

# Moving Forward: T1 Diabetes Research Then, Now and Beyond

Jude Restis and Laura Tremblay, MD | Research Information Volunteers

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### **Impact of Research**

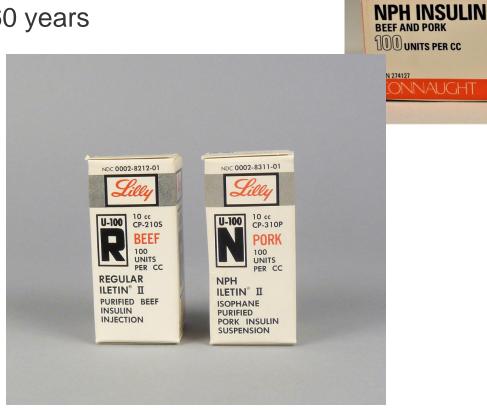
- For 100 Years all we had to treat T1 Diabetes was insulin.
- Now we have the 1<sup>st</sup> approved therapy to delay or stop the progression of T1

What happened in-between? What did research get us?



#### **Insulin - Then**

Insulin from Beef and Pork was all that was available for the first 60 years





NPH INSULIN

BEEF AND PORK

10cc

#### **Insulin - Now**

#### **Human Insulin**







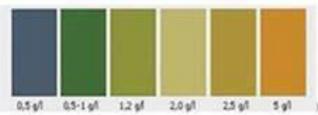


# **Glucose Testing - Then**

#### **Urine Glucose Testing**











# **Glucose Testing - Then**

**Urine - Glucose Testing** 







## **Glucose Testing - Then**

1st Blood Glucose Testing





# **Glucose Testing - Now**

**Glucose Testing** 



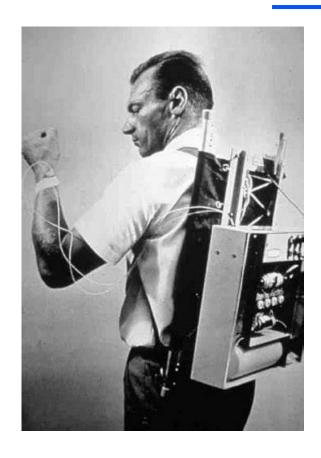






# **Insulin Pumps - Then**

Pumps







# **Insulin Pumps - Now**

#### Pumps







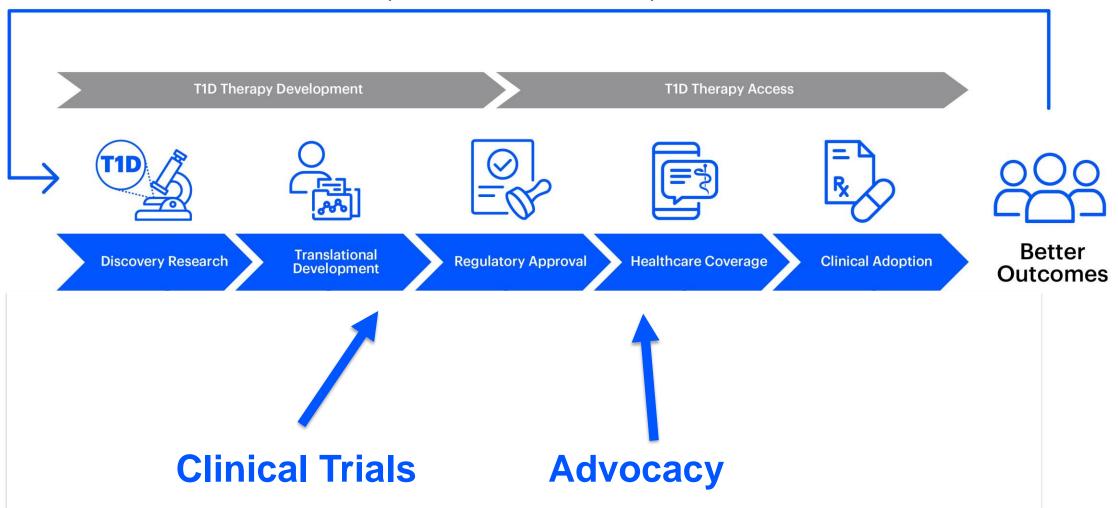
#### **Impact of Research**

 All of these life improving devices and therapies were made possible by research and the support of participants in clinical trials



