



Innovative Medicines Initiative

IMIDIA

IMPROVING BETA-CELL FUNCTION AND IDENTIFICATION OF
DIAGNOSTIC BIOMARKERS
FOR TREATMENT MONITORING IN DIABETES

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efpia

DIABETES

Definition



Diabetes is a metabolic disease characterized by higher than normal blood glucose levels (hyperglycemia) in the fasting state and/or after meals

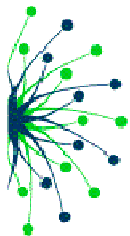
Type 1 and type 2 diabetes are the two main types of diabetes

Type 1 diabetes

- 10 à 15 % of diabetic patients
- Destruction of β cells (autoimmune process)
- Onset in young subjects
- Main feature: weight loss polyphagia, polyuria, asthenia, glycosuria
- Imperatively requires insulin therapy

Type 2 diabetes

- 85 à 90 % of diabetic patients
- No destruction of β cells but a decrease in functional β cells mass
- Onset during maturity and in the elderly
- Very progressive
- In theory insulin therapy is not required



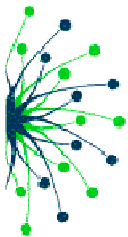
Micro (retinopathy, nephropathy)- and macro-angiopathic complications (cardio and cerebrovascular diseases)

Life span

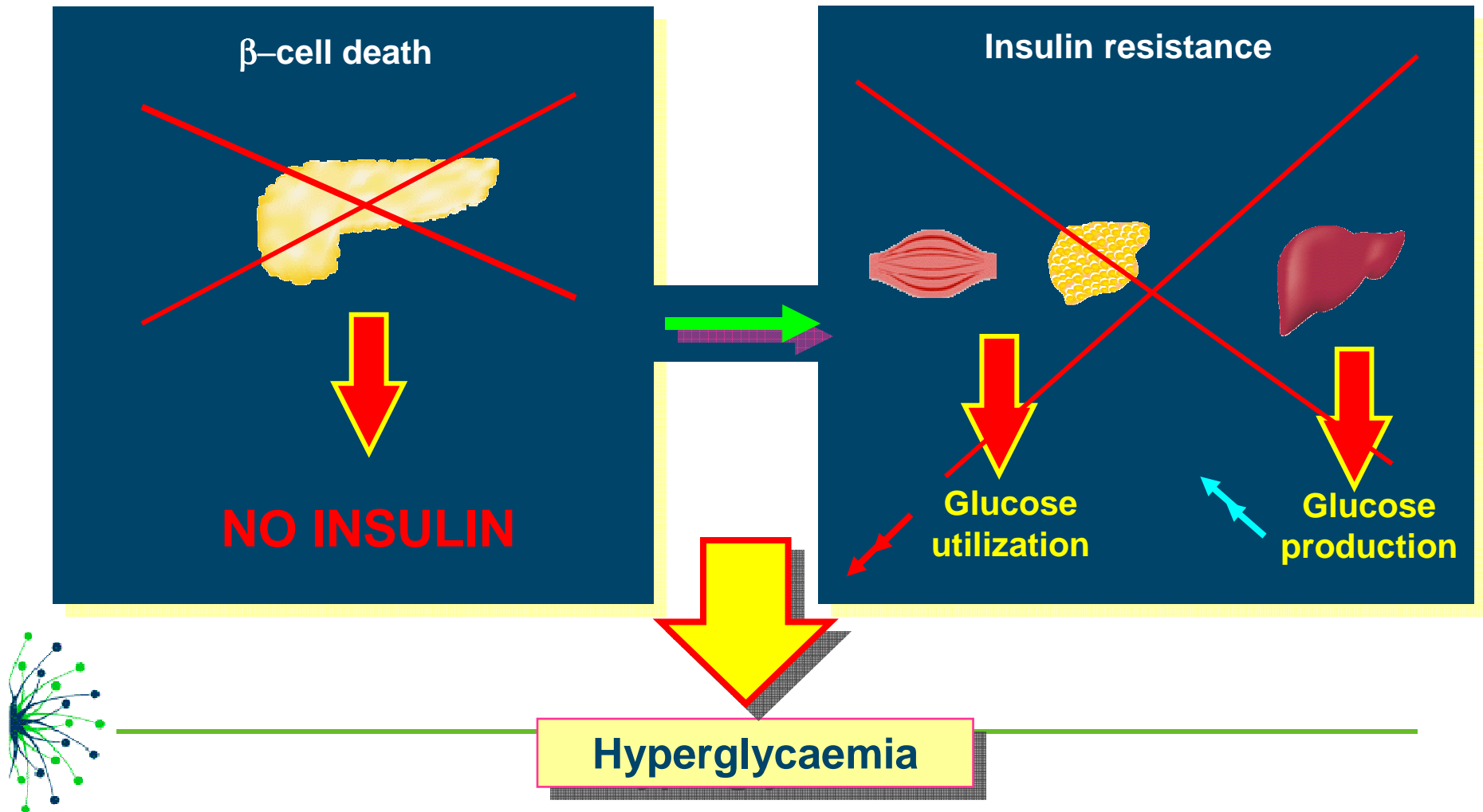
Diabetes – a pandemic disease of the 21st century



