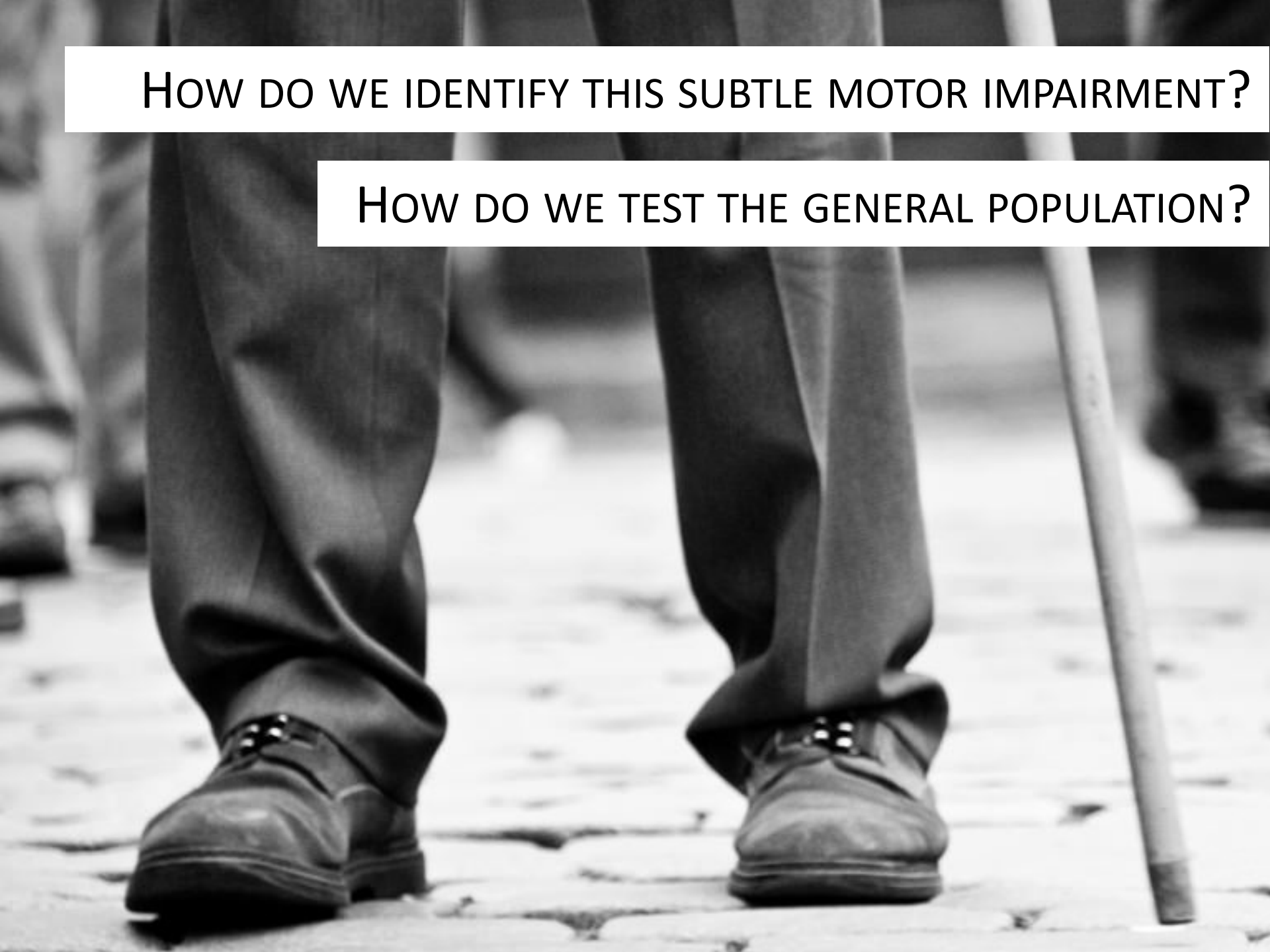
No cure exists, only symptoms are treated



The second most frequent
neurodegenerative disorder in the western world

PARKINSON'S DISEASE





HOW DO WE IDENTIFY THIS SUBTLE MOTOR IMPAIRMENT?

HOW DO WE TEST THE GENERAL POPULATION?

We have the neurological
information at our
fingertips





and we collect it all the time...



and we collect it all the time...



neuro

Q

W

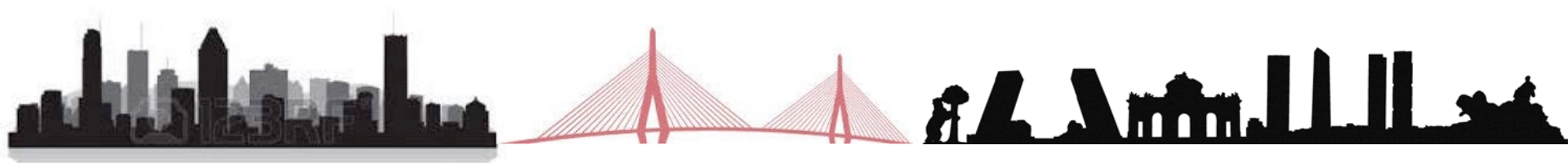
E

R

T

Y

*A transparent tool to detect prodromic symptoms
in Parkinson's Disease*



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i+12
Instituto de Investigación
Hospital 12 de Octubre

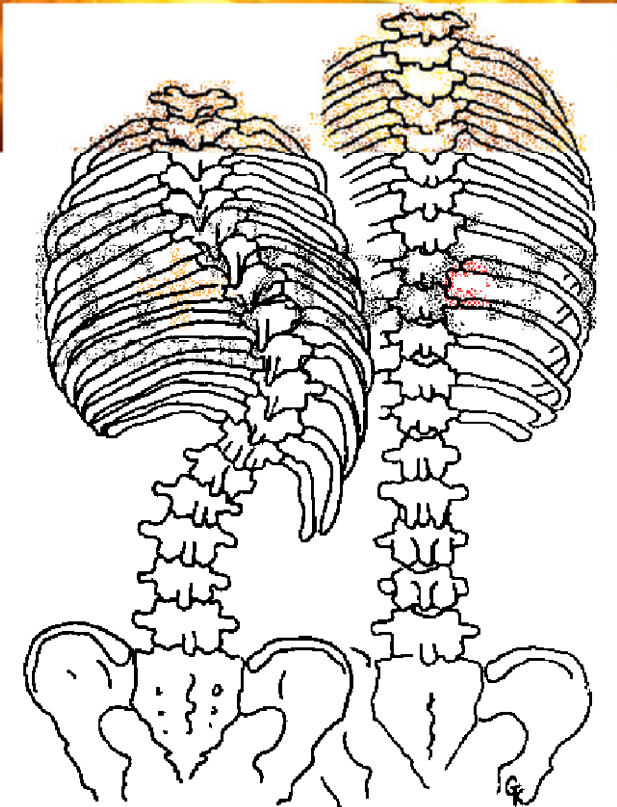
IdISSC
INSTITUTO de
INVESTIGACIÓN SANITARIA
Hospital Clínico San Carlos



earlybird.io

ESCOLIO

UN NUEVO PARADIGMA TERAPEUTICO PARA LA ESCOLIOSIS



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2-3% of adolescents
suffer some degree of
SCOLIOSIS



