SYNDROME ADDS NOTHING TO THE
CARE OF THE PATIENT WITH OR AT
RISK OF TYPE 2 DIABETES MELLITUS
AND CVD

Con: KGMM Alberti

THE "ORIGINAL" SYNDROME X

- Resistance to insulin-stimulated glucose uptake
- · Glucose intolerance
- Hyperinsulinaemia
- Increased VLDL-triglyceride
- · Decreased HDL-cholesterol
- · Hypertension

SYNONYMS

- Syndrome X METABOLIC SYNDROME
- Metabolic Syndrome X
- · Plurimetabolic Syndrome
- · Dysmetabolic Syndrome, etc.
- · Insulin Resistance Syndrome

N.B.

- » Subtle shift from insulin resistance
- » More focus on obesity

METABOLIC SYNDROME

A cluster of risk factors for diabetes and cardiovascular disease

- greater than by chance alone

THE PROBLEM

Diagnostic criteria

Major Definitions of the MetS

WHO EGIR NCEP

*prerequisite

IR	Clamp*	F-insulin*	
IGT	\checkmark		
IFG	≥6.1	6.1-6.9	≥6.1
High-BP	140/90	140/90	130/85
T <i>G</i> ≥1.7	\checkmark	\checkmark	\checkmark
Low-HDL	<0.9 (1.0)	<1.0	<1.04 (<1.29)
Waist		≥94 (≥80)	>102 (>88)
WHR/BMI	√		
Albuminuria			

IDF CONSENSUS MEETING 2004

<u>Aims</u>

- 1. To bring together individuals with different viewpoints
- 2. To establish a unified working diagnostic tool
- 3. To highlight areas where more knowledge is needed
 - N.B. Not just yet another definition

Underlying risk factors for Metabolic Syndrome

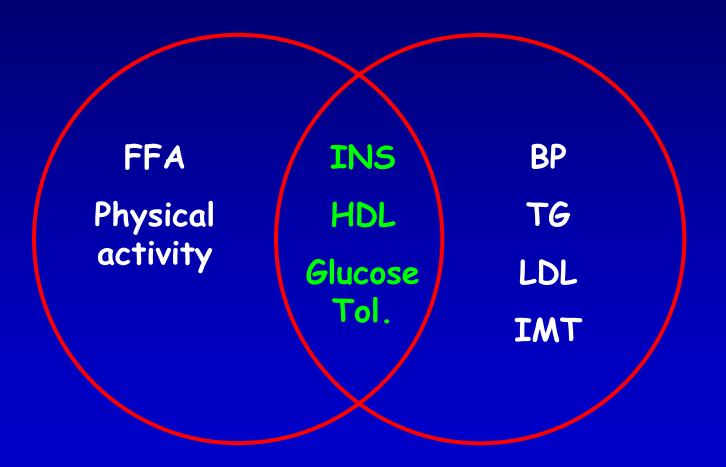
Which is dominant?

Obesity (esp. abdominal obesity)

or Insulin resistance

RISC STUDY CORRELATIONS

(Ferranini et al)



Insulin Resistance

Abdominal Obesity

Loadings of 13 age-adjusted variables related to CVD on 4 factors rotated and extracted with factor analysis in Finnish men

	Metabolic			
Factors	syndrome	Factor 2	Factor 3	Factor 4
Variance explained, %	21	14	11	8
Body mass index, kg/m ²	<u>0.76</u>	-0.28	0.18	-0.05
Waist-to-hip ratio	<u>0.69</u>	-0.18	0.19	0.00
Fasting serum insulin, mU/L	<u>0.74</u>	-0.20	-0.05	-0.17
Fasting glucose, mmol/L	<u>0.45</u>	-0.15	0.33	-0.06
Serum triglycerides, mmol/L	<u>0.58</u>	0.01	<u>-0.42</u>	0.02
HDL cholesterol, mmol/L	<u>-0.48</u>	-0.01	0.67	0.03
Systolic blood pressure, mmHg	<u>0.36</u>	-0.10	0.35	0.18
Smoking, cigarettes/day	0.14	<u>0.78</u>	0.08	-0.01
Alcohol, g/wk	0.20	0.29	0.57	0.17
LDL cholesterol, mmol/L	0.21	0.14	<u>-0.34</u>	0.59
Ischemic heart disease in family	-0.01	-0.14	0.06	<u>0.76</u>
Fibrinogen, g/l	0.26	<u>0.60</u>	0.07	0.01
White blood cells, x 109/mL	0.28	<u>0.72</u>	-0.03	-0.08

Variables with loadings on factors ≥ 0.30 are in bold and underlined.

