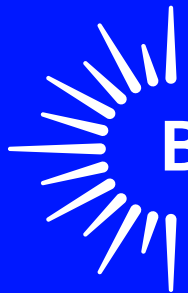


Who should we screen,
at what age,
and how often?

Brigitte Frohnert, MD, PhD
Anette-Gabriele Ziegler, MD



Breakthrough T1D™

Formerly JDRF



Conflicts of interest

BF: none

AGZ: Serves or has served on the advisory panel for Provention Bio, Sanofi, and Novo Nordisk, and DMC for Provention Bio, Sanofi, and ITB-Med. Received education grants and research grants from Sanofi. Holds patent on screening for genetic risk.

Recommendations on population to be screened

Where **policy** and **infrastructure** is in place, IAb screening for early-stage T1D should be offered to the **general population**. [C]

Risk of Developing **Stage 3 T1D** by 20 y of Age

Can they be identified at stage 1 or 2?

First-degree relatives

1/20 (5%)

**10 – 15%
of all cases**

General population

**1/300 (0.3%)
USA**

**85 – 90%
of all cases**

Pathways to be screened

Clinical decision-making

- Capillary or venous draw
- Screening at pediatric clinics



Research (informed consent)

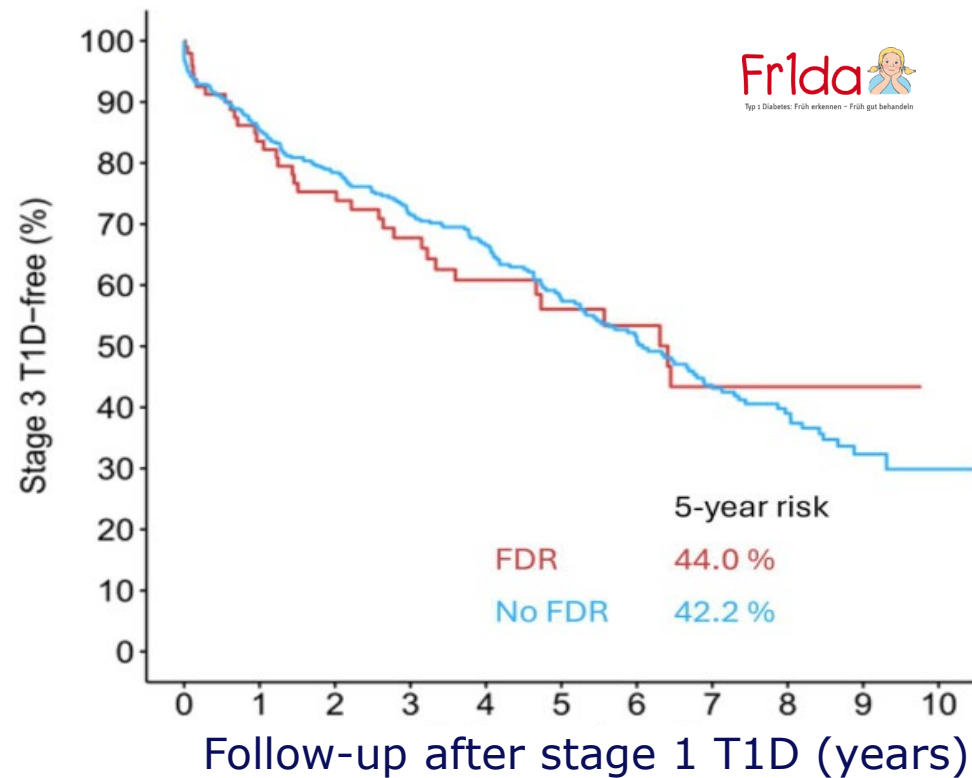
- Laboratory or a finger poke kit to mail



Recommendations on population to be screened

Targeted screening of individuals who are at increased risk due to personal or family history of autoimmune disease may be a practical starting point for screening, **but the ultimate goal is general population screening.** [C]

