### An Update in Management of Type I Diabetes

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## **Diabetes Definition**

- Fasting blood sugar > 126 mg/ dl
- Random glucose in symptomatic patient of  $\geq$  200 mg/dl
- 2 hr OGTT sample > 200 mg/dl
- Hemoglobin A1c <u>></u> 6.5 %

# Pathophysiology of Type I

- Autoimmune islet cell injury, profound insulin deficiency
- Genetic
  - 5 % FH type I
  - HLA
- Environmental

# Epidemiology-Type I

- 2 nd most common chronic disease after asthma
- 1 in 250 children by age 18
- 50-70% present under age 20 and 30-50% over age 20
- Incidence is increasing

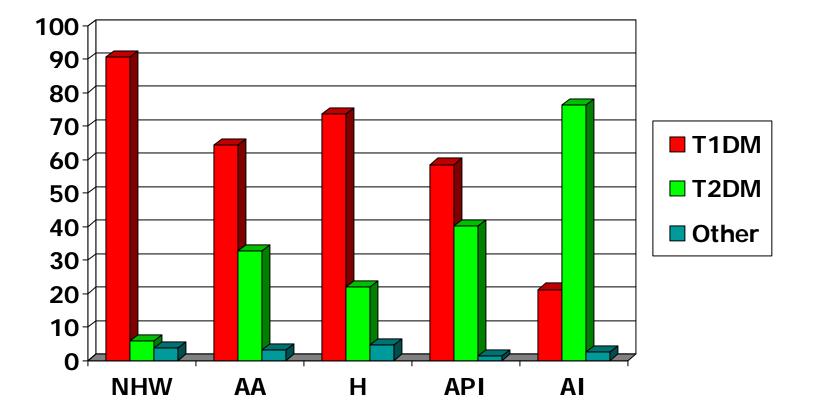
# **Clinical Presentation**

- Polyuria, polydipsia, nocturia
- Weight loss
- Candida perineal infection
- Diabetic ketoacidosis- DKA
  - Fruity odor to breath
  - Nausea, vomiting, abdominal pain
  - Chest pain, difficulty breathing, mental status changes

### Differentiation of Type I vs II

	<u>Type I</u>	<u>Type II</u>
Clinical presentation	Usually fairly rapid	Usually indolent
Age/ PE	Any age <u>&gt;</u> 20 % obese	Usually over 10 yrs 80 - 90% obese Acanthosis
Diabetic Ketoacidosis	35-45 %	5-25 %
Ethnicity	More common in Caucasian	More common in African American, Hispanic, American Indian

# Type of Diabetes by Ethnicity



Pediatrics 2006; 118:1510

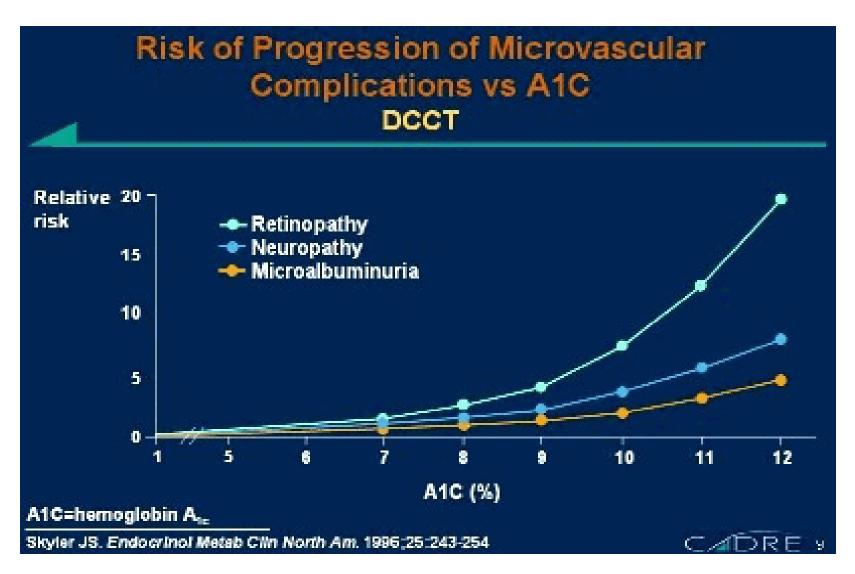
# Take Home Point

- Half of all Type I diabetics presents after childhood, and with the increasing prevalence of obesity a Type I diabetic may have the phenotype of a Type II diabetic
- All new diabetics should have anti-pancreatic antibodies tested
  - GAD 65, IA-2 and insulin antibodies