## **Getting Therapies to the T1D Community**

Cycle of Next Generation T1D Therapies





#### **JDRF Research Priorities**



Global Universal Screening



Disease Modifying Therapies



**Cell Therapies** 





Training of Researchers and Clinicians



# **Screening**



#### What Is Screening?

- T1D is an autoimmune disease
- This means that the body makes antibodies against itself
- In T1D, 5 islet autoantibodies (AABs) have been identified
- The presence or absence of those AABs helps doctors and researchers estimate a person's risk of developing T1D in the future

#### Why Screen?

- Help prevent diabetic ketoacidosis (DKA) at diagnosis
- Intervene early to delay a clinical diagnosis of T1D and the need for insulin
- Identify people who may benefit from earlyintervention clinical trials
- Give individuals and families time to prepare

#### **How to Screen?**

- Screening involves a simple blood test
- There are several different options for screening, including:
  - TrialNet
  - Autoimmunity
     Screening for Kids
     (ASK)
  - Enable Biosciences
  - Commercial Labs



## T1D Risk Screening Available TODAY until 1pm!

#### **Visit the TrialNet Table**

#### Who can be screened:

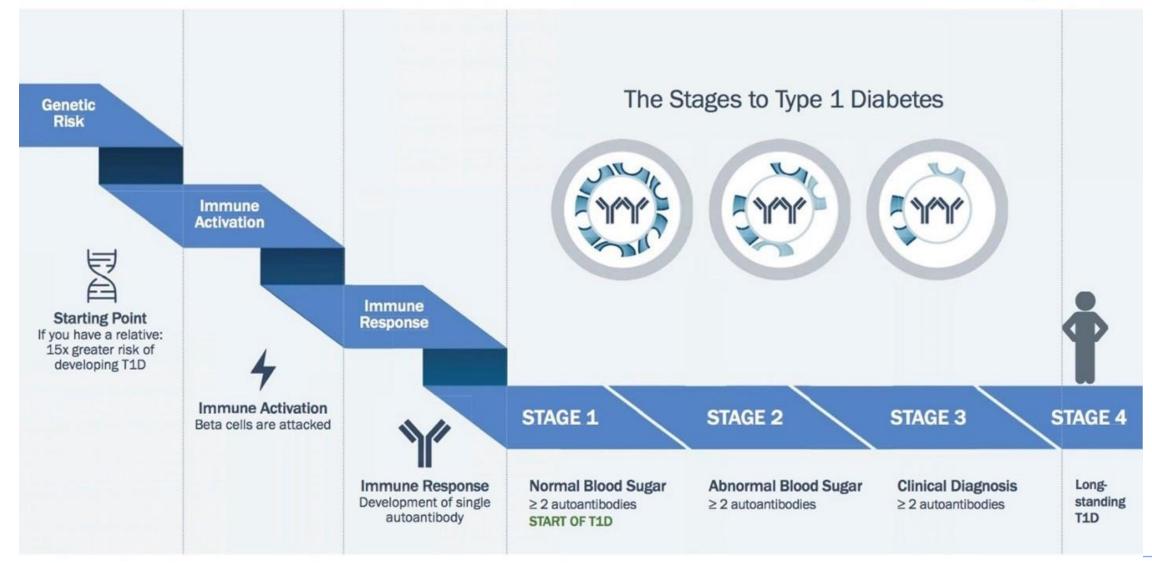
- Anyone age 2.5 through 45 years with a sibling, child or parent with T1D.
- Anyone age 2.5 through 20 years with a cousin, uncle, aunt, niece, nephew, grandparent or half-sibling with T1D.