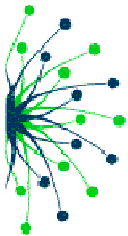
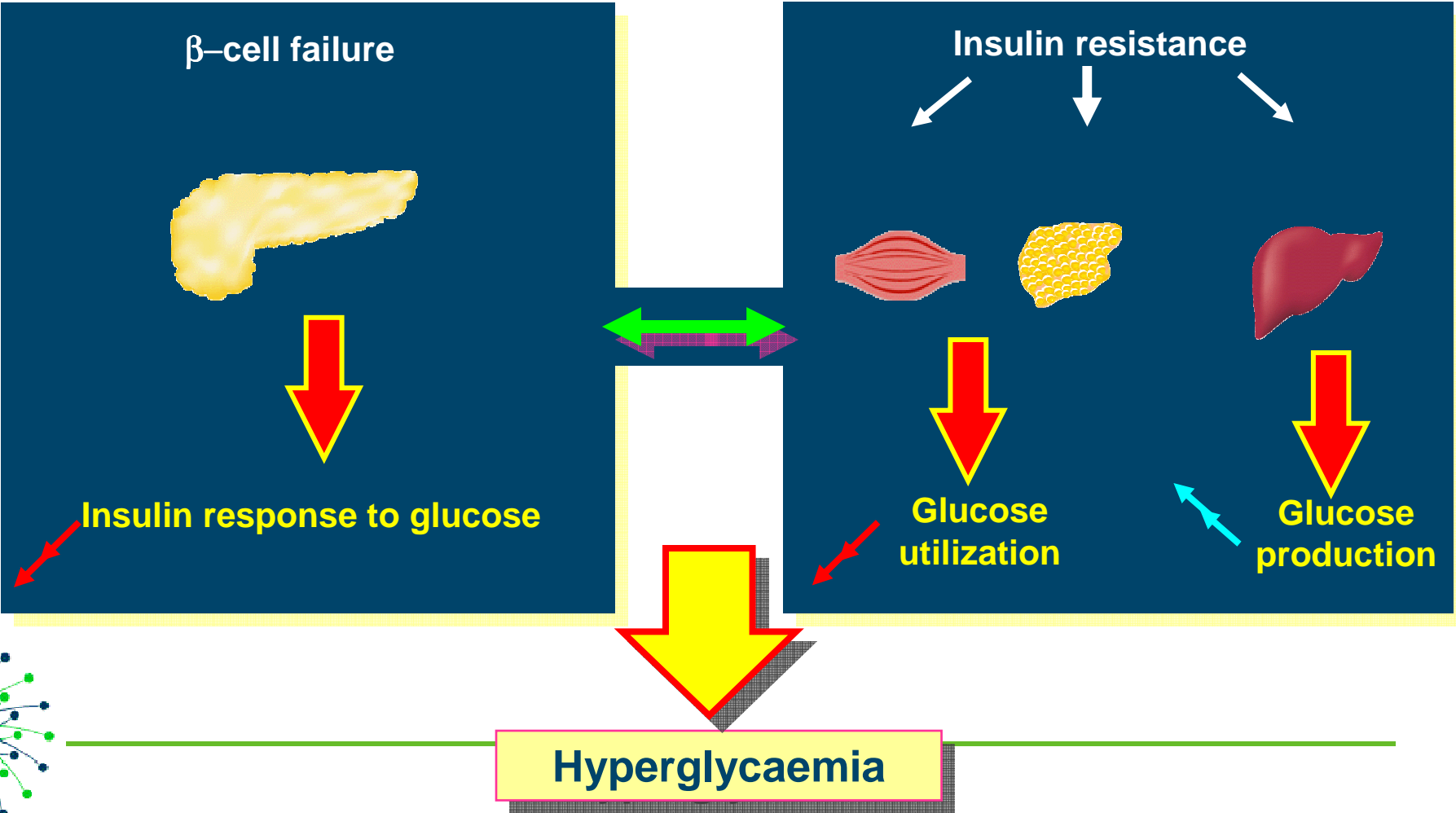
- Number of patients:
 - 2010: 285 million people worldwide
 - 2030: 439 million people worldwide
(in particular spreading to the younger population)
- Pancreatic β -cells:
 - A complete or relative decrease in insulin secretion by pancreatic beta-cells underlies the development of, respectively, type 1 and type 2 diabetes.



PATHOPHYSIOLOGY OF TYPE 1 DIABETES A SCHEMATIC REPRESENTATION



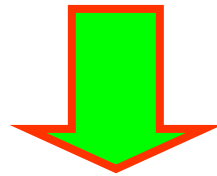
PATHOPHYSIOLOGY OF TYPE 2 DIABETES A SCHEMATIC REPRESENTATION



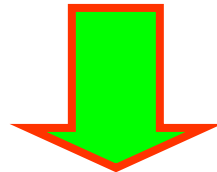


THE AIMS OF TREATMENT OF TYPE 2 DIABETES

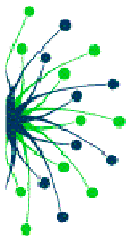
To prevent early death and improve quality of life



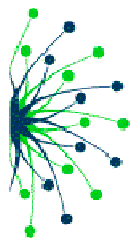
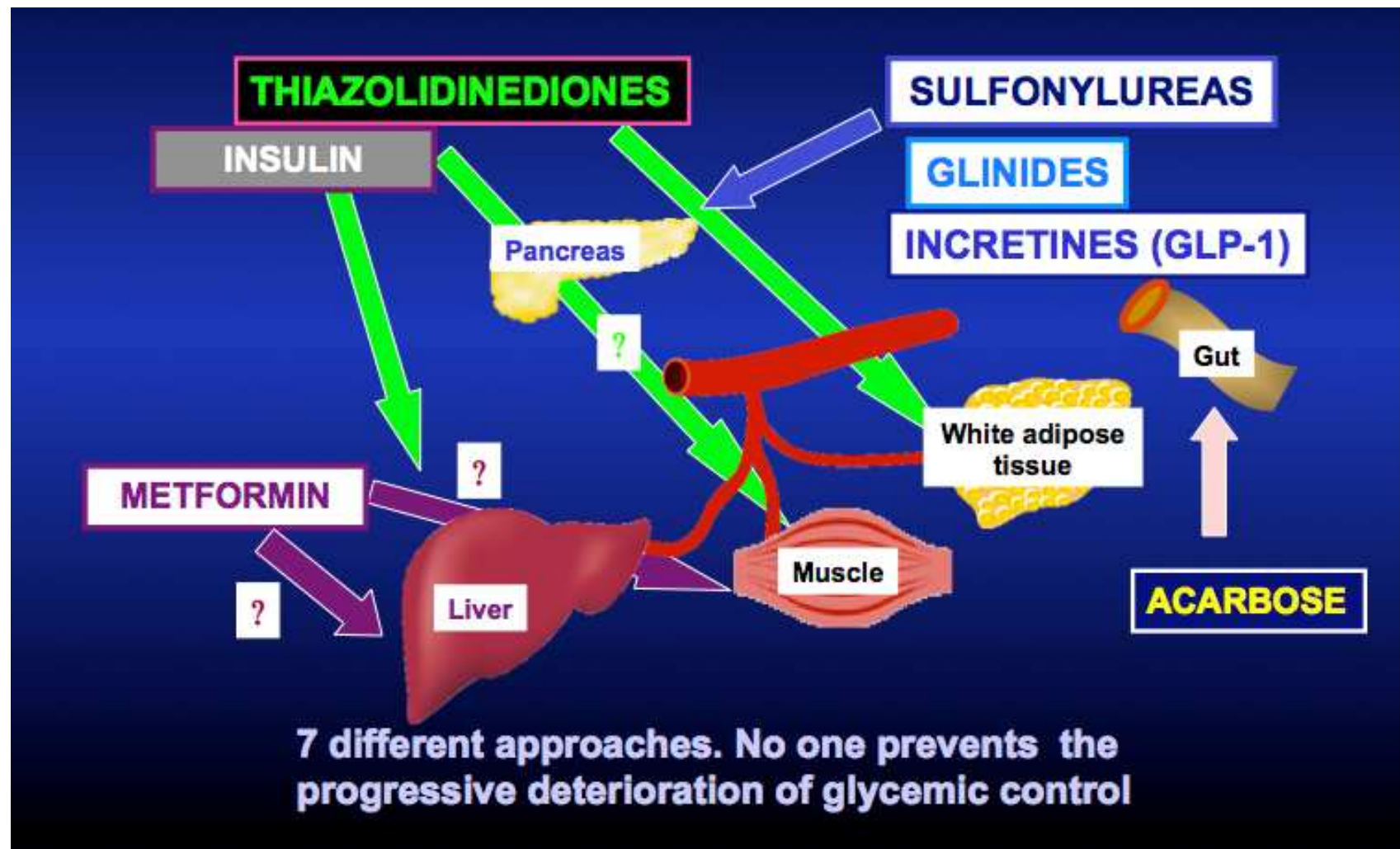
To prevent micro- and macro vascular complications