Attack the cause

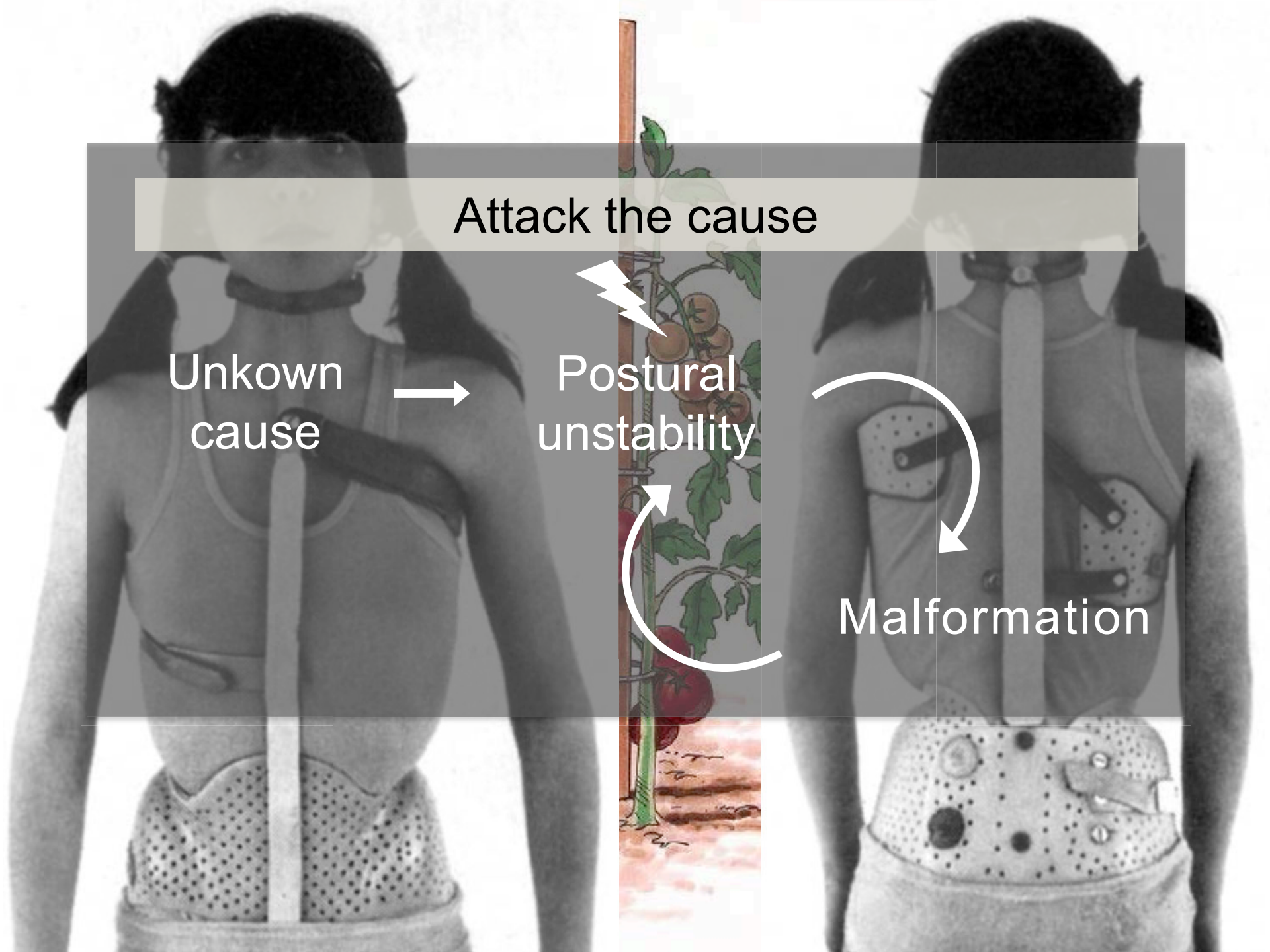
Unkown
cause



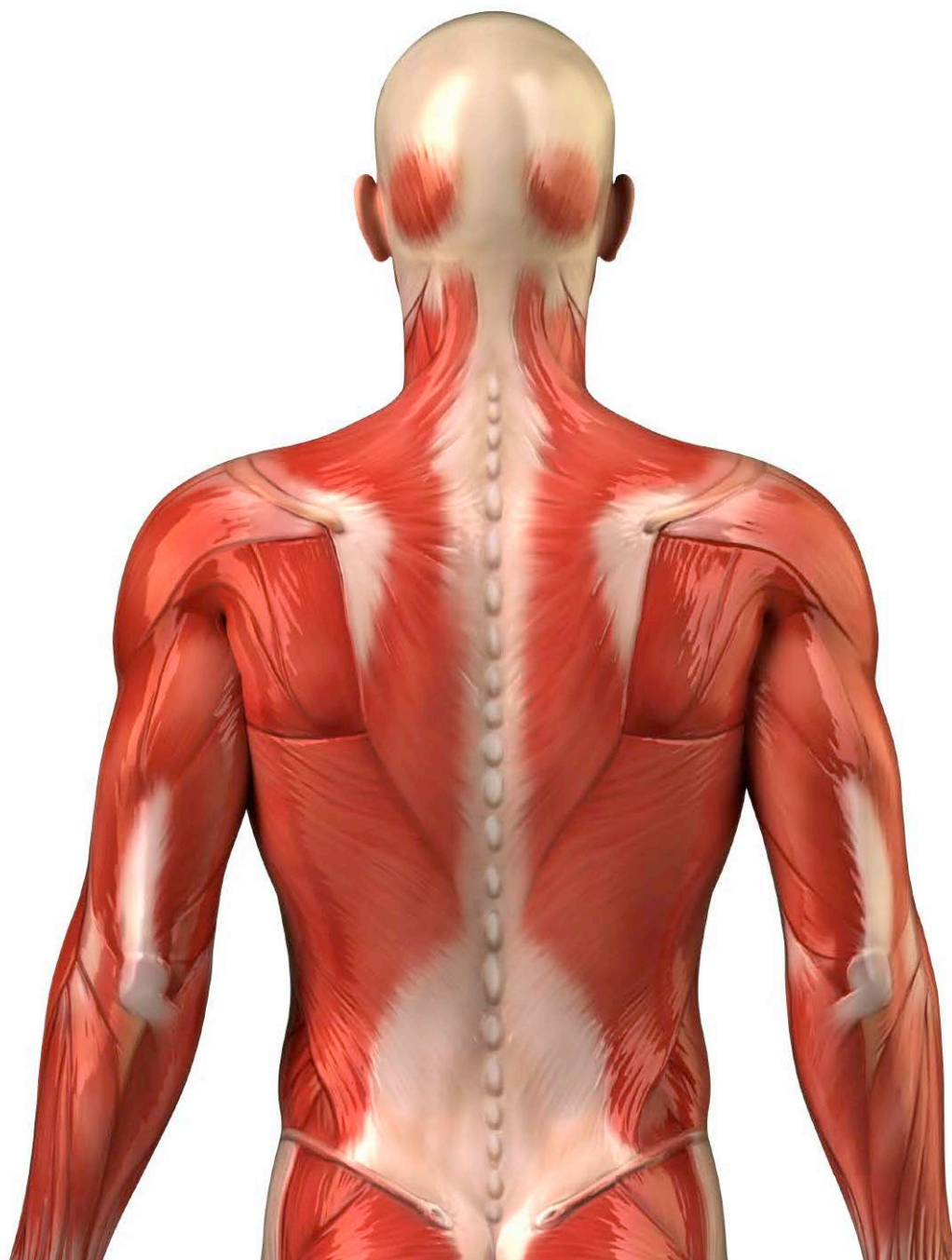
Postural
unstability



Malformation









*"Congratulations,
Maria. Your
posture is almost
perfect"*



Orientation
sensor



Carlos S. Mendoza

PhD

Aurélien Bourquard

PhD

Luca Cignea PhD



Alicia Trallero, DP



asanaweartech.com

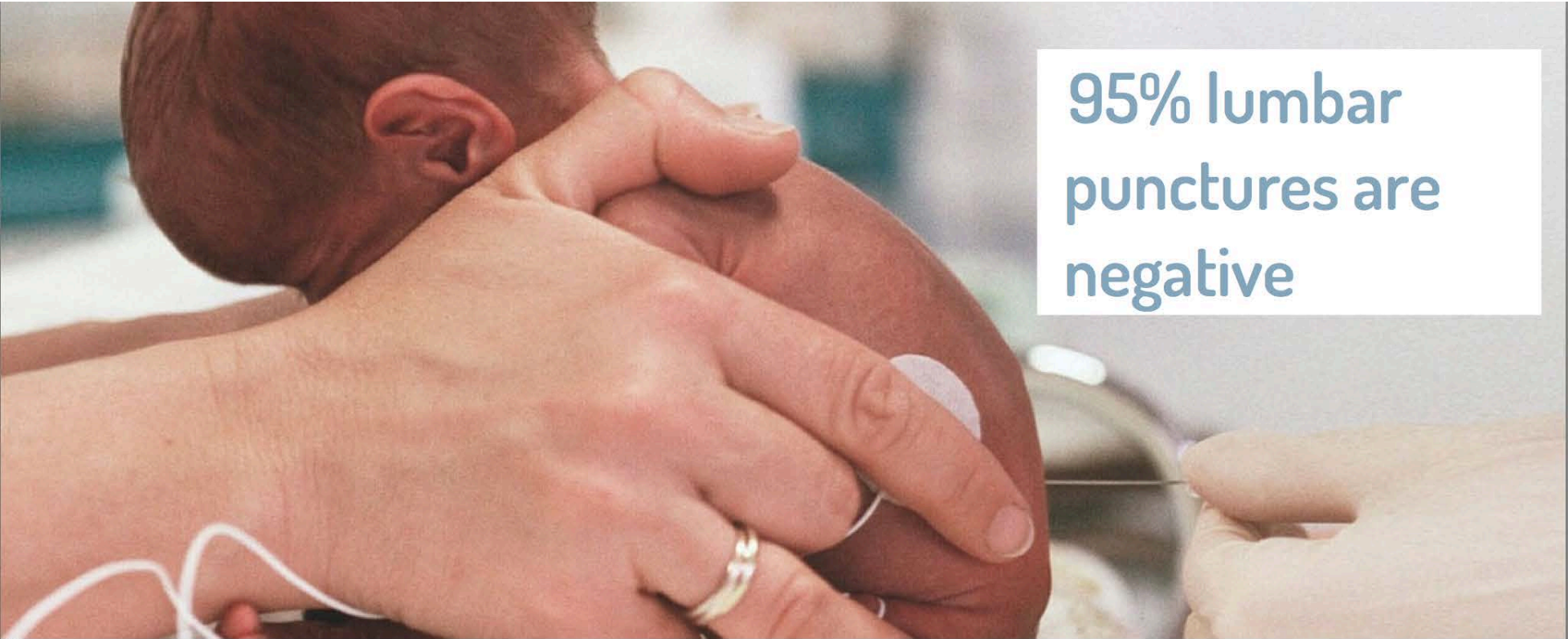


164 newborns die
every day



Bacterial Meningitis is a potentially lethal infection provided it is promptly detected and treated, difficult in the asymptomatic baby

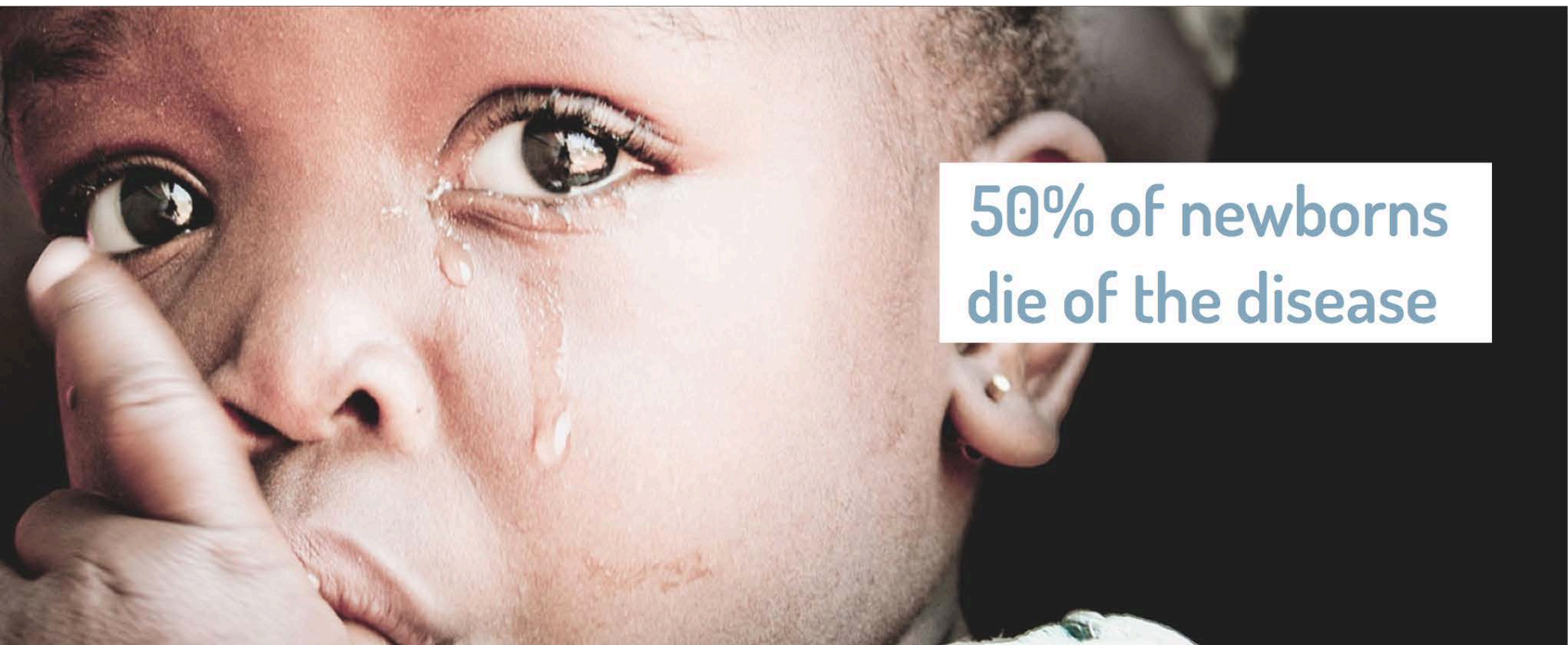
NEWBORN
SOLUTIONS



95% lumbar
punctures are
negative

Lumbar puncture (LP) is the only to draw a sample of cerebrospinal fluid (CSF).
If CSF white blood cell count (WBC) is high, medication is immediately administered
while waiting for diagnostic bacteriologic results, available after 48h

NEWBORN
.SOLUTIONS



50% of newborns
die of the disease

In under-resourced countries, the lack of laboratory facilities to analyse the sample leads to increased misdiagnosis and mortality

NEWBORN
.SOLUTIONS

Neosonics

The first device to non-invasively screen for infant meningitis

Non-invasive measurement of CSF WBC through the infant fontanel, the area of the head where bones are not fully closed.



Non-invasive

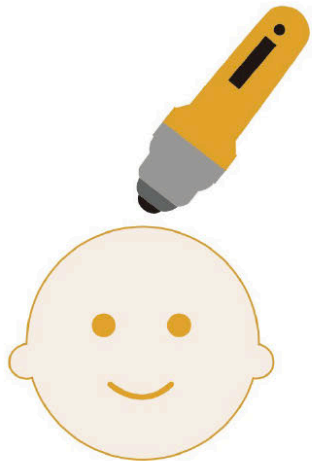
Real time

At the push of a button

Treatment monitoring

Cost-effective

PLACE, PUSH, DETECT



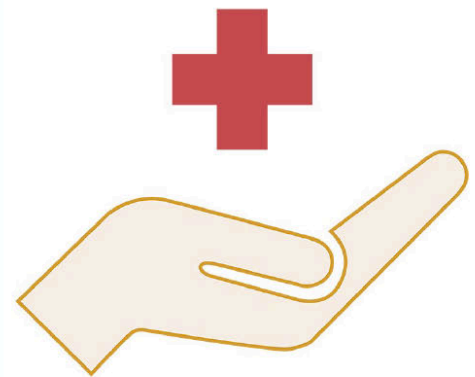
PLACE

on the infant fontanel,
the head area where
bones are not yet closed



PUSH THE BUTTON

to get the cerebrospinal
fluid (CSF) white blood
cell (WBC) count



DETECT

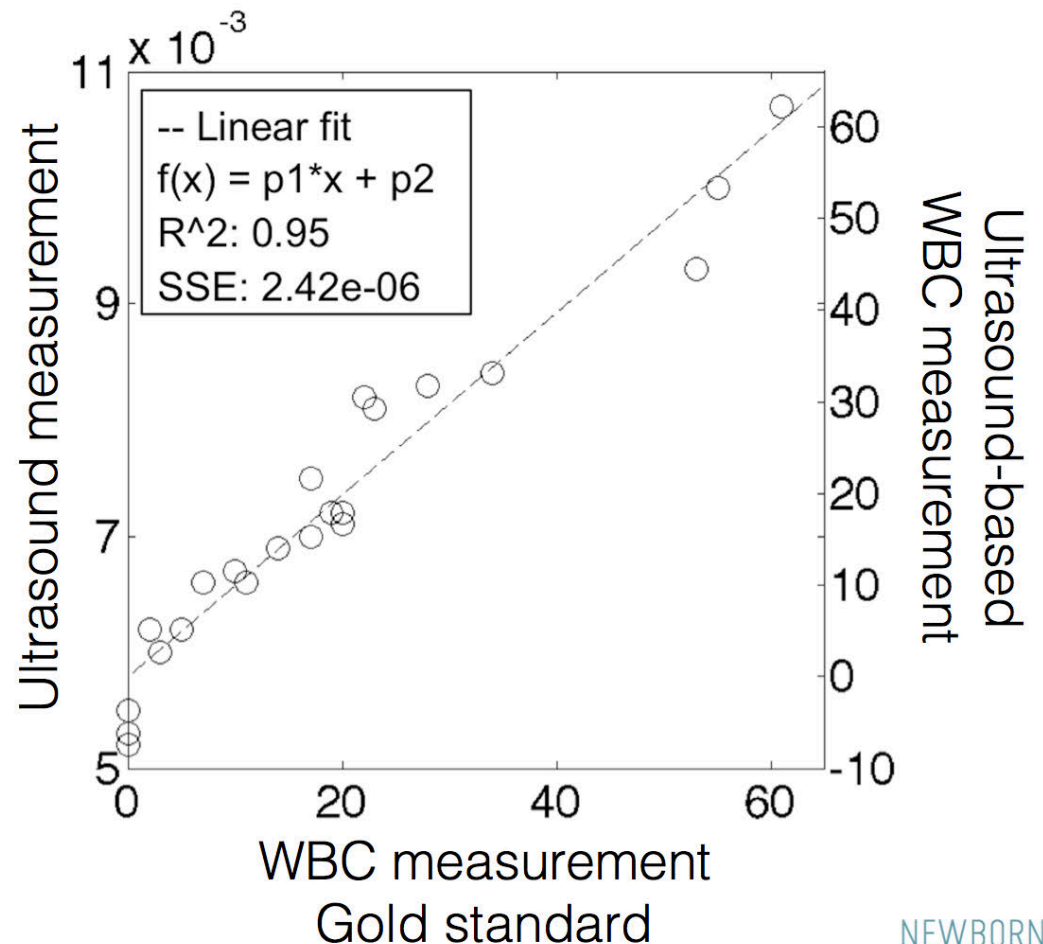
abnormal CSF WBC count to
immediately trigger medication
or to follow-up patient's
response to treatment