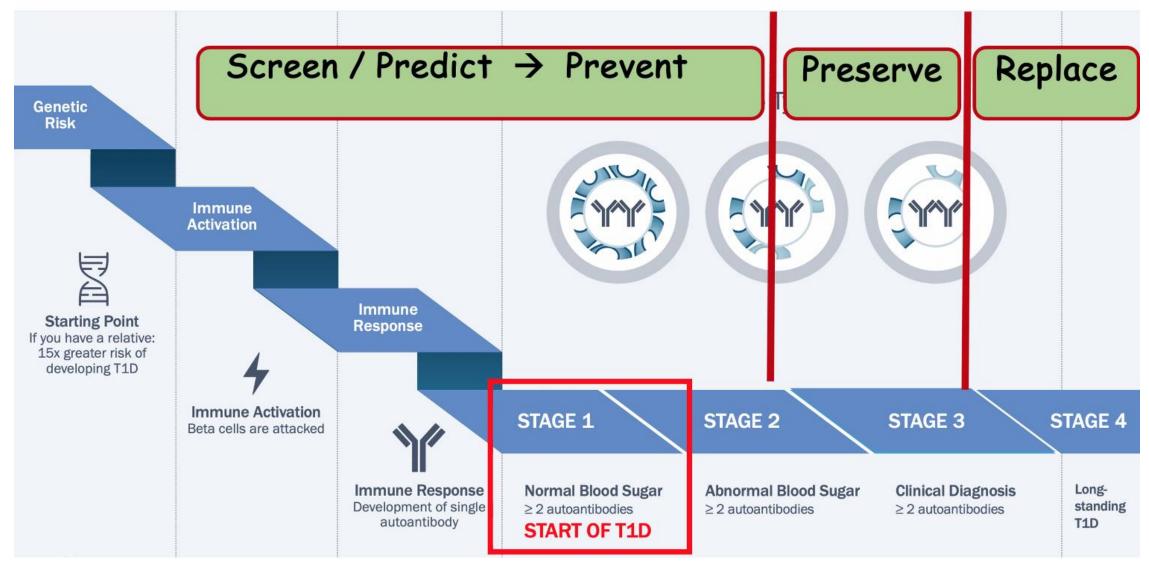
#### T1D Disease Progression







#### **T1D Disease Progression**





# **Disease-Modifying Therapies**



# Multiple Disease Modifying Therapies Currently in Clinical Trials

#### It's an Unprecedented Exciting Time!

Stage 1 Stage 2 Stage 3 Stage 4 Hydroxychloroquine Teplizumab Teplizumab Replacement Therapies Abatacept Low Dose ATG Verapamil Future Regeneration Therapies Oral Insulin Iscalimab **ATG** Baricitinib Ustekinumab Abatacept L. Lactis **Tolerizing Antigens** Diamyd **JAK Inhibitors** 



# Tzield<sup>TM</sup> approved!

- Teplizumab (Tzield), is approved for use in delaying T1D in high-risk individuals aged 8 and above:
  - 2+ islet autoantibodies
  - Abnormal glucose control
- PROTECT Trial also showed that Tzield benefits children/teens with newonset T1D, slowing the loss of beta cells and prolonging insulin production
- JDRF led research into Tzield for over 30 years

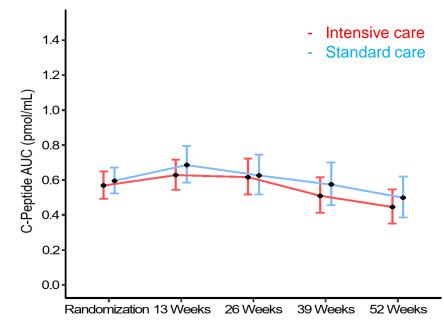


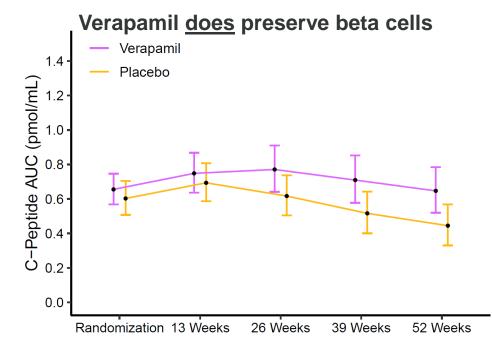


## **CLVer Trial: Verapamil**

- Can AP systems and/or verapamil preserve beta cells (i.e. extend the honeymoon period) in the new-onset population?
- The results are in!
  - AP systems did not preserve beta cells but did provide exceptionally good glucose control in the new onset period
  - Verapamil preserved beta cells—meaning they make more insulin for longer!
  - JDRF will continue to pursue research into verapamil, including using it with other drugs and looking at whether verapamil can be used with people at high-risk, before they develop T1D

