

SMP-Symposium vom 28. August 2023

Milchprodukte und das metabolische Syndrom

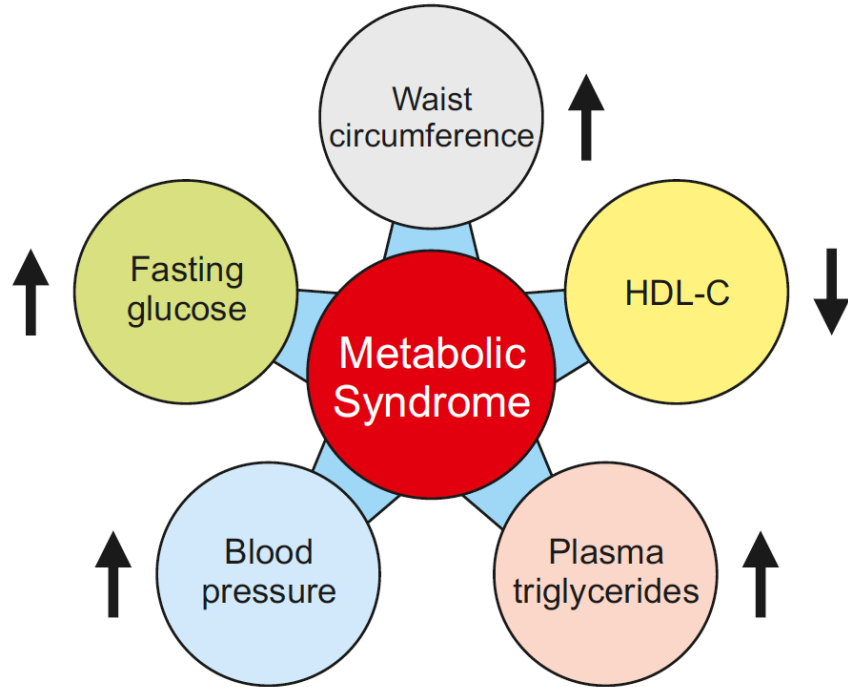
Nicolai Worm

www.nicolai-worm.de

Definition des metabolischen Syndroms (IDF)

	Europäische Männer	Europäische Frauen
Taillenumfang	≥ 94 cm	≥ 80 cm
Plus mindestens zwei der folgenden Faktoren		
Triglyceride	≥ 150 mg/dl	≥ 150 mg/dl
HDL	< 40 mg/dl (0,9 mmol/l)	< 50 mg/dl (1,1 mmol/l)
Systolischer Druck oder (diastolischer Druck)	≥ 130 mg/dl (≤ 85 mmHg)	≥ 130 mg/dl (≤ 85 mmHg)
Nüchternblutglukose	≥ 100 mg/dl	≥ 100 mg/dl

Das metabolische Syndrom



Das metabolische Syndrom: per Definition keine Krankheit, sondern eine Ansammlung einzelner metabolischer Risikofaktoren wie abdominale Adipositas, Bluthochdruck, Hyperglykämie, Hypertriglyzeridämie und niedrige HDL-Cholesterinkonzentration.

Metabolic Syndrome and Its Association with Nonalcoholic Steatohepatitis

KEY POINTS

- The presence of insulin resistance is almost universal in patients with nonalcoholic fatty liver disease (NAFLD) and/or metabolic syndrome (MetS).
- Insulin resistance, and lipotoxicity, are systemic processes that can affect multiple organs, including the liver. Genetic and epigenetic factors determine which organs are affected, and thus defining the specific phenotype of the patient.
- Nonalcoholic fatty liver disease can exist in the absence of the cardiometabolic risk factors that define the metabolic syndrome.
- Targeting insulin resistance in clinical practice should be a major target, including interventions to achieve weight loss, and medications that are insulin sensitizers.

The close relationship between the metabolic syndrome (MetS) and nonalcoholic fatty liver disease (NAFLD) is well described in the literature.^{1,2} Indeed, due to this strong association, NAFLD has oftentimes been referred to as the hepatic expression of the MetS, and recently suggested that NAFLD should be renamed metabolic-associated (dysfunction) fatty liver disease (MAFLD).³ Moreover, a recently proposed

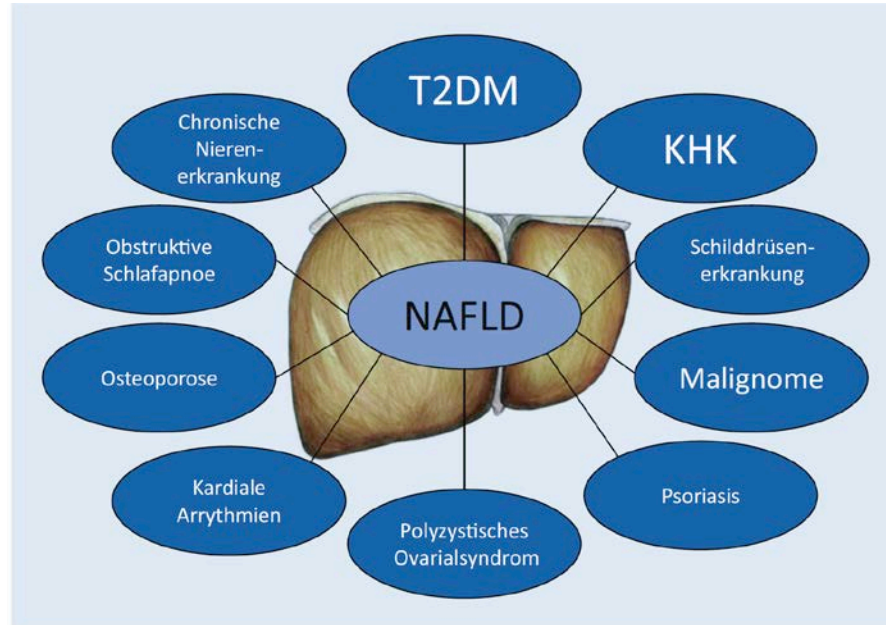
Nichtalkoholische Fettlebererkrankung

Hepatische Manifestation des metabolischen Syndroms



Elke Roeb

Schwerpunkt Gastroenterologie, Zentrum für Innere Medizin, Justus-Liebig-Universität & Universitätsklinikum Gießen, Gießen, Deutschland



Warum ist der BMI kein MetS-Kriterium?

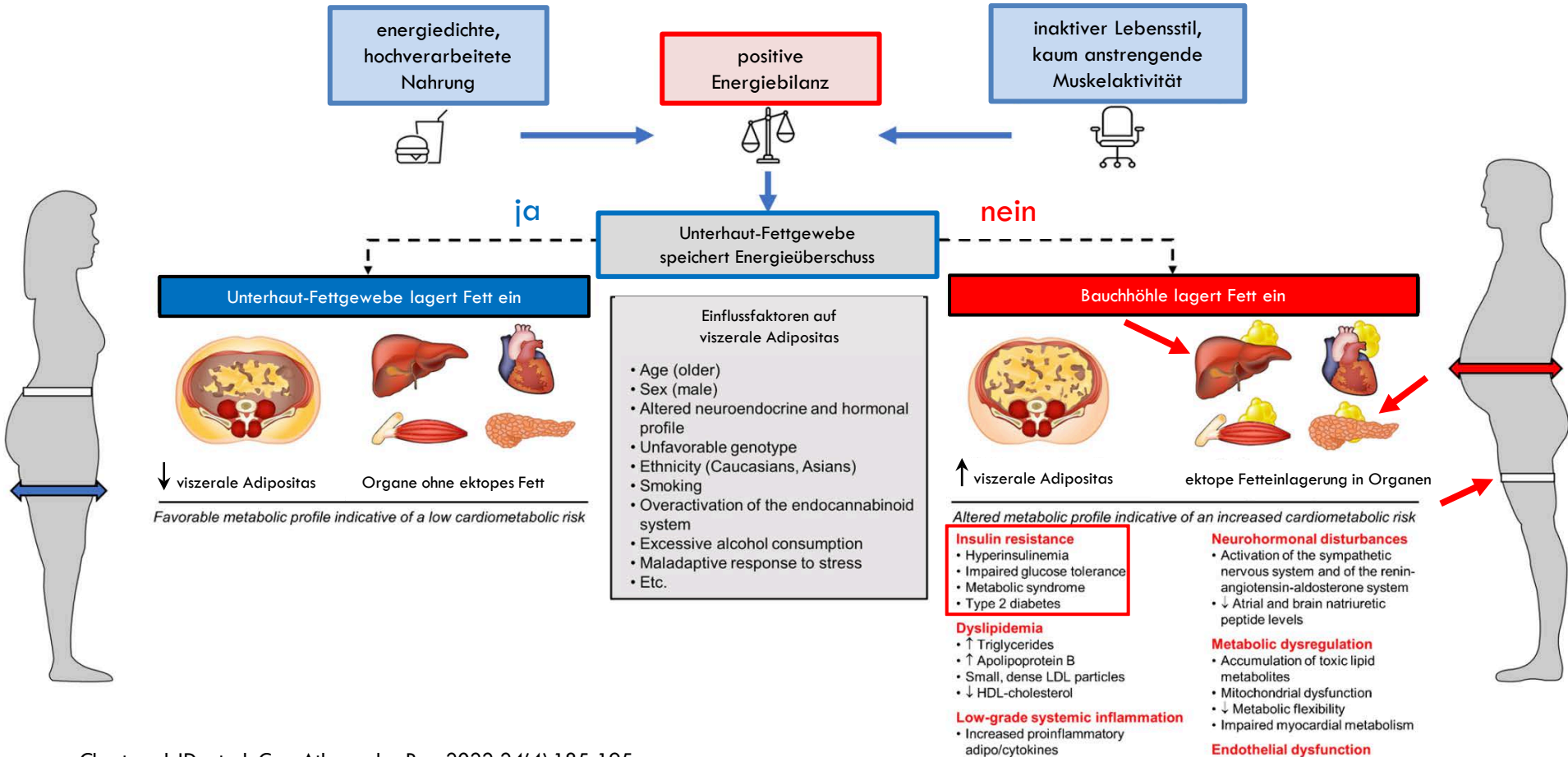


BMI = 30



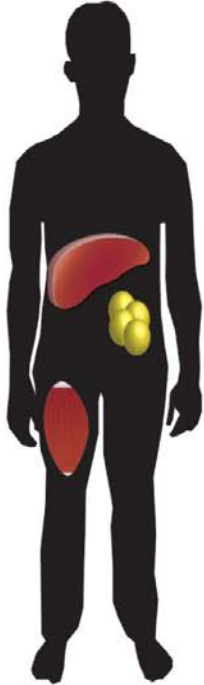
BMI = 30

Metabolisch-vaskuläres Risiko durch ektope Fett

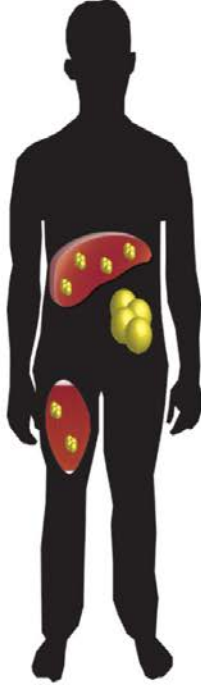


Metabolisches Risiko: Ektopes Fett – nicht „Übergewicht“!

