



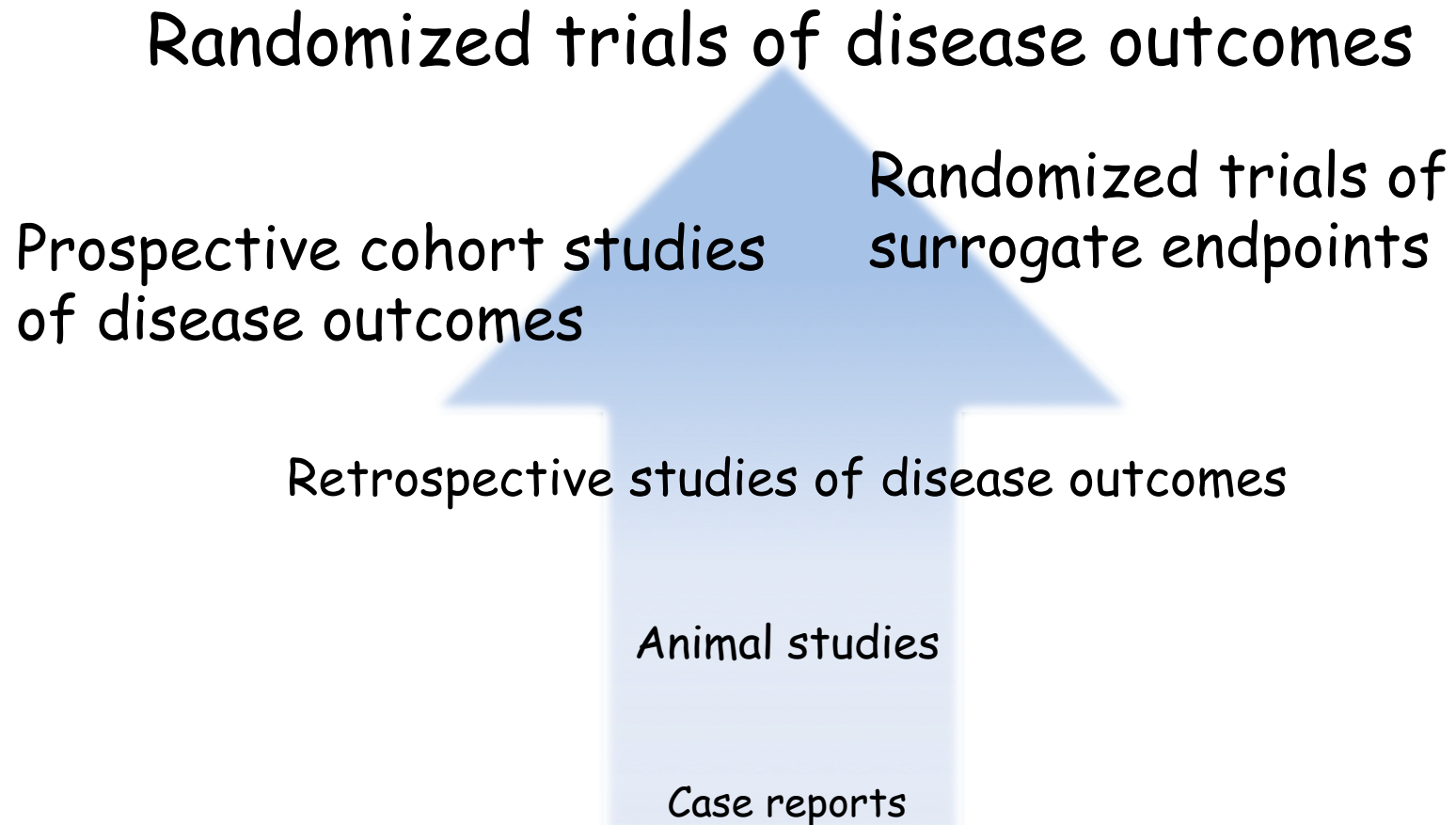
# WHOLE GRAINS AND HEALTH: THE LATEST RESEARCH

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# Hierarchy of Study Designs for Evaluating Strength of Evidence for Disease Risk

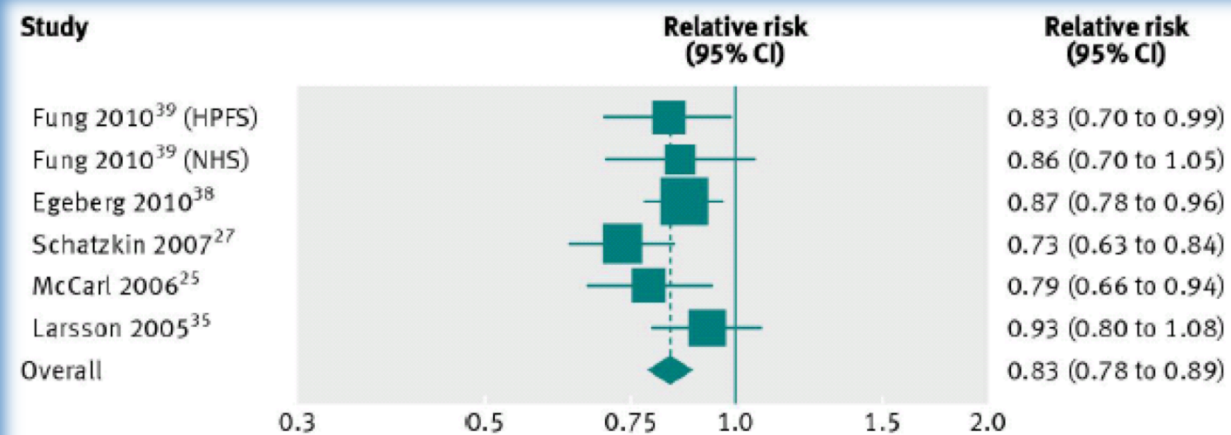


# Prospective human evidence relating whole grains to disease risk

- Prospective cohort studies of disease outcomes
- Randomized trials of surrogate endpoints
- Randomized trials of disease outcomes