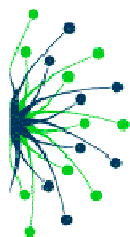
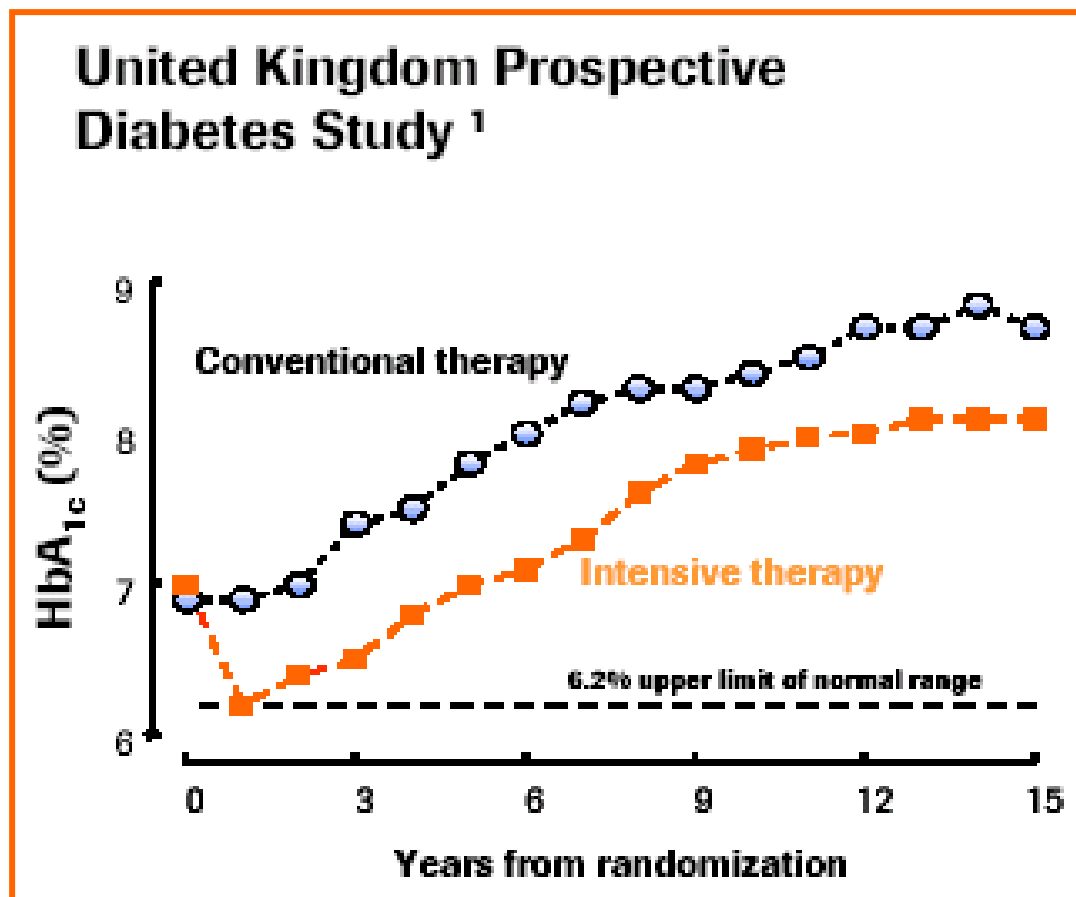
Optimal glycaemic control