Children from the general population have similar rate of disease progression compared to relatives



- Children from the general population have similar rate of disease progression compared to relatives and people at genetic risk: **BABYDIAB, DIPP, DAISY: 43.5% at 5-year follow-up.**(JAMA 2013, JAMA 2020, Diabetologia 2022, Diabetes Care 2025)
- Children with no family history of T1D have a higher risk of DKA (Diabetes Care 2022)
- Children from the general population benefit from early detection (JAMA 2022, Diabetologia 2023)

Many autoimmune conditions in a parent are associated with increased risk of T1D in the child

Bidirectional association between T1D and other autoimmune diseases

Lancet 2023

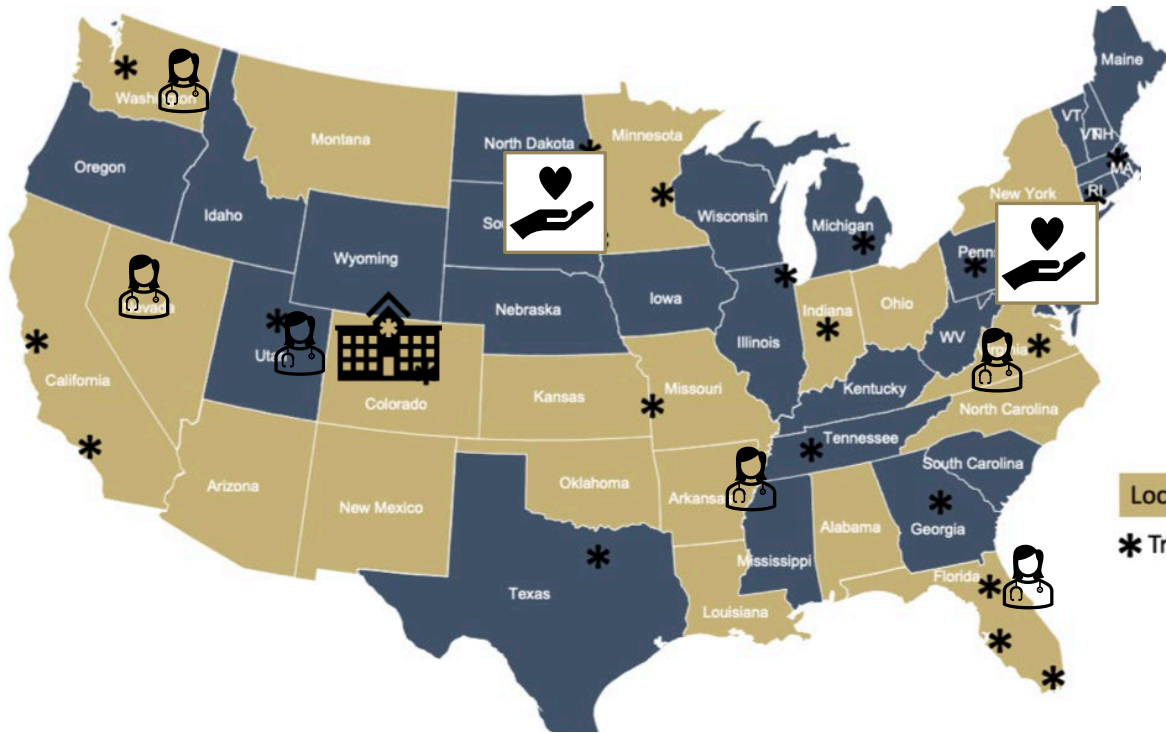
- Celiac disease
- Addison's disease
- Autoimmune thyroiditis
- Polymyalgia rheumatica
- Myasthenia gravis
- Multiple sclerosis
- Vitiligo
- Pernicious anemia
- Systemic lupus erythematosus (SLE)

Recommendations on population to be screened

Screening awareness communications should emphasize the benefits of timely detection of early-stage T1D.[B]

Prior to starting general population screening for early-stage T1D, there must be a developed infrastructure for:

- (i) confirmation of positive results; and
- (ii) monitoring for those with early stage T1D, including referral pathways between primary and secondary care, and between pediatric and adult services, as needed.[C].