Results

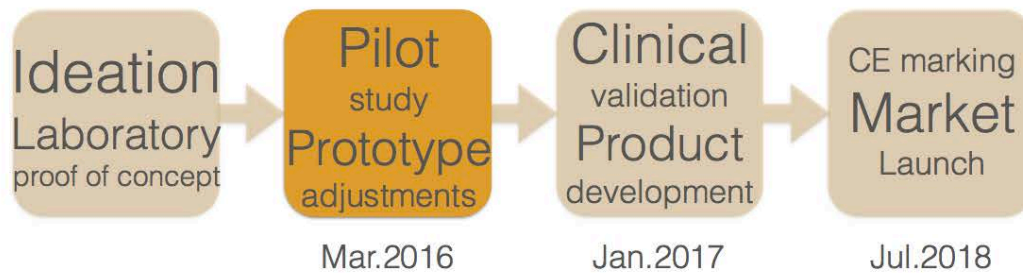
Scientific publication

Journal of Ultrasound
in Medicine and Biology
Approved Jan. 2016

Precise detection of any
WBC concentration in the
meningitis diagnostic range

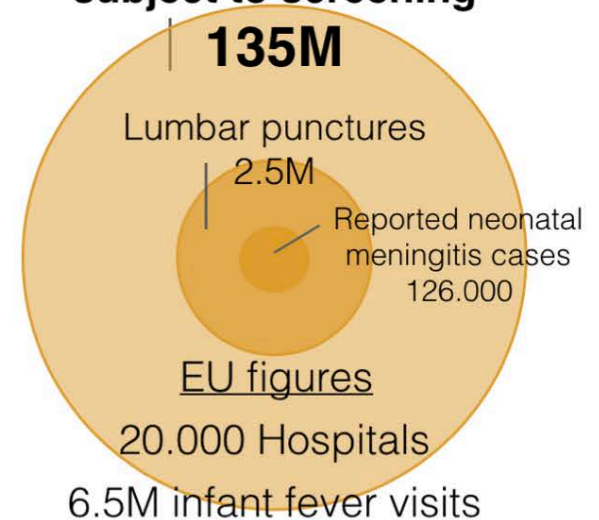


Project status



Market

**Infants with meningeal signs
subject to screening**



Team



Javier Jiménez
Founder and CEO
MIT trainee | EAE MBA
Biomedical Image Processing



Pablo García
CFO
Over 20 years as CFO
Business Expert, Economist



Luis Elvira
Principal Investigator, CSIC
23 years expertise in ultrasounds
Madrid, Spain



Adelina Pellicer
Director of Neonatology
Hospital La Paz
Madrid, Spain



Fernando Cabañas
Director of Paediatrics
Hospital Quirón
Madrid, Spain



Quique Bassat
Paediatrician, Epidemiologist
ISGlobal, Global Health
Barcelona & Mozambique

Patent



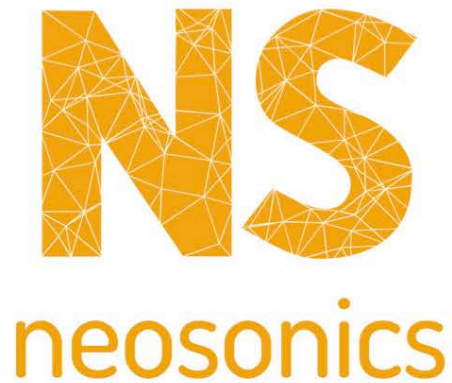
Industrial partner



Clinical partners



NEWBORN
SOLUTIONS



The first hand-held device to non-invasively screen for infant meningitis

Contact information

javier.jimenez@newborn.solutions

+34 626 536 144

www.newborn.solutions

NEWBORN
.SOLUTIONS



Leucemia y linfoma

>130,000
diagnósticos
estimados¹

44,000
muertes¹

>1 millón de
personas

viviendo con
ello¹

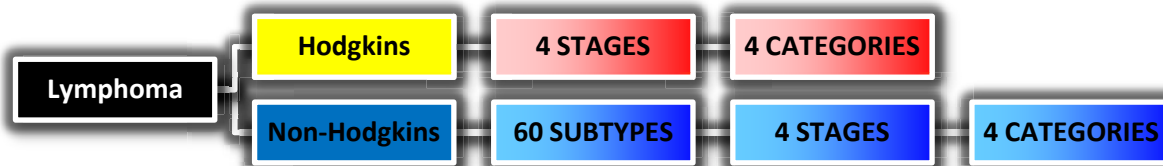
**El diagnóstico preciso de la
clase y subtipo de la
enfermedad es crítico para
un tratamiento correcto y
eficiente^{1,2}**



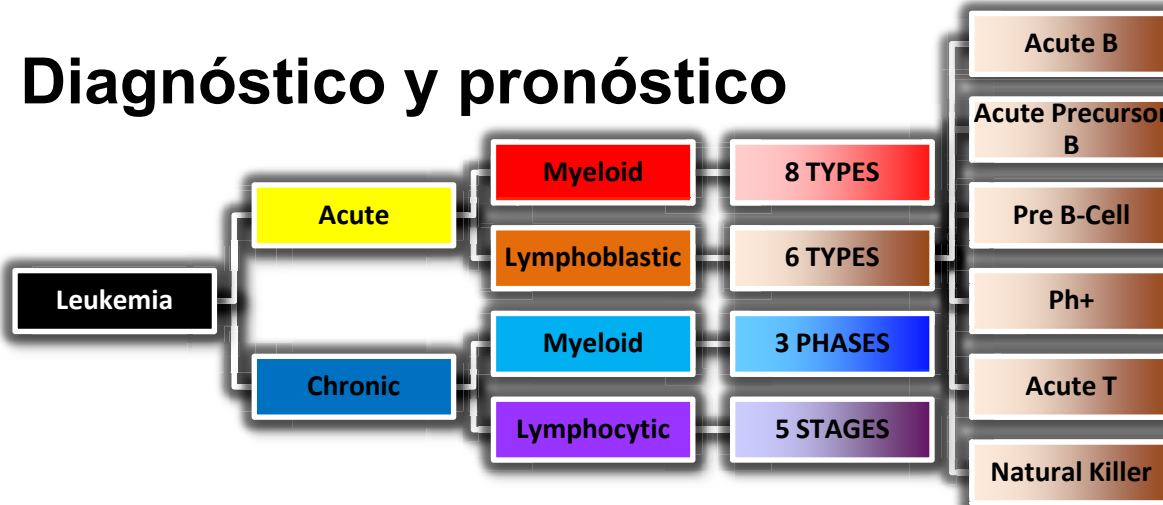
*USA Statistics, 2014

[1] Leukemia and Lymphoma Society Fact Sheet, 2014
[2] National Comprehensive Cancer Network. Guidelines. 2014

Disease Classification



Diagnóstico y pronóstico

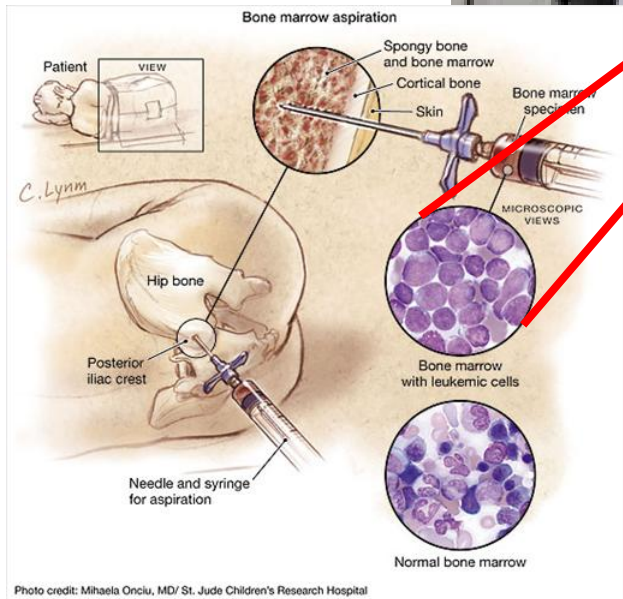


Treatment Plan



Amplio espectro de tipos de enfermedad y fenotipos que deben ser detectados antes del tratamiento

Fenotipado cualitativo: Microscopía



Muestras de sangre/marcado
Identificación de morfología, tipo celular, fenotipo

Cualitativo

Baja eficiencia

Fenotipado cualitativo: Citometría de flujo

Processed Cells



Flow Cytometer



Cell Characterization

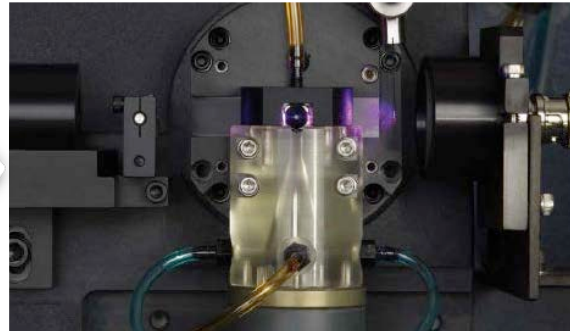


High throughput
Análisis cuantitativo

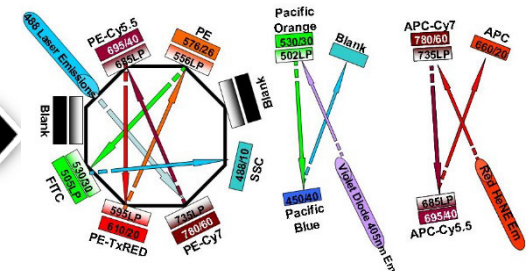
Laser Illumination



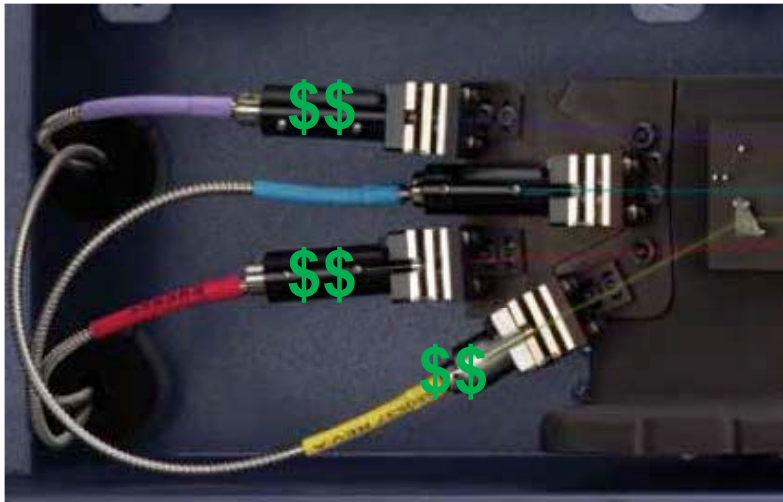
Fluidics System



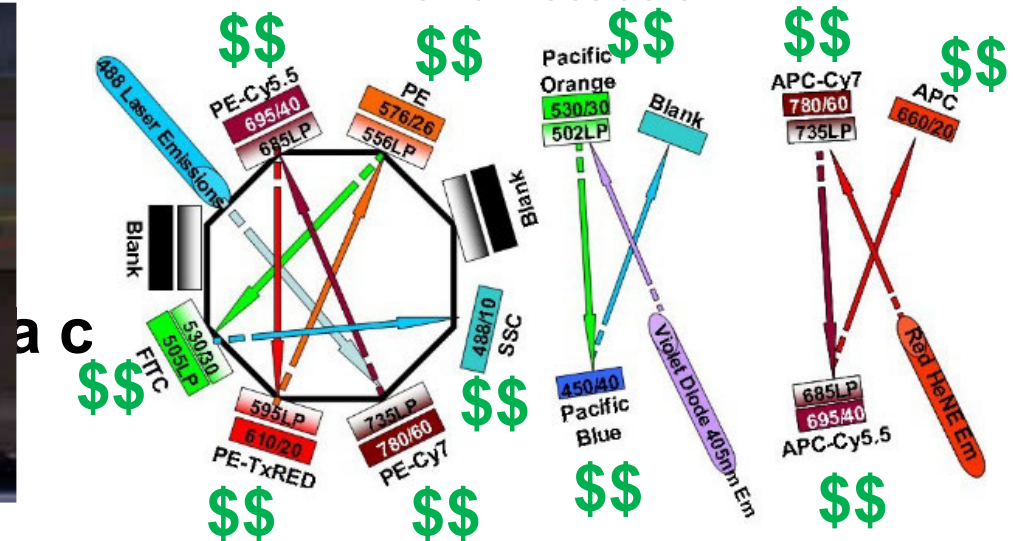
Cell-by-Cell
Event Detection



Laser Illumination



Cell-by-Cell Event Detection



Multi (8+) color flow cytometry requires extensive components for illumination, flow, and detection:

- High cost
- High complexity

Presencia baja de la citometría



**Alto coste,
complejidad**

Poco disponible para el usuario final

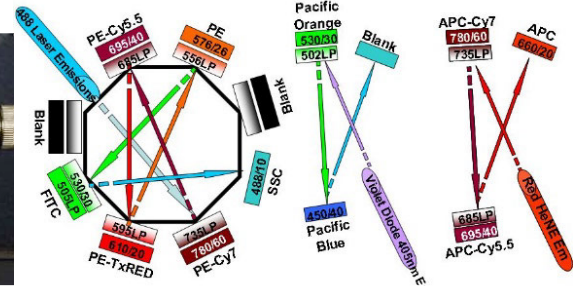
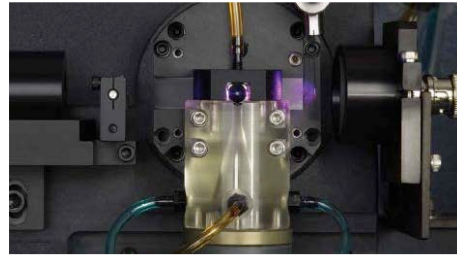
>75% de hospitales sin citómetro de flujo