A SINGLE FACTOR UNDERLIES THE METABOLIC SYNDROME

A confirmatory Factor Analysis

Pladevall et al. Diabetes Care 2006; 29: 113-122

Major Definitions of the MetS

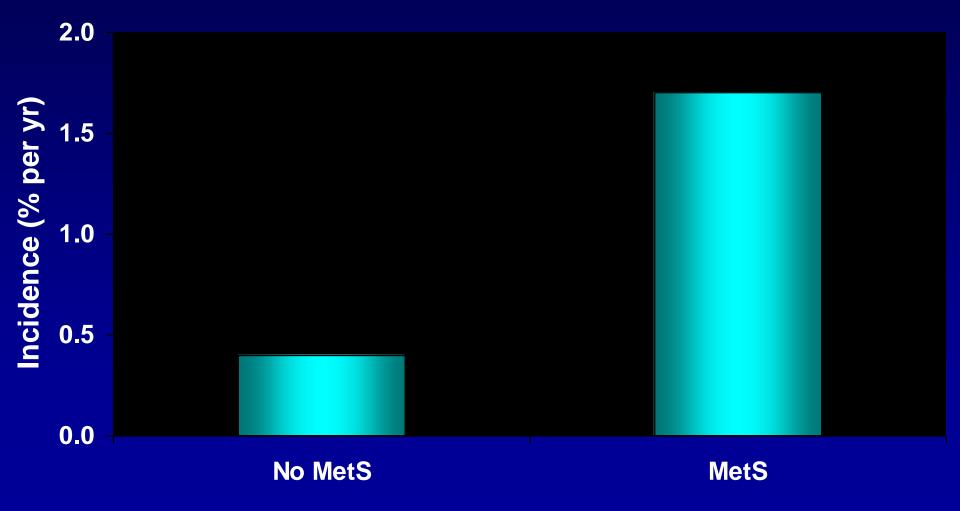
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Low-HDL	<0.9 (1.0)	<1.0	<1.04 (<1.29)	<1.04 (<1.29)
Waist		≥ 94 (≥ 80)	>102 (>88)	≥ 94 (≥ 80)*
WHR/BMI Albuminuria	√			

*prerequisite

Does the metabolic syndrome give indication of risk?

The Metabolic Syndrome increases the Risks of Diabetes 4 times





Metabolic Syndrome as a Risk Condition (Framingham Heart Study)

Cardiovascular disease

- Men: RR 2.50 x increased
- Women: RR 1.58 x increased

Type 2 Diabetes

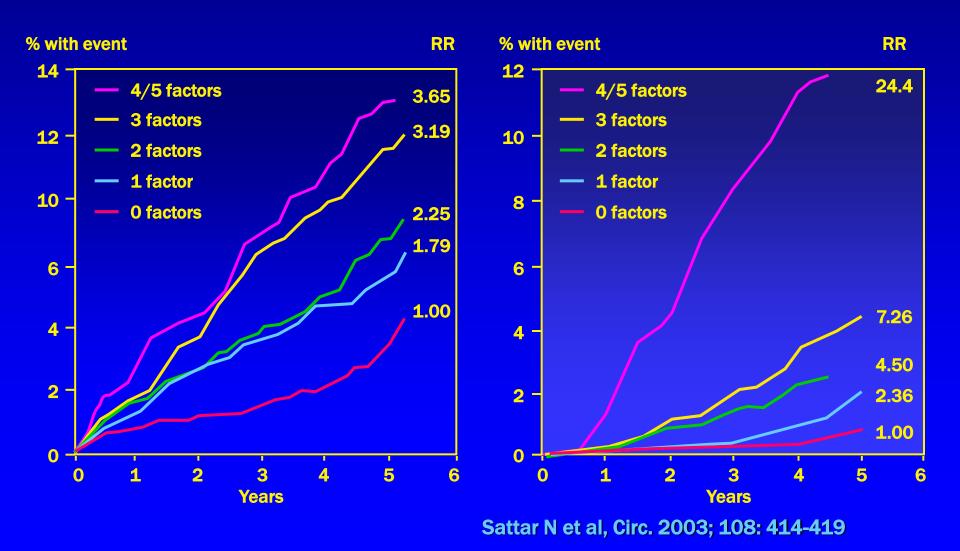
- Men: RR 4.76 x increased
- Women: RR 5.66 x increased