# Intensive Control Guideline

Age	A1c	Preprandial
		Blood Sugar
<u>≤</u> 6	7.5-8.5 %	100-200
6-12	<u>&lt; 8 %</u>	80-180
13-19	<u>&lt;</u> 7.5 %	70-150
Adult	< 7 %	70- 130

## **Therapeutic Rationale**



# Type I Therapy

- Exercise increases glucose transport at the cellular level
- Both sports and daily activities
   should be encouraged



- Carbohydrate (CHO) <u>not</u> under 130 grams/d
  - Best sources are fruits , vegetables, whole grains, legumes and low-fat dairy
- Total CHO guided by current age, gender, BMI and physical activity level
- Saturated fat less than 7%, limit trans fats, cholesterol under 200 mg/d
- Protein 15-20 % total energy needs

What Can I Eat?

#### My Starter Meal Plan – Women

1. Select and Circle Your Activity Level		
Sedentary	Sit most of day, do little walking, watch TV/read in evening	
Moderately Active	Exercise 20 minutes 2-4 times/week – walking around office, climbing stairs	
Active	Exercise 30 minutes 5+ times/week – physical labor, lifting	

2. Select and Circle Your Weight Goal				
	Weight Loss Weight Maintenance			
Activity Level	Carb grams	Calories	Carb grams	Calories
Sedentary	140	1400	160	1600
Moderately Active	160	1600	180	1800
Active	180	1800	200	2000

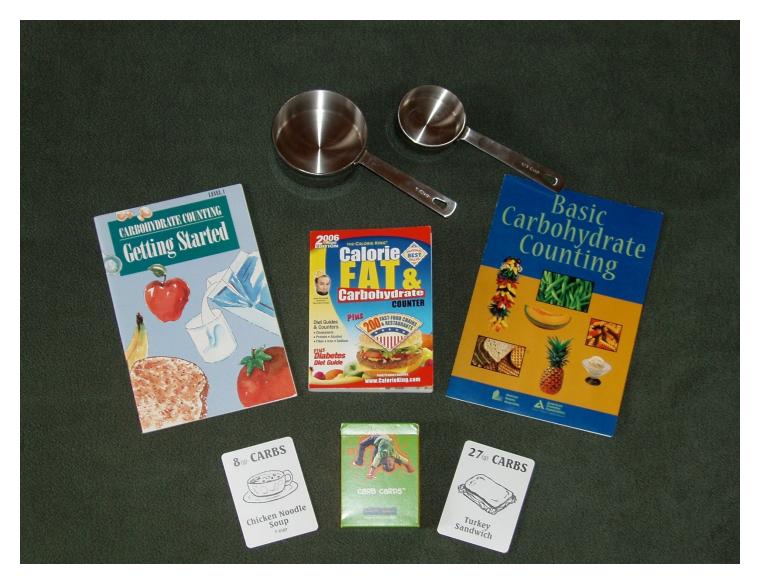
Example of How To Spread Carbs Throughout Meals						
Carb Grams Breakfast Lunch Dinner						
140	45	45	45			
160	45	60	60			
180	60	60	60			
200	60	60	75			

\* Remember to include snacks in total daily carbs if you choose to snack.

3. My Starter Meal Plan (Write-In Total Daily Carbs & Carbs At Each Meal)							
Total Carb Grams	ns Breakfast Lunch Dinner						
* Remember to include snacks in total daily carbs if you choose to snack.							
Snack(s)							

- CHO nomenclature
  - Most common to calculate grams of total CHO
  - Some use a <u>CHO</u>
    <u>"serving or choice</u>"
    description where
    each serving is equal
    to 15 grams of CHO

Serving Size 8 fl oz (24 Servings Per Container Amount Per Serving Calories 130 Calories fro	8
Total Fat 0g	0%
Sodium Omg	0%
Potassium 450mg	13%
Total Carbohydrate 30	g 10%
Sugars 24g	
Protein 2g	
Vitamin C 120% • Calcium	35%
Thiamin 6% • Riboflav	vin 4%
Niacin 2% • Vitamin	B <sub>6</sub> 6%
Folate 15% • Magnes	ium 6%
Not a significant source of saturate cholesterol, dietary fiber, vitamin A " Percent Daily Values are bas 2,000 calorie diet.	and iron.