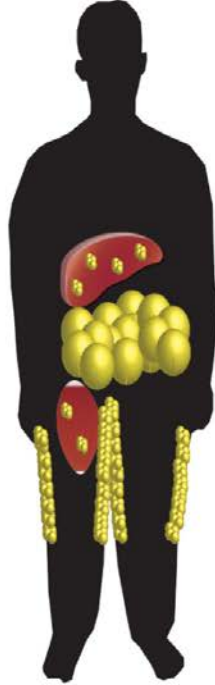
Lean,
insulin sensitive



Lean,
insulin resistant



Obese,
insulin resistant



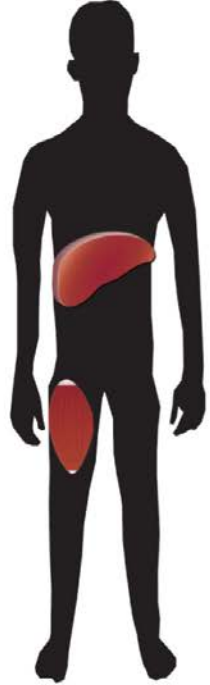
Obese,
insulin sensitive



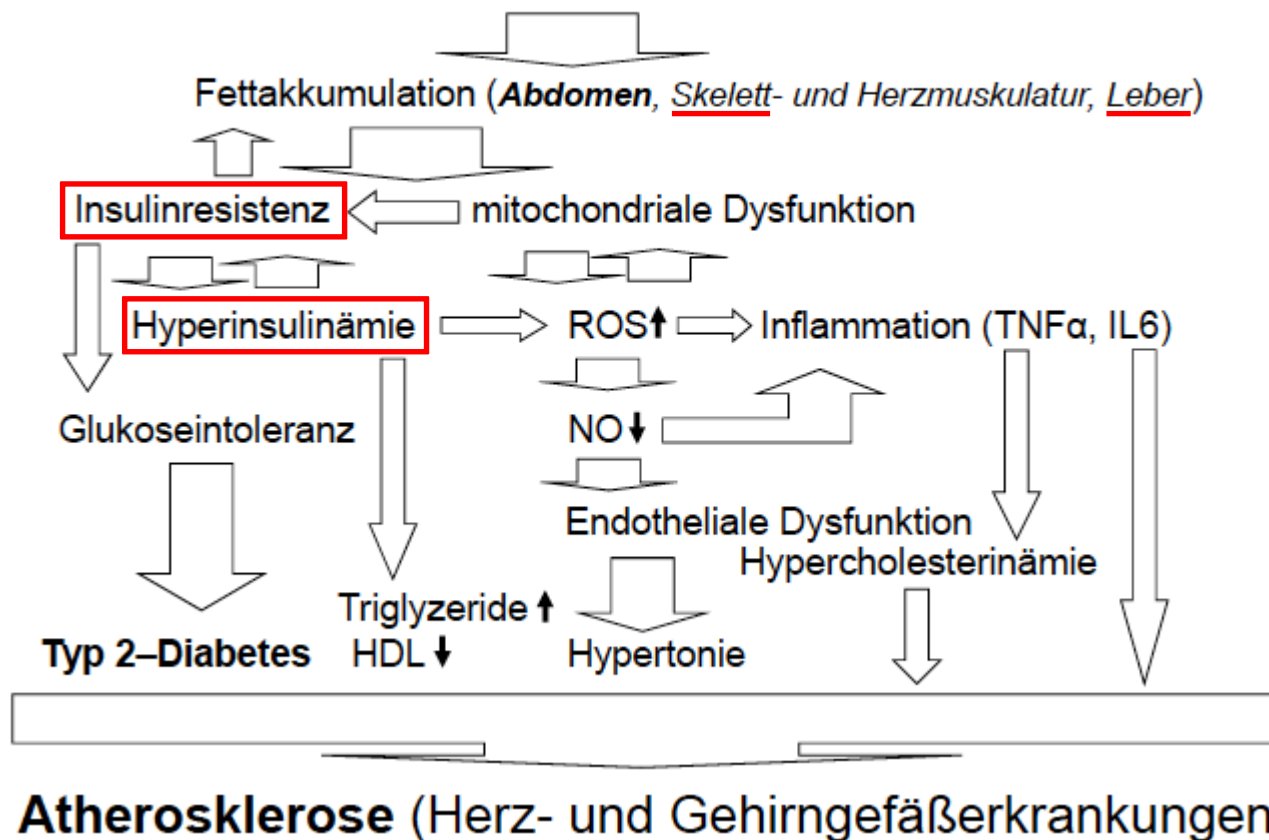
Lipodystrophic,
insulin resistant

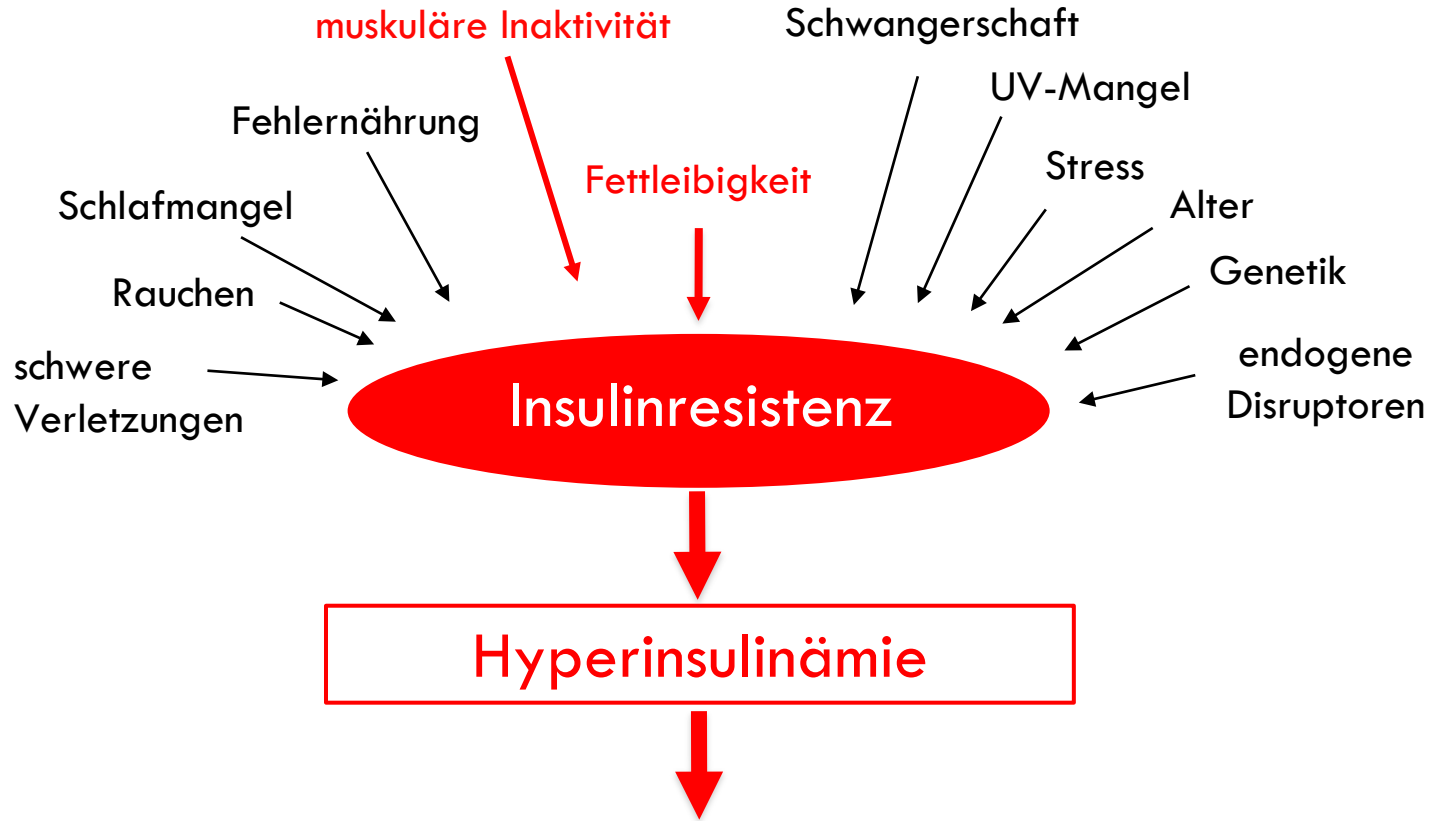


Lipodystrophic, leptin Rx,
insulin sensitive



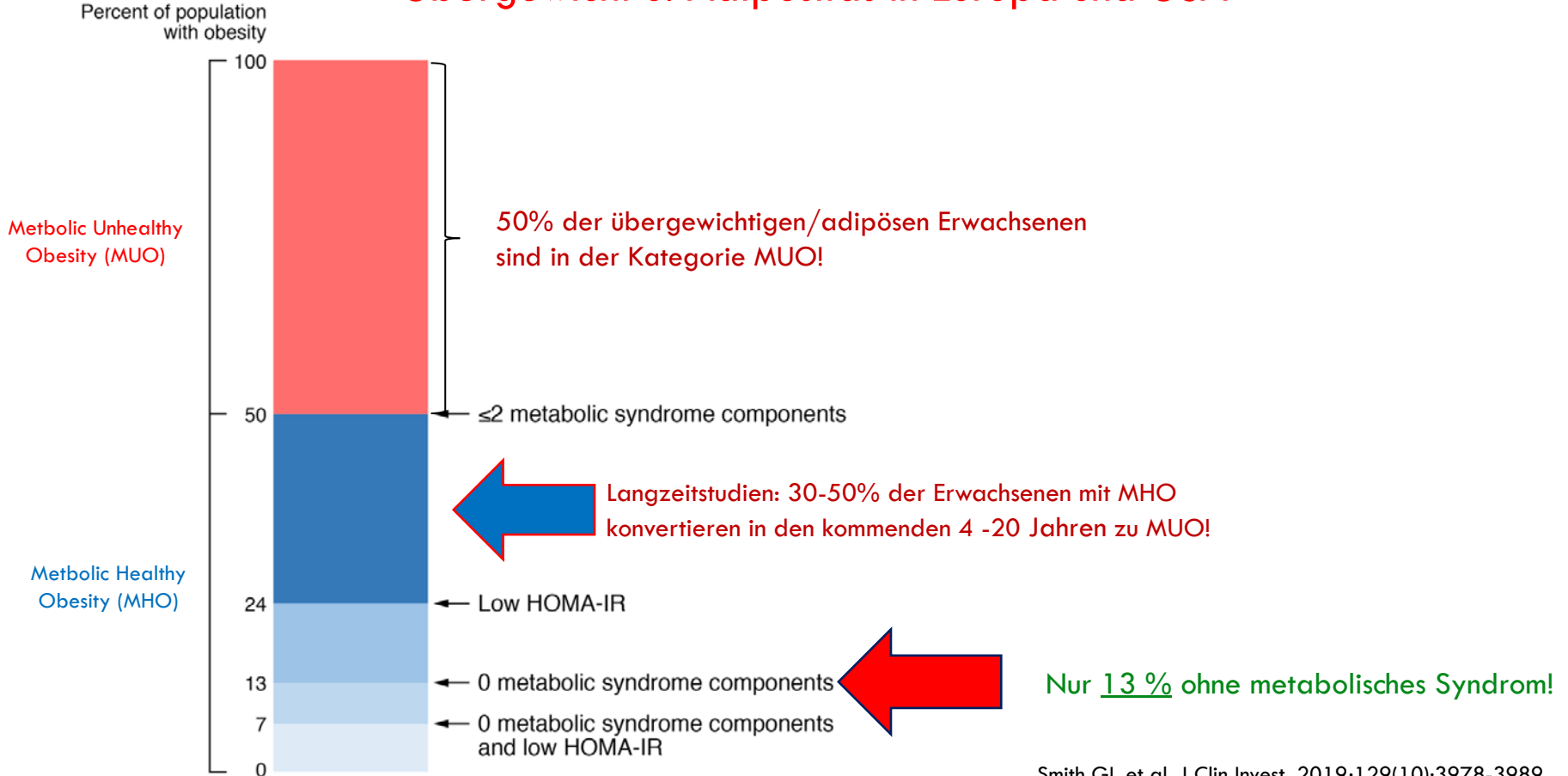
Akkumulation von ektopem Fett fördert Insulinresistenz und Hyperinsulinämie



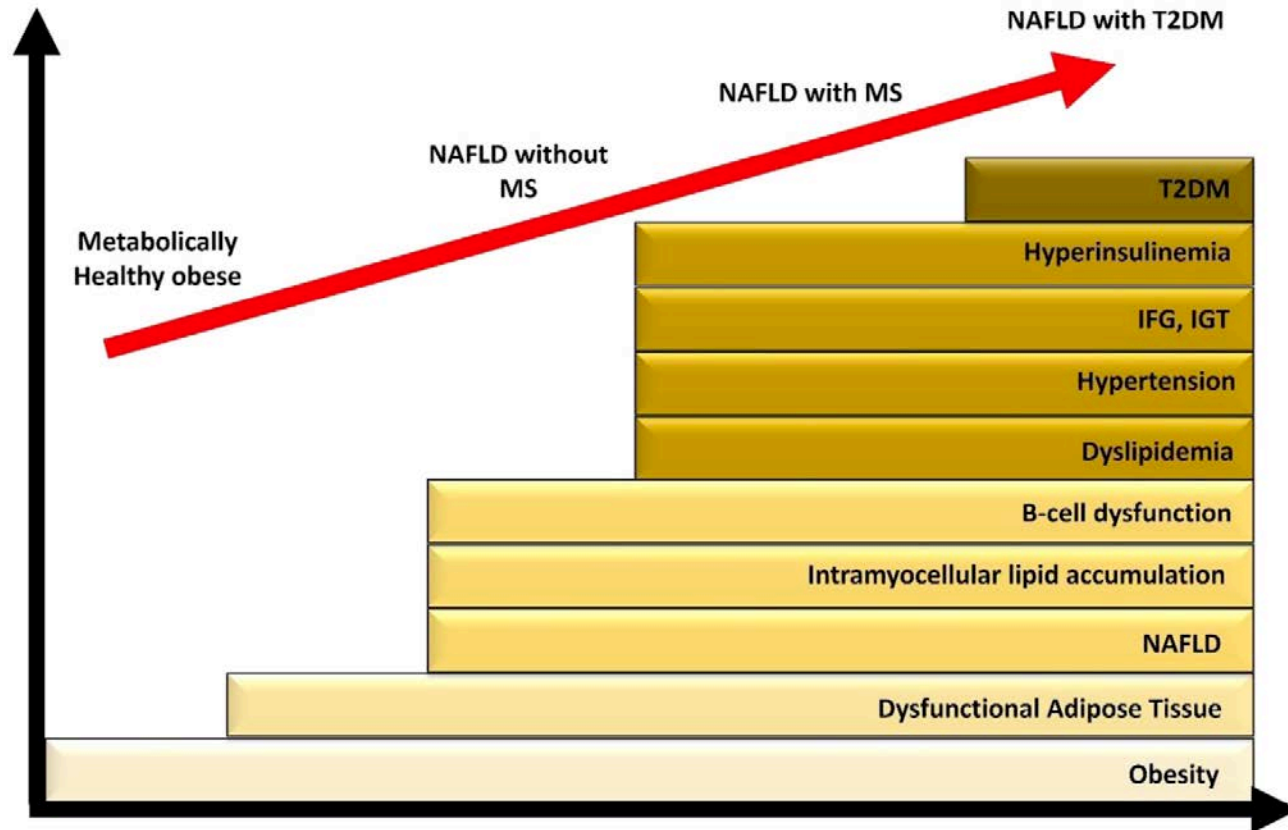


Metabolisches Syndrom, Fettleber, Diabetes, kardiovaskuläre Erkrankungen, Demenz, sexuelle Dysfunktion, Psoriasis und andere Zivilisationskrankheiten