# Whole grain intake and relative risk of colorectal cancer: a systematic review of prospective studies

Aune et al BMJ. 2011

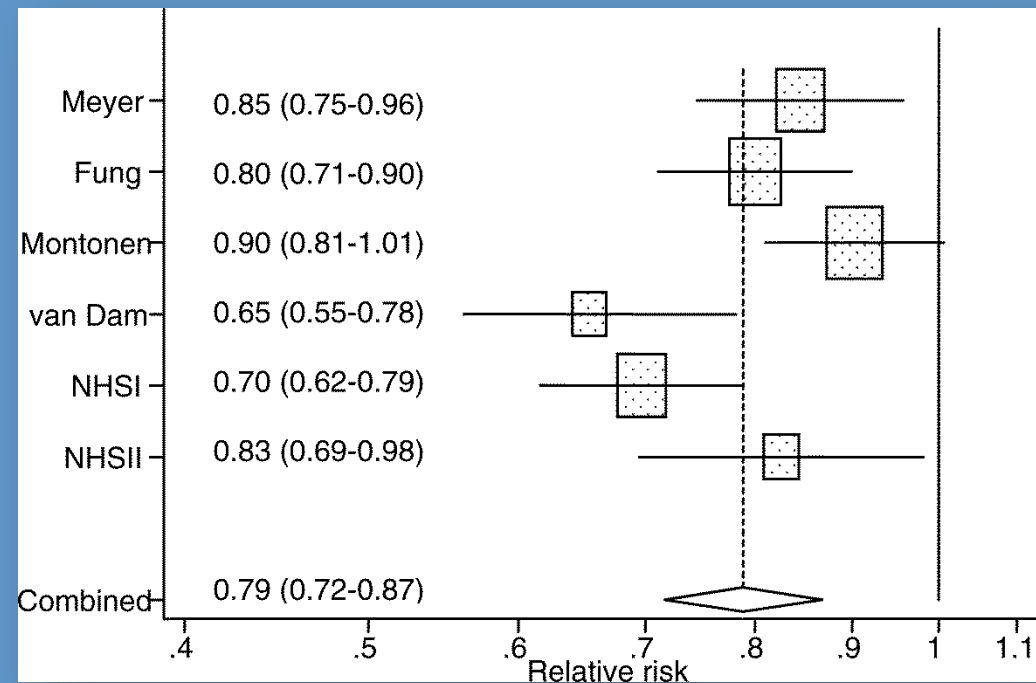


Multivariate-adjusted relative risk of colorectal cancer in high vs low whole grain intake categories. Bars (and diamond) indicate 95% confidence interval. The size of the squares corresponds to the weight of the study in the meta-analysis.

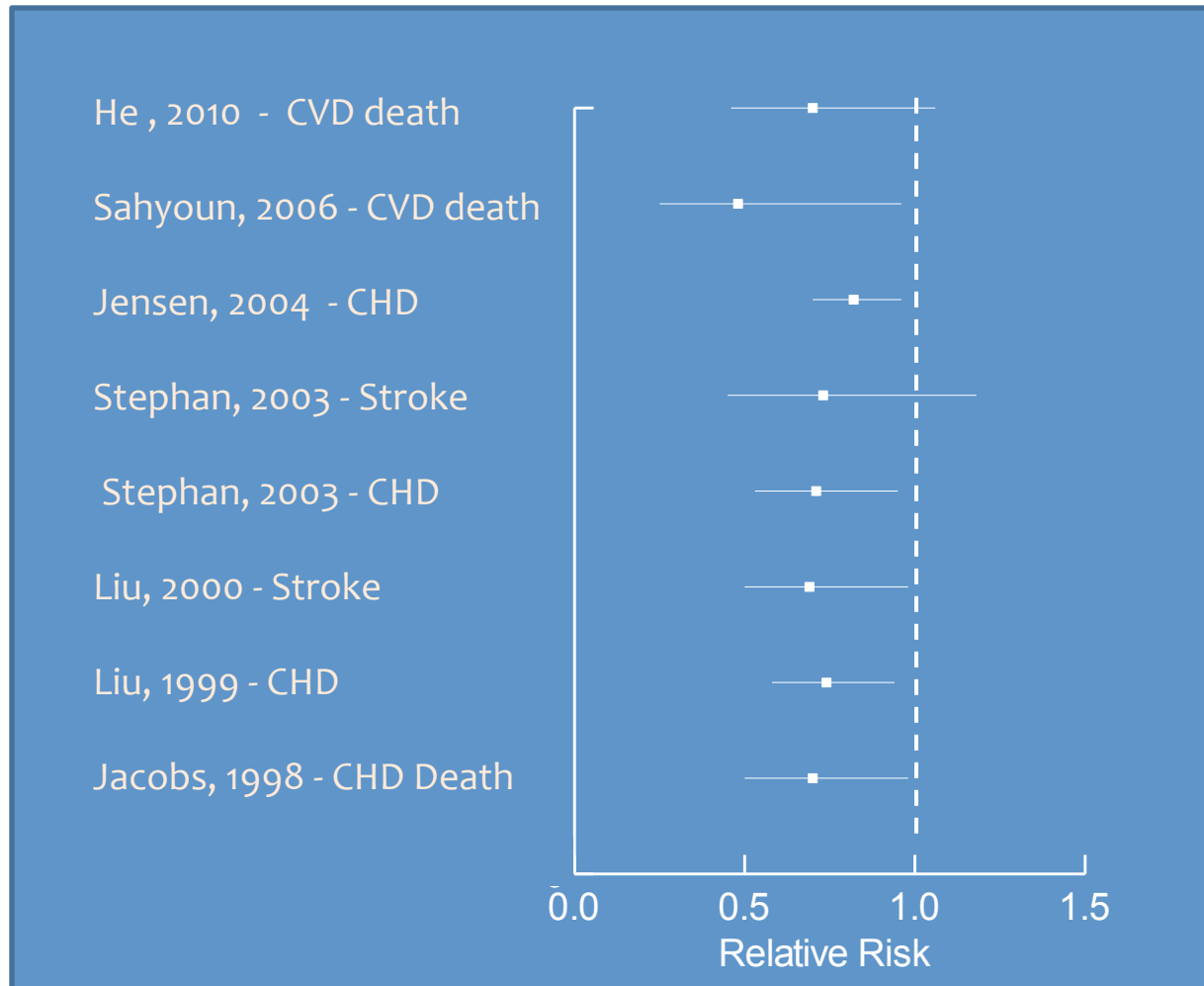
# Whole grain and type 2 diabetes incidence - meta-analysis de Munter et al. PLOS Medicine 2007;4:1385

Multivariate-adjusted relative risk of type 2 diabetes for a two serv/d increment in whole grain intake.