#### Tight glucose control does not preserve beta cells



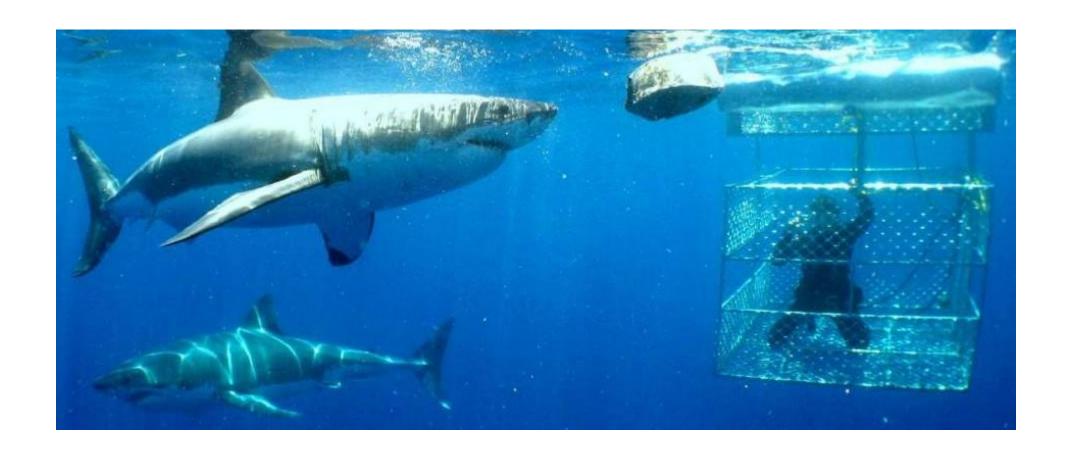




# **Cell Therapies**

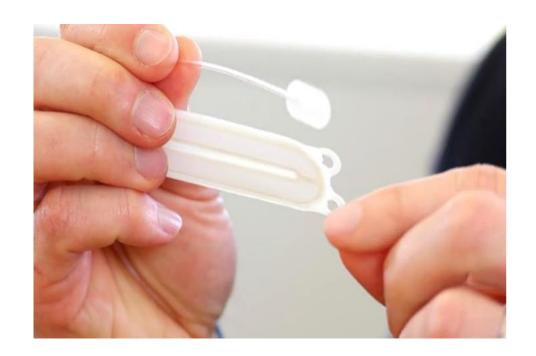


# **Encapsulation Concept**

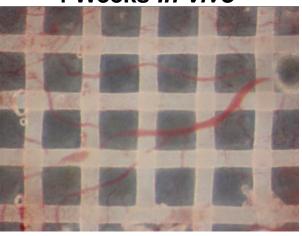




# Beta Cell Replacement



4 Weeks In Vivo



18 Weeks In Vivo



8 Weeks In Vivo





## **Vertex Progress**

Clinical trial for VX-264, an encapsulated cell therapy, to begin in US

2023

JDRF-funded research moving Vertex cell therapies closer to commercialization

Melton funds Semma
Therapeutics to turn these cells
into a curative therapy for T1D

**2015** 

Vertex announces the first person to receive the therapy has a 100% reduction in insulin required

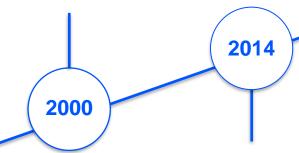
2022

VX-880 receives Fast Track

Designation from the FDA

**2021** 

JDRF awards a grant to Douglas Melton, Ph.D., to create insulin-producing beta cells from stem cells



The T1D fund makes a catalytic investment in Semma, which is then acquired by Vertex Pharmaceuticals

2017

The Melton lab successfully makes insulin-producing beta cells



#### **Aspect and Novo Team up**



- Aspect Biosystems has developed a novel 3D printer that can be used to produce implants made from various biomaterials and cells of interest.
- JDRF began providing support to Aspect in early 2022.
- In April, Novo Nordisk and Aspect announced a \$2.6 billion deal that will combine Aspect's technology with Novo Nordisk's expertise in differentiating stem cells and manufacturing capabilities.
- The Partnership's initial focus will be on creating an implantable encapsulated islet therapy for T1D, designed to maintain normal blood-sugar levels without the need for immunosuppression.