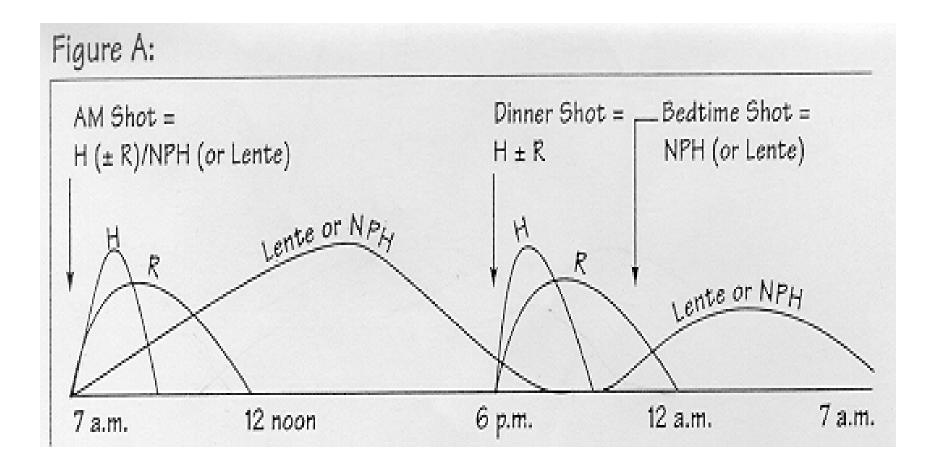


- Misleading labels
  - 1 oatmeal raisin cookie = 17 grams CHO
  - 1 sugar–free oatmeal cookie = 16 grams
    CHO
  - 1 fat free oatmeal raisin cookie = 25 grams
    CHO

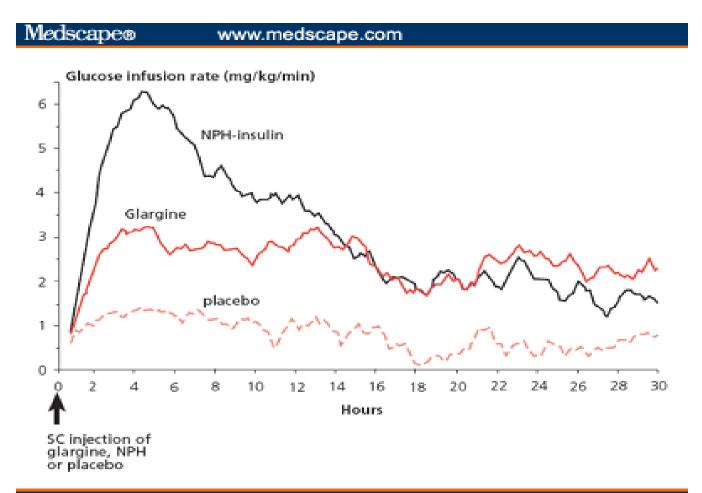


 Underlying principle of medical nutrition therapy in diabetes is <u>estimating</u> <u>carbohydrate intake</u> and matching this intake to medication

