# CGM: Clinical need and Physiology

Jens Sandahl Christiansen
MD, DMSc, FRCPI
Aarhus University Hospital
Denmark



#### Conflicts of interest

None to report



#### Outline of presentation

- Association between glucose levels and complications
- The need to monitor glucose variability
- Feasibility of monitoring glucose levels



# Micromethods for blood glucose determination

Bang: Wiesbaden, 1913
Hagedorn and Norman Jensen,
Copenhagen, 1918
(100μl, 30-60 min)







# Glycaemic control vs late diabetic complications

M. Siperstein, JCI; 47:1973, 1968

R. Østerby, Acta Med Scand; 574:1, 1975

J. Pirart, Diabete Metab; 3:97, 1977

Steno I study, Lancet; i:121, 1982

DCCT, N Engl J Med; 329:977, 1993

UKPDS, Lancet; 352:837, 1998

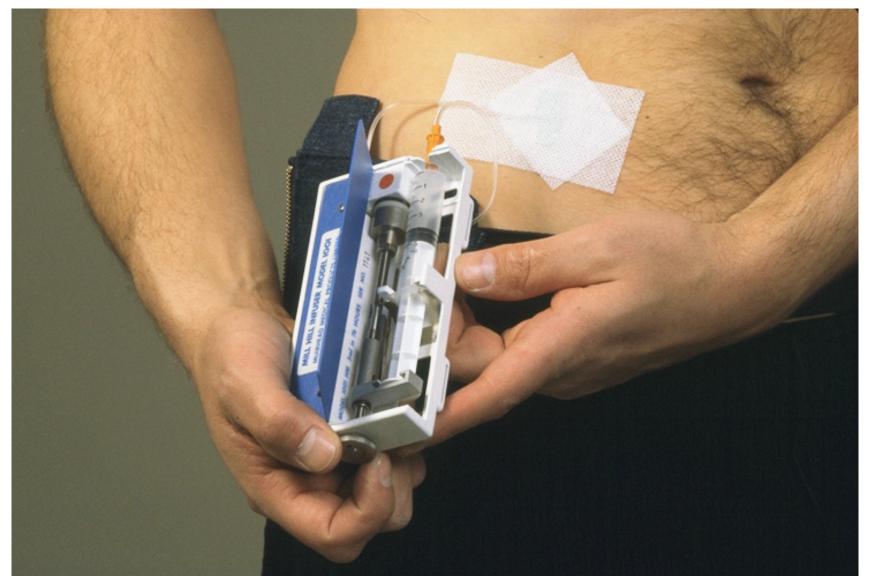














#### Steno I study, Lancet; i:121, 1982

DE MEDICINSKE AFDELINGERS
BIBLIOTEK
BIBLIOTEK
ARHUS KOMMUNEHOSPITAL
SOOD ARHUS C

The Lancet · Saturday 16 January 1982

EFFECT OF 6 MONTHS OF STRICT METABOLIC CONTROL ON EYE AND KIDNEY FUNCTION IN INSULIN-DEPENDENT DIABETICS WITH BACKGROUND RETINOPATHY

Steno Study Group\*

Steno Memorial Hospital and Department of Ophthalmology, Gentofte Hospital, DK-2820 Gentofte, Denmark

Summary The effect of 6 months of strict metabolic

microangiopathy in rats<sup>3</sup> and dogs,<sup>4</sup> and diabetic glomerulopathy in rats has been reversed by islet transplantation.<sup>5,6</sup> Eschwege et al.<sup>7</sup> found that patients treated with multiple insulin injections developed microaneurysms at a slower rate than those treated with one injection a day. Short-term strict metabolic control reduces urinary albumin excretion in insulin-treated diabetics,<sup>8</sup> but no long-term studies have been done.

We present the first report, based on a prospective randomised study of the effects of long-term control achieving near-normal blood-glucose levels on certain aspects

> Diabetes Technology Society

#### DCCT, N Engl J Med; 329:977, 1993

## The New England Journal of Medicine

©Copyright, 1993, by the Massachusetts Medical Society

Volume 329

**SEPTEMBER 30, 1993** 

Number 14

#### THE EFFECT OF INTENSIVE TREATMENT OF DIABETES ON THE DEVELOPMENT AND PROGRESSION OF LONG-TERM COMPLICATIONS IN INSULIN-DEPENDENT DIABETES MELLITUS

THE DIABETES CONTROL AND COMPLICATIONS TRIAL RESEARCH GROUP\*

**Abstract** *Background*. Long-term microvascular and neurologic complications cause major morbidity and mortality in patients with insulin-dependent diabetes mellitus (IDDM). We examined whether intensive treatment with the goal of maintaining blood glucose concentrations close to the normal range could decrease the frequency and severity of these complications.