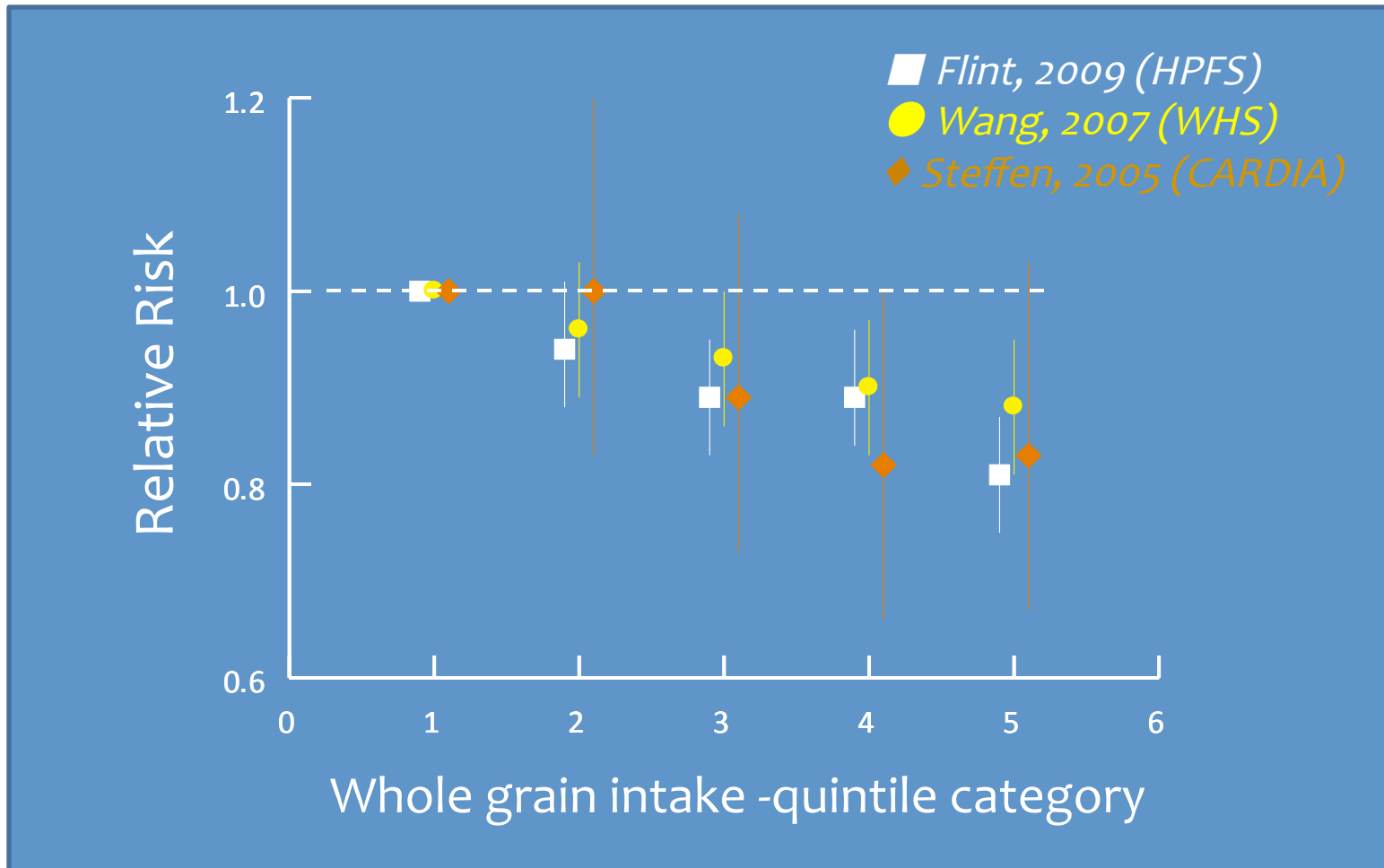
Bars (and diamond) indicate 95% confidence interval. The size of the squares corresponds to the weight of the study in the meta-analysis.