# **Classic Insulin Regimen**

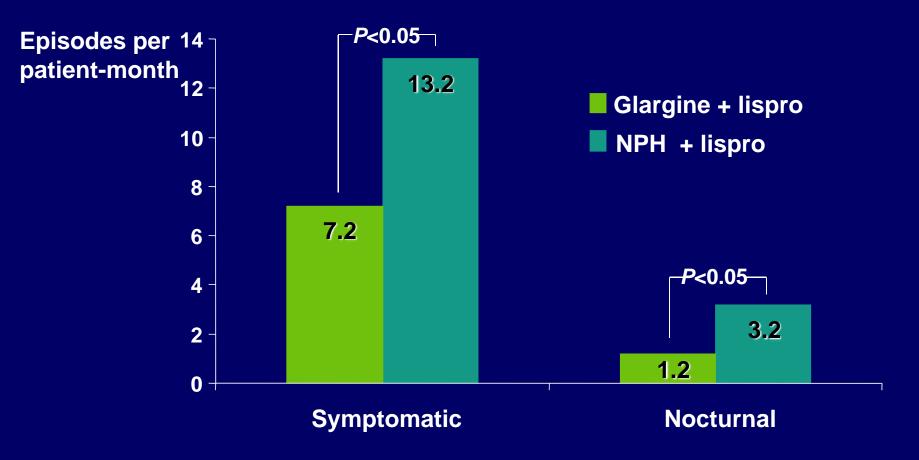


### **Basal Insulin**



### **Basal Insulin and Hypoglycemia**

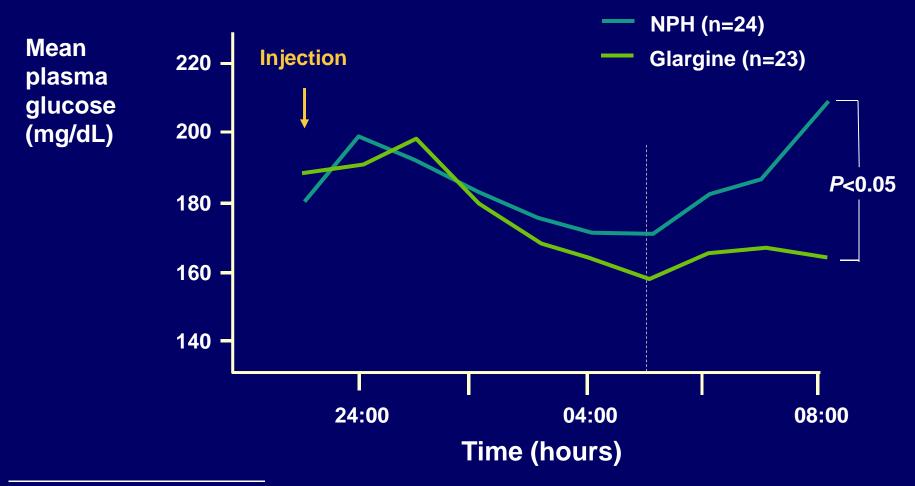
### **Nocturnal and Symptomatic Hypoglycemia**



Porcellati F et al. *Diabet Med.* 2004;21:1213-1220

### Glargine vs NPH Insulin Dawn Phenomenon

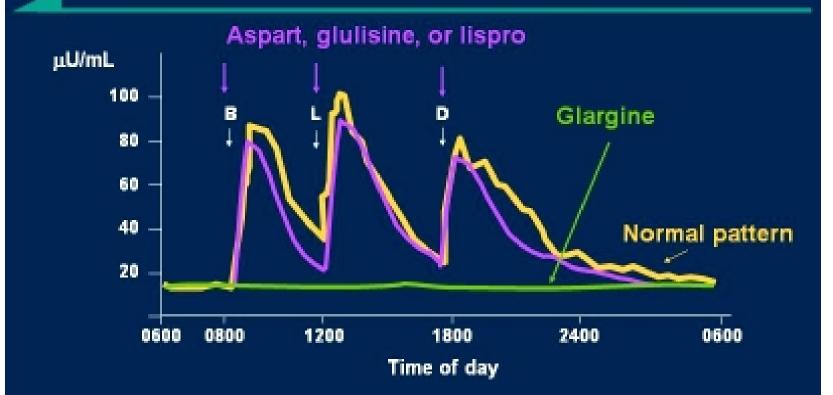
**47 Patients With Type 1 Diabetes** 



Rosenstock J et al. *Diabetes Care.* 2000;23:1137-1142

## **Basal -Bolus Therapy**

### Basal-Bolus Insulin Treatment Long- and Rapid-Acting Insulin Analogues



B=breakfast; L=lunch; D=dinner



# **Basal -Bolus Therapy**

TABLE 1: Main characteristics of rapid and long-acting insulin analogues.