interval, 62 to 85 percent), as compared with conventional therapy. In the secondary-intervention cohort, intensive therapy slowed the progression of retinopathy by 54 percent (95 percent confidence interval, 39 to 66 percent) and reduced the development of proliferative or severe nonproliferative retinopathy by 47 percent (95 percent confidence interval, 14 to 67 percent). In the two cohorts



#### UKPDS, Lancet; 352:837, 1998

ARTICLES

#### Articles

Intensive blood-glucose control with sulphonylureas or insulin compared with conventional treatment and risk of complications in patients with type 2 diabetes (UKPDS 33)

UK Prospective Diabetes Study (UKPDS) Group\*

#### **Summary**

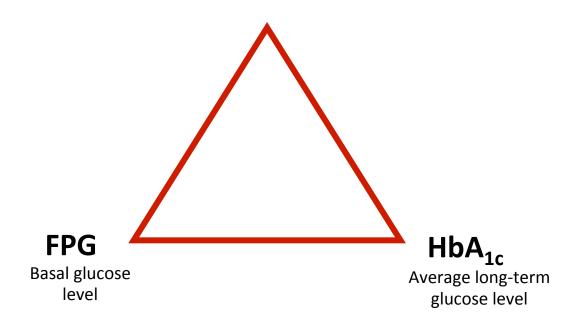
Background Improved blood-glucose control decreases the progression of diabetic microvascular disease, but the effect on macrovascular complications is unknown. There is concern that sulphonylureas may increase cardiovascular mortality in patients with type 2 diabetes and that high insulin concentrations may enhance atheroma formation. We compared the effects of intensive blood-glucose control with either sulphonylurea or insulin

Findings Over 10 years, haemoglobin  $A_{1c}$  (Hb $A_{1c}$ ) was 7.0% (6·2–8·2) in the intensive group compared with 7.9% (6·9–8·8) in the conventional group—an 11% reduction. There was no difference in Hb $A_{1c}$  among agents in the intensive group. Compared with the conventional group, the risk in the intensive group was 12% lower (95% Cl 1–21, p=0·029) for any diabetes-related endpoint; 10% lower (–11 to 27, p=0·34) for any diabetes-related death; and 6% lower (–10 to 20, p=0·44) for all-cause mortality.



#### 'Glucose triad' of diabetes management

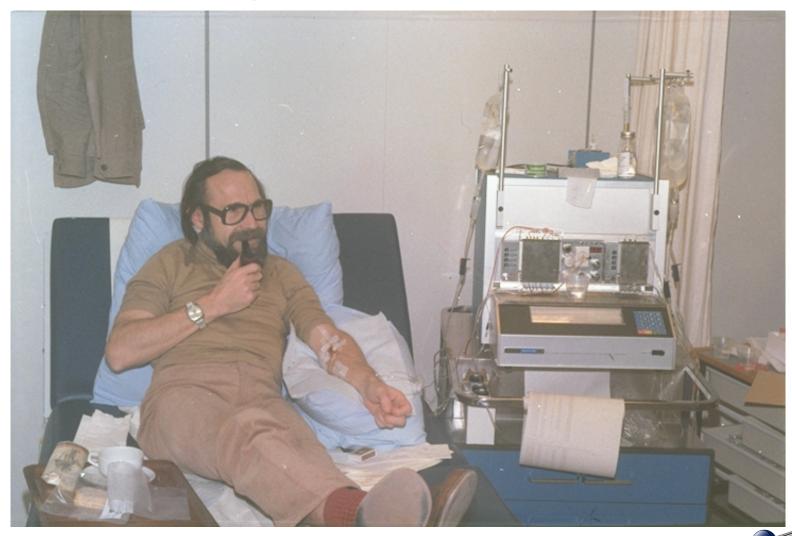
#### Postmeal glucose



HbA<sub>1c</sub> = glycated haemoglobin FPG = fasting plasma glucose



### ... using an artificial beta-cell



Diabetes Technology Society

#### Outline of presentation

- Association between glucose levels and complications
- The need to monitor glucose variability
- Feasibility of monitoring glucose levels



#### The clinical importance of CGM

- Postprandial glucose
- Hypoglycaemia(nocturnal)



#### Key questions assessed by IDF

- 1) Is postprandial hyperglycaemia harmful?
- 2) Is treatment of postprandial hyperglycaemia beneficial in improving clinical outcomes and glycaemic control?
- 3) Which therapies are effective in controlling postprandial plasma glucose?
- 4) What are the targets for postprandial glycaemic control and how should they be assessed?