Where to turn when your patient or child has screened positive for type 1 diabetes or celiac.



Barbara Davis Center Faculty

Local Expert(s)
* TrialNet centers



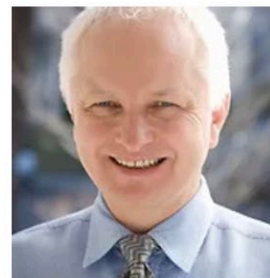
Kimber Simmons
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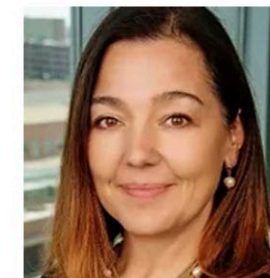
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Ask the Experts

FOR EARLY T1D ANSWERS AND GUIDANCE

Recommendations on age and cadence of screening (Youth)

General population screening:

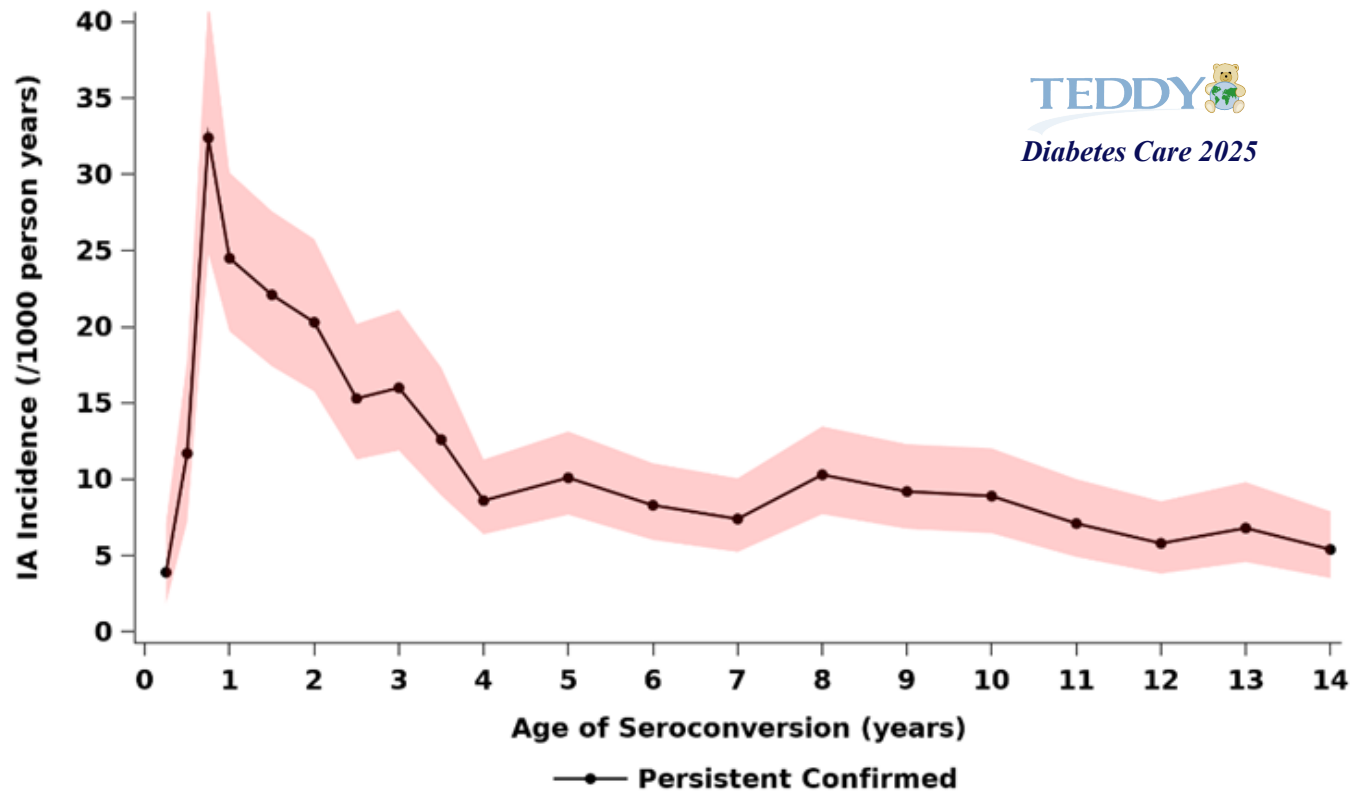
- at age 2-4 years and, if IAb negative, again
- at age 6-8 years, and
- at age 10-15 years