Analogue	Trade name/manufacturer	Onset (min)	Peak (hrs)	Duration (hrs)
Long-acting analogues				
Glargine	Lantus/Sanofi-Aventis	4–6 hrs	No peak	>24 hrs
Detemir	Levemir/Novo Nordisk	4-6 hrs	8–10 hrs	$\sim \! 17  hrs$
Rapid-acting analogues				
Lispro	Humalog/Eli Lilly	15–30 min	0.5–2.5 hrs	36.5 hrs
Aspart	Novorapid/Novo Nordisk	10–20 min	1-3 hrs	3–5 hrs
Glulisine	Apidra/Sanofi-Aventis	10–15 min	1-1.5 hrs	3–5 hrs

# Bolus insulin – Sliding scale

Blood Glucose (mg/dl)	Breakfast Bolus	Lunch Bolus	Supper Bolus	Snack Bolus
<70	Treat low blood sugar with 15 grams carbohydrate, recheck in 15 minutes, retreat until above 70, then take insulin in 70-90 row.			
70 - 90	2	2	4	0
91 - 130	4	4	6	1
131 - 150	5	5	F	2
151 - 200	6	6	8	3
201 - 250	Ŧ	F	9	4

AM LONG ACTING INSULIN TYPE\_\_\_\_\_ DOSE\_\_\_\_\_TAKE BEFORE BREAKFAST PM LONG ACTING INSULIN TYPE\_<u>Lantus</u> DOSE\_\_\_\_\_T<u>AKE BEFORE SUPPER</u> T**X**KE AT BEDTIME Bolus Insulin Calculation Advanced CHO counting

 1 unit per X grams CHO consumed is estimated based on age, weight, activity level

- Example
  - 1 unit per 30 gram
  - If 90 grams is eaten at the meal
  - Short acting insulin for food is 3 units

# Bolus Insulin Calculation-Advanced

- Correction for high blood sugar if necessary
- Example
  - Blood sugar is 300 mg /dl
  - Target blood sugar is 150
  - Correction factor 50 = 1 unit of insulin will lower blood sugar by 50 mg/dl
  - 300 actual -150 target , then divided by CF of 50 = 3 units for the high blood sugar

# Insulin Pump Therapy

- Only contain short acting insulin
- Programmable basal rate
- Have bolus calculator feature that guides dosage when CHO grams entered and blood sugar entered
- An important safety feature is the <u>insulin</u> on board calculation
  - Factors duration of insulin action into dose calculation to prevent insulin over dosage

## Animas Ping

I always feel close to you.



### OneTouch<sup>®</sup> Ping.<sup>™</sup> Meet the brains of this outfit.

Even when we're 9.84 feet apart.

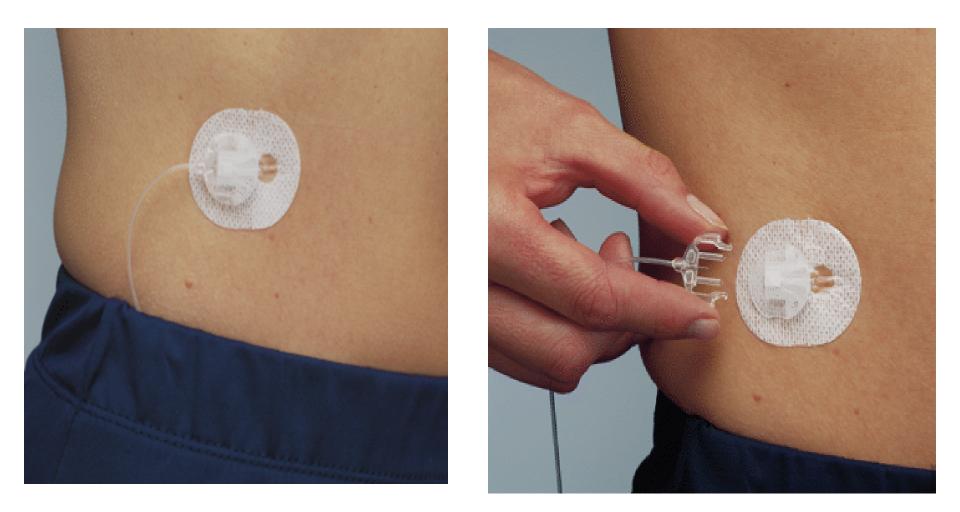




### MiniMed Paradigm



### Infusion Sets



### <u>OmniPod</u>



# <u>Continuous Glucose Monitoring-</u> <u>CGM</u>

- Measure interstitial glucose each 5 minutes
- Must calibrate with fingerstick blood sugars
- Replace sensor each 3-7 days
- High and Low blood sugar alarms
- Blood sugar trend arrows
- To dose insulin a fingerstick blood sugar is required