## Question 1: Is postprandial hyperglycaemia harmful?



#### Is postprandial hyperglycaemia harmful?

#### **IDF Recommendation:**

Postprandial hyperglycaemia is harmful and should be addressed



# Question 2: Is treatment of postprandial hyperglycaemia beneficial in improving clinical outcomes and glycaemic control?



## Is treatment of postprandial hyperglycaemia beneficial?

#### **IDF Recommendation:**

Implement treatment strategies to lower postprandial plasma glucose in people with postprandial hyperglycaemia





Diabetes Technology Society

#### The clinical importance of CGM

- Postprandial glucose
- Hypoglycaemia(nocturnal)



## Management of glycaemia

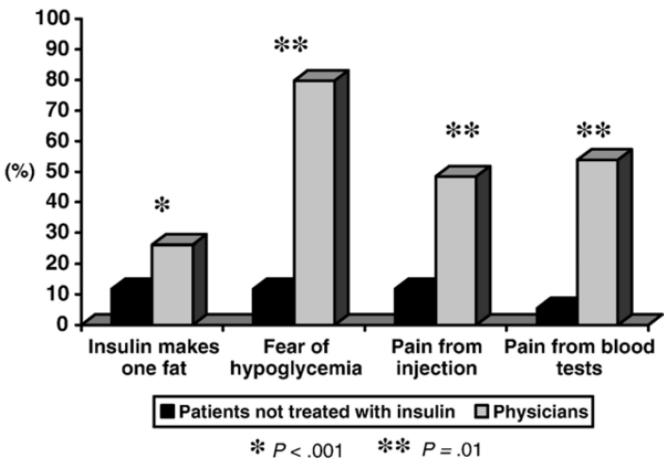
## The journey between Scylla and Charybdis





Diabetes Technology Society

#### Barriers for insulin treatment



Journal of Diabetes and Its Complications 21 (2007) 220-226



#### Outline of presentation

- Association between glucose levels and complications
- The need to monitor glucose variability
- Feasibility of monitoring glucose levels



#### What and where to monitor?



#### Vietnamese potbellied pig





## Muscle



## **SCAT**



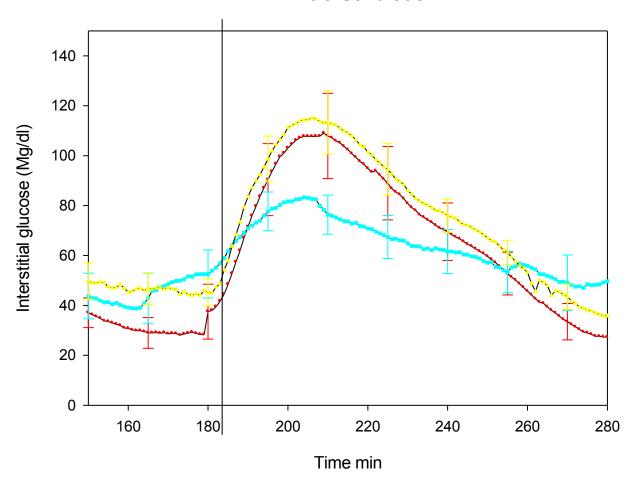


## CNS





#### In Vitro Calibration



**CNS** 

**SCAT** 

**MUSCLE** 

20 g iv glucose Challenge, Diabetes, 54:1635, 2005



# SCAT is clinically appropriate in continuous monitoring of glucose



- Financial
- Technological
- Organisational



#### **Technological**

- Robustness
- Reliability



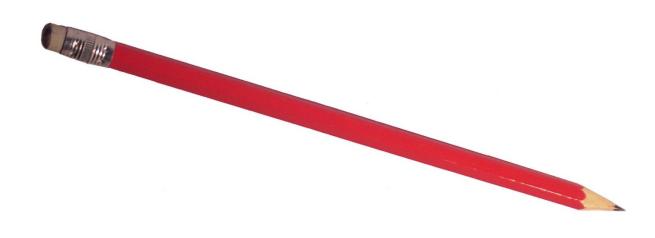
#### Optimal vs. pragmatic solution

For the first manned space shuttle flight, the Americans developed an ink-pen that could not leak in a zero gravity environment.

The costs for this development were \$ 1 million.