Prävalenz von Insulinresistenz / Metabolischem Syndrom bei Übergewicht u. Adipositas in Europa und USA



Von anfangs noch „gesunder“ Fettleibigkeit über Fettleber und metabolischem Syndrom zum Typ-2-Diabetes

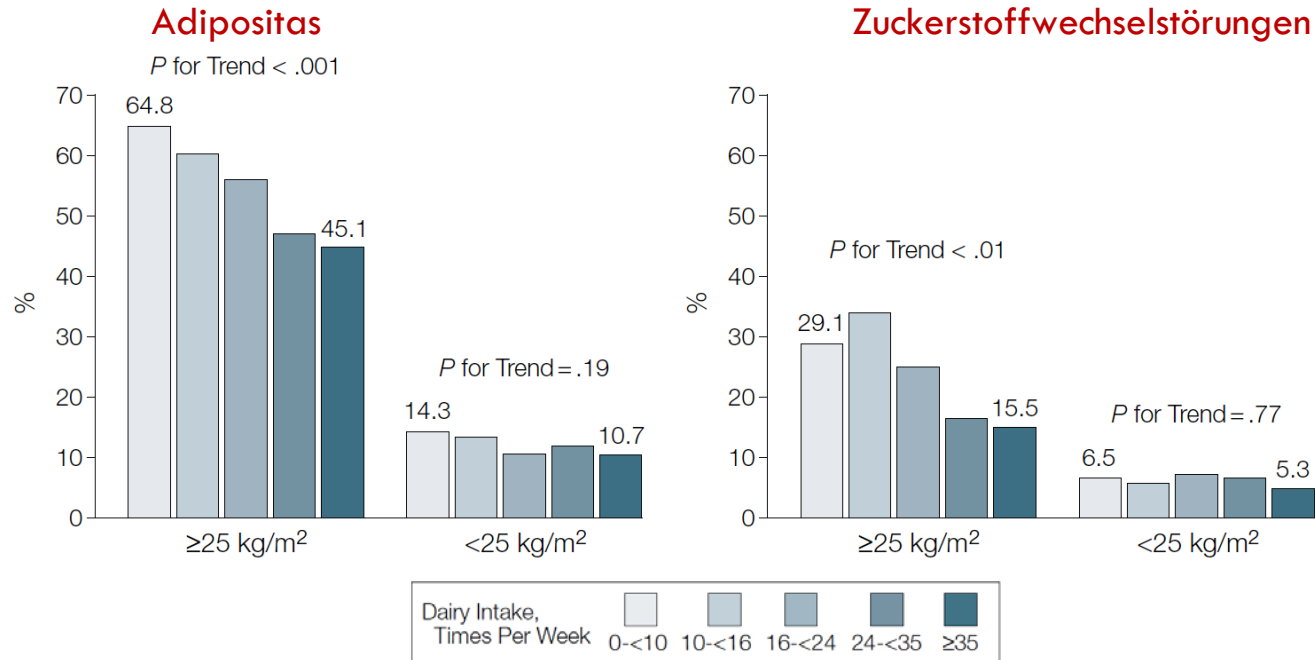


Beobachtungsstudien: Milchprodukte und MetS

Dairy Consumption, Obesity, and the Insulin Resistance Syndrome in Young Adults

The CARDIA Study

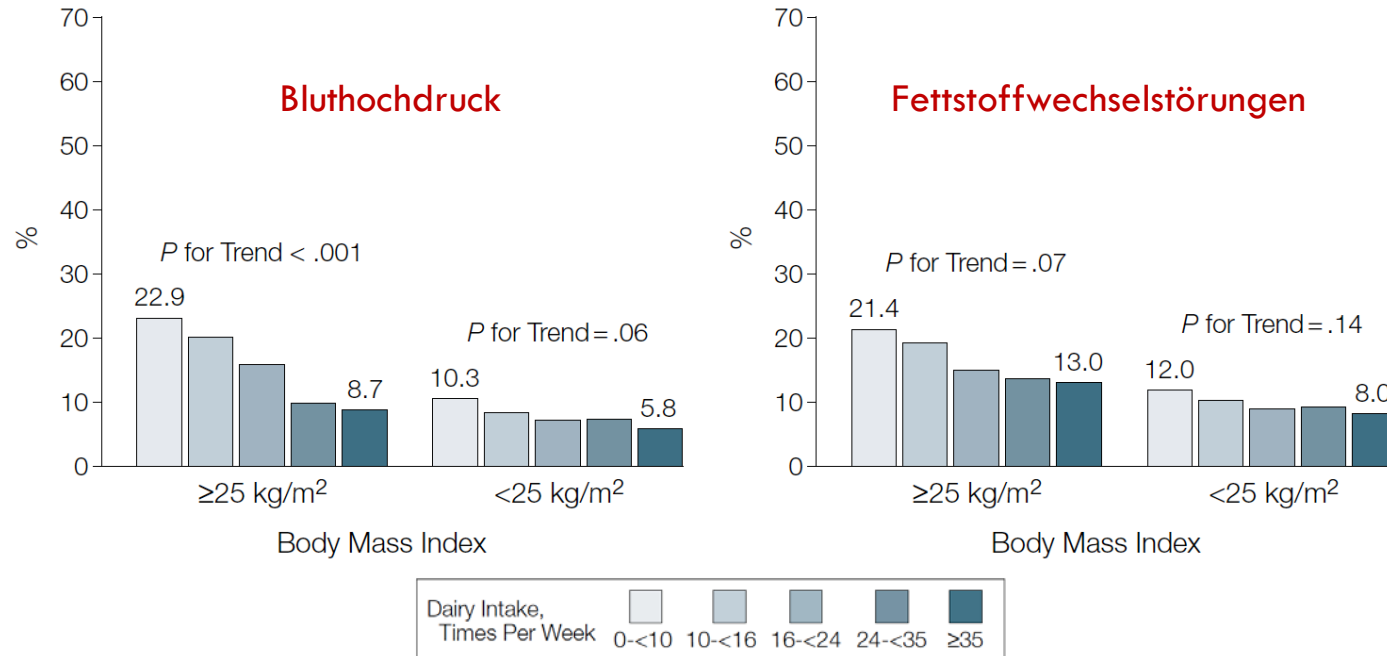
3.157 US-AmerikanerInnen, Alter: 18-30 Jahre; Followed up: 1985/86 – 1995/96.



Dairy Consumption, Obesity, and the Insulin Resistance Syndrome in Young Adults

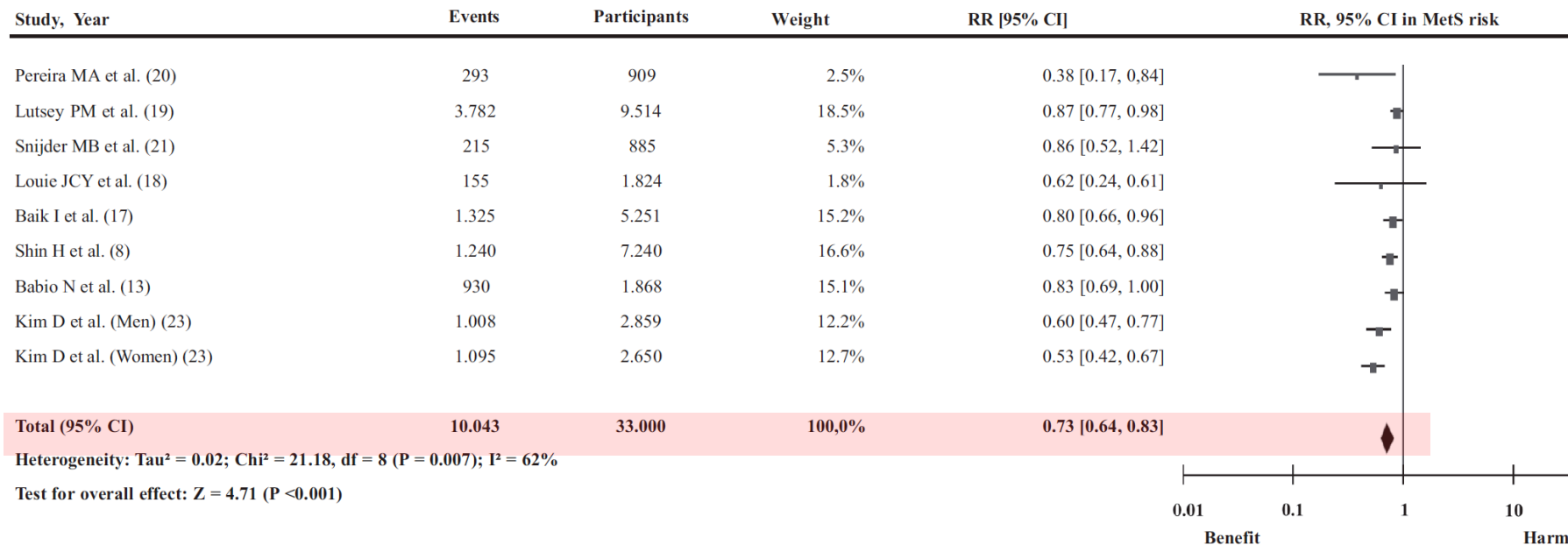
The CARDIA Study

3.157 US-AmerikanerInnen, Alter: 18-30 Jahre; Followed up: 1985-86 – 1995-96.



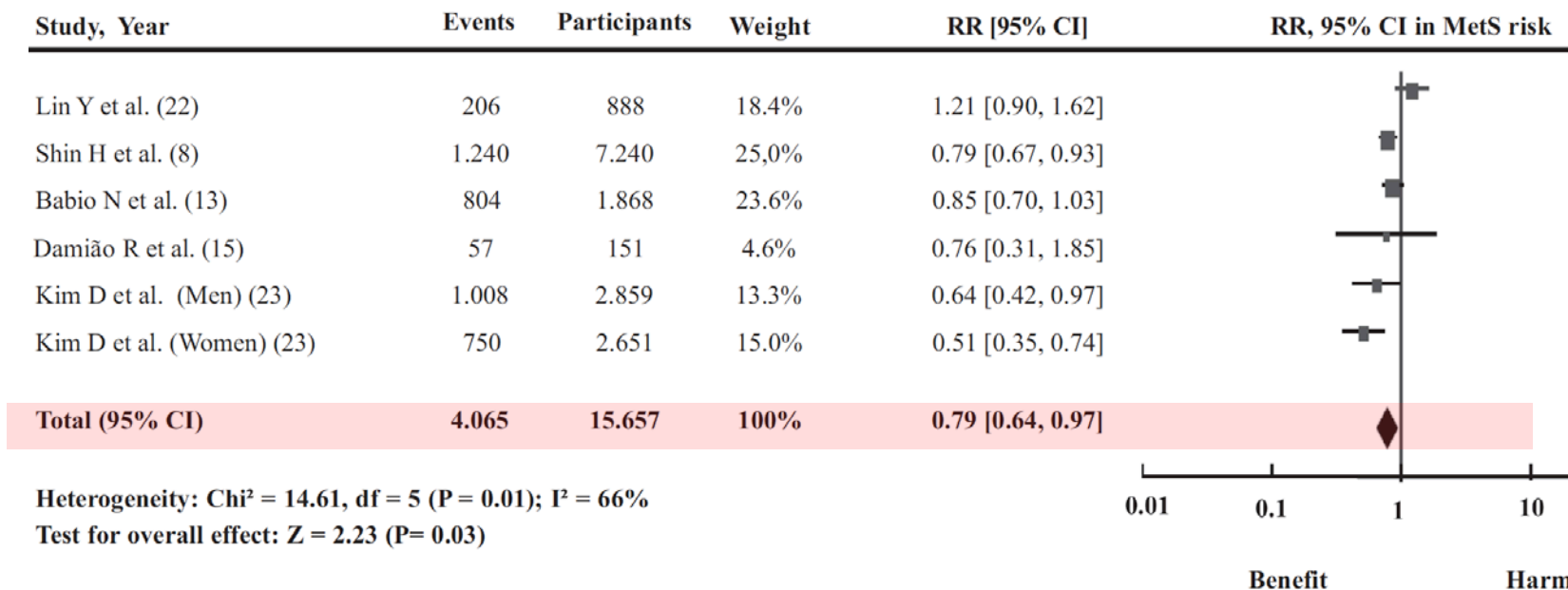
Dairy Product Consumption in the Prevention of Metabolic Syndrome: A Systematic Review and Meta-Analysis of Prospective Cohort Studies