is recommended for detecting individuals with early-stage T1D.

Children who have never been screened should catch up by joining this schedule at the earliest opportunity. [B]

For any screening, the age ranges indicated can be adapted to the screening opportunities provided within established public health activities.[E]

Islet Autoantibodies Develop Throughout Childhood



Frída
Typ 1 Diabetes: Fröh erkennen - Fröh gut behandeln
Lancet Diabetes and Endocrinol 2024

First screen at 2 yrs

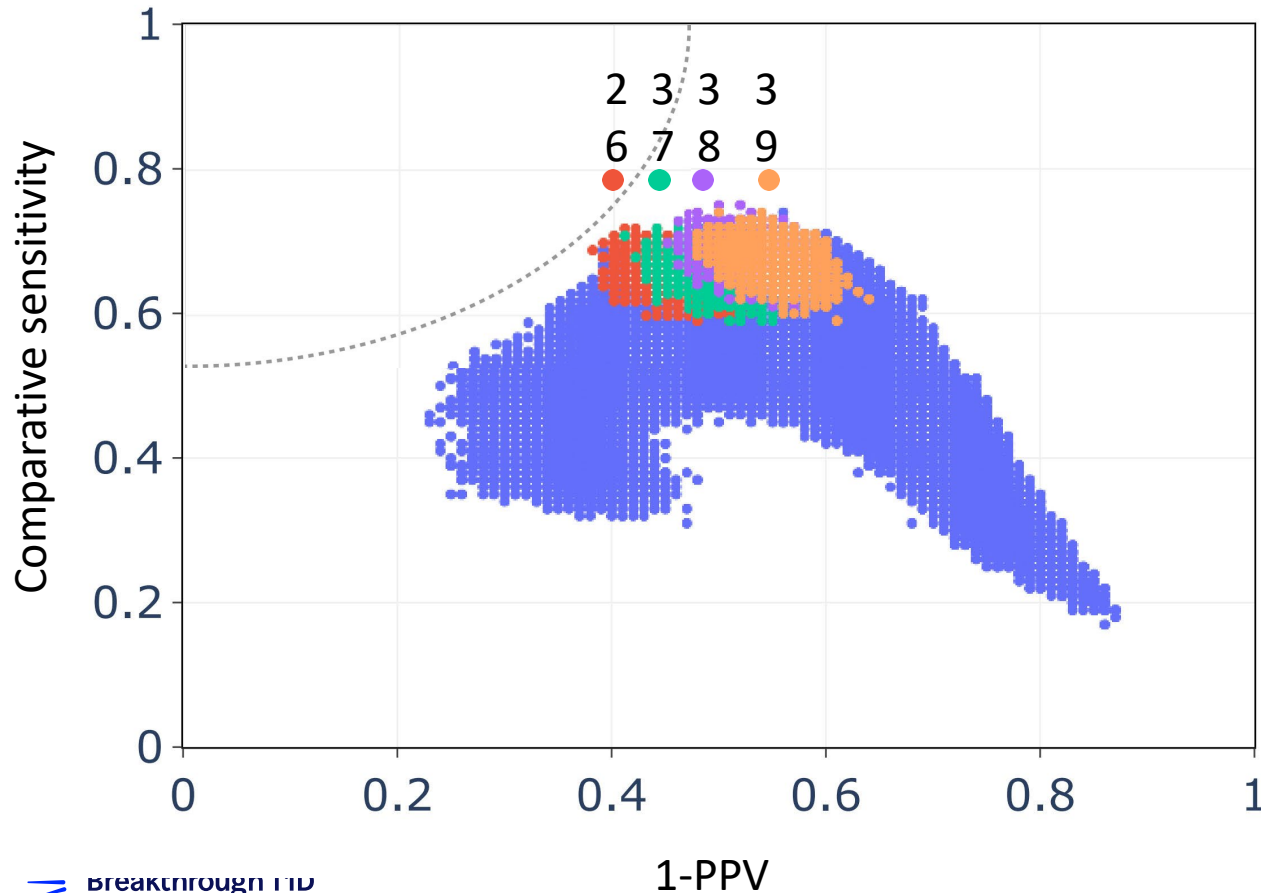
0.21%

Rescreen age 6 yrs

0.19%

Screening for islet autoantibodies at 2 and 6 years can detect 82% of future T1D cases

From 10,000 run bootstrap set for each screening-age pair



Test ages, y	Screening outcome	T1D dx, y	Sensitivity
2	≥ 1 autoAb	2 - 15	54% (50-58)
2 and 6	≥ 1 autoAb	2 - 15	82% (79-86)
2 and 6	≥ 2 autoAb	2 - 15	63% (59-67)

Islet Autoantibodies Develop Throughout Childhood:

2-3 Screens Are Optimal For High Sensitivity

Sensitivity

One screening: age 3-4 years (BABYDIAB, TEDDY, T1DI): **40% - 54%**

Two screenings: age 2-3 and 6-7 years (TEDDY, T1DI, Fr1da): **65% - 82%**

Three screenings: age 2-3 and 6-7 and 10 years (T1DI, Fr1da): **80% - 87%**

BMJ open 2016

Diabetes Care 2019

Lancet Diabetes and Endocrinol 2022

Lancet Child Adolesc Health. 2023

Diabetologia 2025

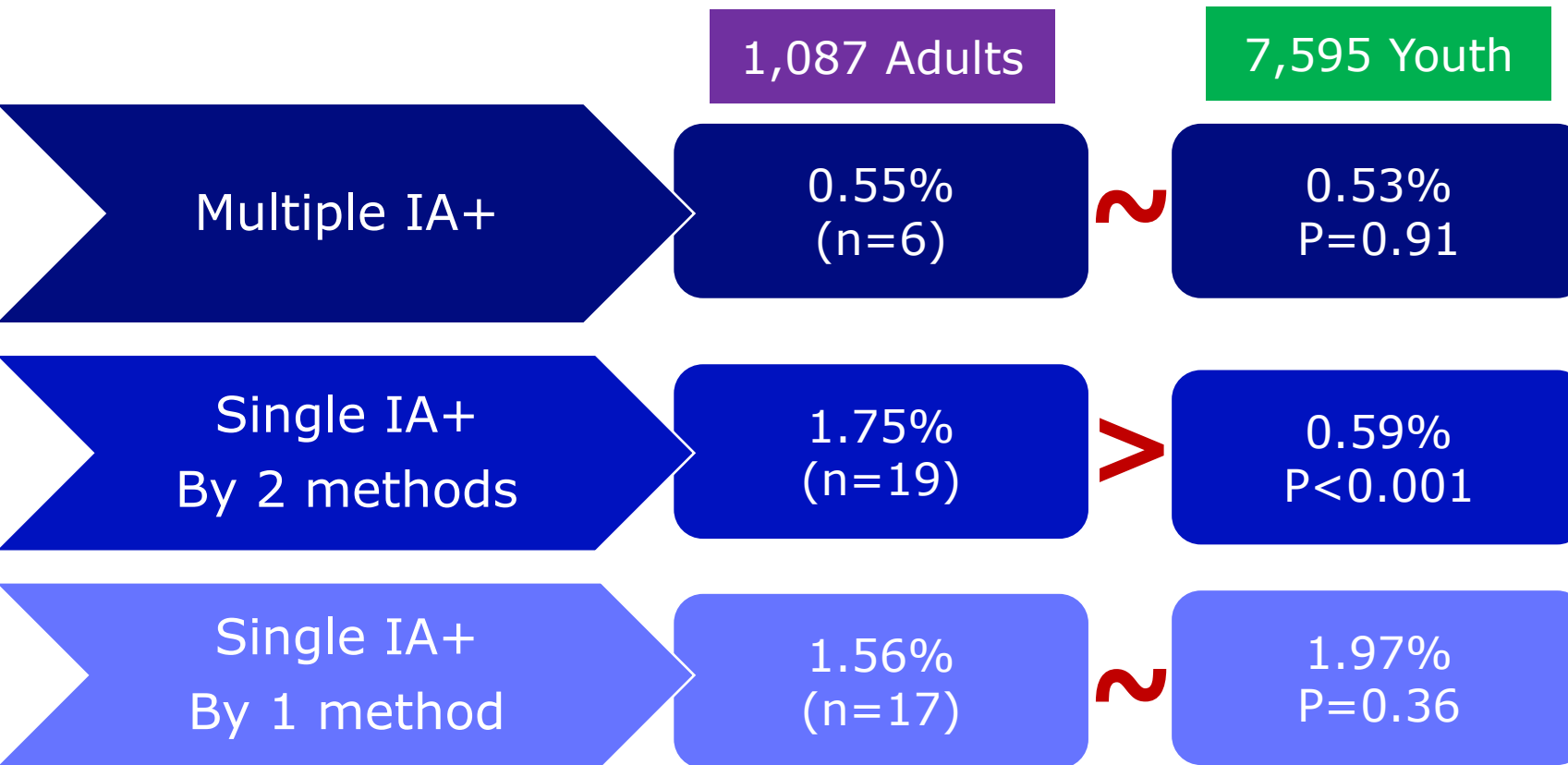
DOM 2025

Recommendations on age and cadence of screening (Adults)

Less is known about early-stage T1D progression in the general population older than 15 years.

If not done earlier, a one-time screening after 15 years of age may be considered in conjunction with established public health activities. [E]