# Prospective studies of whole grain intake and incident CVD

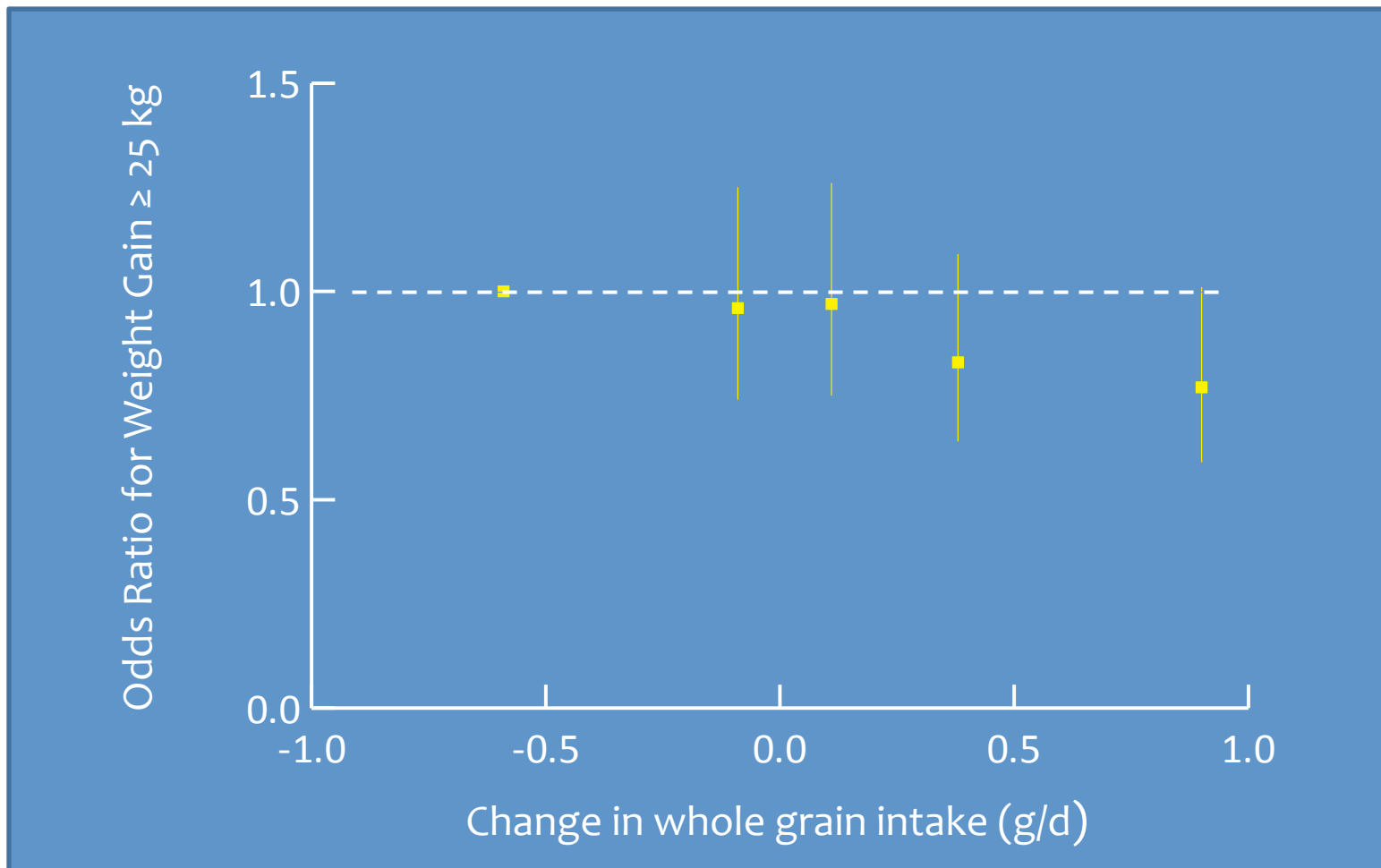


# Whole grain intake and incidence of elevated blood pressure



# Whole grain intake and development of obesity

Adapted from Liu et al. AJCN 2003

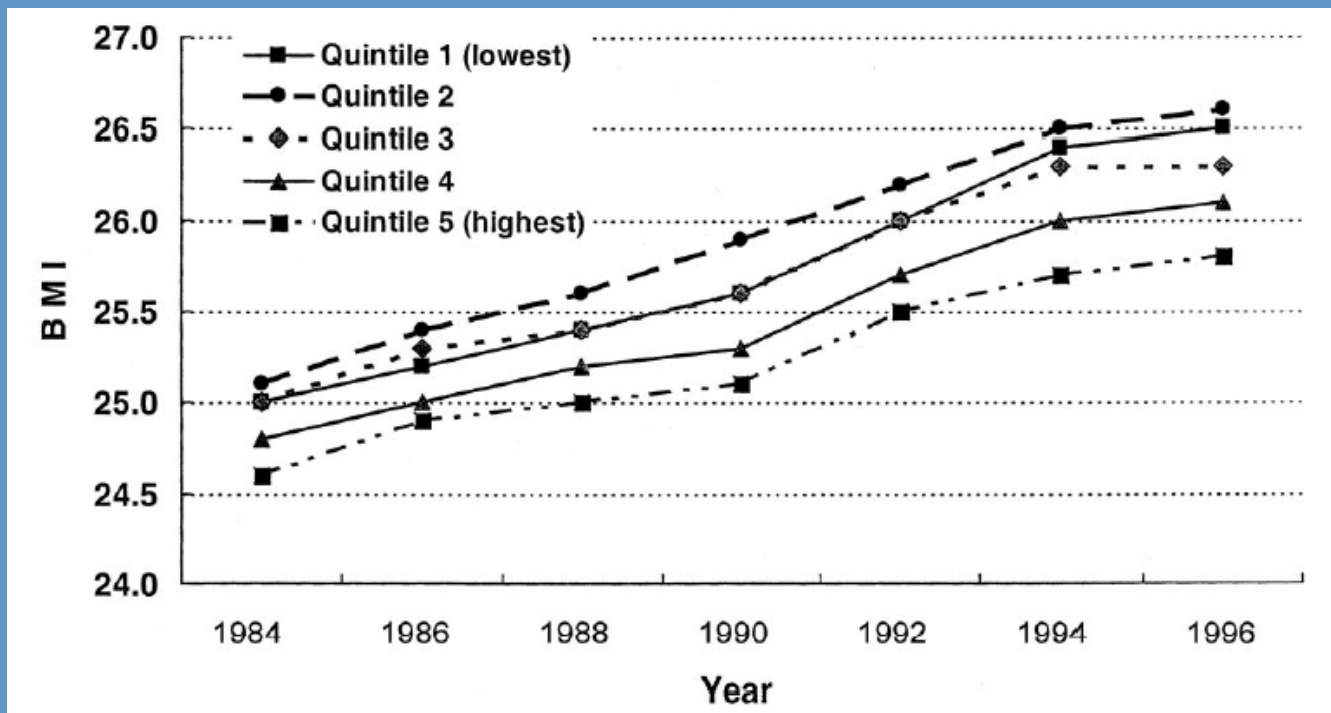




# Whole grain intake and Change in BMI

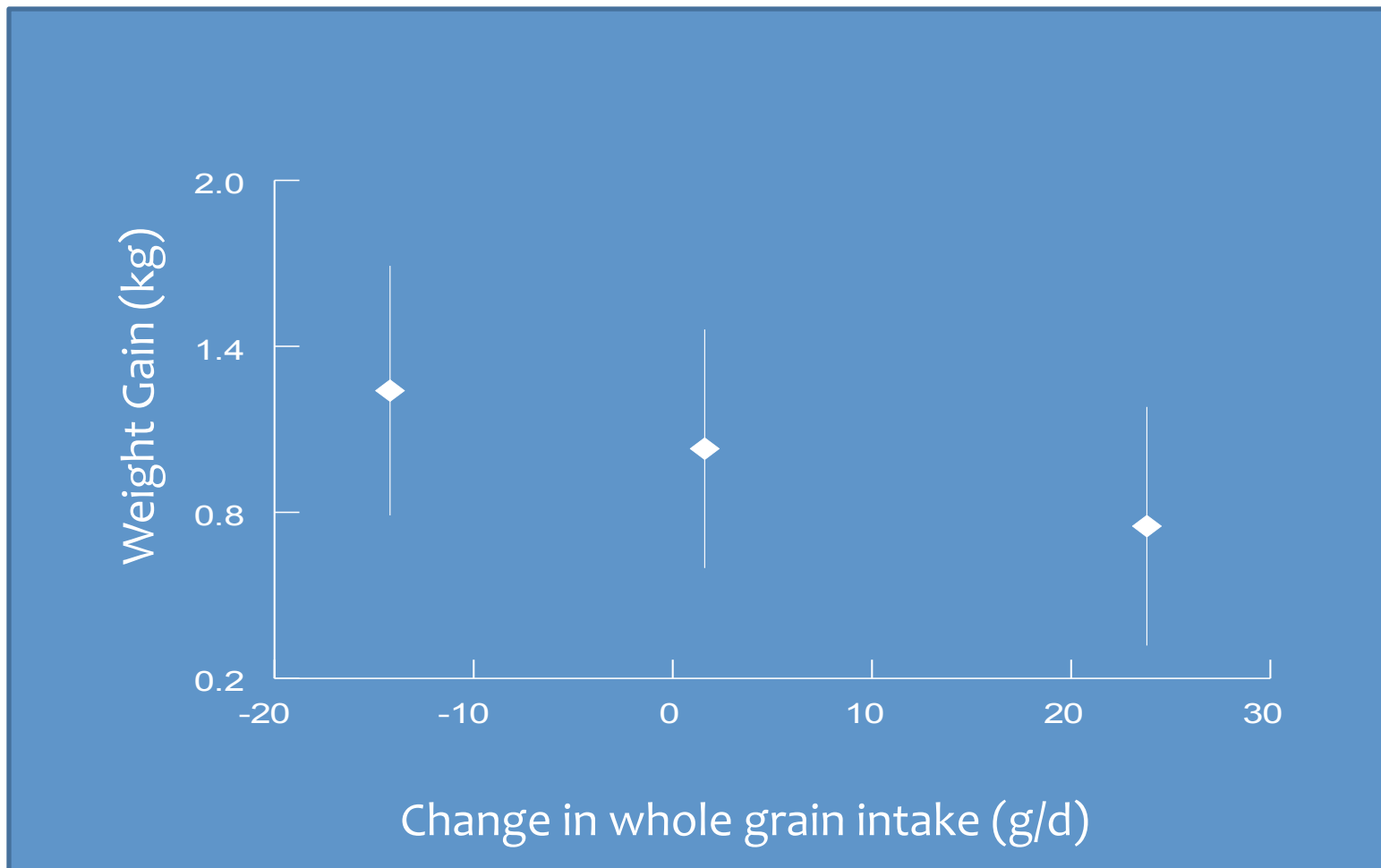
Adapted from Liu et al. AJCN 2003

Mean BMI from 1984 to 1996 according to quintiles of whole-grain intake