El paciente debe acudir a un centro especializado

Retraso en el diagnóstico

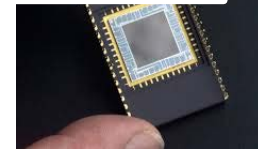
Retraso en el diagnóstico comercial

Degradación de la muestra



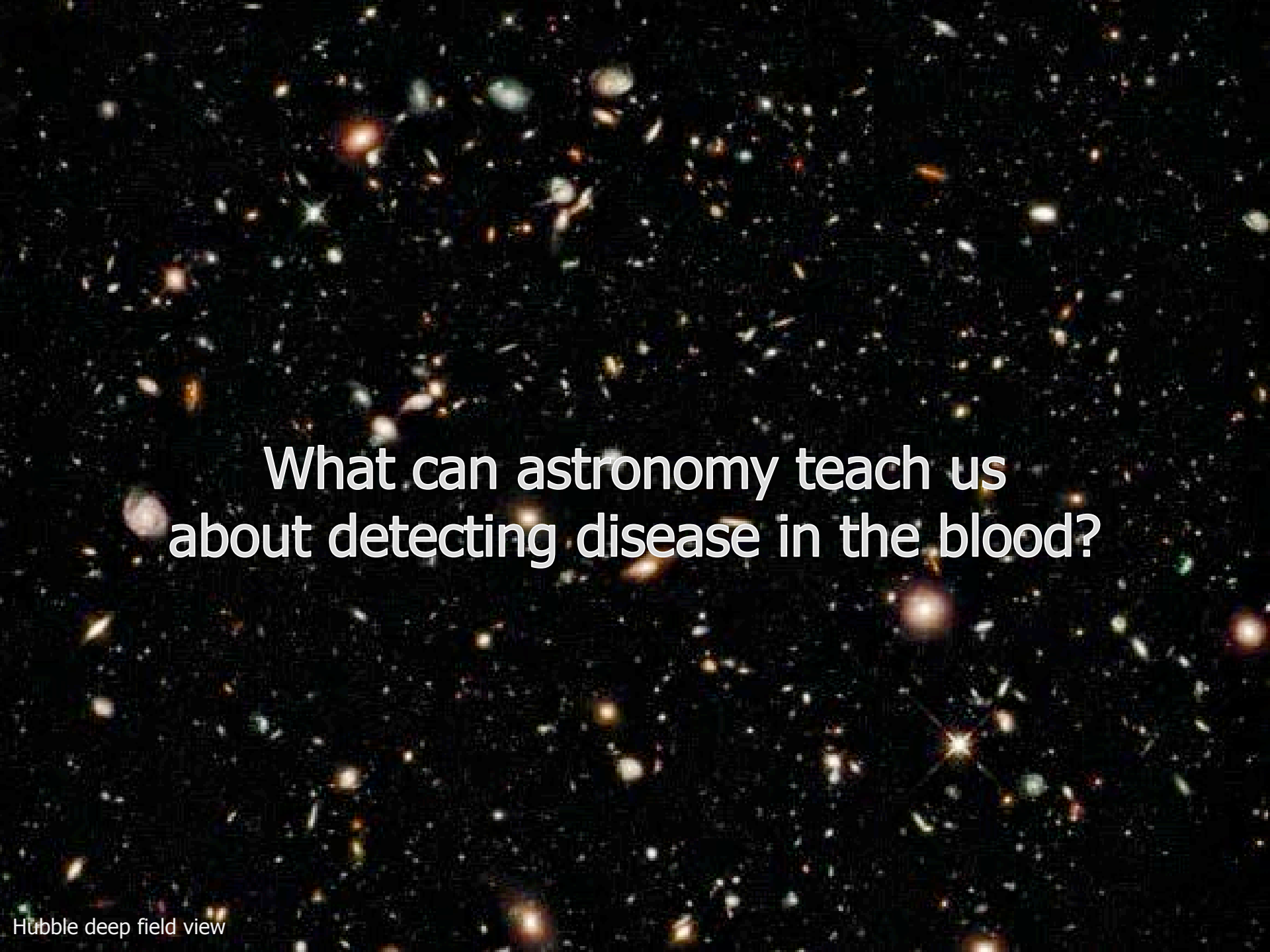
¿Por qué no usamos un microscopio para hacer citometría sin flujo de alto rendimiento?

Citron

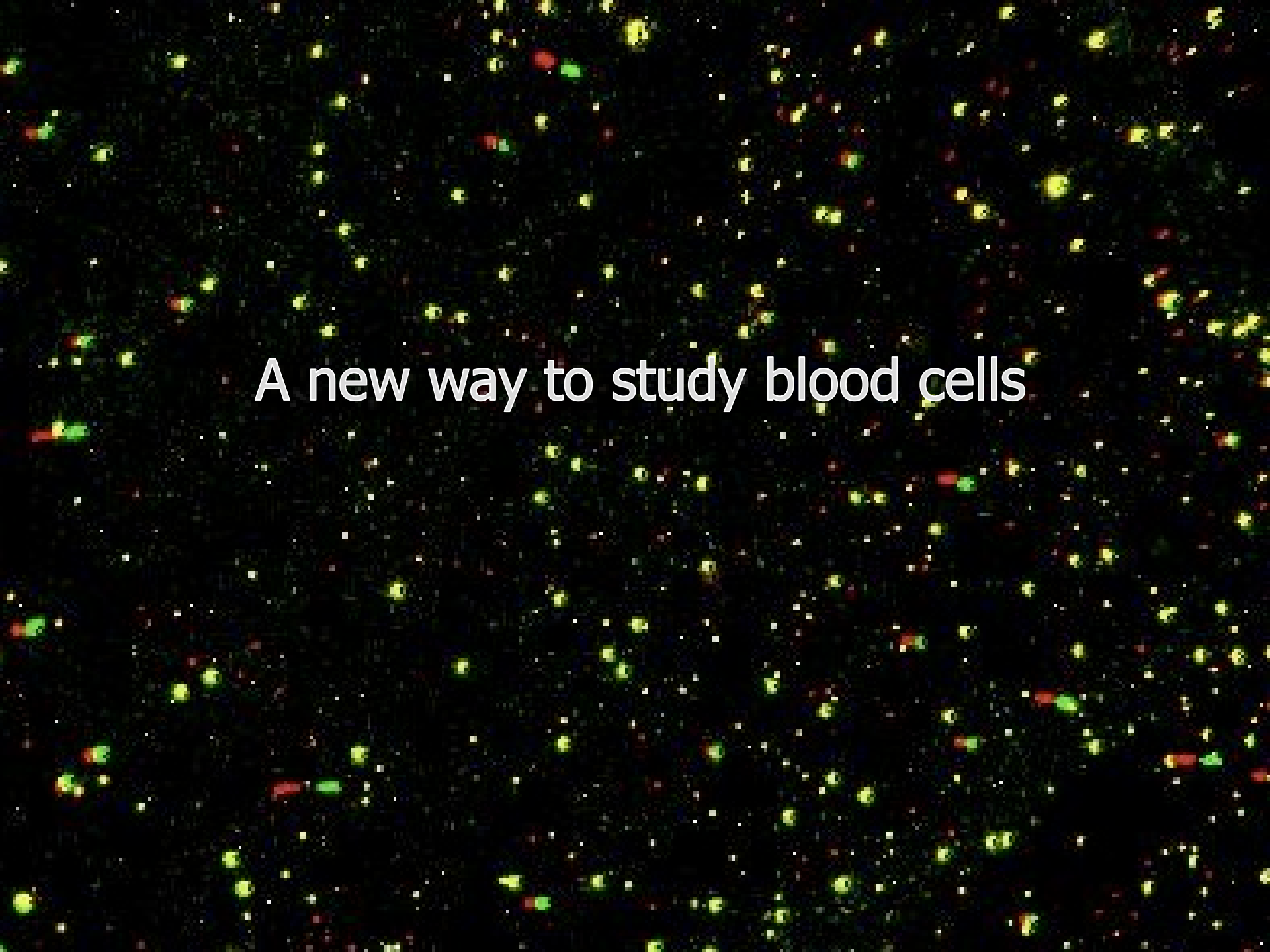


Los avances en electrónica, procesamiento de imagen, y LEDs de alta potencia permiten **la imagen cuantitativa de alto rendimiento** sensible y asequible

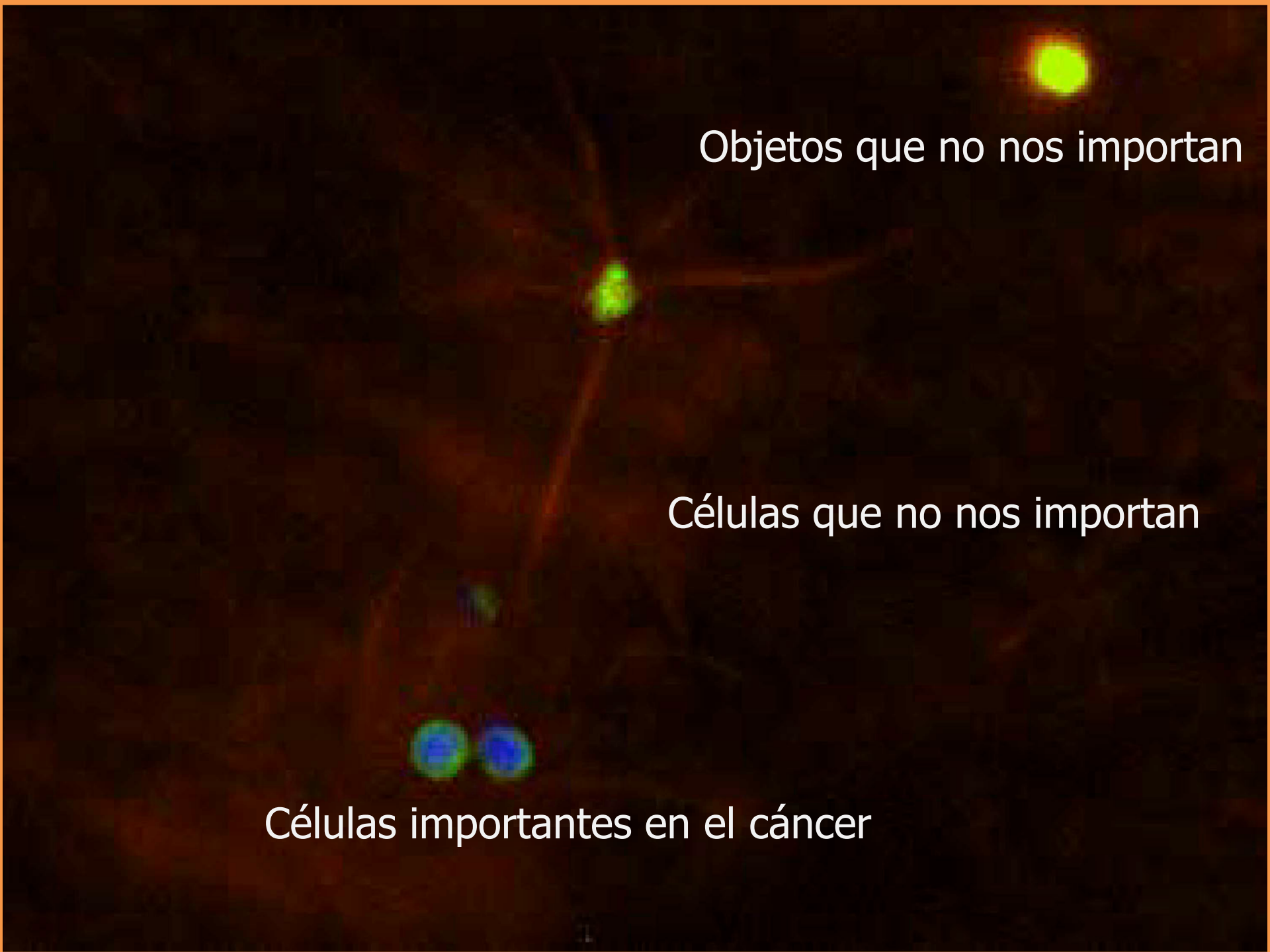


The background of the slide is a deep-field image from the Hubble Space Telescope, showing a vast field of distant galaxies. The galaxies are of various shapes and sizes, some appearing as bright, smooth ellipses and others as more complex, irregular structures. They are scattered across a dark, black background, with some showing distinct colors like orange, red, and blue. The overall effect is a sense of immense scale and depth in the universe.

What can astronomy teach us
about detecting disease in the blood?