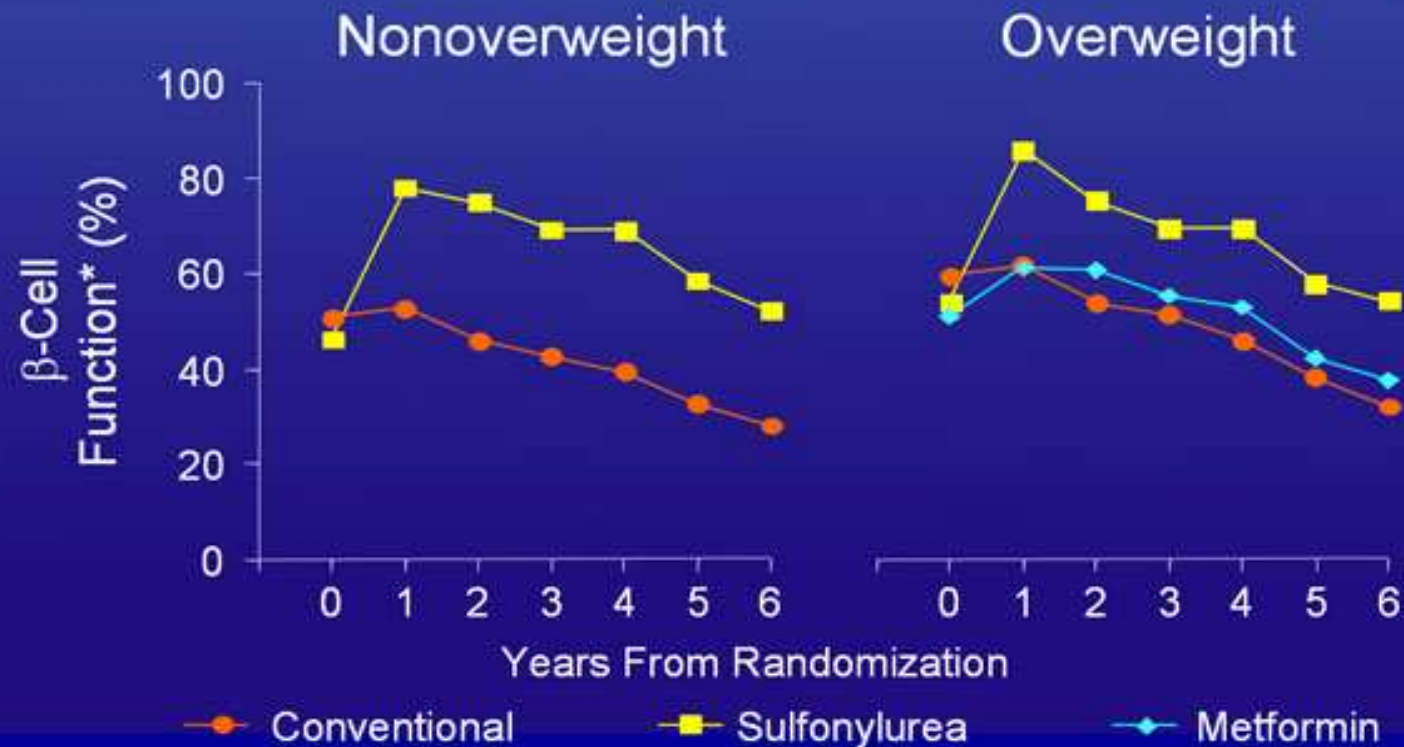
MAIN AGENTS USED IN THE TREATMENT OF TYPE 2 DIABETES



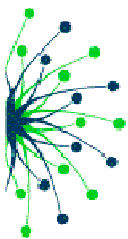
PROGRESSIVE DETERIORATION OF GLYCEMIC CONTROL IN TYPE 2 DIABETES



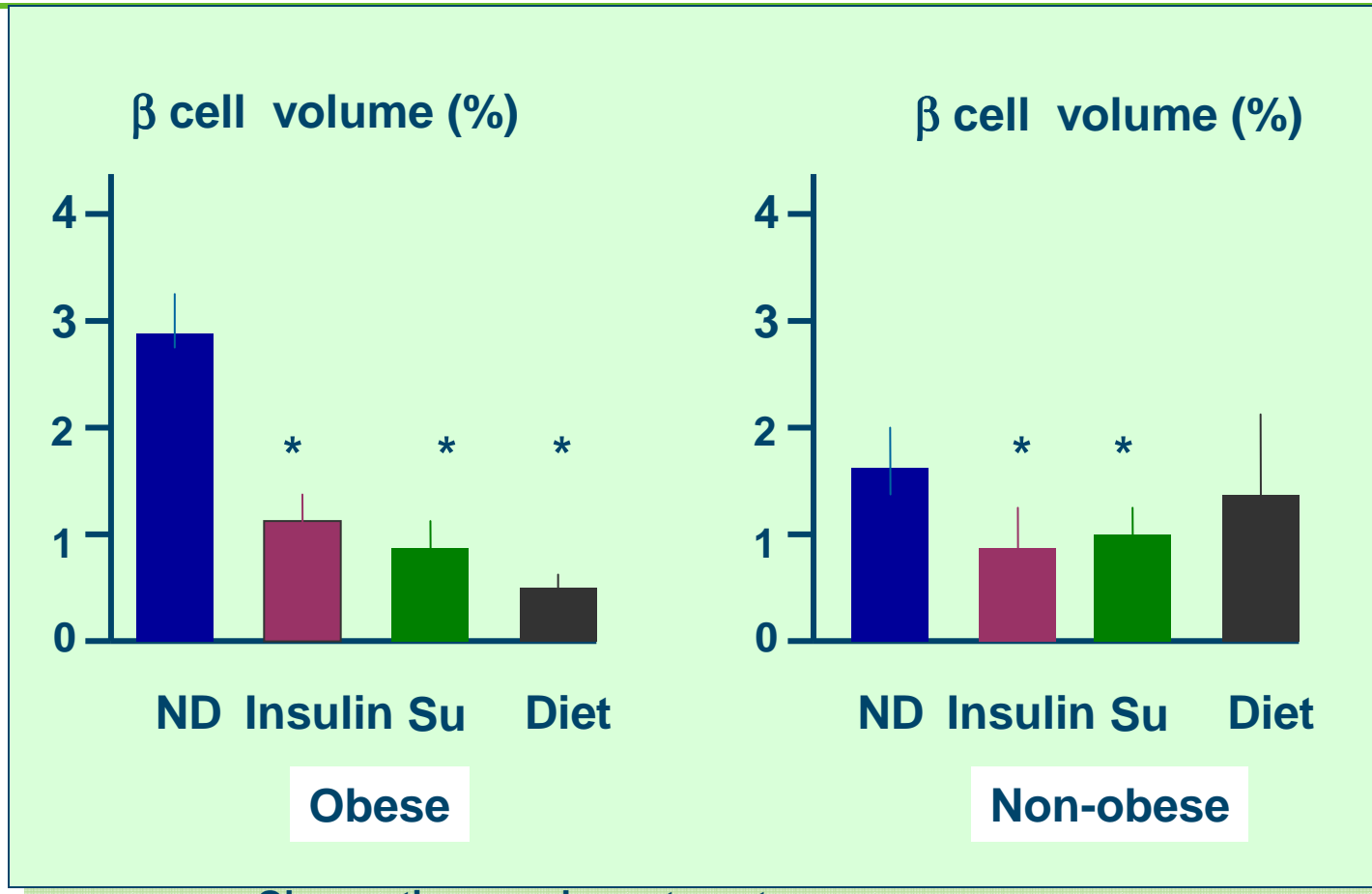
Conventional Therapies Do Not Influence β -Cell Failure



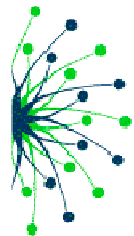
UKPDS Group. *Lancet*. 1998;352:854-865.
 UKPDS Group. *Diabetes*. 1995;44:1249-1258.