That's how the Russians solved the problem:





#### Organisational

The difference between RCT and everyday clinical reality



#### Organisational

- Instruction (time needed)
- Calibration (in-house, out patient)
- Maintenance



#### **Financial**

 Development is needed to provide low cost monitoring to 3. world







# Beneficial effects upon Morbidity and Mortality still needs to be finally demonstrated



#### American Sociological Review 2013 78: 26

# Egalitarianism, Housework, and Sexual Frequency in Marriage

American Sociological Review 78(1) 26–50
© American Sociological Association 2012
DOI: 10.1177/0003122412472340
http://asr.sagepub.com



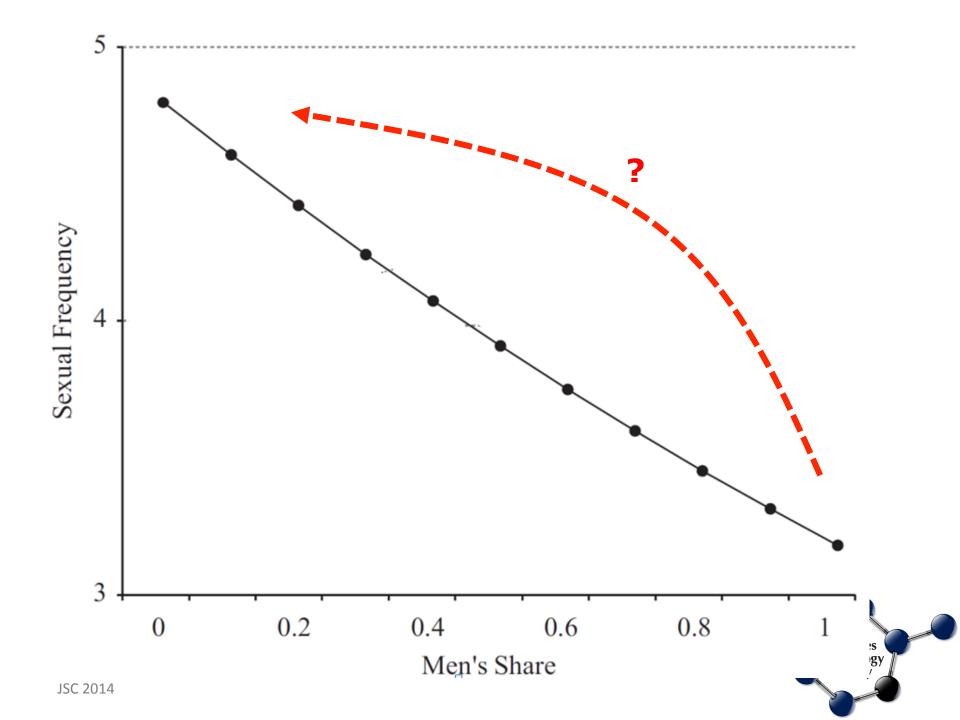
Sabino Kornrich,<sup>a</sup> Julie Brines,<sup>b</sup> and Katrina Leupp<sup>b</sup>

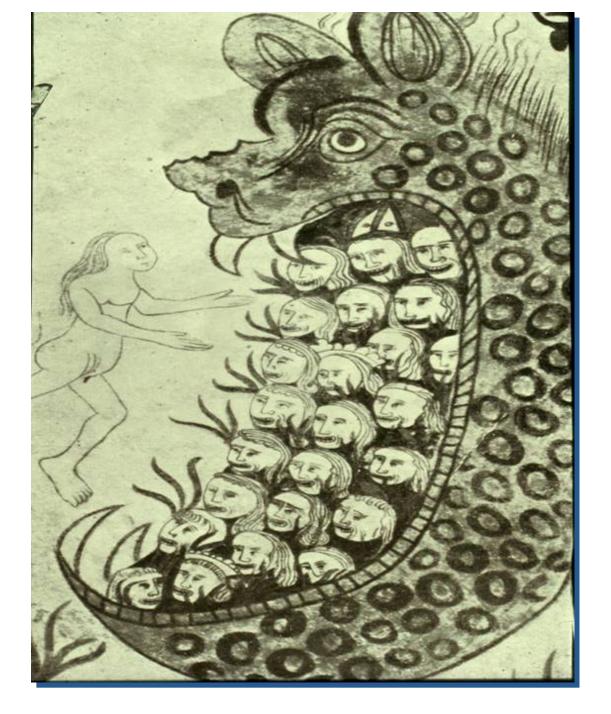
#### **Abstract**

Changes in the nature of marriage have spurred a debate about the consequences of shifts to more egalitarian relationships, and media interest in the debate has crystallized around claims that men who participate in housework get more sex. However, little systematic or representative research supports the claim that women, in essence, exchange sex for men's participation in housework. Although research and theory support the expectation that

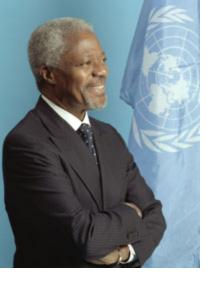
This study investigates the links between men's participation in core (traditionally female) household tasks and sexual frequency.











"It is my aspiration that health will finally be seen not as a blessing to be wished for, but as a human right to be fought for."

Kofi Annan





#### THANK YOU FOR YOUR ATTENTION!!