Colorado population screening: Rate of multiple IA+ similar in adults and youth

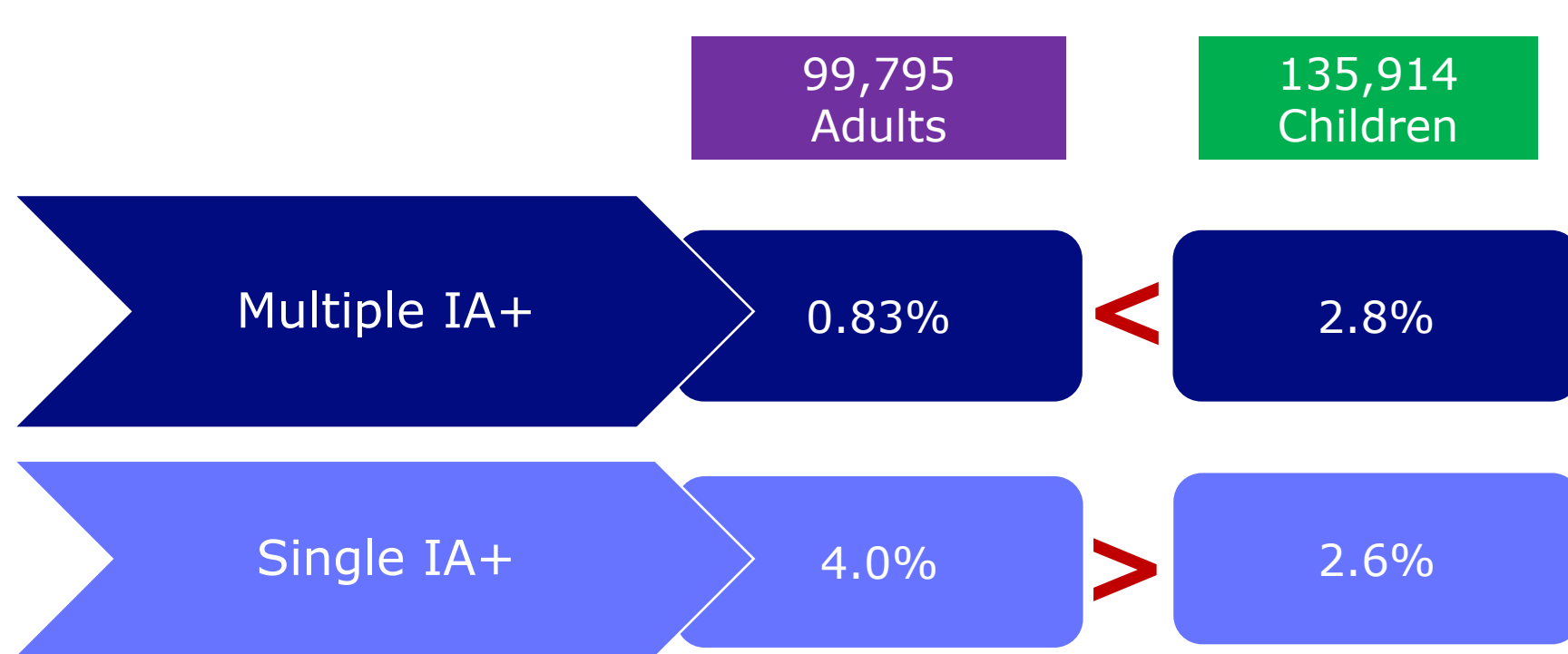


- 4 antibody screen:
 - IAA, IA2A, GADA, ZnT8A
- Adult mean age 41y (20-64)
- Youth mean age 9.3y (1-18)
- Frequency matched:
 - FDR