#### MiniMed REAL-Time Paradigm



#### **MiniMed REAL-Time Guardian**

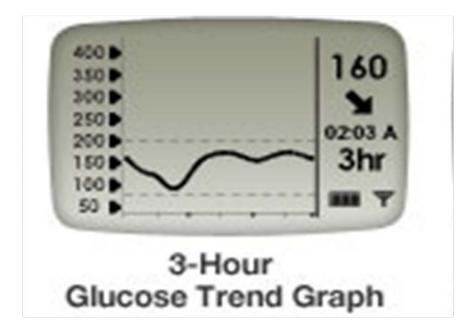


DexCom



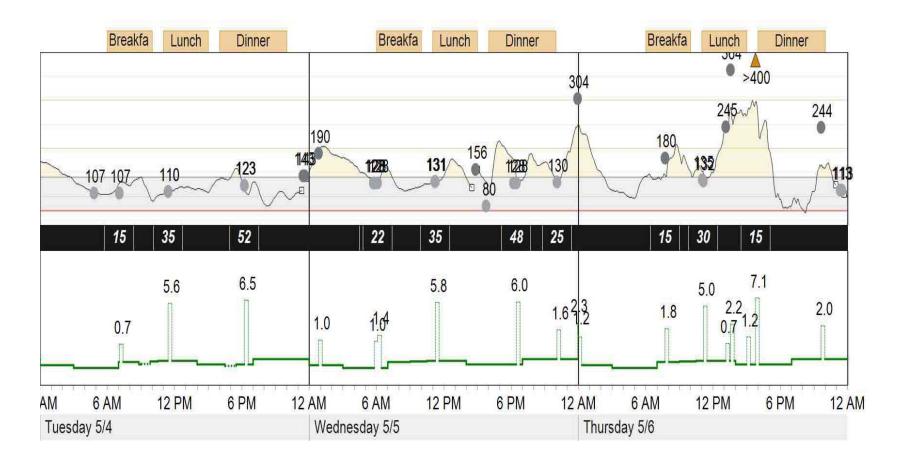
## <u>Displays</u>



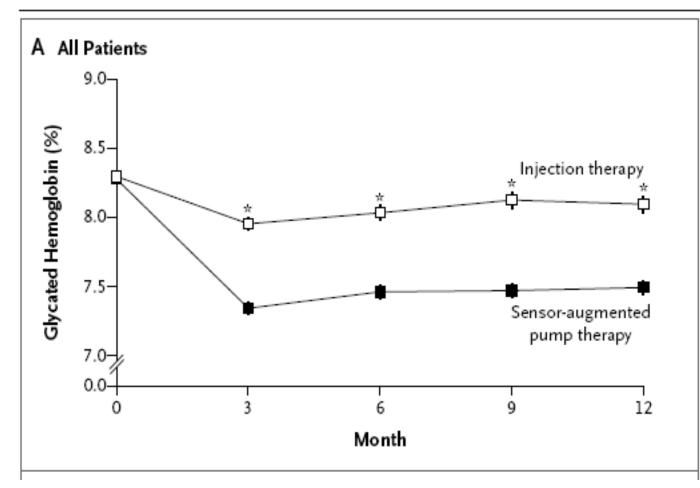


# Download Data to review for Blood Sugar Patterns

**Sensor & Meter Overview** 



### Sensor Augmented Pump Therapy



NEJM July 2010

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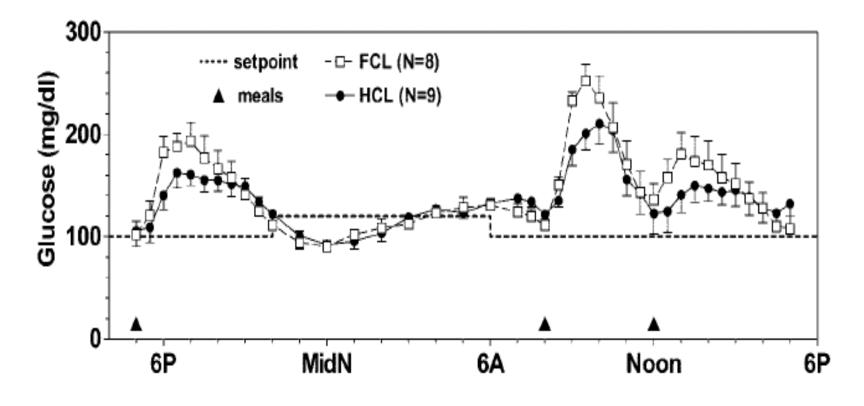
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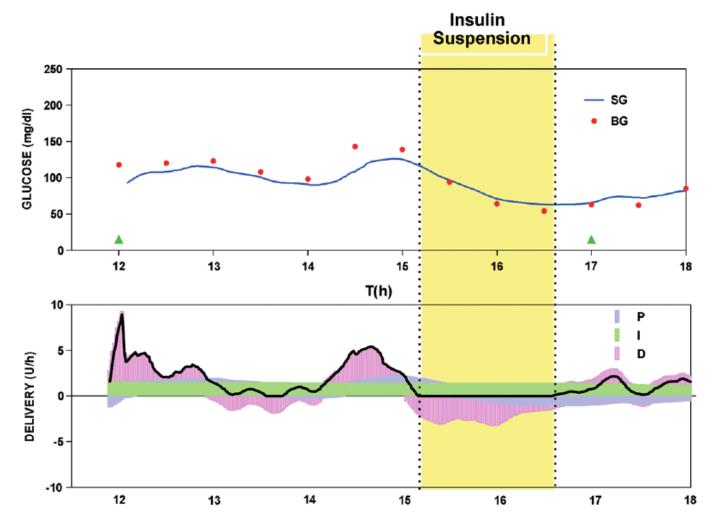
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# <u>Artificial Pancreas Project-</u> <u>Closed Loop</u>



Diabetes Care may 2008

# Artificial Pancreas Project-Closed Loop



Diabetes Technology Therapy April 2009

### Type I Diabetes Immunotherapy

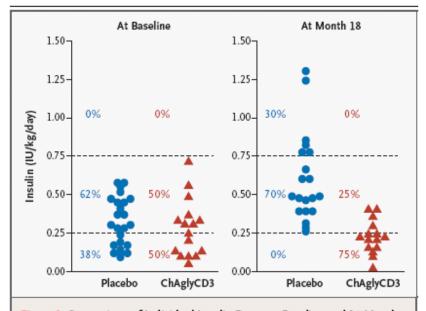
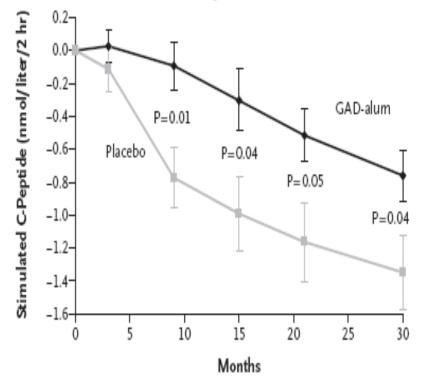