# Whole grain intake and 8 year weight change

Adapted from Koh-Banerjee et al. *AJCN* 2004;80:1237

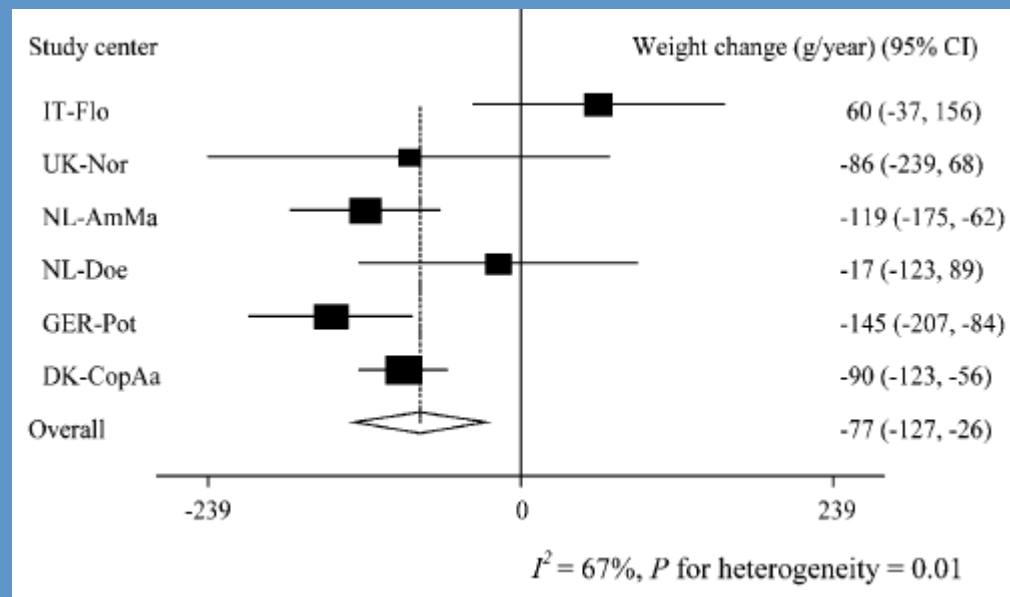


# Cereal fiber intake and weight gain

Du et al. *AJCN* 2010;91:329

Prospective cohort study of 89,432 Europeans (EPIC) who were free of cancer, CVD, and diabetes followed for an average of 6.5 y.

The values presented are regression coefficients for a 10-g/d cereal fiber intake. Bars (and diamond) indicate 95% confidence interval. The size of the squares corresponds to the weight of the study in the meta-analysis.