 - Race/ethnicity
 - Sex

TrialNet Screening of relatives:

Adults were less likely mIA+, more likely sIA+

Diabetes Care 2025



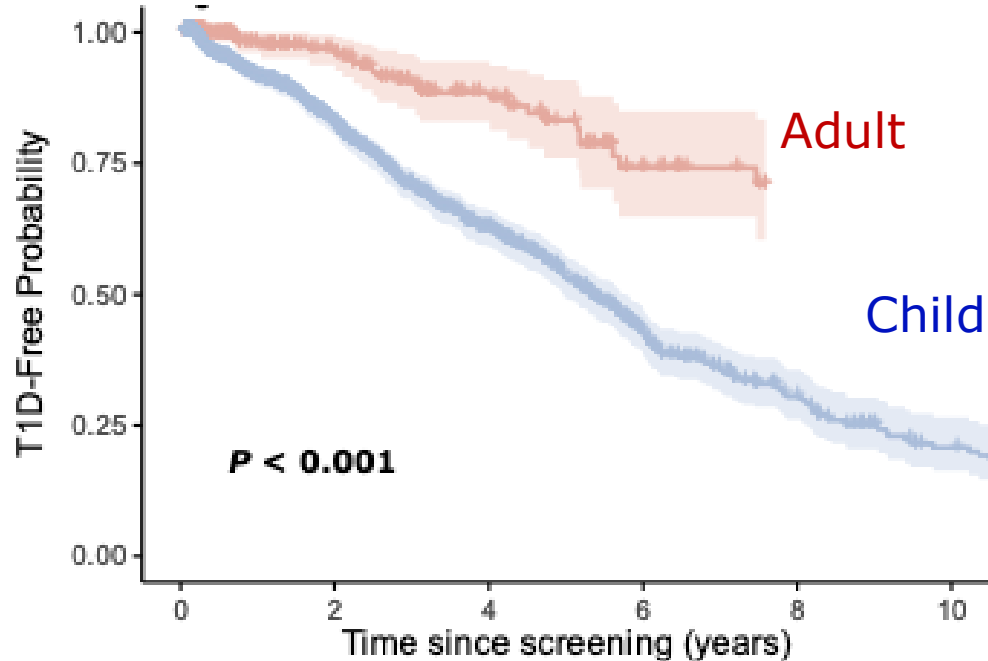
- 4 antibody screen:
 - IAA, IA2A, GADA, ZnT8A
- Adults more likely first-degree