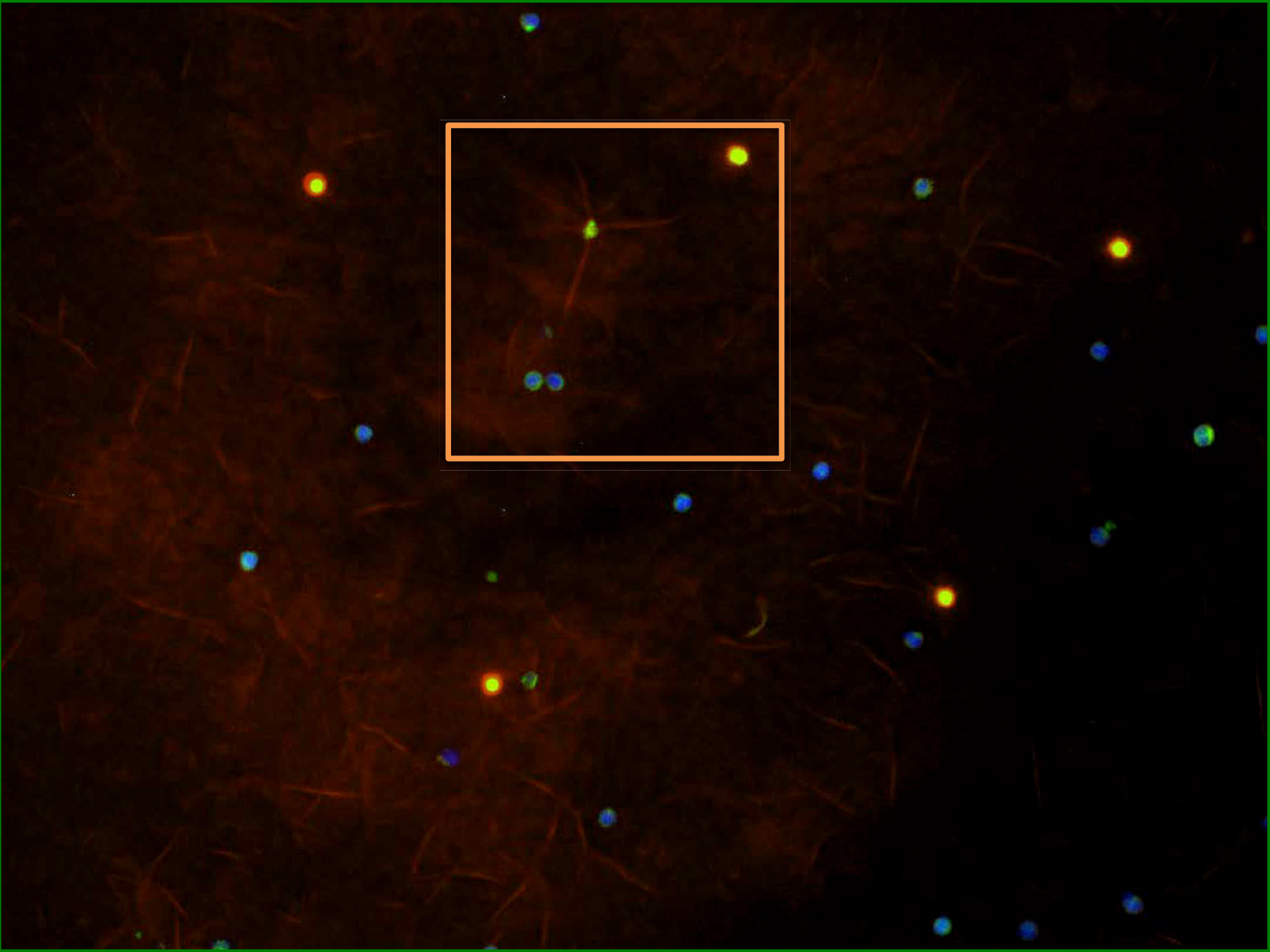
A new way to study blood cells

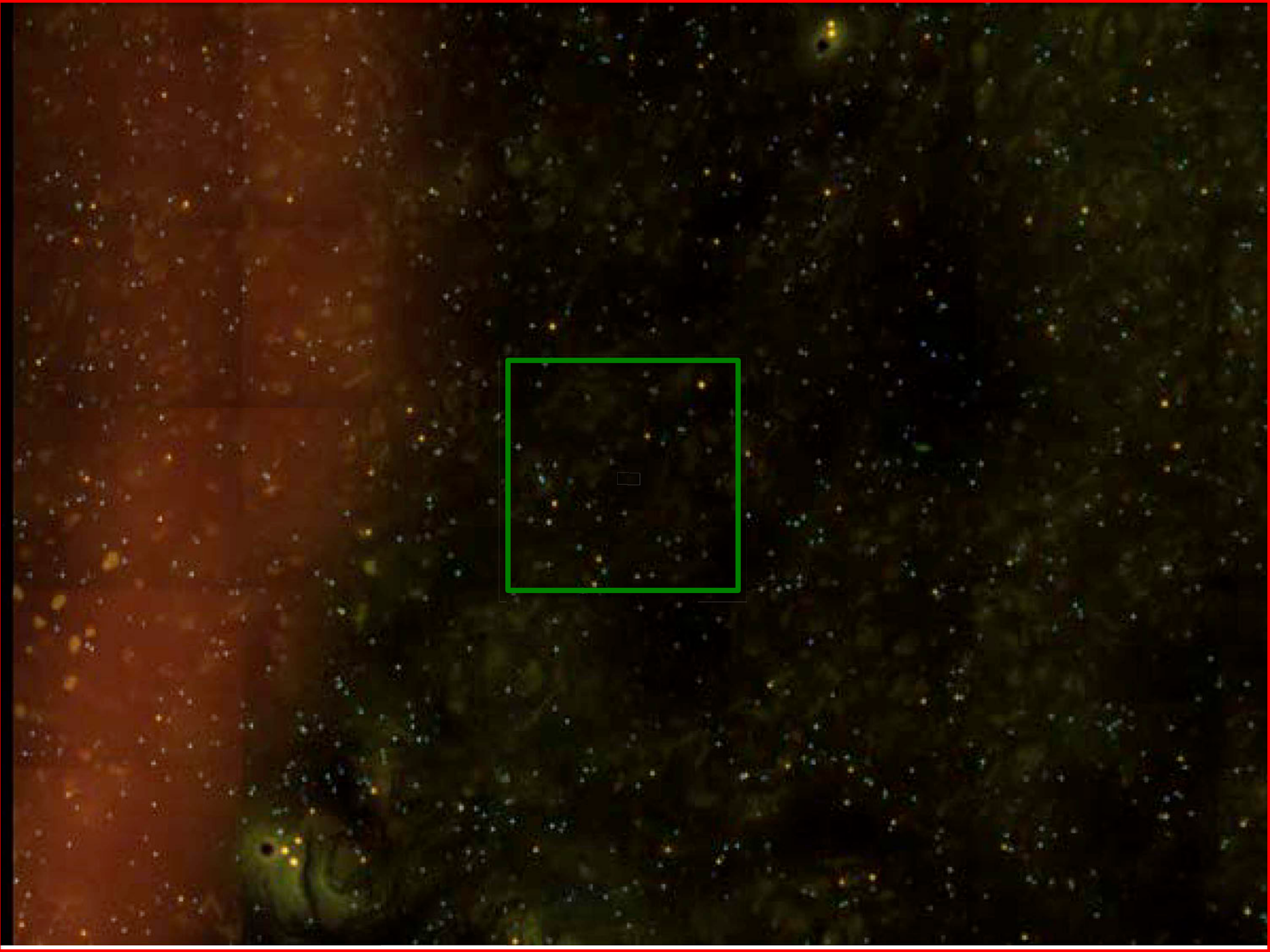


Objetos que no nos importan

Células que no nos importan

Células importantes en el cáncer

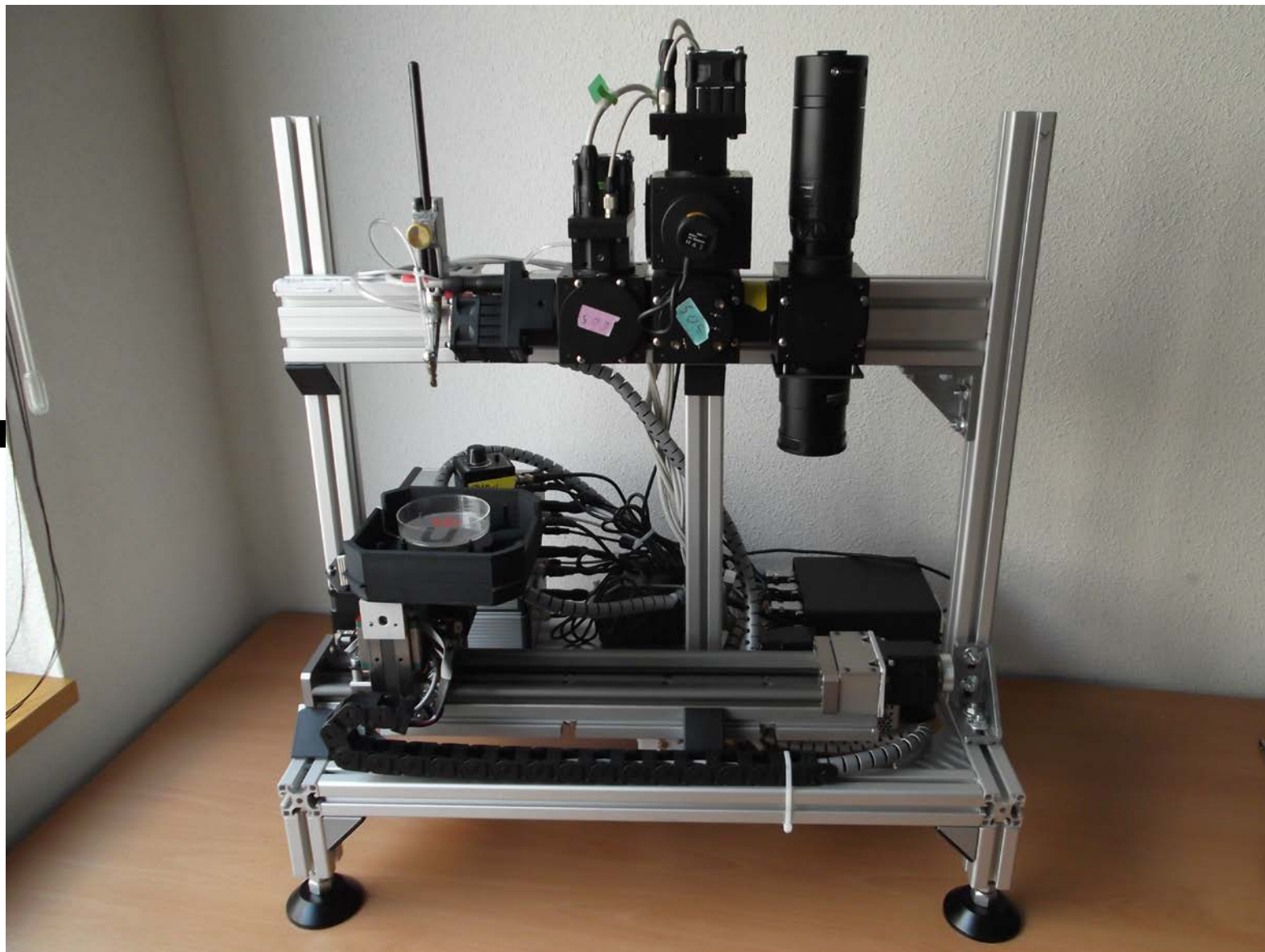


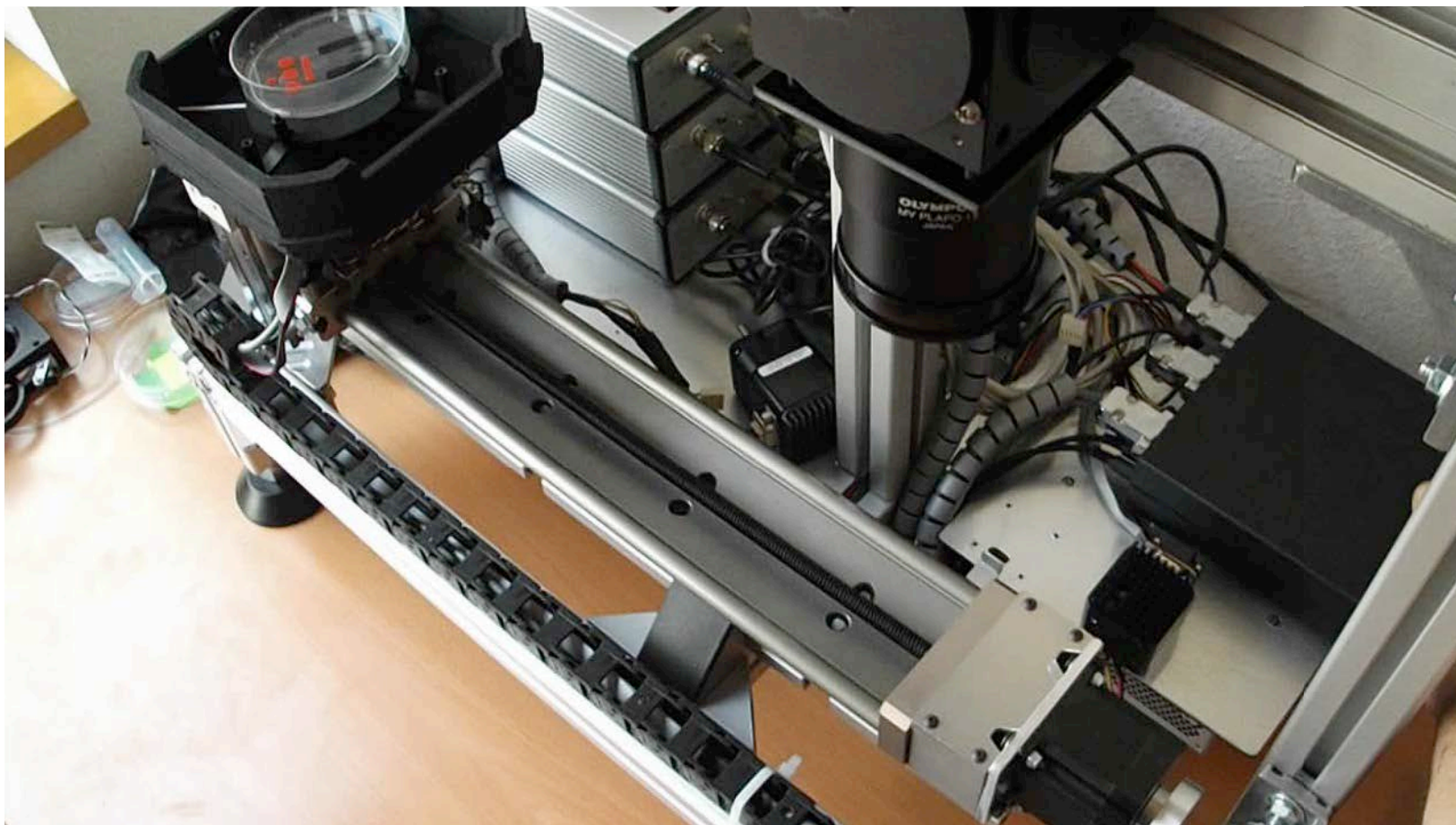


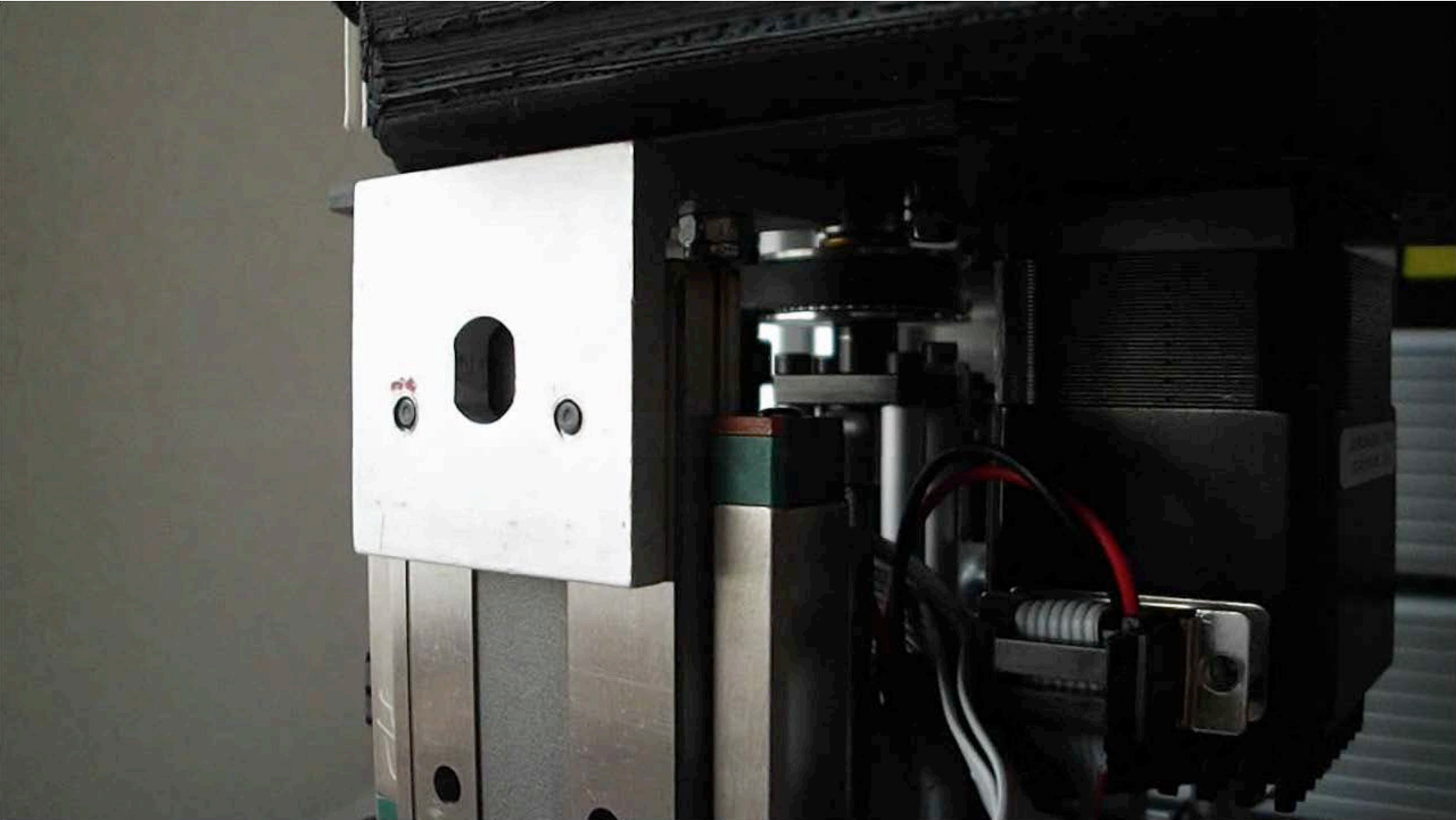
¡1.000.000 de células
detectadas en segundos!



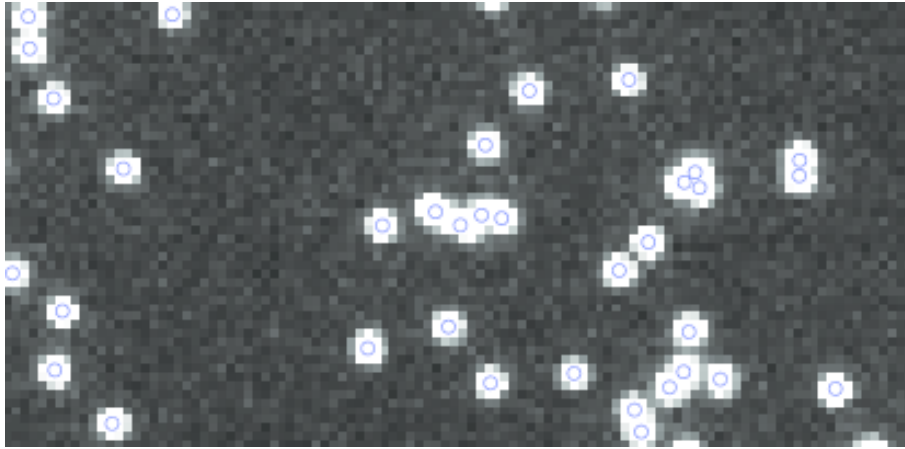
Un



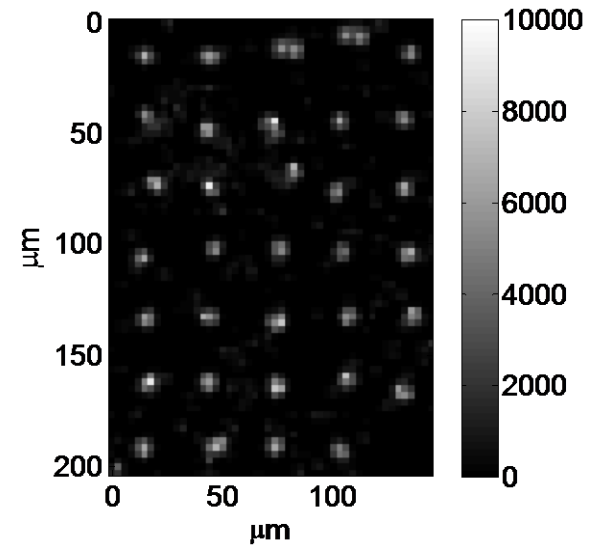