β -CELL MASS IN DIABETIC AND NON DIABETIC SUBJECTS

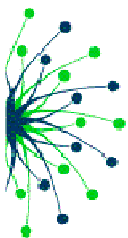
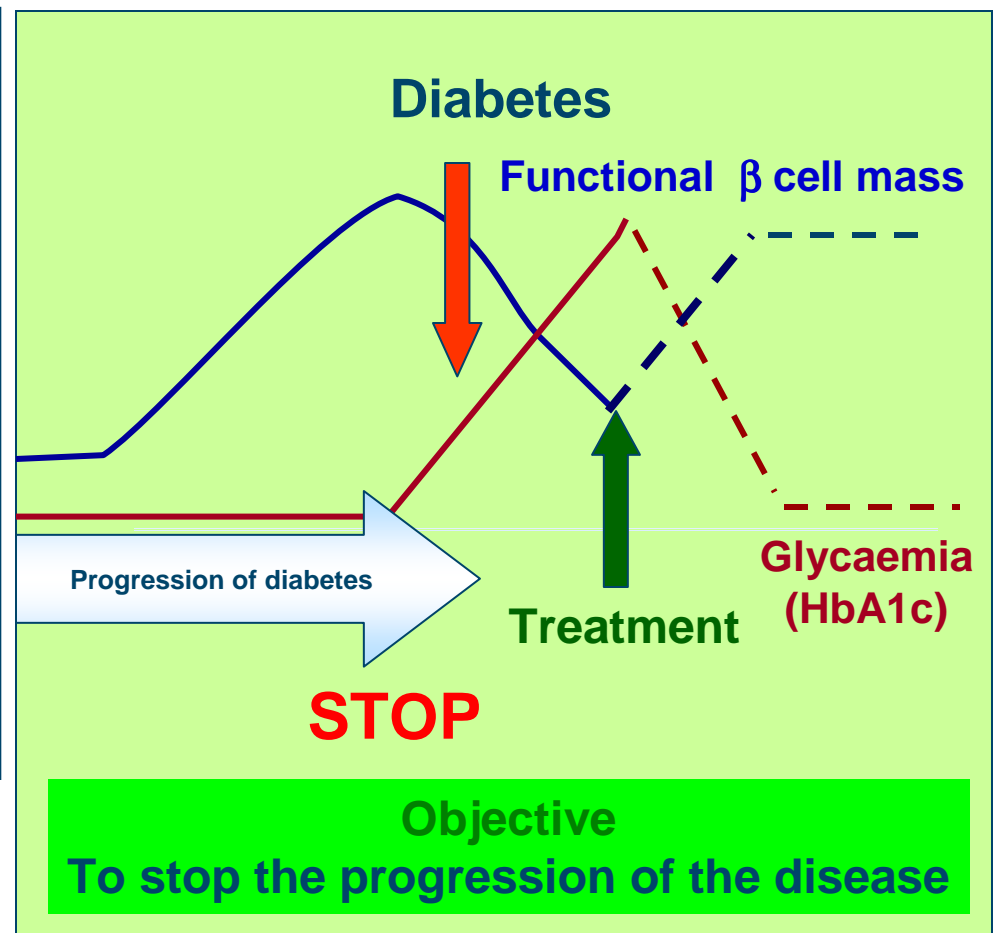
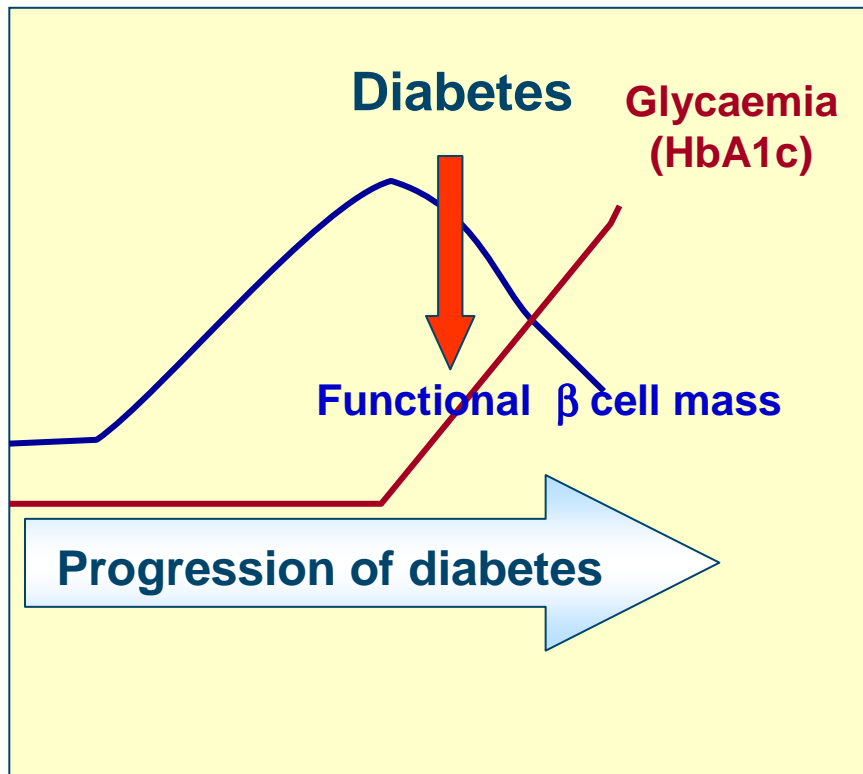


- Observations made post-mortem
- No information on the time-course of the β cell mass