Meta-Analyse: 11 prospektiven Kohortenstudien Milch/Milchprodukte und Inzidenz des MetS



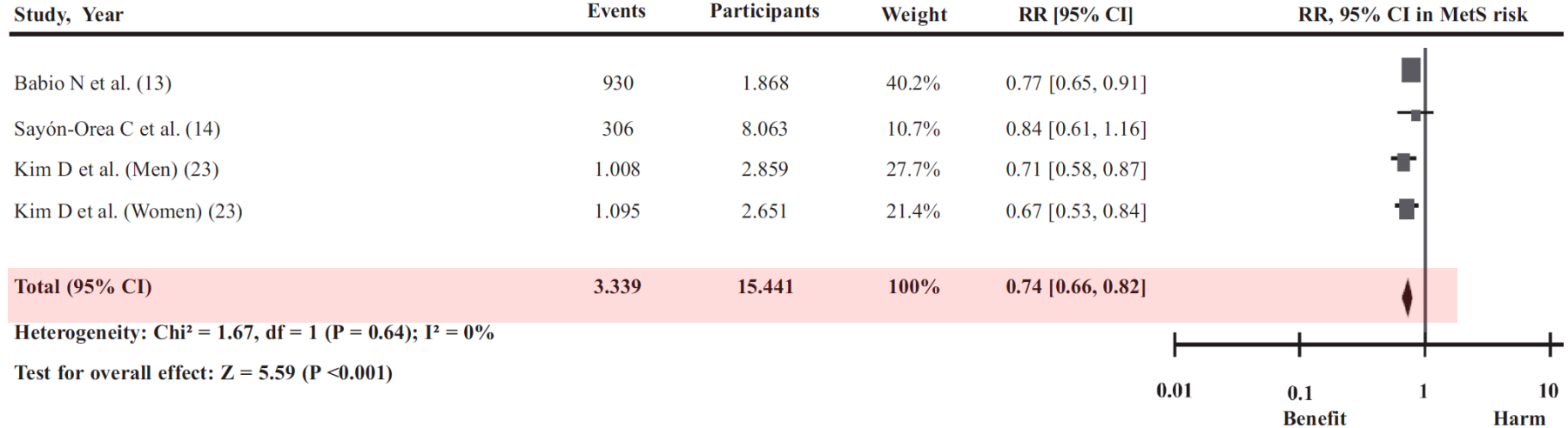
Dairy Product Consumption in the Prevention of Metabolic Syndrome: A Systematic Review and Meta-Analysis of Prospective Cohort Studies

Meta-Analyse: 11 prospektiven Kohortenstudien Milch-Konsum und Inzidenz des MetS



Dairy Product Consumption in the Prevention of Metabolic Syndrome: A Systematic Review and Meta-Analysis of Prospective Cohort Studies

Meta-Analyse: 11 prospektiven Kohortenstudien Joghurt-Konsum und Inzidenz des MetS



Dairy Consumption and Risk of Metabolic Syndrome: Results from Korean Population and Meta-Analysis

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Meta-Analyse von 12 Kohortenstudien und
25 Querschnittsstudien mit n = 398.877

Dosis-Wirkungs-Analyse

Subgroups	No. of Studies	Relative Risk (95% CI)
Milk	20	0.83 (0.78–0.89)
Study design		
Cohort	7	0.83 (0.72–0.97)
Cross-sectional	13	0.83 (0.77–0.90)
Sex		
Men	7	0.83 (0.75–0.92)
Women	7	0.79 (0.69–0.90)
Geographical region		
America	6	0.86 (0.78–0.95)
Asia	10	0.80 (0.72–0.89)
Europe	3	0.87 (0.45–1.71)
Yogurt	12	0.89 (0.83–0.95)
Study design		
Cohort	6	0.84 (0.71–0.98)
Cross-sectional	6	0.93 (0.87–0.99)
Sex		
Men	4	0.86 (0.72–1.02)
Women	4	0.91 (0.81–1.02)
Geographical region		
America	3	0.71 (0.42–1.22)
Asia	6	0.91 (0.84–0.998)
Europe	2	0.78 (0.67–0.91)
Cheese	8	0.98 (0.86–1.11)
Study design		
Cohort	4	1.03 (0.87–1.22)
Cross-sectional	4	0.91 (0.74–1.14)
Geographical region		
America	3	1.07 (0.93–1.25)
Asia	2	0.92 (0.71–1.20)
Europe	2	1.03 (0.65–1.64)

	Studies	Dose	Relative Risk (95% CI)
Total dairy	6	400 g/day	0.71 (0.59–0.85)
Milk	5	200 g/day	0.85 (0.79–0.93)
Yogurt	5	200 g/day	0.63 (0.53–0.75)
Cheese	3	50 g/day	0.99 (0.73–1.35)

Dairy product consumption and risk of non-alcoholic fatty liver disease: A systematic review and meta-analysis of observational studies

^a Department of Agricultural, Food and Nutritional Science, University of Alberta, Edmonton, Alberta, Canada

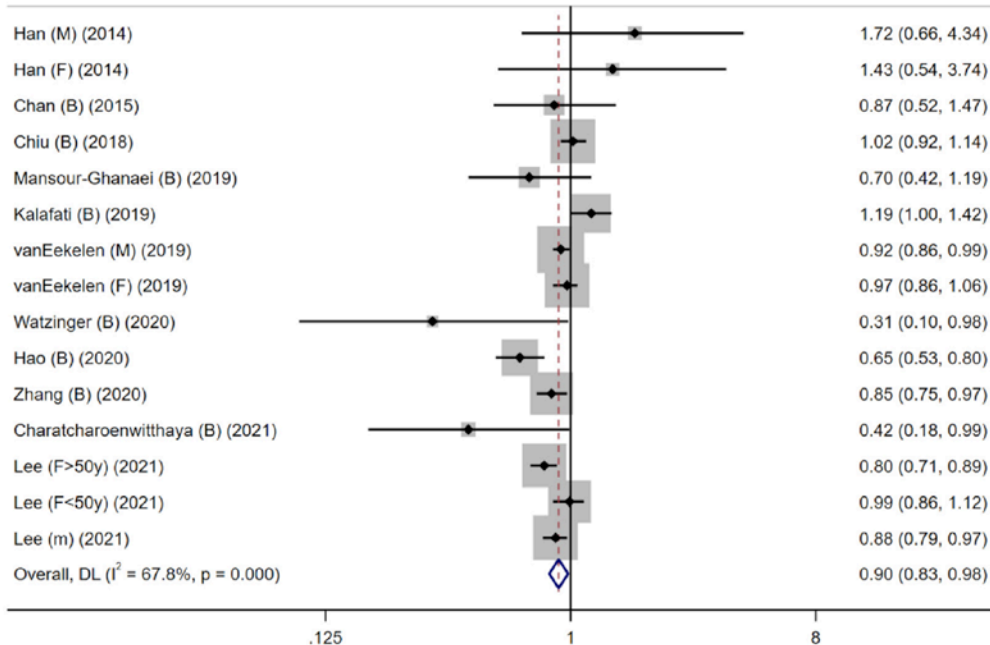
^b Department of Cell Biology, University of Alberta, Edmonton, Alberta, Canada

^c Alberta Health Services, Edmonton, Alberta, Canada

^d School of Public Health, University of Alberta, Edmonton, Alberta, Canada

^e Department of Physiology, University of Alberta, Edmonton, Alberta, Canada

OR: NAFLD Risiko für Milch/Milchprodukte



Meta-Analyse von 11 Beobachtungsstudien zur Inzidenz von NAFLD, n=43.649, 11.020 Fälle;

Milchprodukte: OR=0,90; 95% KI: 0,83-0,98

Fetteiche Milchprod.: OR=0,38; 95% KI: 0,19-0,75

Milch: OR=0,86; 95% KI: 0,78-0,95

Joghurt: OR=0,88; 95% KI: 0,82-0,96

Käse: OR=1,01; 95% KI: 0,82-1,25

Dairy products consumption and the risk of hypertension in adults: An updated systematic review and dose–response meta-analysis of prospective cohort studies

^a Department of Biostatistics and Epidemiology, School of Health, Isfahan University of Medical Sciences, Isfahan, Iran

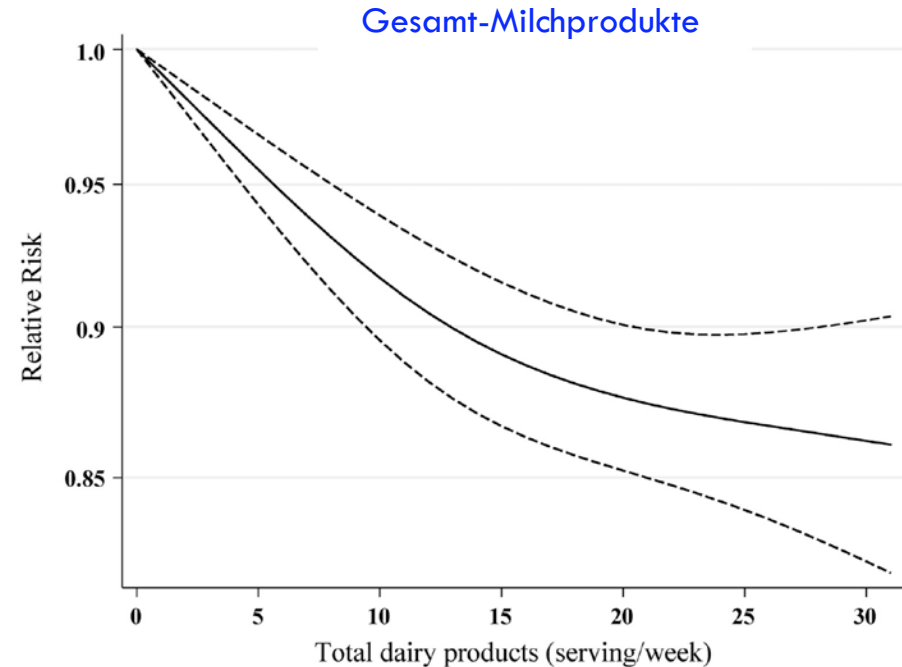
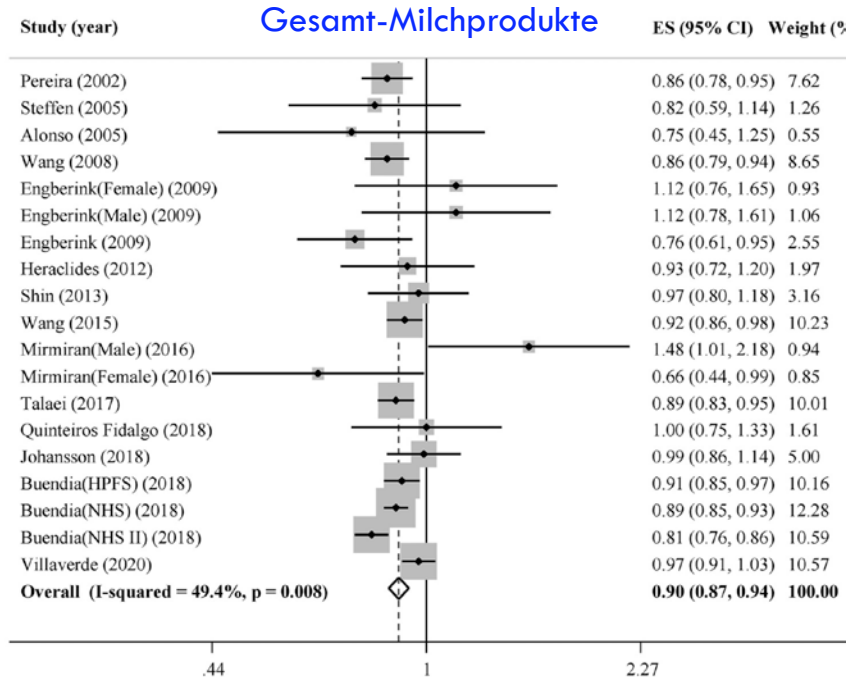
^b Cardiac Rehabilitation Research Center, Cardiovascular Research Institute, Isfahan University of Medical Sciences, Isfahan, Iran