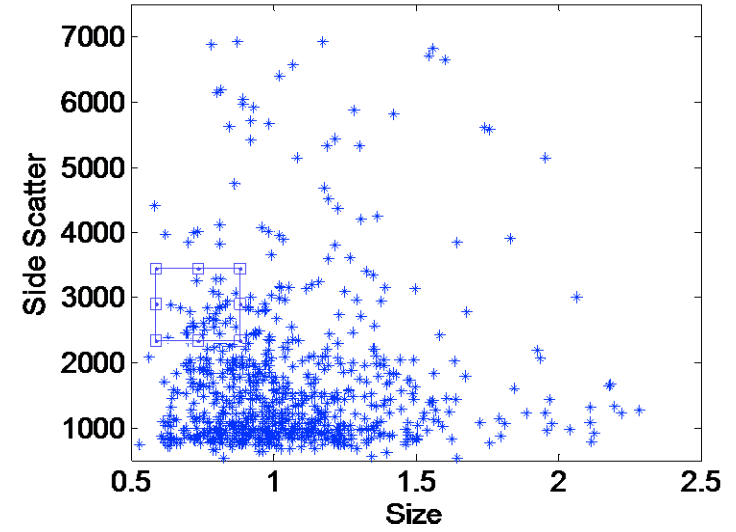
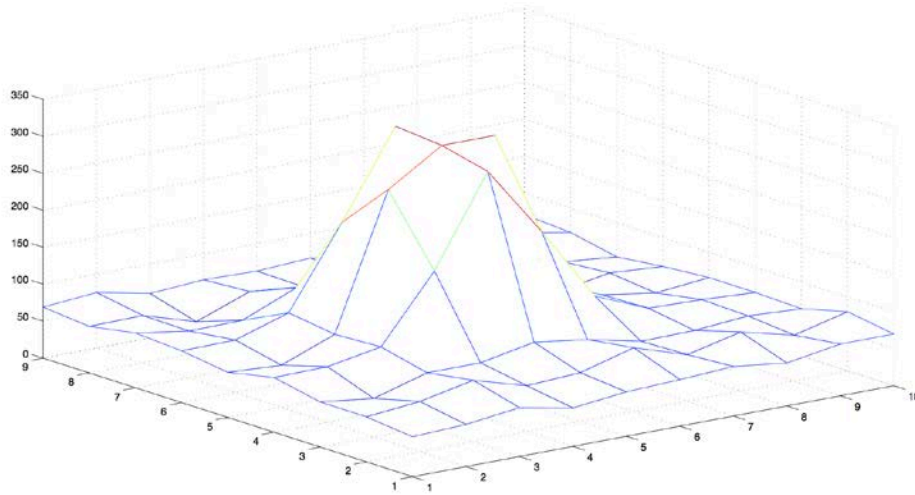








Análisis de imagen



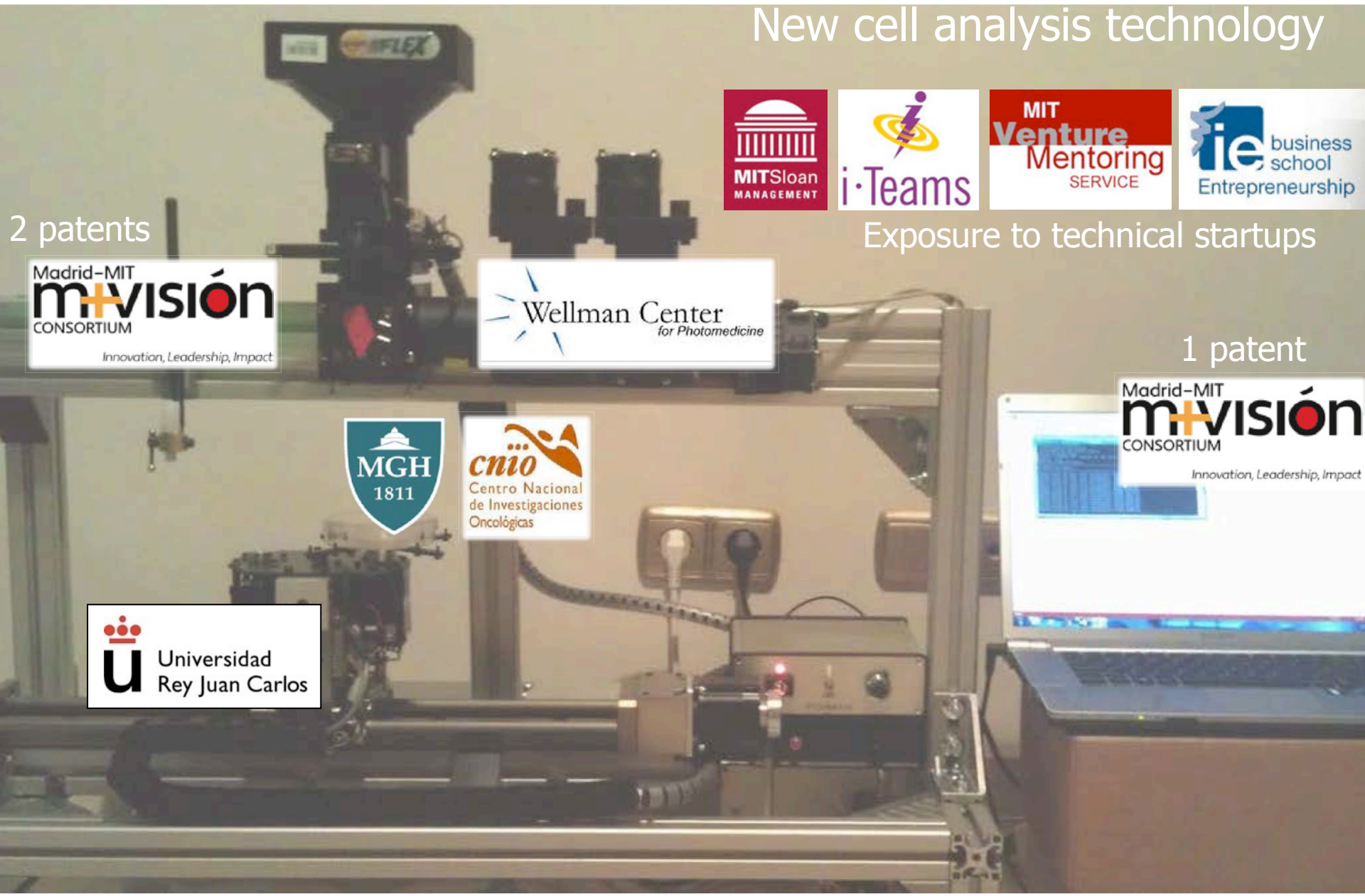
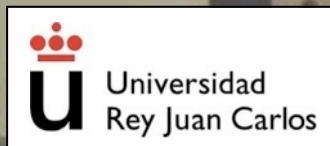
New cell analysis technology

2 patents



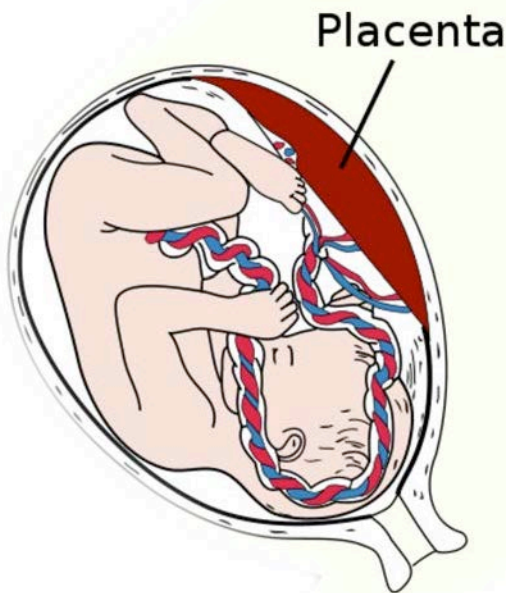
Exposure to technical startups

1 patent



4% de todos los fetos en riesgo de morir en el útero debido a un transporte insuficiente de oxígeno y nutrientes a través de la placenta