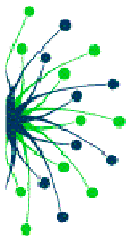
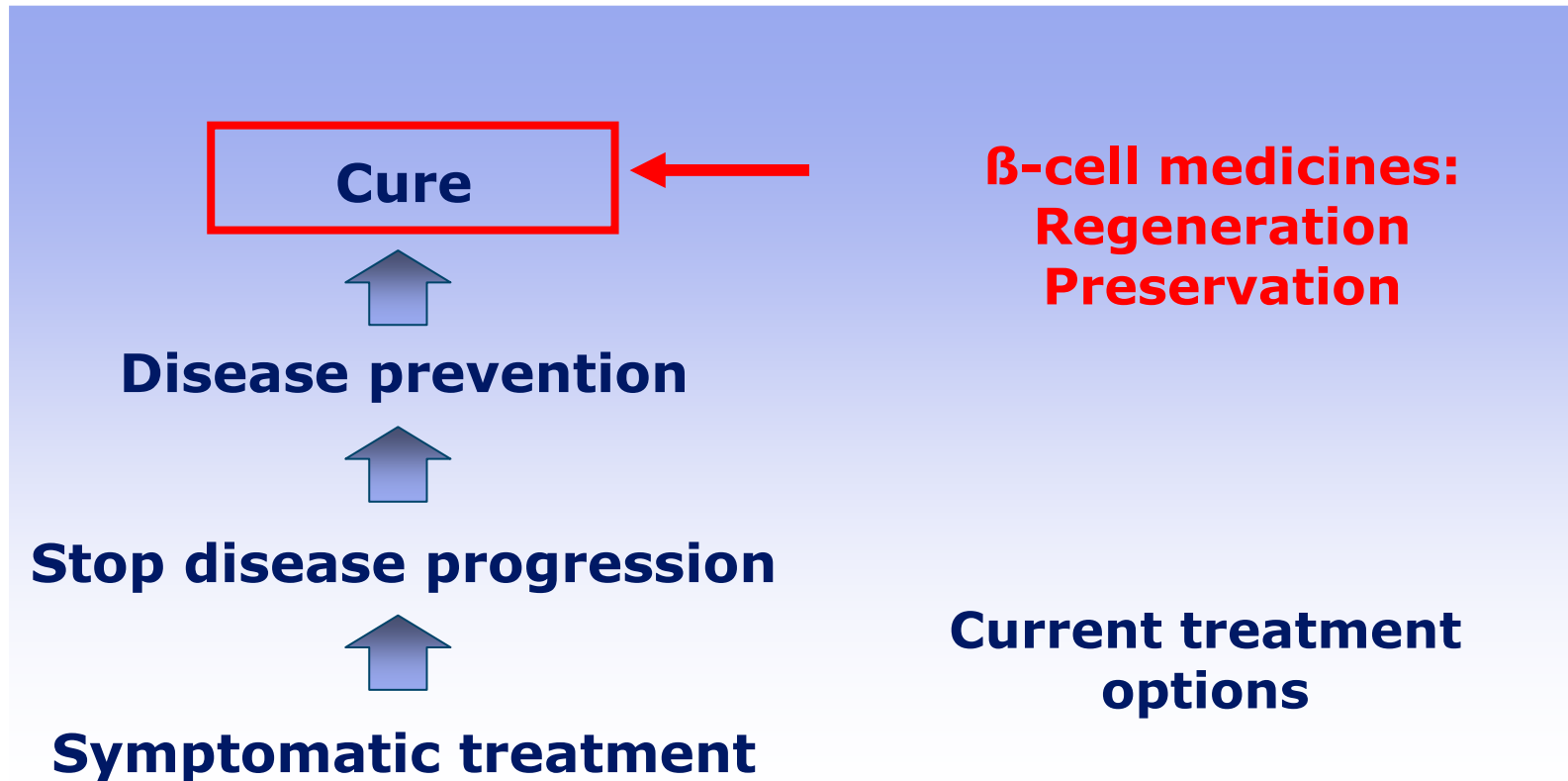


The requirement of non-invasive methods for the measurement of β cell mass

Relationship between functional β cell mass and glycemic control during the progression of diabetes



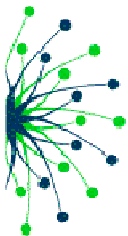
A Paradigm Shift in Diabetes Therapy: from symptomatic Treatment to Cure



IMIDIA will address key bottlenecks for the development of these new therapies



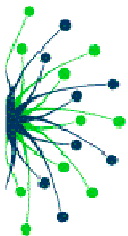
- Novel tools for the study of:
 - human beta-cell development, function and survival;
 - human beta-cell functional modulation by potential therapeutic compounds;
 - in vivo beta-cell imaging.