^c Shahid Motahari Hospital, Foadlshahr, Isfahan, Iran

^d Centre for Intelligent Healthcare, Coventry University, Coventry, CV1 5FB, UK

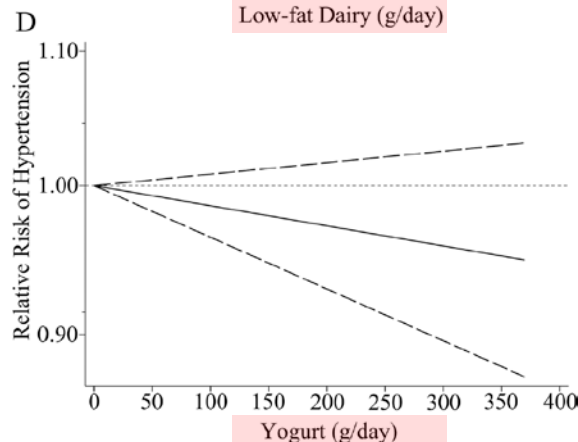
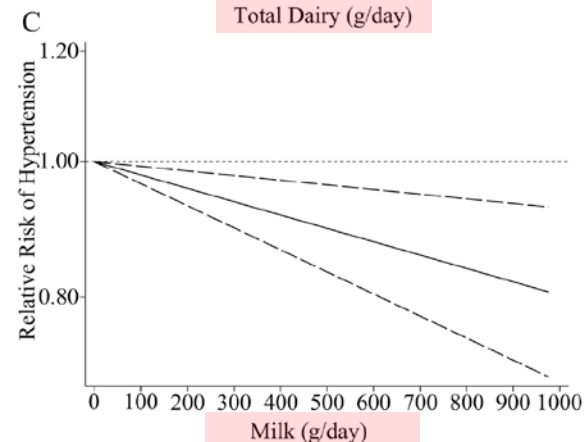
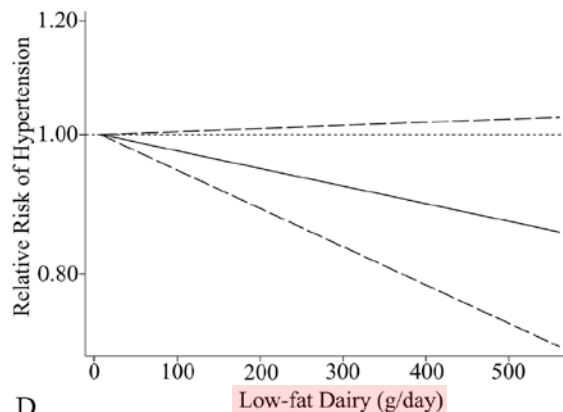
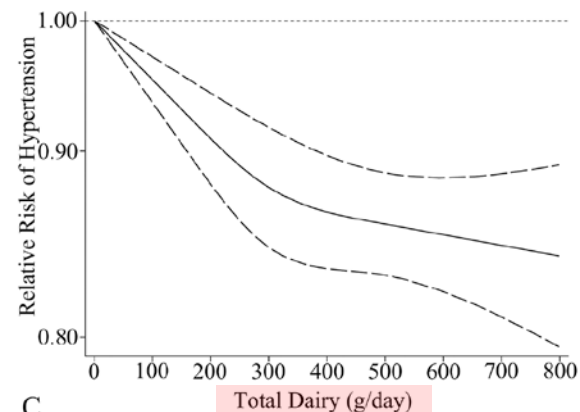
^e Isfahan Cardiovascular Research Center, Cardiovascular Research Institute, Isfahan University of Medical Sciences, Isfahan, Iran

Meta-Analyse von 16 prospektive Kohortenstudien: Inzidenz von Bluthochdruck (Relatives Risiko)



Consumption of Dairy Products and the Risk of Overweight or Obesity, Hypertension, and Type 2 Diabetes Mellitus: A Dose-Response Meta-Analysis and Systematic Review of Cohort Studies

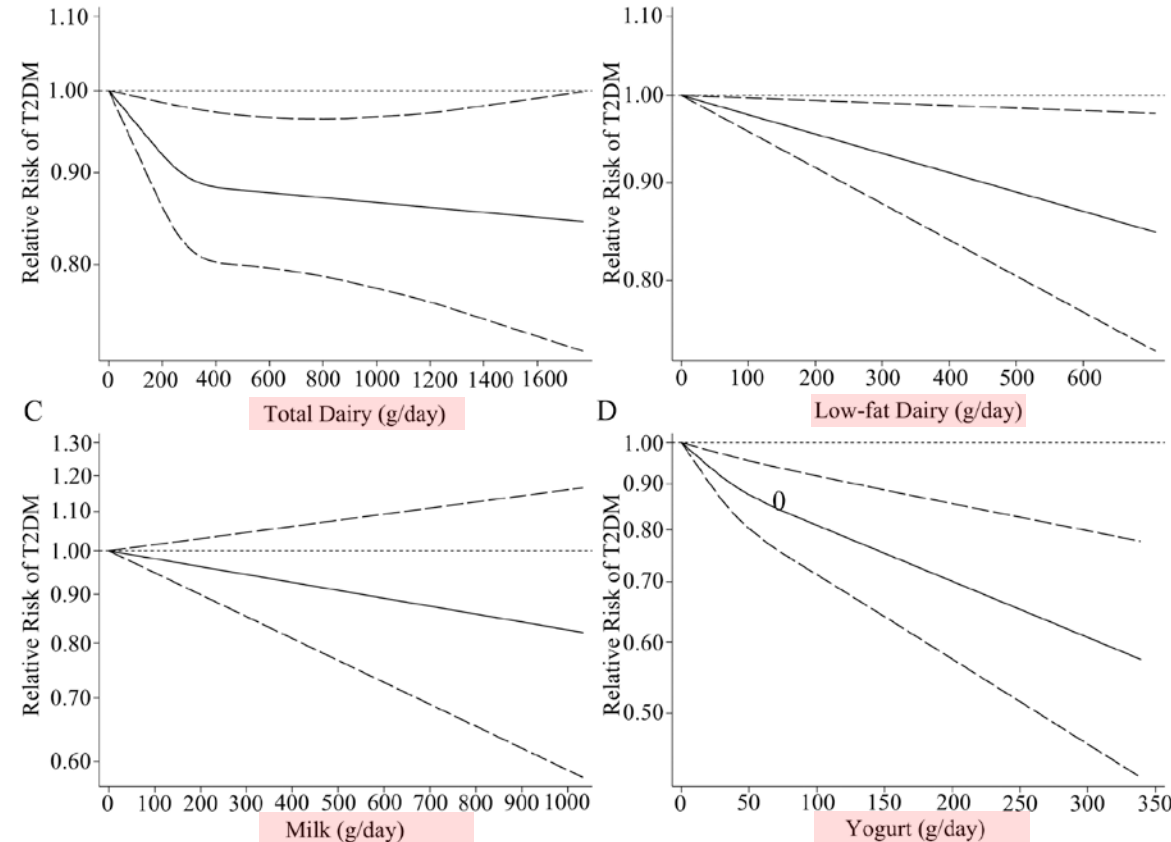
¹Department of Biostatistics and Epidemiology, School of Public Health, Shenzhen University Health Science Center, Shenzhen, Guangdong, People's Republic of China; ²Department of Epidemiology and Biostatistics, College of Public Health, Zhengzhou University, Zhengzhou, Henan, People's Republic of China; ³Department of Medical Record Management, Shenzhen Qianhai Shekou Free Trade Zone Hospital, Shenzhen, Guangdong, People's Republic of China; ⁴Department of Endocrinology, Shenzhen Qianhai Shekou Free Trade Zone Hospital, Shenzhen, Guangdong, People's Republic of China; ⁵Guangdong Provincial Key Laboratory of Regional Immunity and Diseases, Shenzhen University Health Science Center, Shenzhen, Guangdong, People's Republic of China; and ⁶Department of General Practice, The Affiliated Luohu Hospital of Shenzhen University Health Science Center, Shenzhen, Guangdong, People's Republic of China



Bluthochdruck: Meta-Analysen von 17 prospektiven Kohortenstudien

Consumption of Dairy Products and the Risk of Overweight or Obesity, Hypertension, and Type 2 Diabetes Mellitus: A Dose–Response Meta-Analysis and Systematic Review of Cohort Studies

¹Department of Biostatistics and Epidemiology, School of Public Health, Shenzhen University Health Science Center, Shenzhen, Guangdong, People's Republic of China; ²Department of Epidemiology and Biostatistics, College of Public Health, Zhengzhou University, Zhengzhou, Henan, People's Republic of China; ³Department of Medical Record Management, Shenzhen Qianhai Shekou Free Trade Zone Hospital, Shenzhen, Guangdong, People's Republic of China; ⁴Department of Endocrinology, Shenzhen Qianhai Shekou Free Trade Zone Hospital, Shenzhen, Guangdong, People's Republic of China; ⁵Guangdong Provincial Key Laboratory of Regional Immunity and Diseases, Shenzhen University Health Science Center, Shenzhen, Guangdong, People's Republic of China; and ⁶Department of General Practice, The Affiliated Luohu Hospital of Shenzhen University Health Science Center, Shenzhen, Guangdong, People's Republic of China

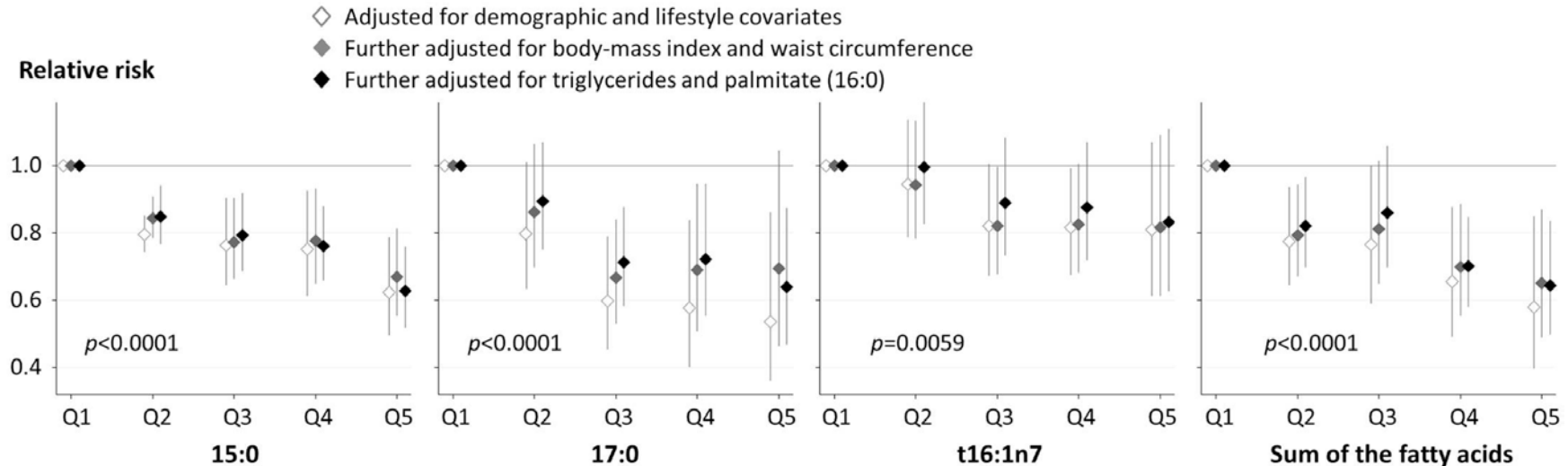


Typ-2-Diabetes: Meta-Analysen von 28 prospektiven Kohortenstudien

Fatty acid biomarkers of dairy fat consumption and incidence of type 2 diabetes: A pooled analysis of prospective cohort studies