# Prospective human evidence relating whole grains to disease risk

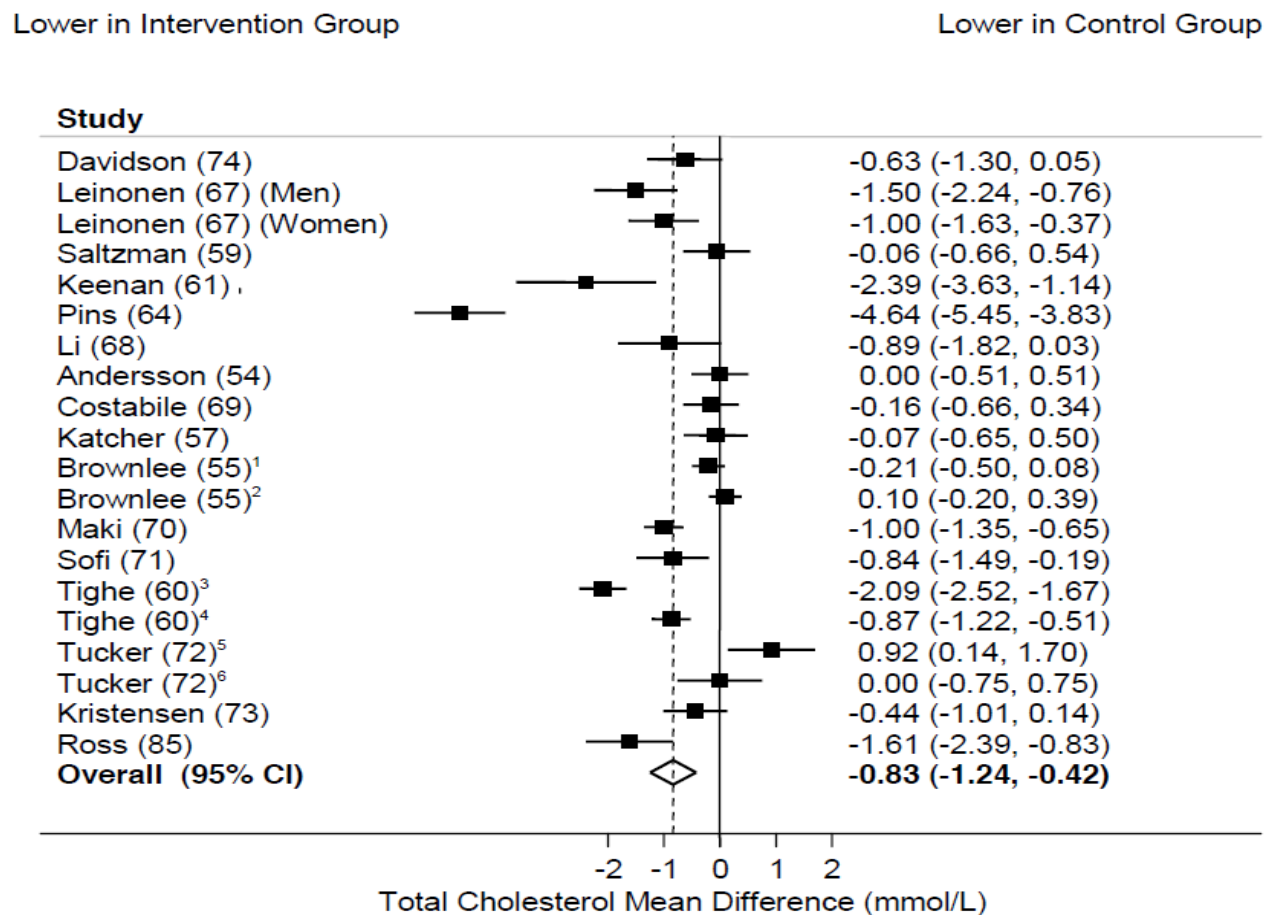
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Mean difference in post-treatment metabolic markers:  
whole-grain intervention groups versus controls\*  
(Ye et al, J Nutr 2012;142: 1304)

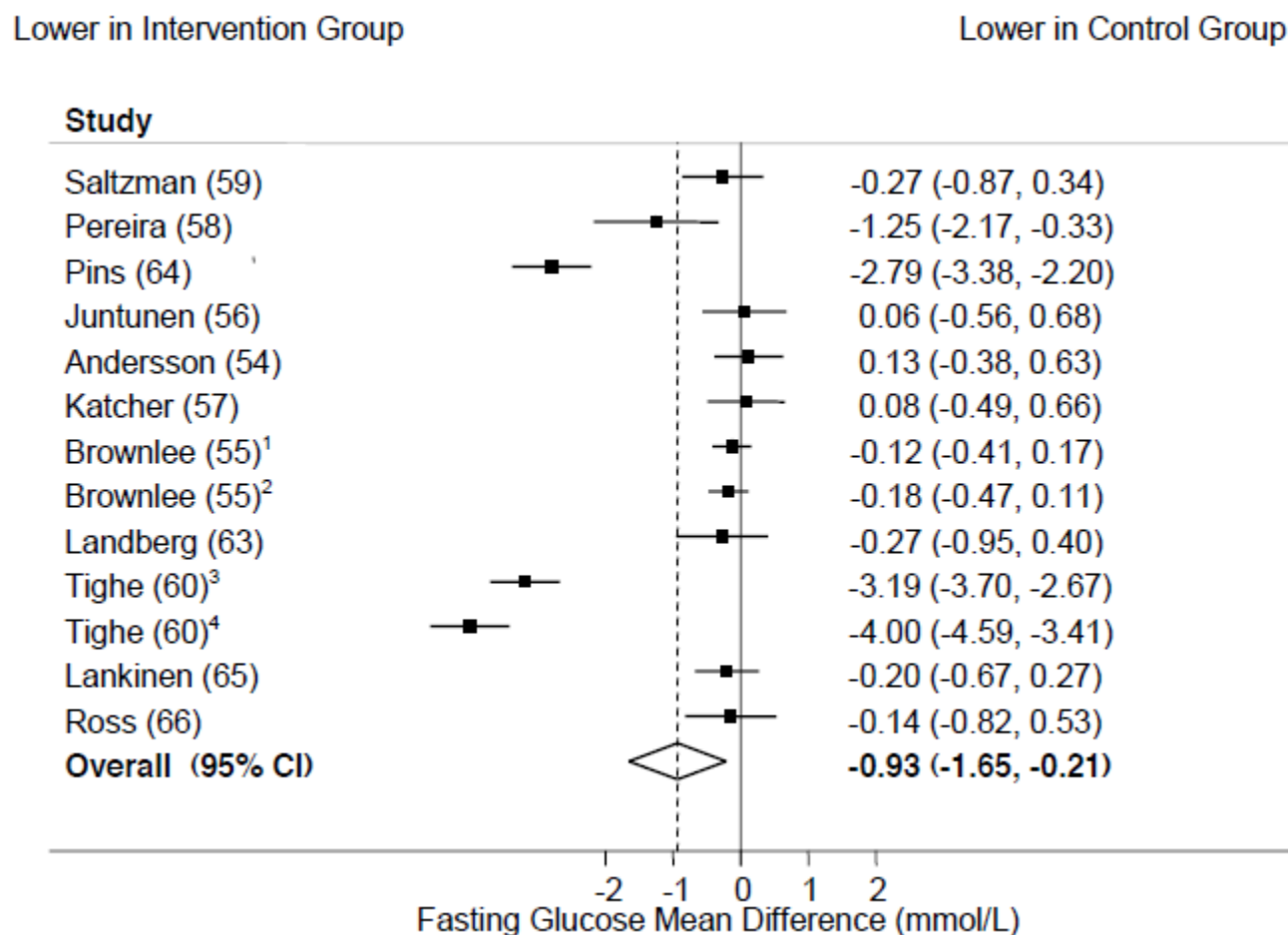
Metabolic biomarkers	Studies, <i>n</i>	Weighted mean difference (95% CI)
Fasting insulin, <i>pmol/L</i>	10	−0.29 (−0.59, 0.01)
Fasting glucose, <i>mmol/L</i>	11	−0.93 (−1.65, −0.21)
Total cholesterol, <i>mmol/L</i>	16	−0.83 (−1.24, −0.42)
LDL-cholesterol, <i>mmol/L</i>	15	−0.72 (−1.34, −0.11)
Systolic blood pressure, <i>mm Hg</i>	6	−0.06 (−0.21, 0.10)
Diastolic blood pressure, <i>mm Hg</i>	6	−0.05 (−0.21, 0.11)
Weight gain, <i>kg</i>	9	−0.18 (−0.54, 0.18)