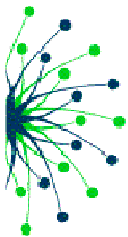
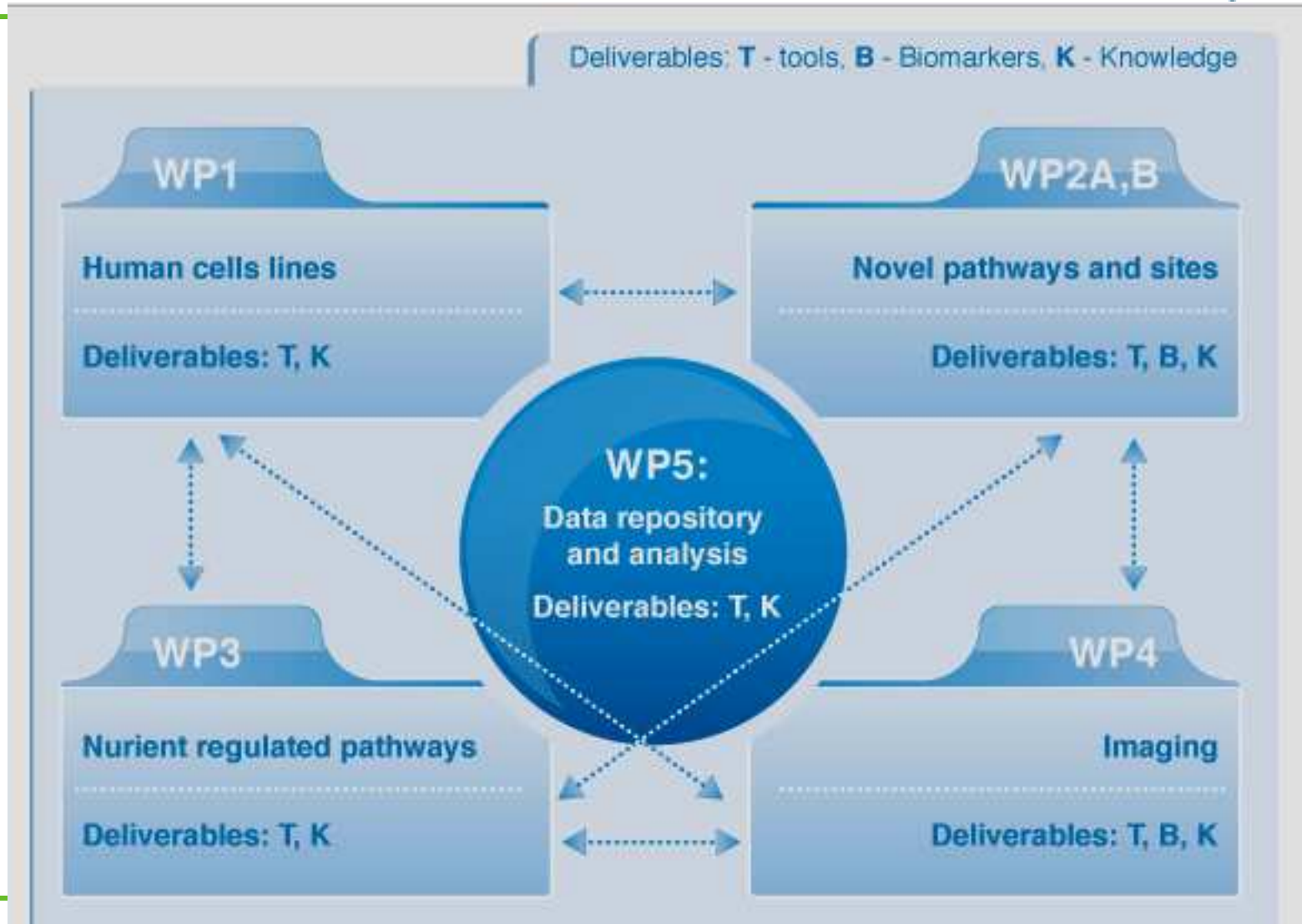
IMIDIA will address key bottlenecks for the development of these new therapies



- Biomarkers:
 - for the diagnosis and prognosis of beta-cell failure;
 - for monitoring diabetes progression and treatment.
- Knowledge:
 - on novel pathways and sites that control beta-cell proliferation, differentiation and apoptosis,
 - on the role of nutrient-regulated pathways in controlling beta-cell mass and function.



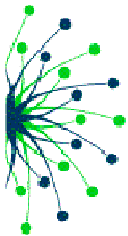
Integrated approach



Expected Outcome – I:



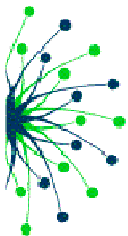
- Disease relevant human islet cell lines:
 - will lead to better models for the development and assessment of diabetes therapies.
- Beta islet cell precursor isolation and purification:
 - will help understanding the birth of beta-cells to help find methods for beta-cell regeneration in diabetes.
- A Systems Biology approach of beta-cell demise in type 2 diabetes:
 - will provide better understanding of the beta-cell pathogenesis;
 - will deliver biomarker candidates for diagnosis, prognosis and assessment of therapeutic efficacy.



Expected Outcome – II:



- A network and technology allowing the isolation of human islet cells from surgical specimen of diabetic and non-diabetic patients.
- Unraveling key pathways and sites that control GLP-1 trophic actions on beta-cells.
- New and unique imaging technologies and novel probes for earlier and better monitoring of beta cell function and mass in humans.



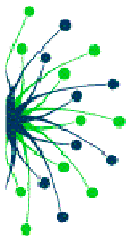
Benefit to the patient:



To monitor specific disease progression and enable improved disease management.

To pave the way for the development of β cell focused therapies via:

- Better biomarkers to monitor therapy benefit in patients
- Better disease centric in-vitro and in-vivo models
- Better understanding to enable focused therapeutic approaches



IMIDIA: Collaboration (Sustainable win-win)



IMIDIA - Collaboration

Data / Results Sharing

Innovation
Focus academia



Application
Focus industry

Generation of
models

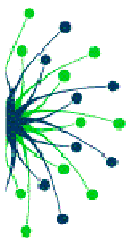
In-vitro Models:

Assessment
Drug Discovery

Imaging Biomarkers:

In-vitro Assessment

Clinical Assessment



Results/achievements so far: 4 months into the project

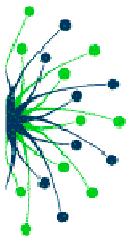


Tools:

- Evaluation of first version of in-vitro models
- Data acquisition process for Systems Biology established
- Biorepository Harmonization Initiated
- Synthesis of first Imaging Molecules Candidates initiated

Biomarker:

- Human assesment study in preparation



Time and money

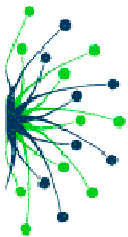


Financing

- IMI funding: € 7.074.760
- EFPIA contribution, mainly in kind: € 15.081.800
- Other contributions
(e.g. unfunded act., act. in the USA) € 3.750.920
- Total project cost: € 25.907.480

Timing:

- Starting date: 01. Feb. 2010
- Duration: 5 years



Leadership team

- Sanofi-Aventis Deutschland W. Kramer
- Servier A. Ktorza
- University of Lausanne B. Thorens

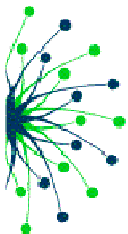
Participants

- AstraZeneca Pharmaceuticals
- Boehringer Ingelheim Pharma
- CEA /Institut d'imagerie Biomédicale
- CNRS UMR 7091
- CNRS-University Paris Diderot
- Dresden University of Technology
- F. Hoffmann-La Roche
- Hannover Medical School (MHH)
- Imperial College London
- INSERM U845
- Lilly Deutschland
- Novo Nordisk A/S
- Novartis Institutes
- SARL ENDOCELLS
- Swiss Institute of Bioinformatics
- University of Geneva
- University of Pisa
- Vrije Universiteit Brussel

[View all participants](#)



*Location of key scientific contacts in Europe



Press Release Issued 14/06/10

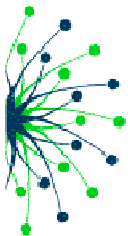


Launch of IMIDIA

An Innovative Medicines Initiative Project for Diabetes

***Academia, biotech and the pharmaceutical industry have
joined forces to fight diabetes.***

Frankfurt, Germany / Lausanne, Switzerland / Paris, France - June 14, 2010.