Figure 2. Comparison of Individual Insulin Doses at Baseline and 18 Months in Patients with an Initial Secretory Response at or above the 50th Percentile. Circles represent the placebo group, and triangles the ChAglyCD3 group.

NEJM June 2005

E Patients Treated <6 Mo after Diagnosis



NEJM October 2008

# Islet cell transplantation

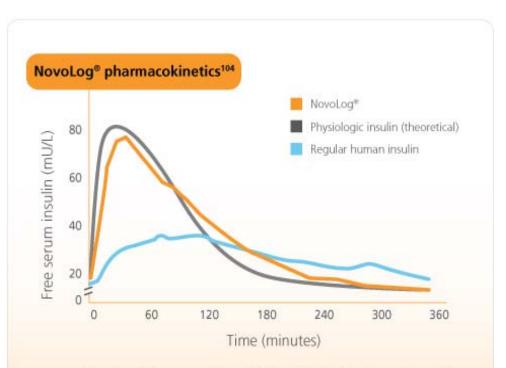
- Initial success to obtain insulin independence with the Edmonton protocol published in 2000
- More than 50 % must restart insulin within 2 years, although residual insulin from transplant did smooth blood sugar control
- Subsequent complete transplant failure
- Limited islet supply
- Immunosuppressive drugs increase cancer risk

# <u>Hypoglycemia</u>

- <u>Minidose Glucagon</u>
  - To treat hypoglycemia when patient is alert but due to nausea or vomiting cannot take oral fluids
  - Less risk of emesis from the Glucagon
  - 15 units measured on an insulin syringe and given just like insulin
- Alcohol
  - Decreases gluconeogenesis