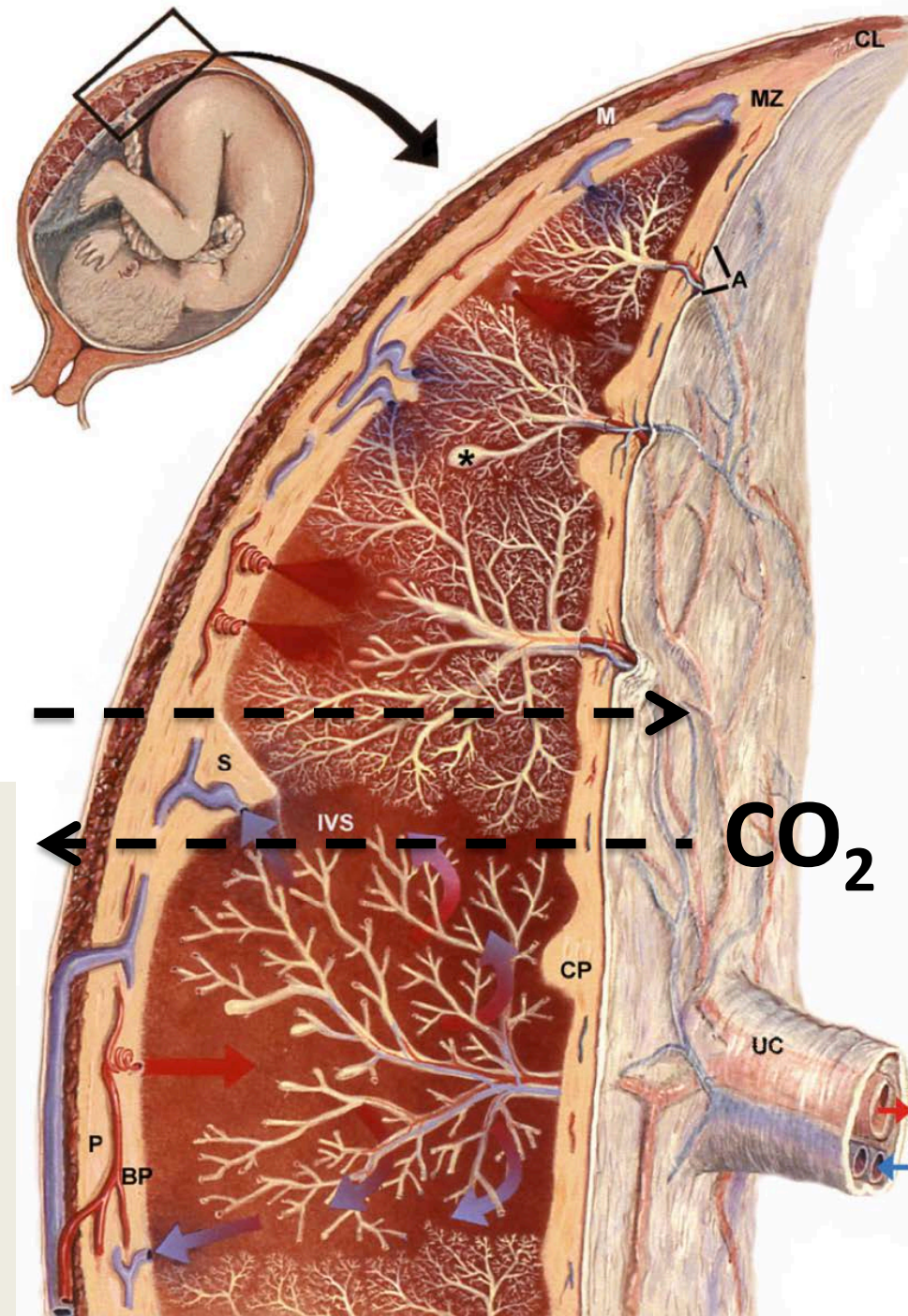
10% de los fetos en riesgo de **crecimiento intrauterino restringido (IUGR)**



Actualmente, **no existe una manera de medir** el transporte placentario.

NO hay forma de detectar **si llega suficiente O_2 y nutrientes** al feto.

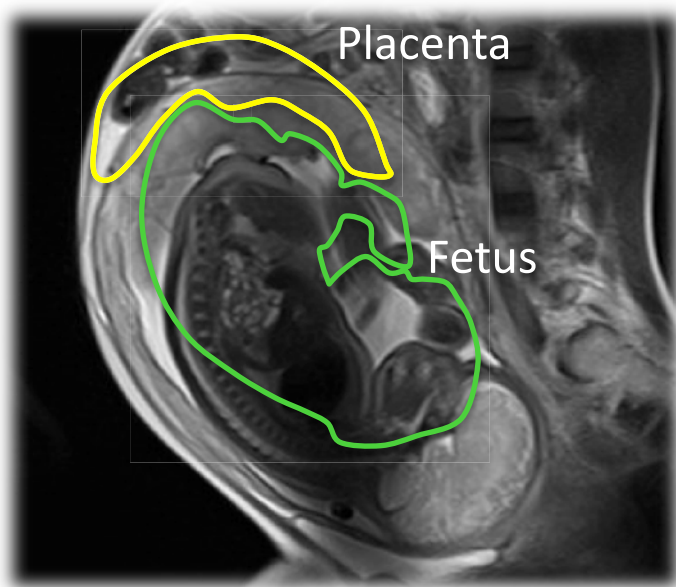
Fig. 4.6 The mature human placenta in situ is composed of the chorionic plate (CP) and the basal plate (BP) surrounding the intervillous space (IVS) as cover and as underside, respectively. The fetally vascularized villous trees project from the chorionic plate into the intervillous space and are directly surrounded by the maternal blood that circulates through the intervillous space. The loose centers of villous trees, arranged around the maternal arterial inflow area, are frequent features. *P* placental bed; *M* myometrium; *CL* chorion laeve; *A* amnion; *MZ* marginal zone between placenta and fetal membranes, with obliterated intervillous space and ghost villi; * cell island, connected to a villous tree; *S* placental septum; *UC* umbilical cord (Source: From Kaufmann and Scheffen (1992), with permission)



O_2

CO_2

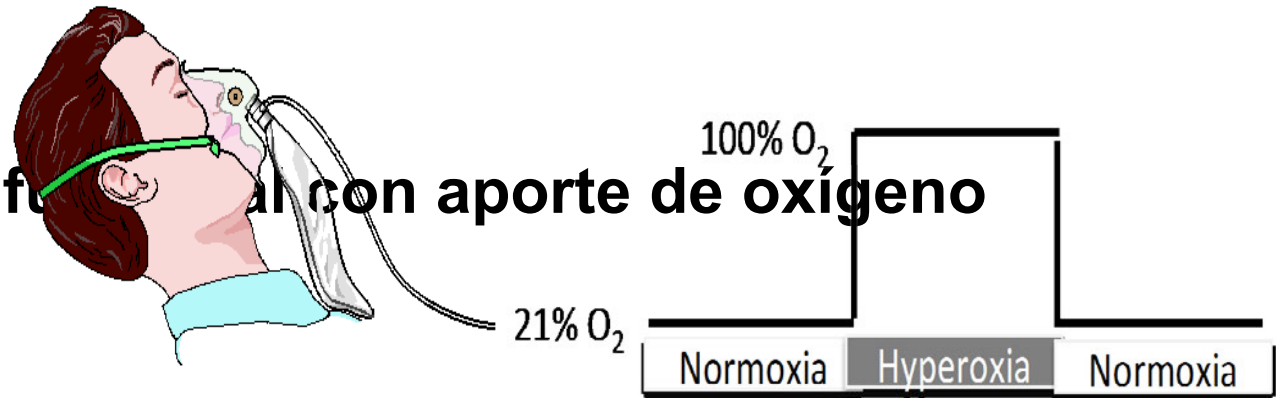
- No mixing between blood on maternal and fetal side
- O_2 carried by hemoglobin in the blood
- transport by diffusion



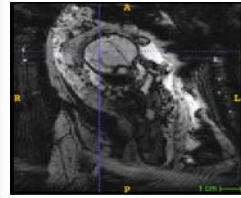
Objetivo:
Desarrollo de técnicas de resonancia para medir el transporte de oxígeno a través de la placenta



M f... al con aporte de oxígeno



1. Data Acquisition

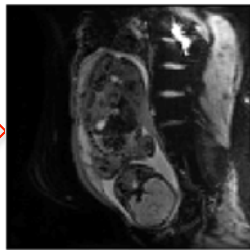
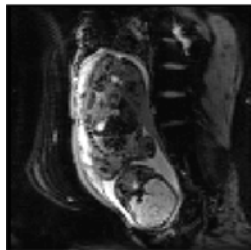


3T Scanner
2D multi slice single-shot GRE-EPI
In plane resolution: $3 \times 3 \text{ mm}^2$
Slice thickness 3 mm,
Interleaved

TR: 6 ~ 7 sec; Total 30 minutes

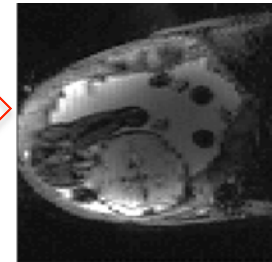
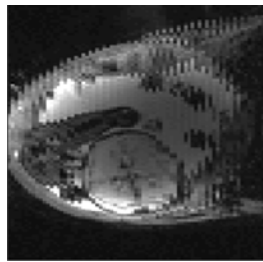
2. Data processing

Signal non-uniformity correction



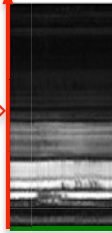
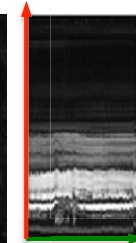
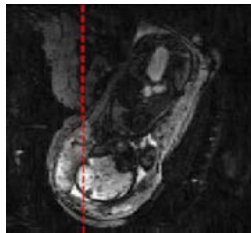
Used N4ITK based correction approach

Volume Correction



Non-rigid transformation using **Elastix^N** with customized parameter file

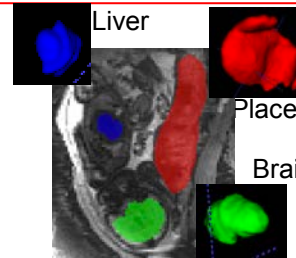
Registration and Transformation



Non-rigid transformation using **Elastix^N**

time

Manual segmentation on reference volume

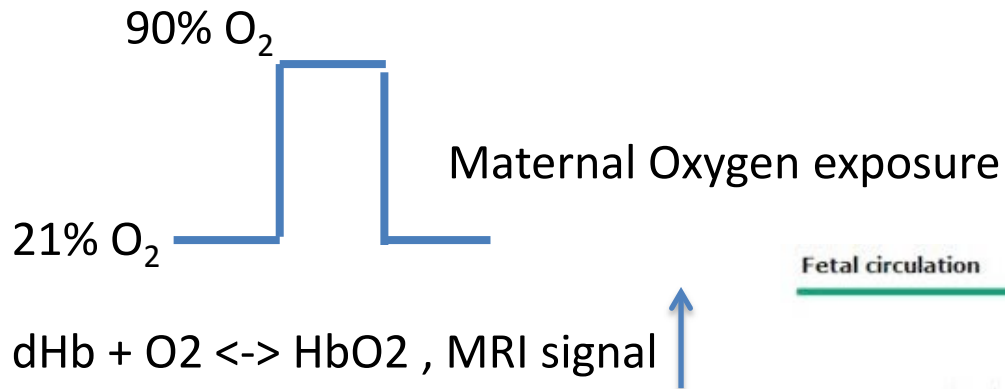


Using ITK-SNAP*

3. BOLD signal change curve generation:



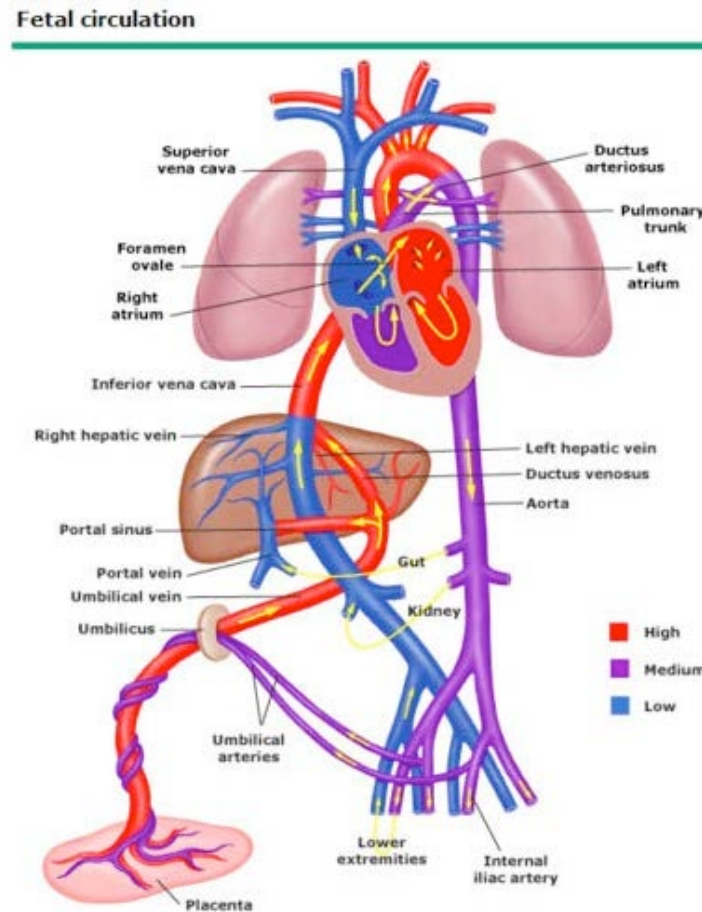
Data Acquisition-BOLD MRI with oxygen exposure