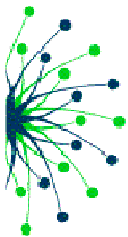


Further information



- www.imidia.org
- Email: [info \[AT\] imidia.org](mailto:info@imidia.org)

www.imi.europa.eu



IMIDIA



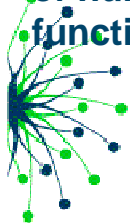
Improving β -cell Function and Identification of Diagnostic Biomarkers For Treatment Monitoring in Diabetes

- Type 1 and type 2 diabetes impose a huge burden to welfare systems
- Relative or complete decrease in insulin secretion underlies the development of type 1 and type 2 diabetes
- Limited therapeutic options ← Limited knowledge β -cell biology (function, survival, pathophysiology)
- Lack of biomarkers for prognostic of β -cell failure



The IMIDIA project

New tools for the study
of human β -cell development
function and survival

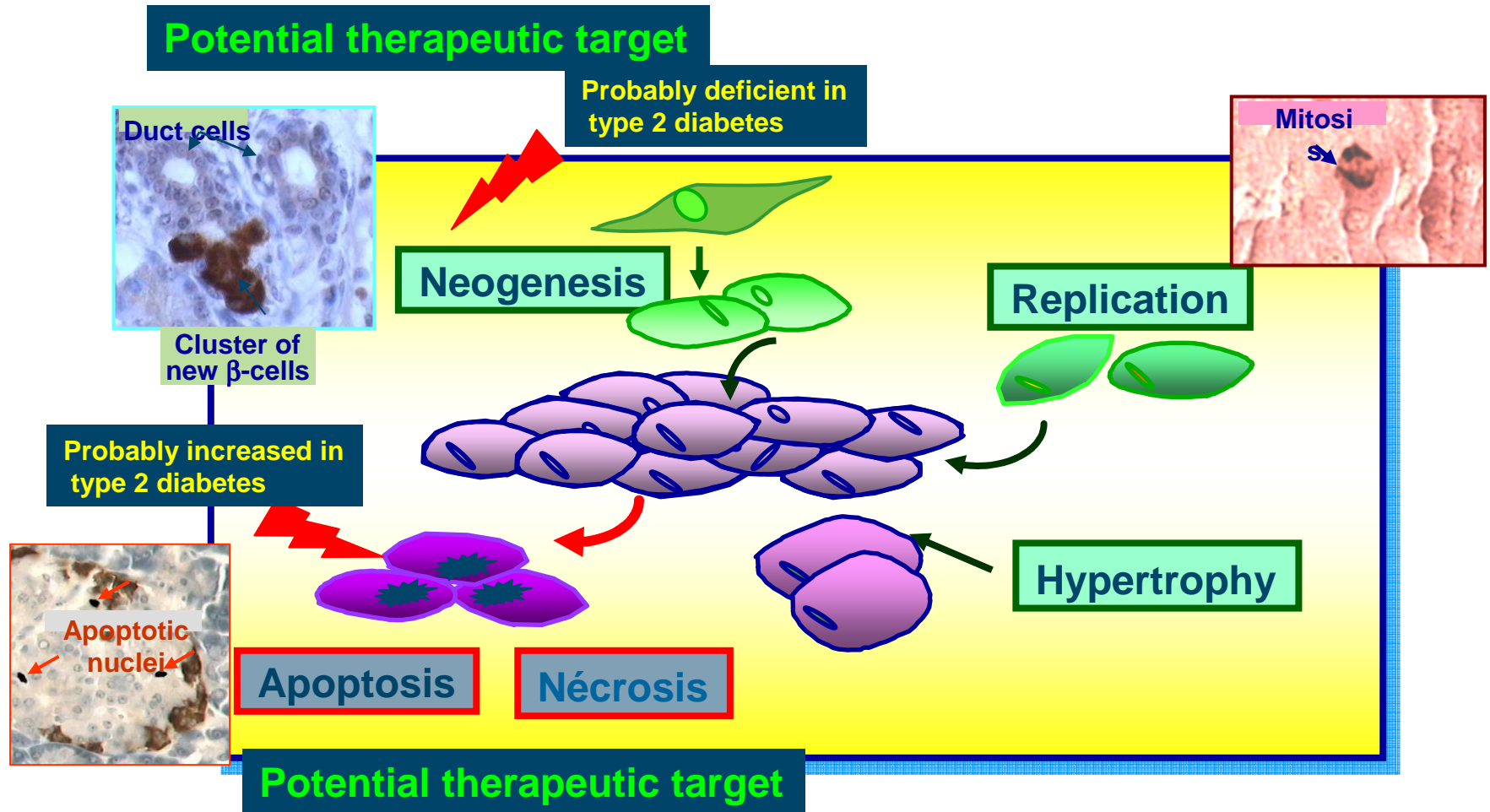


Biomarkers

Knowledge of novel
pathways and sites
that control β -cell
development,
function and survival

Non-Invasive Imaging
of the Human
Endocrine Pancreas

CONTROL OF BETA-CELL MASS



EXPECTED MAIN COMPETITIVE ADVANTAGE



Close interaction between academic teams, pharmaceutical companies and SMEs



Unique conjunction of expertise and will form a strong basis for a successful enterprise



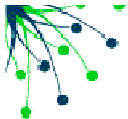
Improvement of industrial competitiveness and Public Health in Europe

- **Content advantage**

- All results transparent to all project participants during the project
- “Validation” of new technologies / tools from academia during the project by participating industry

- **Time advantage**

Access to generated IP within the project “foreground” for “research use”



The Path to innovative Diabetes Therapies: enhancing functional β -Cell Mass