\*Based on 21 RCT's that directly investigated the effects of whole-grain interventions on one or more metabolic intermediate risk factors.

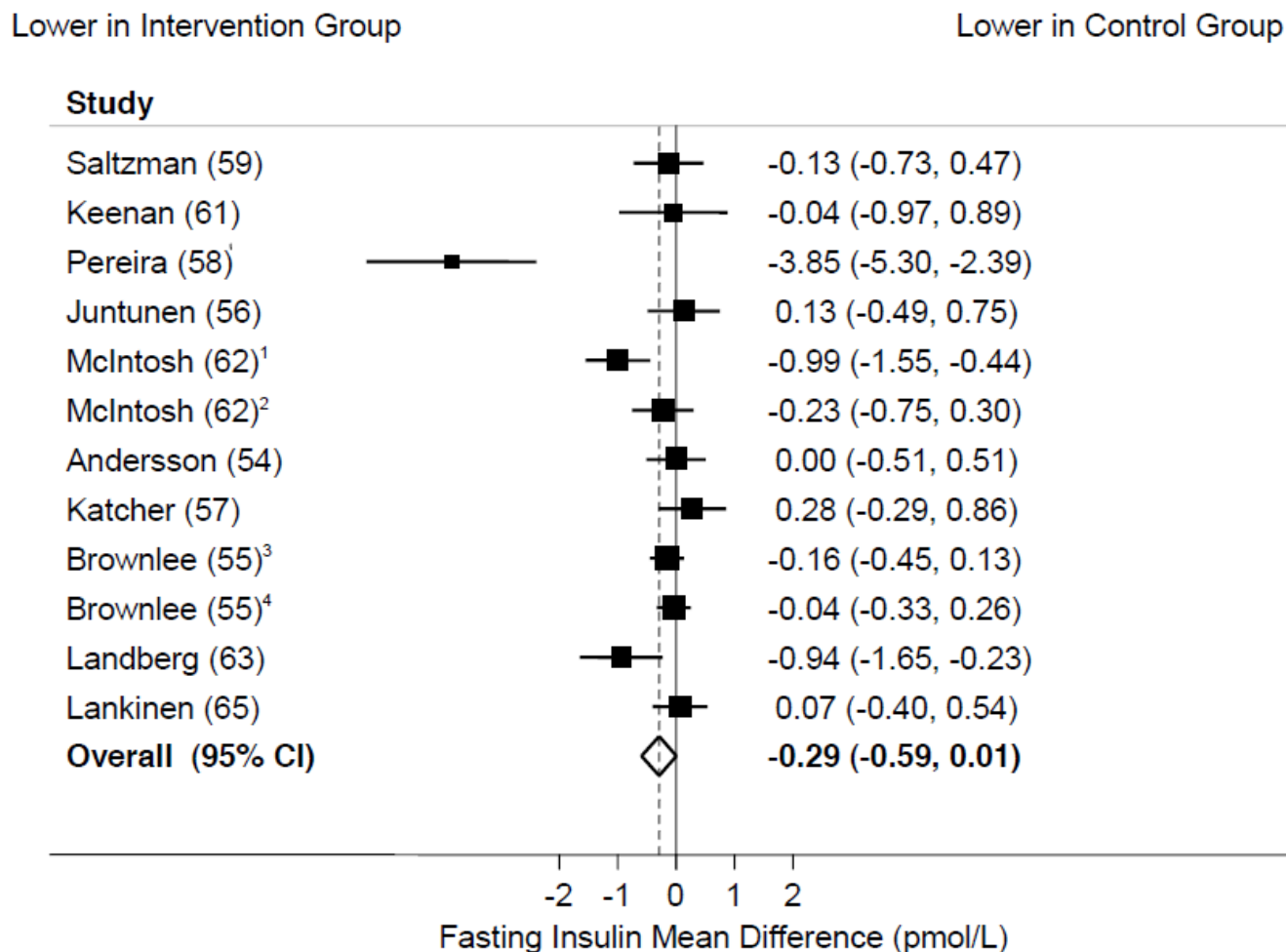
# Mean differences (95% CI) in total cholesterol (mmol/L): whole-grain intervention groups versus controls (Ye et al, J Nutr 2012;142: 1304 – online supporting material)



Mean differences (95% CI) in fasting glucose (mmol/L):  
 whole-grain intervention groups versus controls  
 (Ye et al, J Nutr 2012;142: 1304 – online supporting material)



# Mean differences (95% CI) in fasting insulin (pmol/L): whole-grain intervention groups versus controls (Ye et al, J Nutr 2012;142: 1304 – online supporting material)