Eugenia Martinez Vallejo

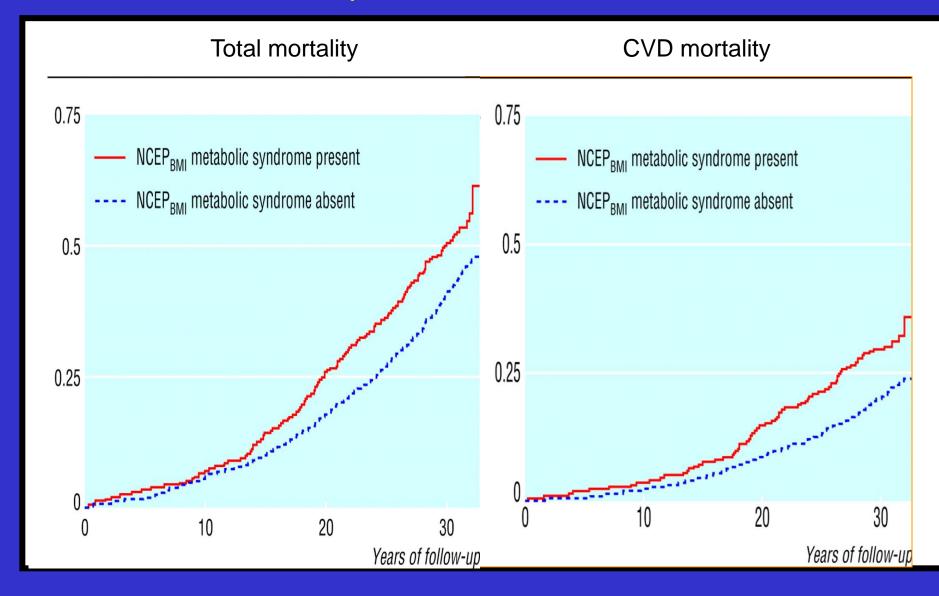
Metabolic Syndrome as a predictor of CHD and Diabetes: WOSCOPS