T1D progression in people older than 15 years.

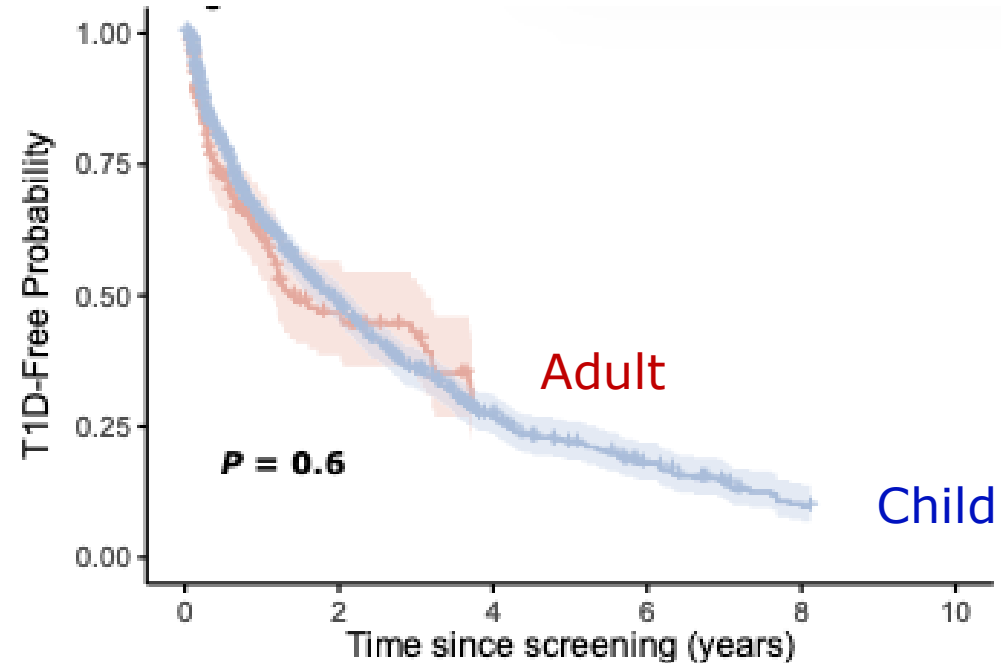
Progression from stage 2 to stage 3 is similar in children and adults with family history of T1D.

Diabetes Care 2025

Progression from Stage 1 to Stage 3



Progression from Stage 2 to Stage 3



T1D prevalence and progression in people older than 15 years.

- Prevalence of islet Abs may be similar or lower
- Little information about positive predictive values to clinical diabetes
(Diabetologia 1999, Diabetes 2023, Diabetes Care 2025, J Endocrine Soc)
- Adults have a considerable prevalence of DKA at diagnosis
(Diabetes Care 2022, Diabetologia 2024)

Conclusions

- ❖ All children should be screened 2-3 times during childhood:
 - ❖ 2-3, 6-7 and 10-15 years
- ❖ Targeted screening of at-risk groups may be a reasonable first step.
- ❖ For those older than 15 years, a one-time screening should be considered
- ❖ Screening program requires infrastructure for:
 - ❖ Confirmation
 - ❖ Monitoring