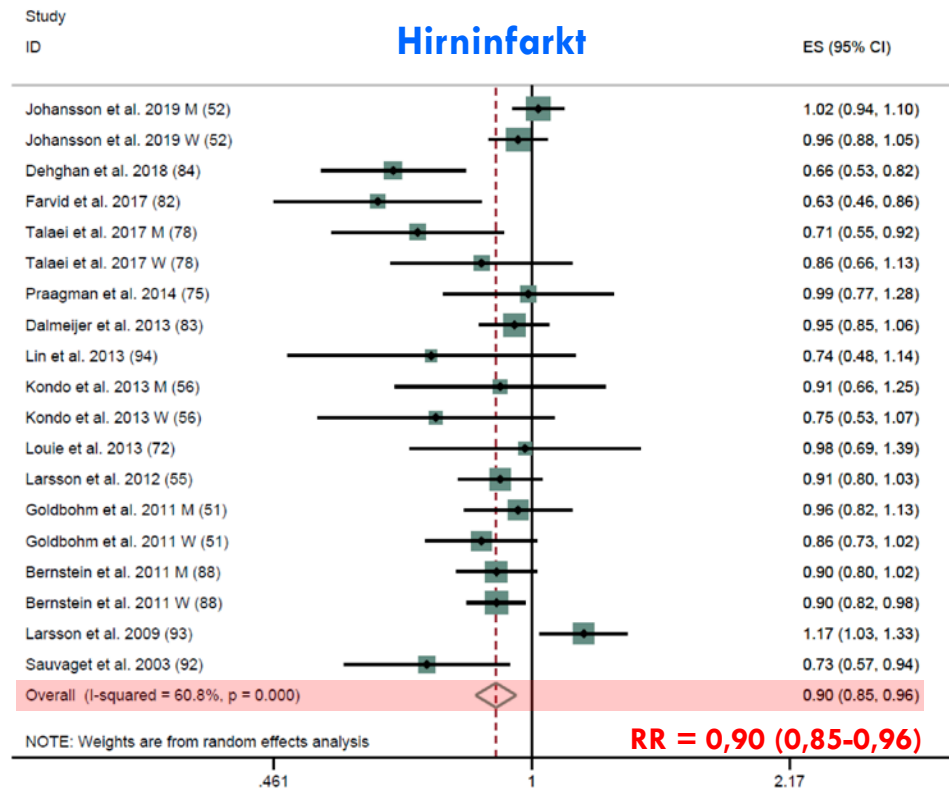
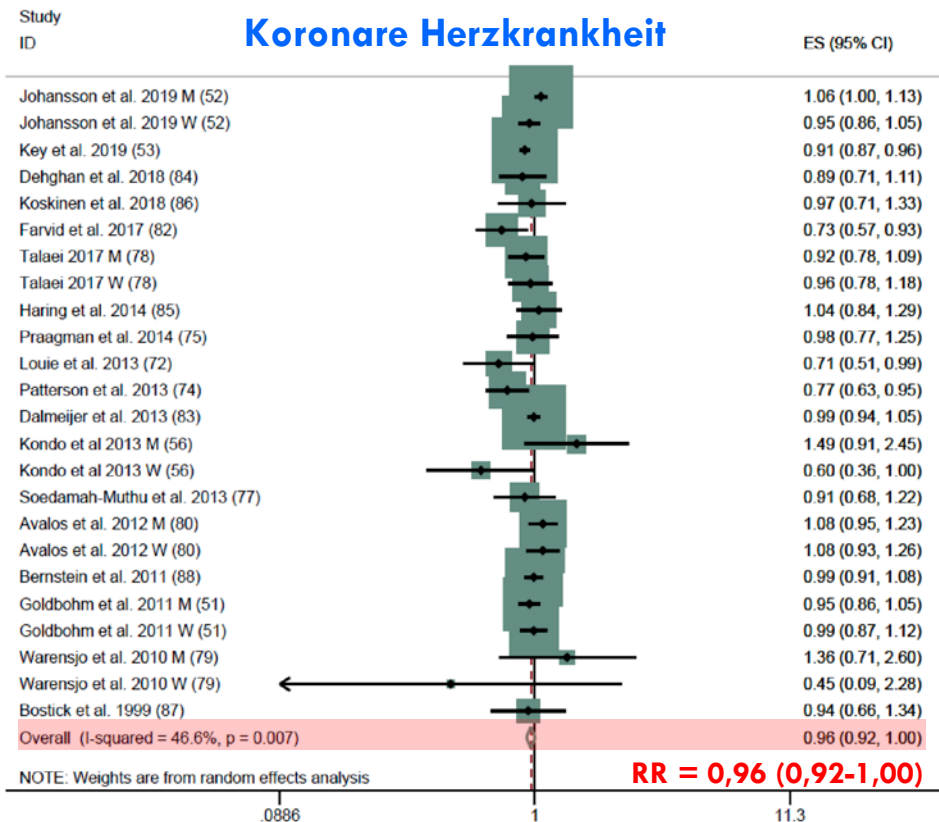
16 prospektive Kohortenstudien aus 12 Ländern (7 USA, 7 Europa, 1 Australia, 1 Taiwan);
n=63.682 aus allen Alters- und BMI-Bereichen; 9 Jahre Follow-up, 15.180 T2D-Fälle;

C15:0, C17:0, C16:1n-7 => Biomarker für den Konsum von Milchfett/Milchprodukten



Konsum von Milch/Milchprodukten und Herz-Kreislauf-Risiko

Meta-Analyse: Höchster vs niedrigster Konsum



Consumption of Dairy Foods and Cardiovascular Disease: A Systematic Review

Übersichtsarbeit: Meta-Analysen von Kohortenstudien zu harten kardiovaskulären Endpunkten – und Meta-Analysen von randomisiert-kontrollierten Diät-Interventionsstudien zur kardiovaskulären Risikofaktoren.

Abstract: Limited consumption of dairy foods and use of low-fat products is recommended for cardiovascular (CV) prevention; however, other features besides fat content modulate their metabolic effects. We analyze updated evidence on the relationship of different dairy products (low/full-fat dairy, milk, cheese, yogurt) with CVD by reviewing meta-analyses of cohort studies and individual prospective cohort studies with CV hard endpoints (CVD/CHD incidence/mortality), together with meta-analyses of randomized controlled trials exploring the effect of dairy on major CV risk factors. The analyses provide evidence that moderate dairy consumption (up to 200 g/day, globally) has no detrimental effects on CV health and that their effect depends more on the food type (cheese, yogurt, milk) than on the fat content. These data expand current knowledge and may inform revision of current guidelines for CVD prevention.

Kontrollierte Experimente

Sucrose-sweetened beverages increase fat storage in the liver, muscle, and visceral fat depot: a 6-mo randomized intervention study¹⁻³

Maria Maersk, Anita Belza, Hans Stødkilde-Jørgensen, Steffen Ringgaard, Elizaveta Chabanova, Henrik Thomsen, Steen B Pedersen, Arne Astrup, and Bjørn Richelsen

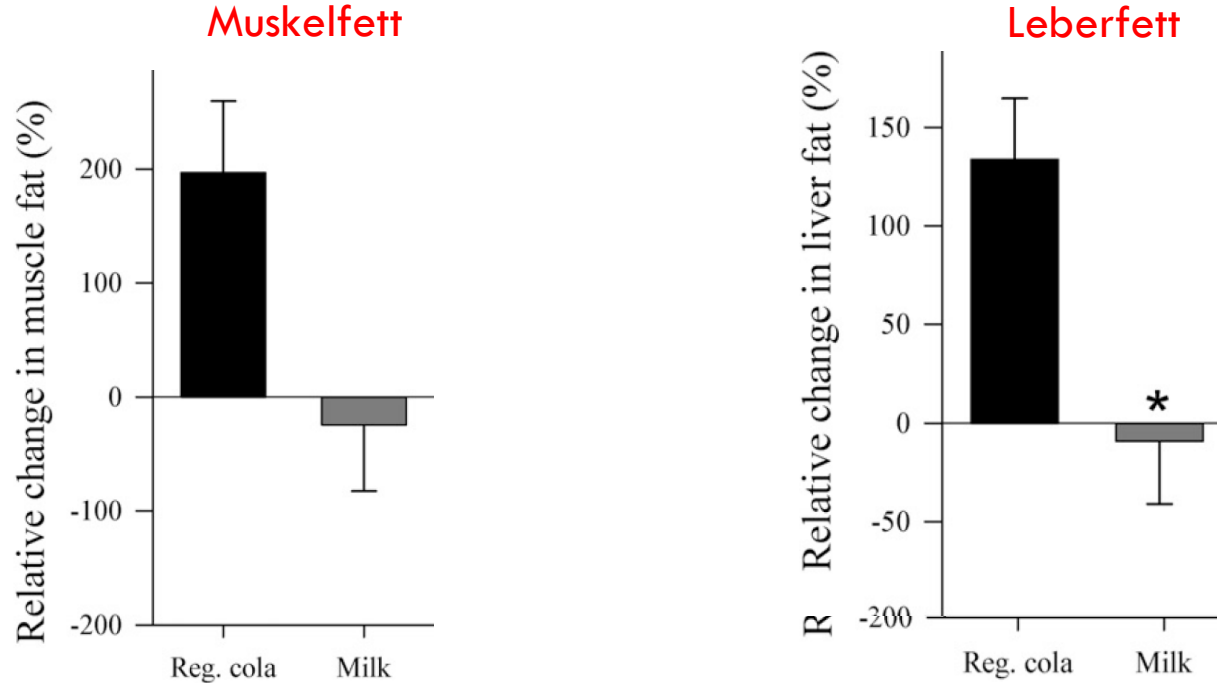
RCT bei 47 Übergewichtigen, 6 Monate täglich 1 Liter Test-Getränk;
Cola mit Saccharose (50% Fruktose/50% Glukose); Cola-Light; Milch mit 1,5% Fett;

	Sucrose-sweetened regular cola	Milk
Carbohydrate (g/100 mL)	10.6	4.7
Protein (g/100 mL)	0	3.4
Fat (g/100 mL)	0	1.5
Energy (kJ/d)	1800	1900
Volume (mL)	1000	1000
Energy density (kJ/g)	1.8	1.9

¹ The subjects drank 1 L of 1 of 4 test drinks daily for 6 mo.

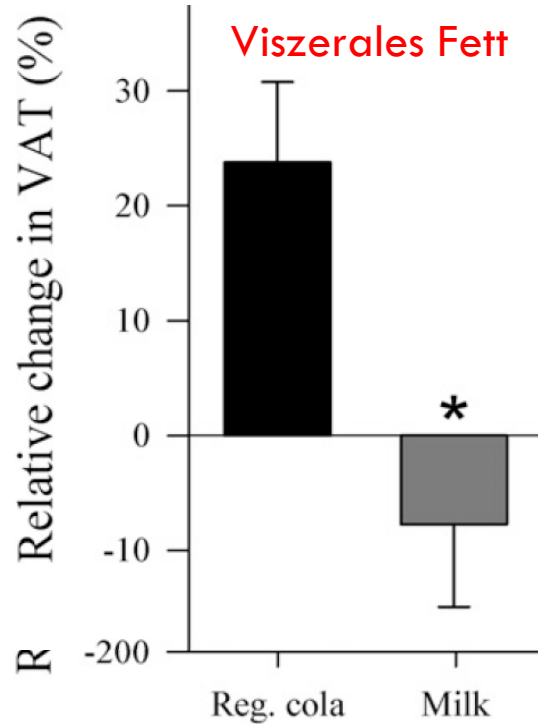
Sucrose-sweetened beverages increase fat storage in the liver, muscle, and visceral fat depot: a 6-mo randomized intervention study¹⁻³

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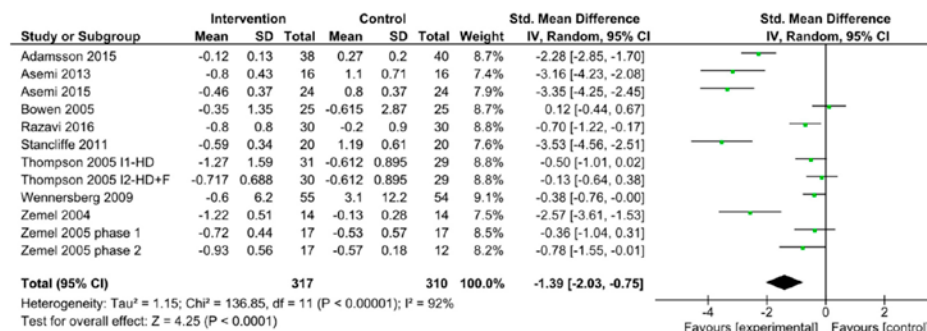
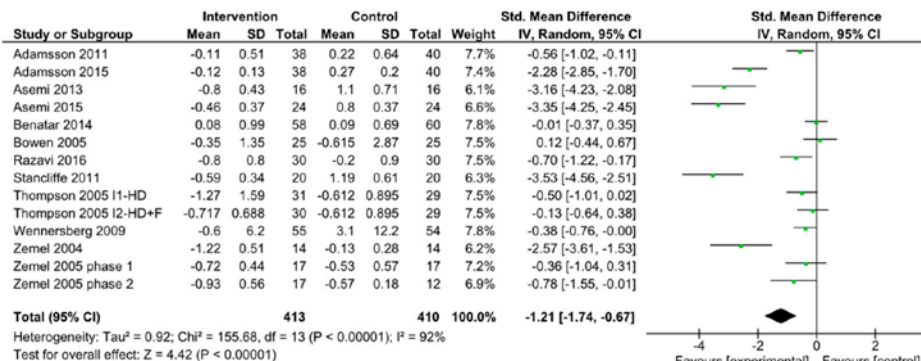