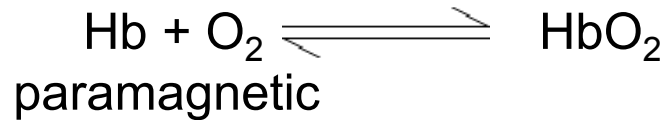
-> Placenta

-> fetal liver

-> fetal brain



OxyMRI – oxygen enhanced MRI based on BOLD effect



Oxygen Paradigm: 10 min normoxia, 10 min hyperoxia, 10 min normoxia
Total time: 30 min

Results - placenta

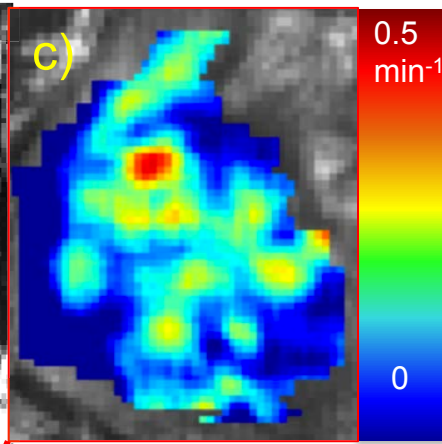
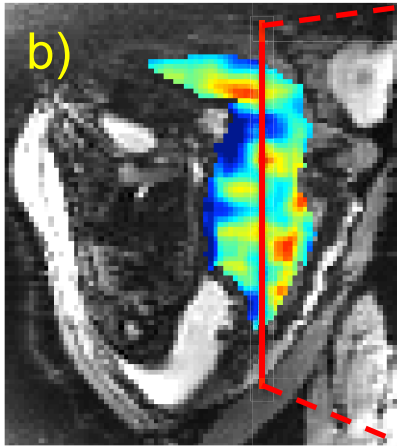
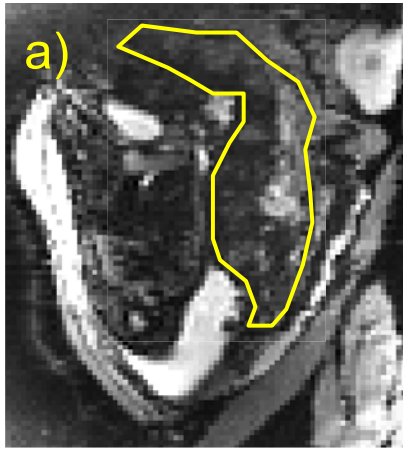
Normoxia

Hyperoxia

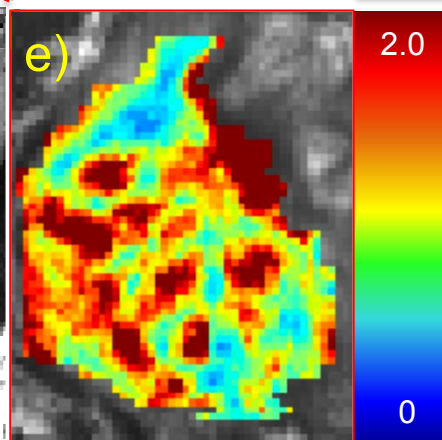
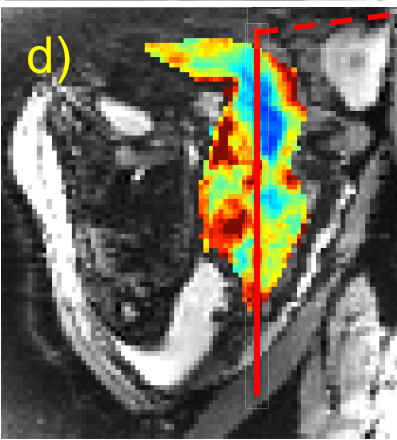
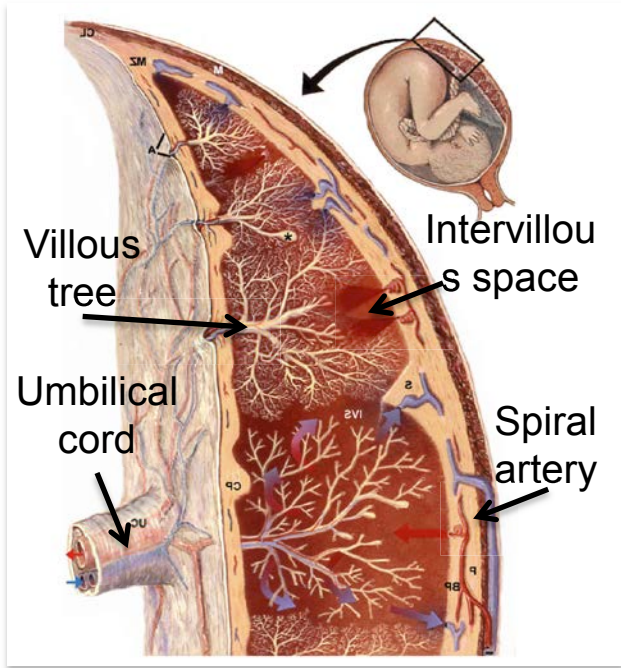
Normoxia



Heterogeneous response of placenta to hyperoxia



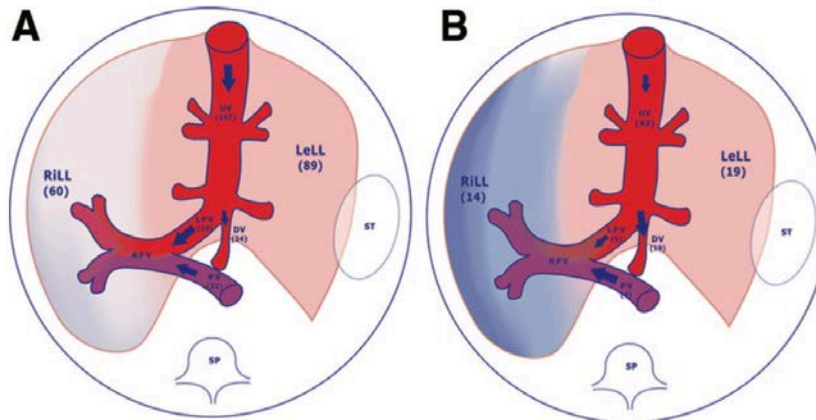
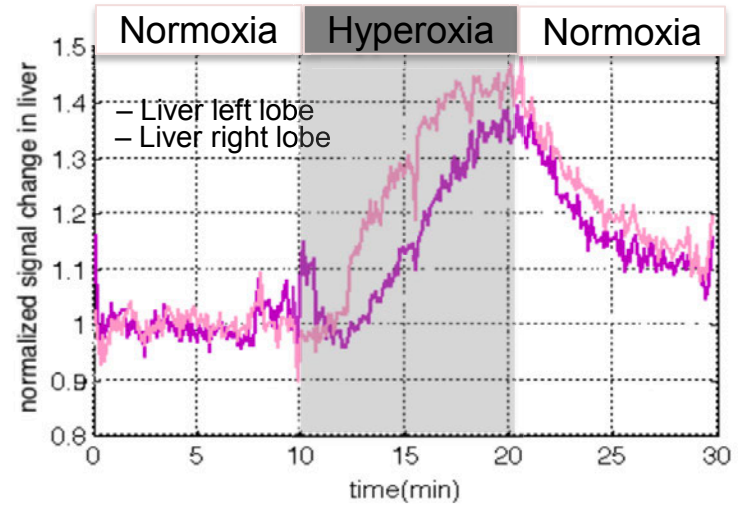
“slope map”



“residual map”

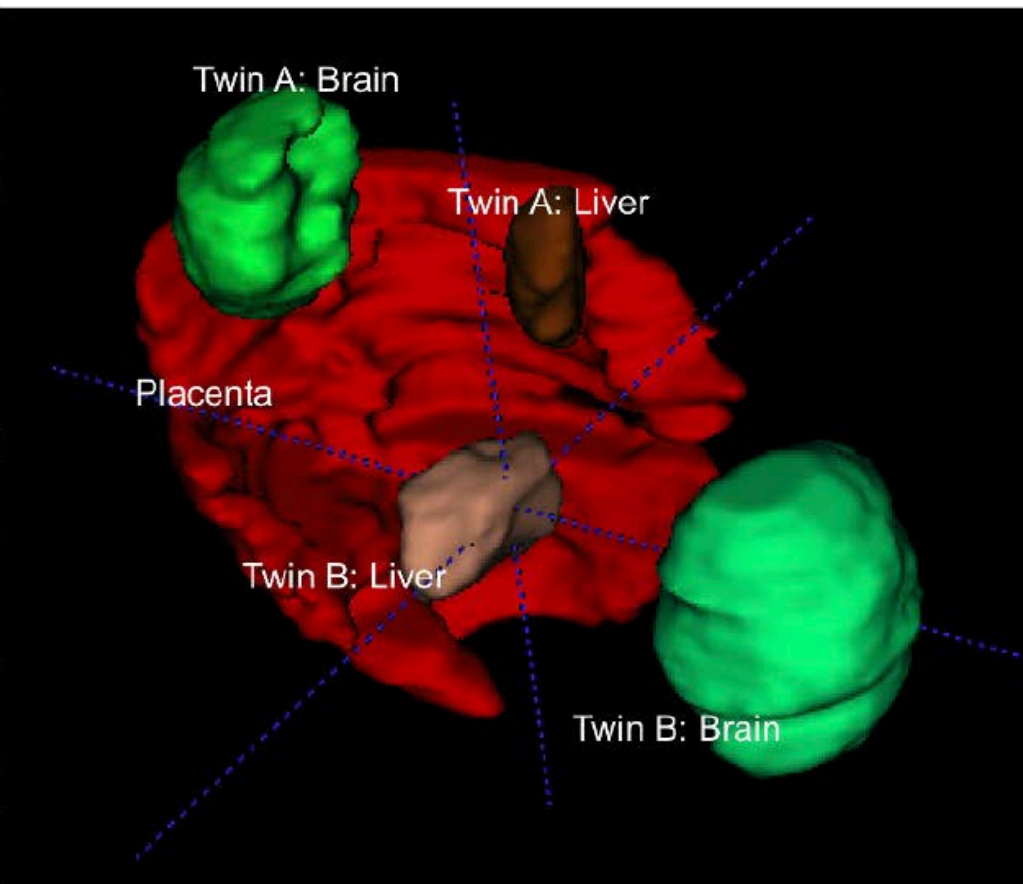
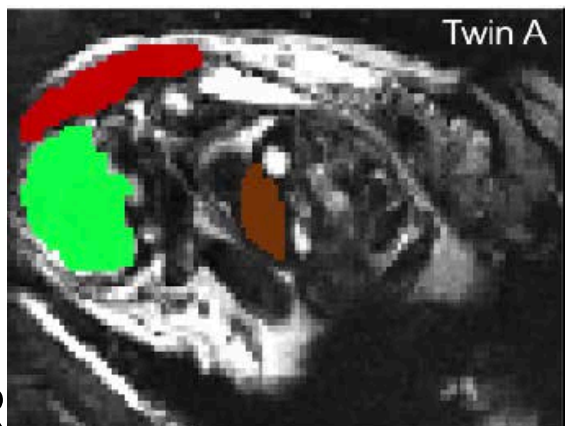
Fetal Liver

IUGR case



A: Healthy
B: Growth restricted

R





Converter: From technology to product

Real-world product development immersion for recent graduates and mid-career innovators

Designed to help bridge the gap between proven technology and readiness for new business startup or outside investment

Teams are supported by mentors in engineering, business, and product development to achieve viability by “de-risking” their technology, market plan, and financial model.



Converter: How it works

- + Teams chosen for two-year term
- + Selected for proven technology and viable strategy
- + Focus on successful exits

Four phases:

- + Project Definition
- + Risk identification
- + Risk mitigation
- + Pitch preparation

Support:

- + Senior mentor
- + Expert review panel
- + Targeted consulting / contractors
- + International community
- + Supplies, travel, indirect cost coverage



Benefits to partners

Scalable, effective strategies to cultivate a technology innovation ecosystem

investment

- + Focus on excellence through elite training
- + Enable change-makers
- + Promote an innovation culture

+ Improve returns on R&D



Improve returns on R&D resource investment

A fraction of R&D investment deployed through the M+Visión system. Objective to drive research toward economic and healthcare impact

- + Identify champions who can inspire and drive innovation projects
- + Projects developed through Catalyst and

Converter programs

- + International collaboration
- + Leverage Boston innovation cluster



Focus on excellence through elite training

Talent development for science- and technology-driven MS, MD, PhD and MBAs—tomorrow's innovation leaders

- + Participants explore and develop their own ideas
- + Training a mix of academic and practice



Enable change-makers

Close collaboration with leaders in positions to effect system-wide change.

- + Focus high impact policy evolution
- + Create environment for cross-institutional collaborations
- + Enable translation-oriented research initiatives



Promote an innovation culture

Short duration, high volume activities that inspire innovators, create connections, and build innovation capacity

- + Hacking Medicine
- + IDEA² Madrid
- + Workshops and courses
- + Conferences and network events



Madrid

mvision
CONSORTIUM

V

mvision.madrid.org

CONSORTIUM

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