CHD Death/Non-fatal MI

Onset of New DM



Metabolic Syndrome and 30 year mortality in 50 year old Swedish men

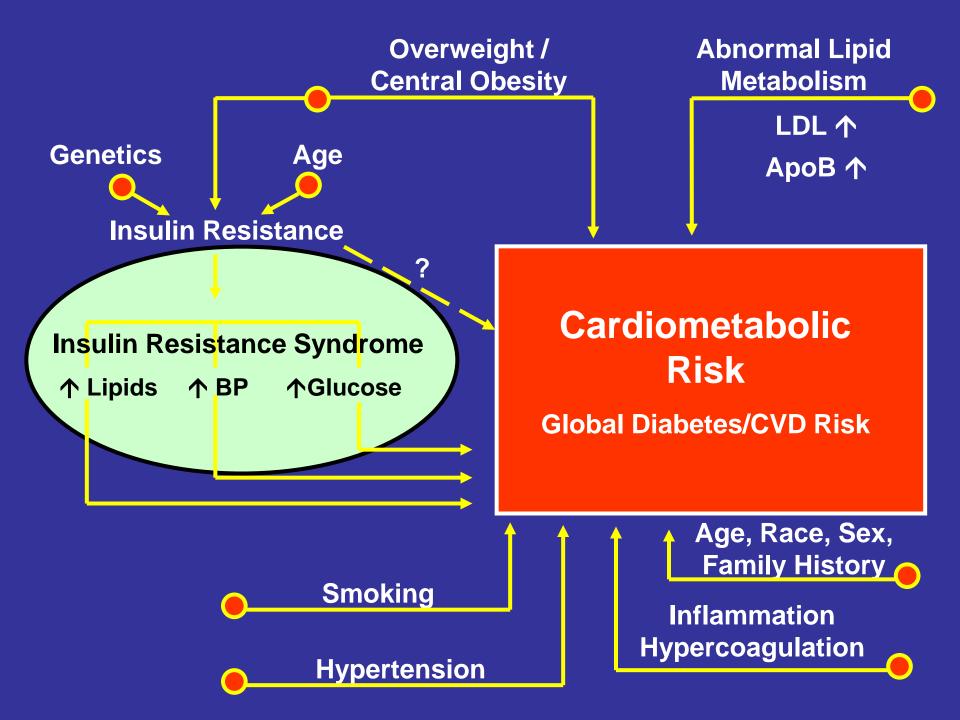


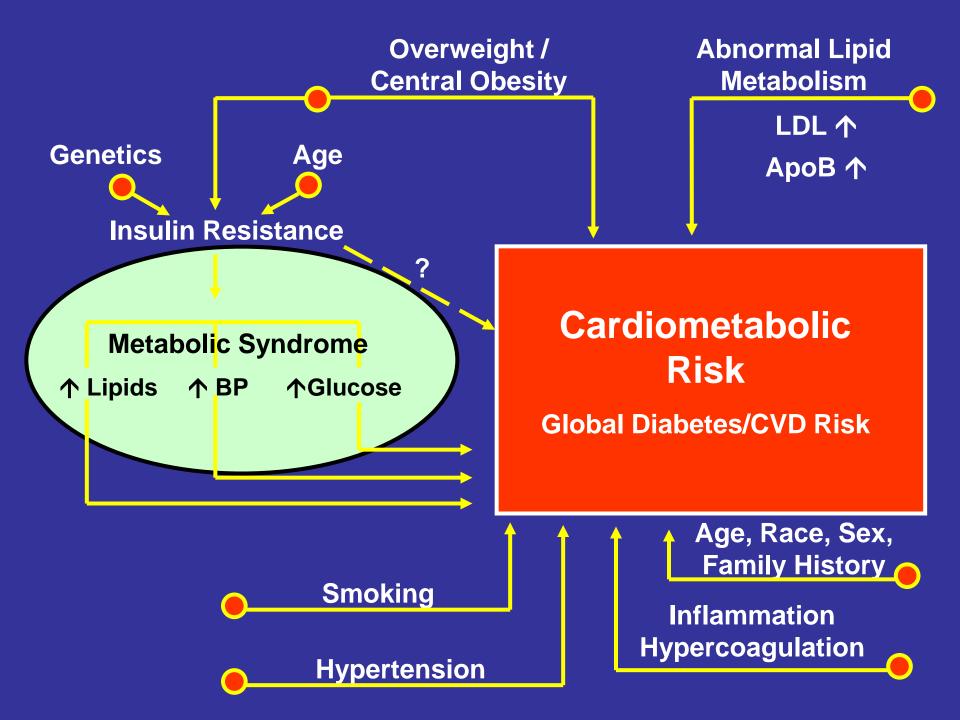
THE METABOLIC SYNDROME

More than its components?

N.B.

Does not give ABSOLUTE risk but picks out those at high RELATIVE risk





IDF POSITION

- A useful way of focusing on subjects at high risk of CVD and diabetes
- A simple diagnostic "set" capable of use in primary care world wide
- Strong need for long term detailed studies of other putative components

The Basic Public Health Screening Tool





THE METABOLIC SYNDROME

"whilst not a precise tool for determining global cardiovascular risk ... identifying those with the metabolic syndrome can help target them for more intensive therapies"

Holman, 2006

WHAT DOES THE METABOLIC SYNDROME ADD?

- Awareness of high risk associated with central obesity, raised glucose, ↑TG, ↓HDL, raised BP
 - particularly in combination
- · Hook to hang screening on for primary care
- · Emphasis on diabetes for cardiologists
- · Emphasis on cardiology for diabetologists
- Stimulus to research

Diagnosis of the Metabolic Syndrome adds nothing to the care of the patient with or at risk of Type 2 diabetes mellitus and CVD

N.B. They are not all patients Yes it does

- in most health care settings