The Effects of Dairy Intake on Insulin Resistance: A Systematic Review and Meta-Analysis of Randomized Clinical Trials

30 randomisiert-kontrollierte Diät-Interventionstudien aufgenommen;

Milchprodukte und HOMA-Index

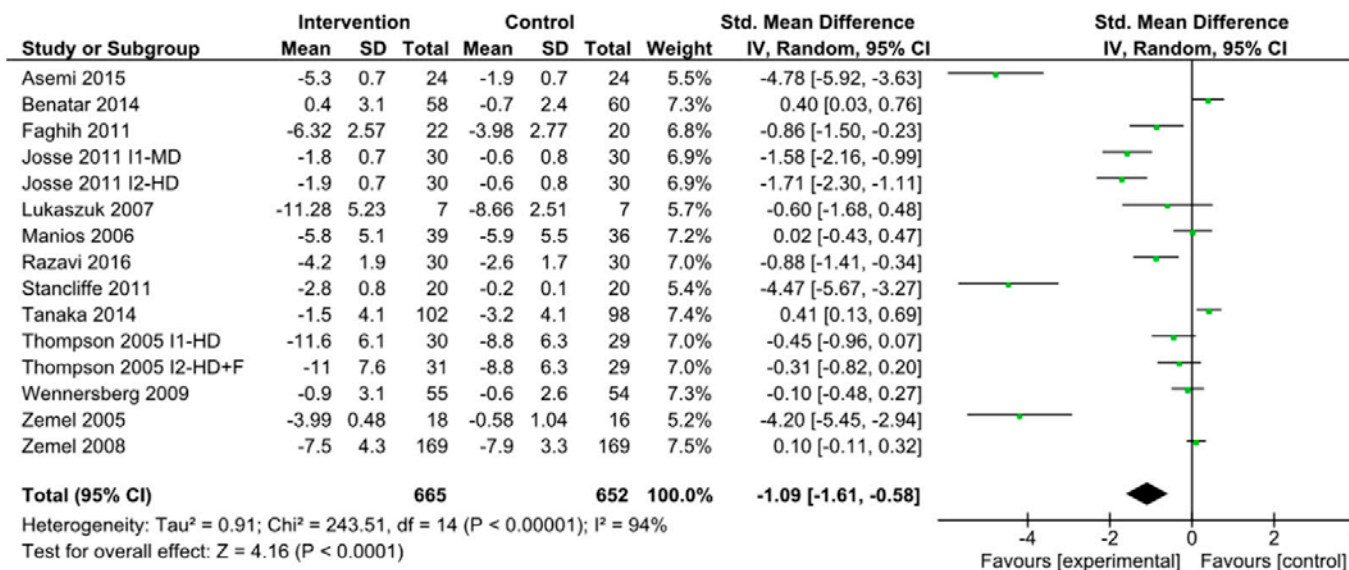
Milchprodukte und HOMA-Index bei BMI>25



The Effects of Dairy Intake on Insulin Resistance: A Systematic Review and Meta-Analysis of Randomized Clinical Trials

30 randomisiert-kontrollierte Diät-Interventionsstudien aufgenommen;

Milchprodukte und Taillenumfang



Milk and Dairy Product Consumption and Inflammatory Biomarkers: An Updated Systematic Review of Randomized Clinical Trials

¹Department of Nutrition, Institute of Basic Medical Sciences, University of Oslo, Oslo, Norway; ²Norwegian National Advisory Unit on Familial Hypercholesterolemia, Oslo University Hospital, Oslo, Norway; ³Department of Biochemistry and Molecular Biology II, School of Pharmacy; and ⁴Institute of Nutrition and Food Technology "José Mataix," Biomedical Research Center, University of Granada, Granada, Spain; ⁵ibs.GRANADA, University Hospital Complex of Granada, Granada, Spain; and ⁶CIBEROBN (CIBER Physiopathology of Obesity and Nutrition CB12/03/30028), Institute of Health Carlos III, Madrid, Spain

Systematische Übersicht von 16 Diät-Experimenten

The most common inflammatory markers analyzed were hs-CRP, IL-1 β , IL-6, MCP-1, and TNF- α (23–29). Three reports included sICAM-1 and sVCAM-1 (23). Adiponectin (27), macrophage inflammatory protein-1 α (MIP-1 α) (25), and retinol-binding protein 4 (RBP-4) (29) were measured, however, with no significant changes or differences between the groups. Three studies, in addition to the study by Pei et al. (18), investigated the effect of the intervention on gene expression levels (24, 26, 27).

In conclusion, the consumption of milk or dairy products did not show a proinflammatory effect in healthy subjects or individuals who were overweight or obese or had other metabolic abnormalities. The evidence from long-term supplementation showed a weak anti-inflammatory effect in both healthy and metabolically abnormal adults. The evidence from acute and short-term interventions is scarce and thus inconclusive. Further studies need to be developed with enhanced designs and better reporting, and the characterization of the dairy products should be included.

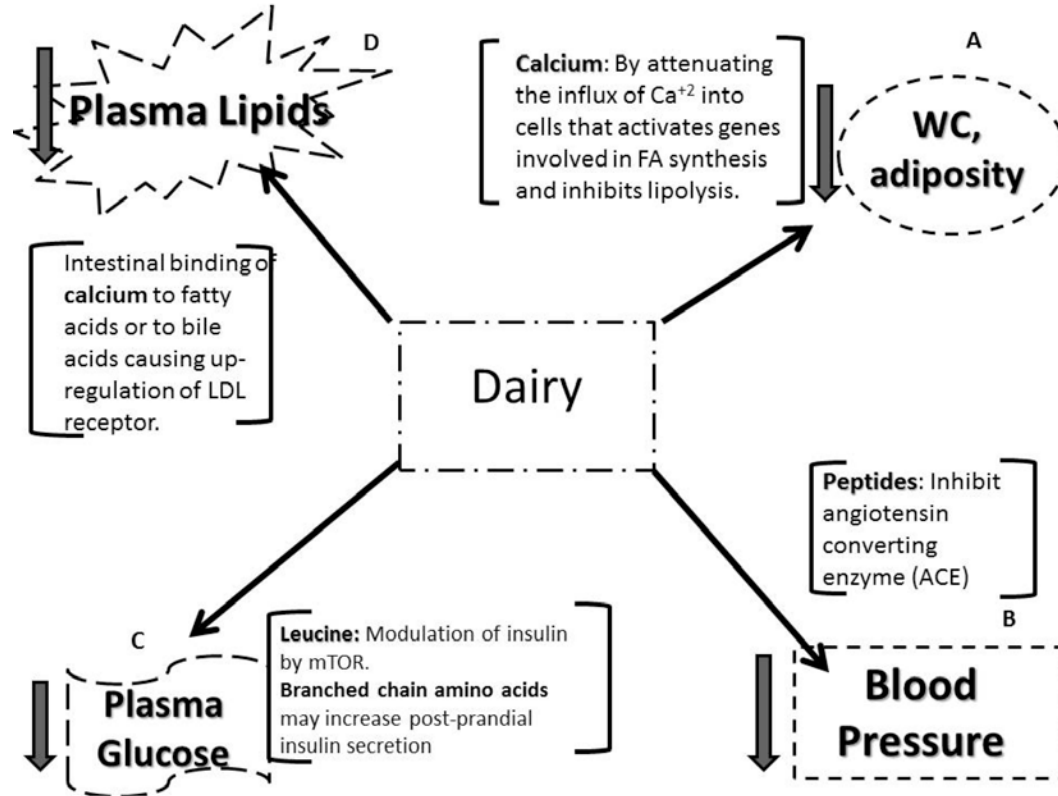
Ergebnis: Keine oder leicht anti-entzündliche Effekte

Mechanismen

Effects of Dairy on Metabolic Syndrome Parameters: A Review

Christine E. Dugan, PhD, and Maria Luz Fernandez, PhD*

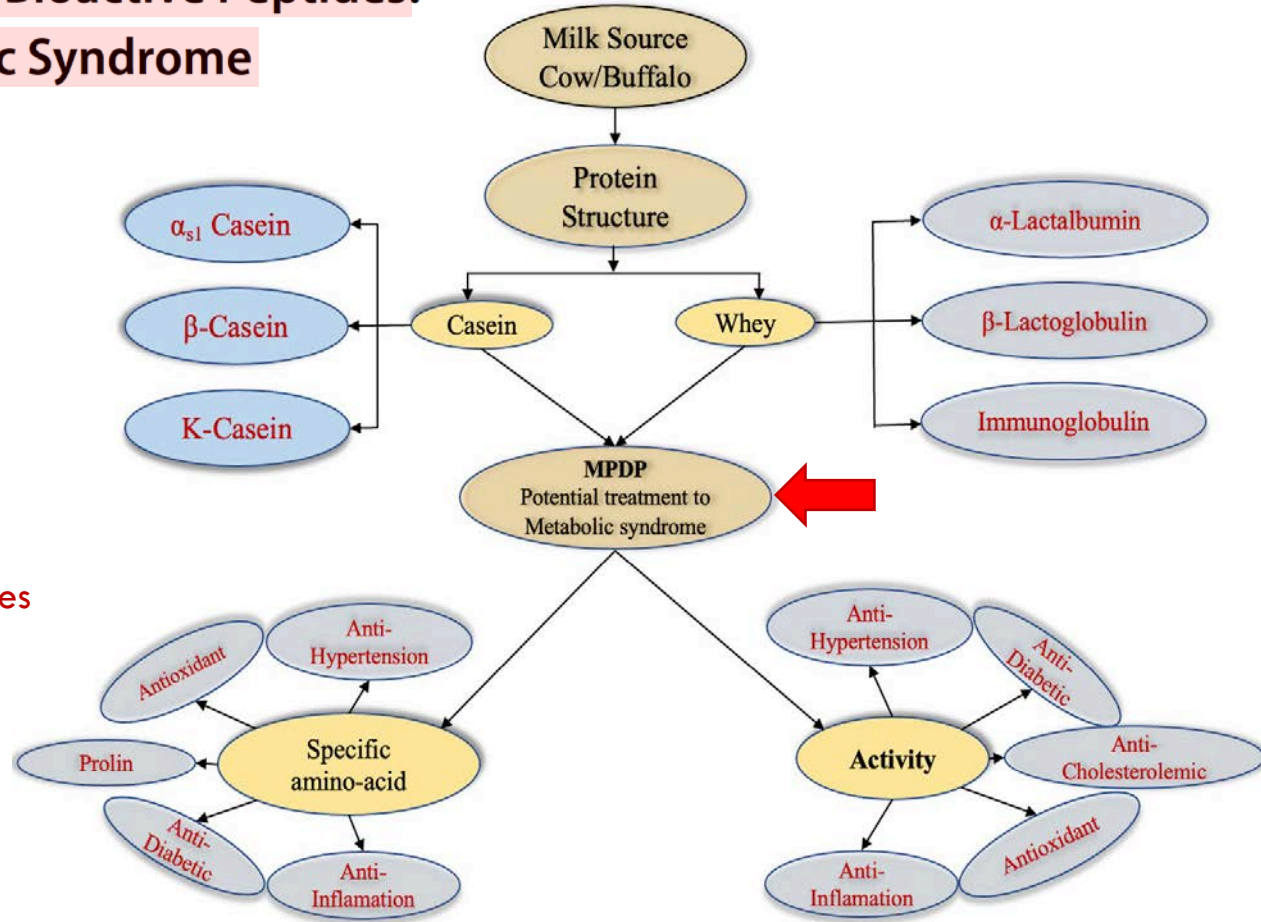
Department of Nutritional Sciences, University of Connecticut, Storrs, Connecticut