### **Hyperglycemia**

- <u>Causes</u>
  - Blood sugar tested
    too close to the last carbohydrate intake and insulin dose
  - Insufficient or missed insulin dose



 NovoLog<sup>®</sup> takes a median of 40 to 50 minutes to reach peak serum insulin concentration

### <u>Ketones</u>



### <u>Ketones</u>

- Urine ketone test
  - Inexpensive, 16 cents
    each
  - Measures

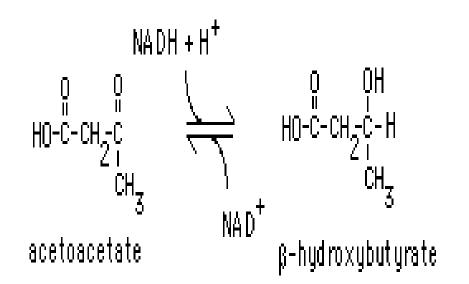
#### acetoacetate

May not reflect current status

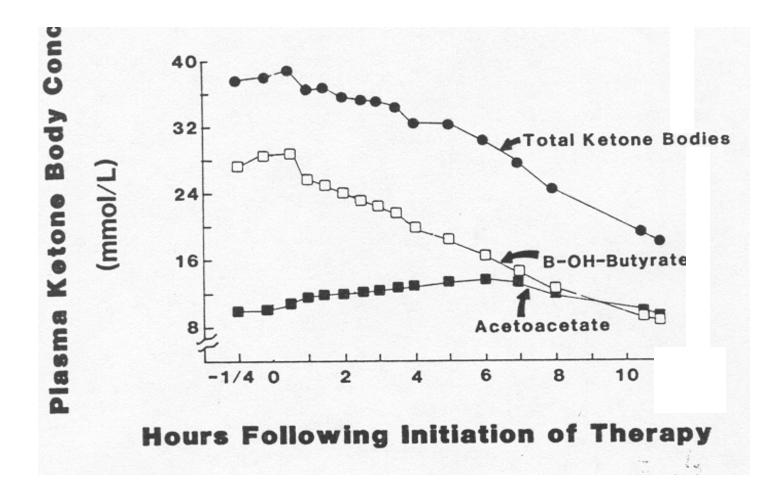
- Blood ketone test
  - Expensive, \$3.75each strip
  - Measures **B**
    - hydroxybutyrate ( BHB )
  - Reflects current status

### <u>Ketones</u>

- When the blood pH is closer to normal Acetoacetate is the <u>major</u> form
- When the blood pH is lower B-hydroxybutyrate is more prevalent
- As blood pH improves Bhydroxybutyrate is <u>converted</u> to acetoacetate



### $\beta$ -hydroxybutyrate is a better indicator of metabolic status when detecting and treating DKA



# Comparison of Blood and Urine

Ket	ton	es

<u>Blood Ketones ( mmol/l)</u>	<u>Urine Ketones</u>
< 0.6 mM	Negative
0.6 mM to 0.9 mM	Trace to small
1.0 mM to 1.5 mM	Moderate
1.6 to 3.0 mM	Large
≥ 3.0 mM	Very Large

### **Diabetic Ketoacidosis Therapy**

- Conservative fluid therapy- initial bolus
  10 cc/kg normal saline , then IV fluid (NS- first 4-6 hr, then ½ NS) at 1.5 times maintenance
- Electrolyte replacement Na, K, P
- Continuous Intravenous low dose
  (0.1unit /kg /hr) regular insulin therapy
- <u>No</u> insulin bolus recommended
- Bicarbonate therapy rarely used unless ionotropic dysfunction. Use is associated with increased risk of cerebral edema

# **Type I Diabetes Therapy**

- This talk has been glucocentric
- However management of other cardiovascular risk factors is crucial
  - Hyperlipidemia
  - Hypertension
  - Smoking cessation
  - Vitamin D deficiency

### **Education**

- PLANNING FOR COLLEGE:
- A Workshop for High School Students with Diabetes and Their Families
- Learn how to prepare yourself NOW for life at college when you graduate.
- March 3, 2011 at 6pm
- Joslin Diabetes Center
- 3229 East Genesee Street (across from Nottingham High School)
- Syracuse, NY 13214 RSVP: 464-8668