JDRF Beta Cell Replacement Consortium: Sharing Data, Maximizing Resources, and Saving Time and Money

Novel

**Biomaterials** 

Over 50 members 2 Core Facilities

**Industry Members** 



















**Optimize Local** mmunosuppression approaches

**Macro Device** Design

#### **JDRF**

Foster collaboration Reagent sharing & protocol standardization Prioritize activities & resources **Cross-team comparisons Identify gaps for RFA** 

Calls

External expertise

**Preclinical to** clinical translation studies

**Standardization Tools** 





# **Improving Lives**



## **Thriving Device Market**



Medtronic 780G



Abbot Freestyle Libre 3



Beta Bionics iLet



Omnipod 5 Approved in kids 2+



Tandem Mobi





#### **Tidepool Loop**



- The FDA Authorized Tidepool Loop on January
   24
- JDRF has prioritized interoperable, or open protocol artificial pancreas systems, working with the FDA to create the regulatory pathways and with industry partners to encourage development
- JDRF and Helmsley Charitable Trust –
   Foundational funders of Loop
- Next step: Partnership with pump companies to incorporate the Loop algorithm



## **Psychosocial Impact**

- There is increasing awareness of the emotional toll of living with T1D
- Research is focusing on developing programs and interventions to help reduce negative mental health effects
- Areas of focus include:
  - Stress on interpersonal relationships (spouse, parent-child)
  - Depression
  - Anxiety
  - Disordered eating



#### **Insulin Therapy Is Not Enough**

#### Among people with T1D in the US:



~70% do not consistently achieve target blood-glucose control levels



**2/3** of adults have obesity or overweight



~1 in 3 will develop kidney disease



~2 in 3 will develop heart disease

- The future of T1D care is multiple drugs taken safely and conveniently for glucometabolic control and long-term health
- JDRF is funding clinical trials in T1D for medications already available for Type 2 Diabetes, such as GLP-1 agonists (e.g. Ozempic) and SGLT-2 inhibitors (e.g. Jardiance). In the T2D population, these medications been shown to reduce heart and kidney complications



## **Clinical Trials**



#### **Golden Age for T1D Trials**





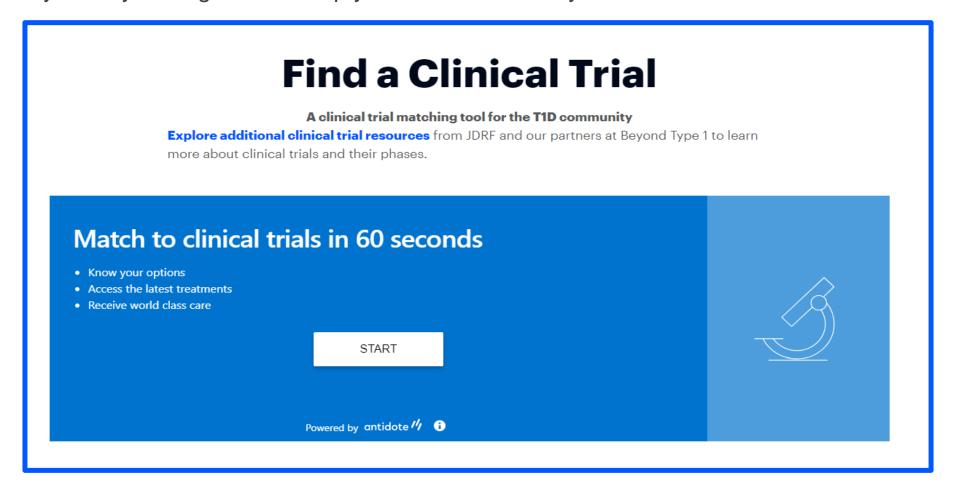
150 trials actively recruiting in the US

70+ JDRF-supported trials around the world



# Fostering a Culture of Clinical Trial Participation

JDRF Clinical Trials Connection tool (jdrf.org/clinical-trials) can match you with T1D trials for which you may be eligible and help you contact the study site to learn more





# **Looking Ahead**



#### **Looking Ahead**

#### **Screening**

- Develop a simple genetic risk test for identifying at-risk individuals
- Develop a protocol for general population screening

#### **DMT**

Develop combination approaches that will stop the immune attack <u>and</u> preserve or regrow beta cells

#### **Cell Therapies**

 Create a JDRF beta cell factory, which will provide greater access for teams working to develop new products

#### **Improving Lives**

Fully close the loop

JDRF Accelerating Breakthroughs in ALL areas



#### **Sneak Peak at FY24**

- There's a LOT to look forward in FY24, including:
- Several DMT clinical trials will be reading out, including:
  - BANDIT JAK inhibitors in newly diagnosed individuals
  - DiViD Combination of two antivirals (pleconaril and ribavirin) in newly diagnosed individuals
- Updates from Vertex VX-880 and VX-264
- Sernova Cell Pouch results
- New devices
  - Ypsomed ACE pump



# THANKYOU