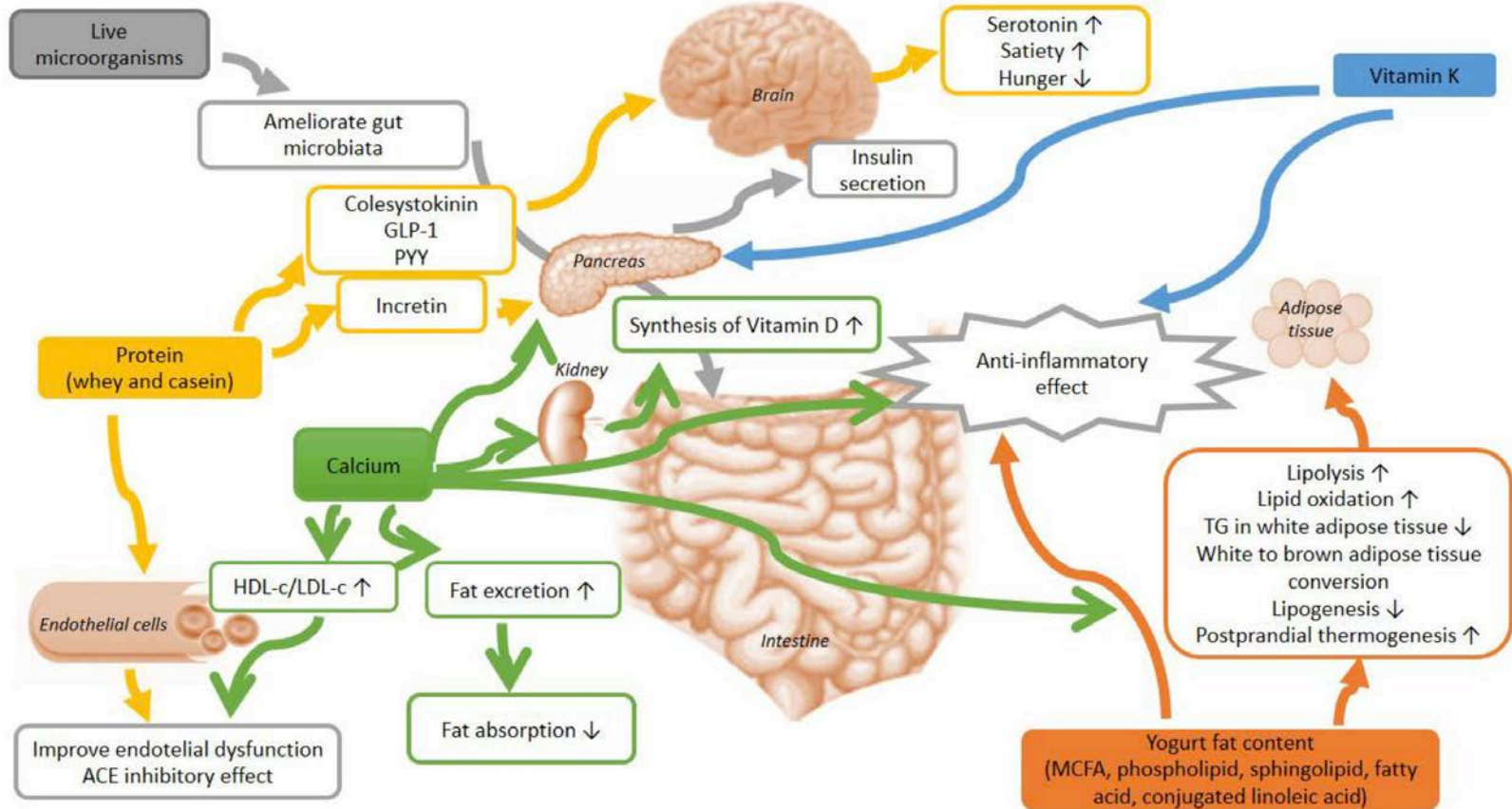
Dairy Milk Protein–Derived Bioactive Peptides: Avengers Against Metabolic Syndrome

- ¹ Institute of Nutrition, Mahidol University, 999 Phutthamonthon 4 Road, Salaya, Nakhon Pathom 73170, Thailand
- ² Department of Nutrition and Dietetics, Central Campus of Technology, Tribhuvan University, Kirtipur, Nepal
- ³ Department of Food Science, College of Food and Agriculture, United Arab Emirates University, Al Ain 15551, United Arab Emirates
- ⁴ Department of Food Science and Nutrition, College of Agriculture and Food Sciences, King Faisal University, Al-Hofuf, P. O. Box 400, Al-Ahsa 31982, Saudi Arabia

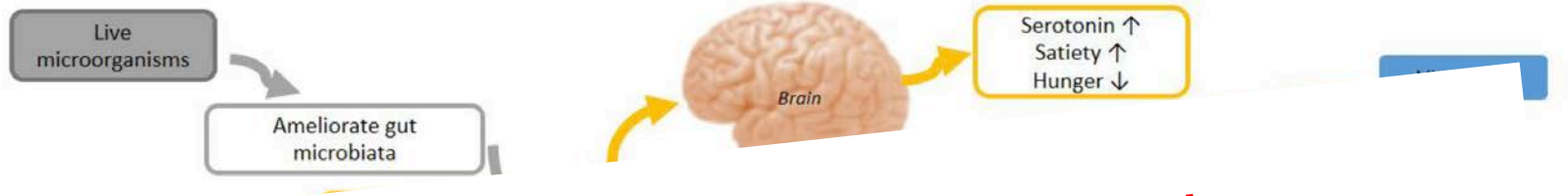
MPDP = Milk Protein–Derived Peptides



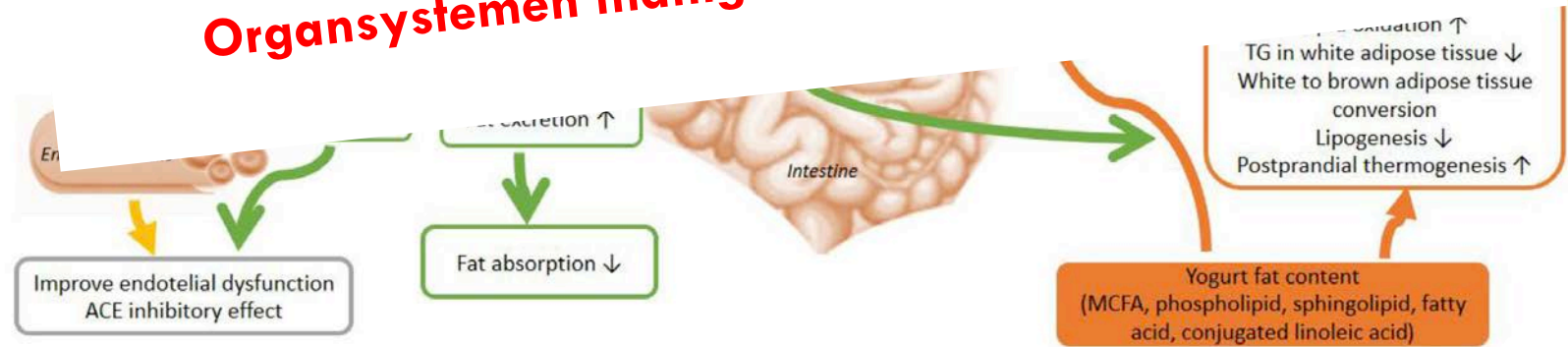
Traditional plain yogurt: a therapeutic food for metabolic syndrome?



Traditional plain yogurt: a therapeutic food for metabolic syndrome?



Die Forschungsarbeiten der letzten Jahre machen klar, dass die verschiedenen Milchbestandteile über komplexe Interaktionen mit den verschiedenen Organsystemen manigfaltige Wirkungen ausüben.



Alles klar?

Saturated fatty acid and *trans*-fatty acid intake for adults and children

WHO guideline



© World Health Organization 2023



These recommendations
high levels of SFA should

Saturated fatty acid and *trans*-fatty acid intake for adults and children

WHO guideline



© World Health Organization 2023



ity in the world. Modifiable risk factors such as use of alcohol are major risk factors. Among saturated and *trans*-fatty acids (TFA) in the diet have the highest CVDs.

Saturated fatty acid intake to 10% of total

to less than 10% of total energy intake

Diet with polyunsaturated fatty acids from plant sources (*conditional recommendation*) including naturally occurring dietary fibre, such as whole grains (*conditional recommendation*).

However, foods containing high levels of intake.

Total fat intake for the prevention of unhealthy weight gain in adults and children

WHO guideline



World Health Organization

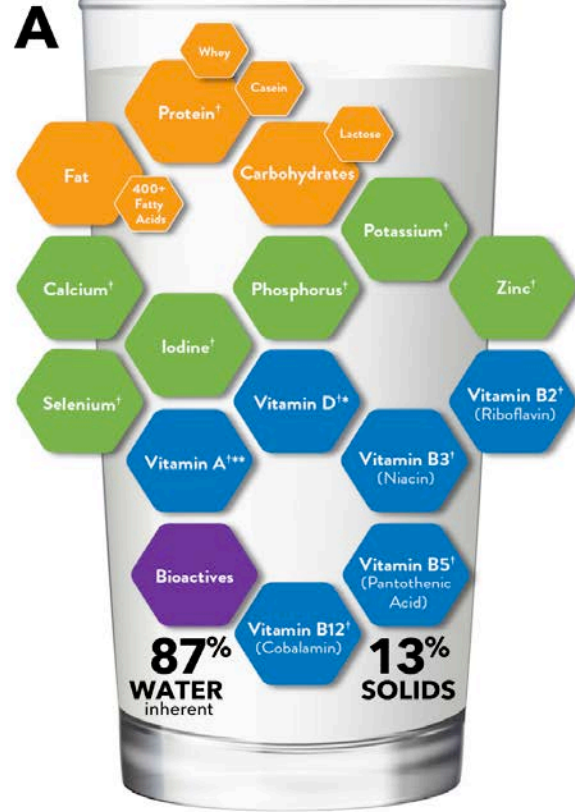
Saturated fatty acid and *trans*-fatty acid intake for adults and children

WHO guideline



World Health Organization

Bestimmt ein Inhaltstoff tatsächlich den Gesundheitswert?



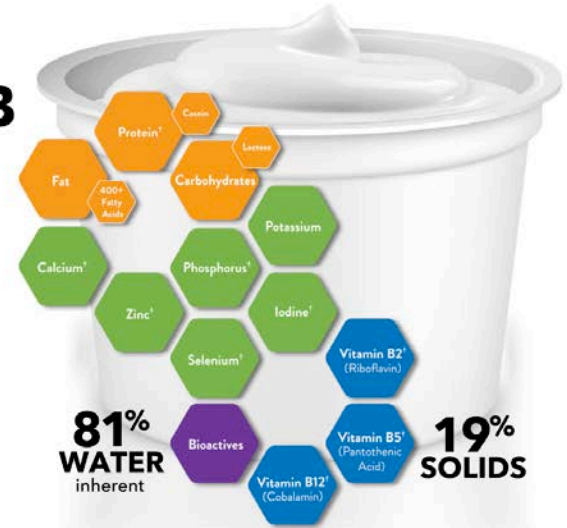
Macronutrients

Minerals

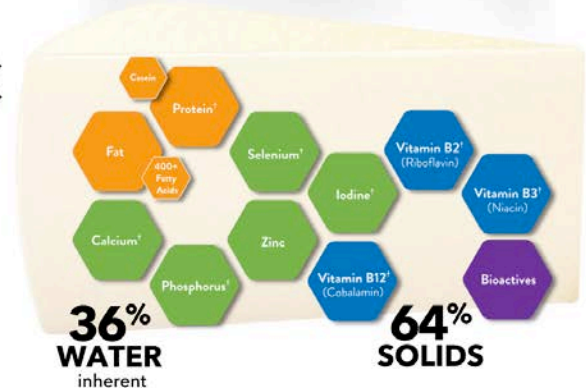
Vitamins

Bioactives

B



C



„a modest amount of meat or whole grains
can be part of a healthy diet.“

New PURE Diet Score (6 healthy components)



LOWER RISK

CARDIOVASCULAR DISEASE (CVD) IN DIVERSE WORLD REGIONS



“PURE” DIET SCORE ASSOCIATIONS
WITH CVD WERE SIMILAR TO
A MEDITERRANEAN DIET SCORE,
AND MUCH STRONGER THAN A
“PLANETARY HEALTH” DIET SCORE

FINDING WERE FROM
DIVERSE GLOBAL NATIONS
INCLUDING LOW
AND MIDDLE INCOME
COUNTRIES

DAIRY WAS
MOSTLY
WHOLE FAT

AVOIDING RED MEAT
DID NOT STRENGTHEN
THE PURE DIET SCORE

Mozaffarian D. Eur Heart J. 2023;44(28):2580–2582

DIE Flexi-CARB-PYRAMIDE für den Gewichtserhalt



Flexi-Carb-Pyramide gestaltet nach Energiegehalt, Nährstoffgehalt, Kaloriengehalt/Öl, Verarbeitungsgrad, nach Worts/Landtags/Hausgehalt © Eva Verlag, 2023

Vielen Dank für Ihr Interesse
und Ihre Aufmerksamkeit



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