# Comparison of intervention vs. observational studies - whole grain and diabetes risk

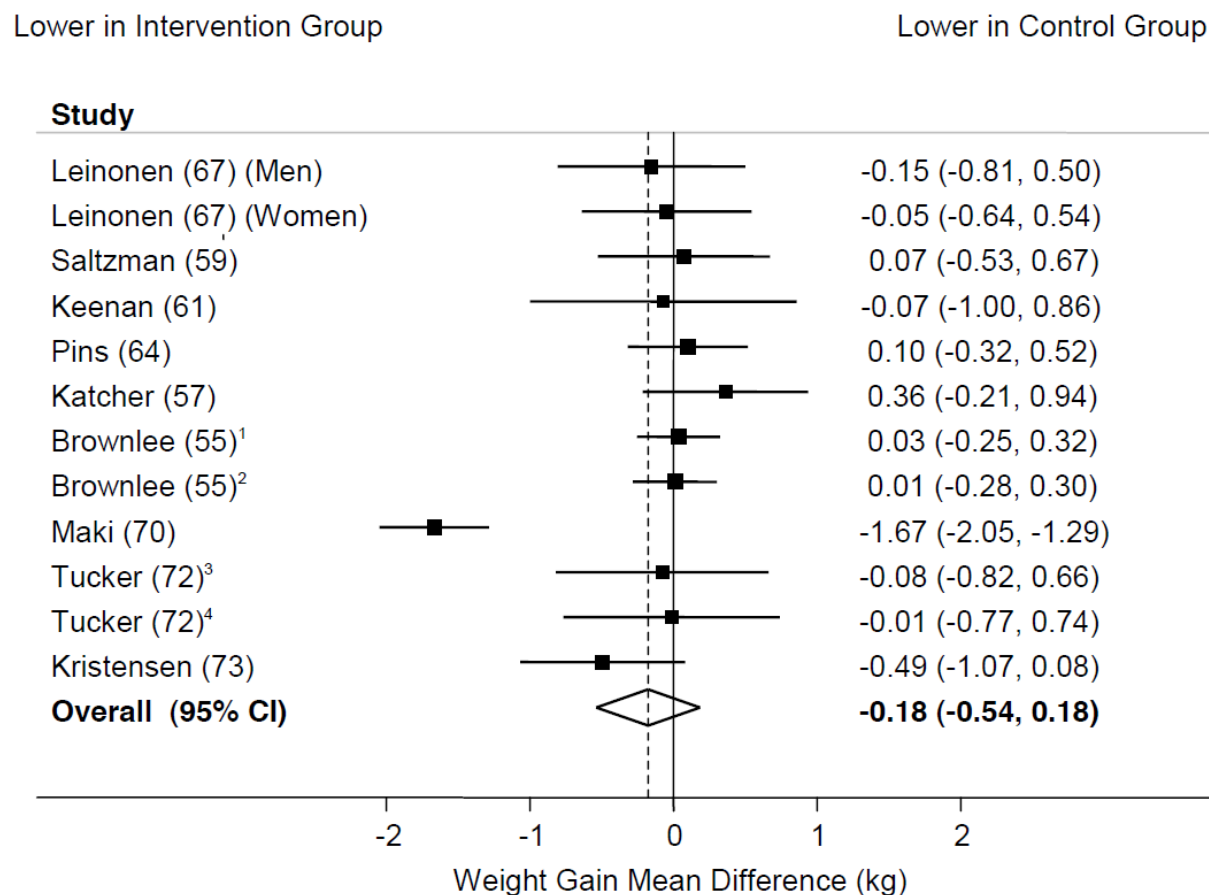
## Katcher et al (2008)

- Randomized, parallel study
- 50 obese adults with metabolic syndrome
- Followed 12 weeks
- Whole grain exposure: 4-7 vs. 0 servings (~64-112 g)/ day as part of hypocaloric diet
- No significant difference between treatment groups for any insulin or glucose measures (fasting, 2 hr OGT, AUC) or for the Insulin Sensitivity Index.

## de Munter et al (2007)

- Cohort study
- 161,737 women without diabetes, CVD or cancer
- Followed 12-14 years
- Whole grain exposure: 31-40 vs. 4-6 gm/d
- 14-25% lower risk of type 2 diabetes

# Mean differences (95% CI) in weight gain (kg) whole-grain intervention groups versus controls (Ye et al, J Nutr 2012;142: 1304 – online supporting material)



1 Dosage: 60 g/day; 2 Dosage: 60-120 g/day; 3 Healthy participants; 4 Hyperglycemic participants

# Whole Grain Interventions in Trials of Physiologic Measures

## Examples selected from 21 trials

- high-fiber rye vs white-wheat breads to make up  $\geq 20\%$  of energy
- dietary advice to avoid whole-grain foods vs advice to obtain all grain from whole grains
- hypocaloric diet (maintenance energy minus 4.2 MJ/d) with or without oats at 45 g/4.2 MJ dietary energy
- oat cereal group (standardized to 5.52 g/day beta-glucan) vs a low-fiber cereal control group (less than 1.0 g/day total fiber)
- standard diet (100% of carbohydrates from rice) vs a barley diet (30% carbohydrate from barley and 70% from rice)
- a diet containing 150 g/d of either whole grain bread made from a variety of old grain grown in Tuscany vs commercially available bread
- two portions per day of whole-grain RTE oat cereal (3 g/day oat  $\beta$ -glucan) vs energy-matched low-fiber foods (control), as part of a reduced energy (500 kcal/day deficit) dietary program

# Prospective human evidence relating whole grains to disease risk

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# Randomized trials of whole grains and disease outcomes

None

# Summary

Current evidence: Do we know more than we think we know?

- ❑ Prospective cohort studies
  - ❑ show consistent findings for most health outcomes
  - ❑ supported by trials of surrogate outcomes for total and LDL cholesterol and fasting glucose.
- ❑ Trials of surrogate outcomes
  - ❑ Comparability a problem because there are no standard interventions
  - ❑ Inconsistency between cohort and trial finding
    - ❑ Lack of consistency between trials (interventions)
    - ❑ Different paradigm/different hypotheses

# Future Directions & Challenges

## Future evidence for whole grains and health

- ❑ Prospective cohort studies
  - ❑ More cohort studies of surrogate endpoints, particularly for weight/waist circumference change
  - ❑ Need to consider types of whole grain/whole grain intake in grams
- ❑ Randomized intervention trials of surrogate endpoints
  - ❑ Focus of future evidence
  - ❑ Standardization interventions for comparability among trials
    - ❑ Dose response effects
    - ❑ Effect of different types of whole grain and interactions between them