



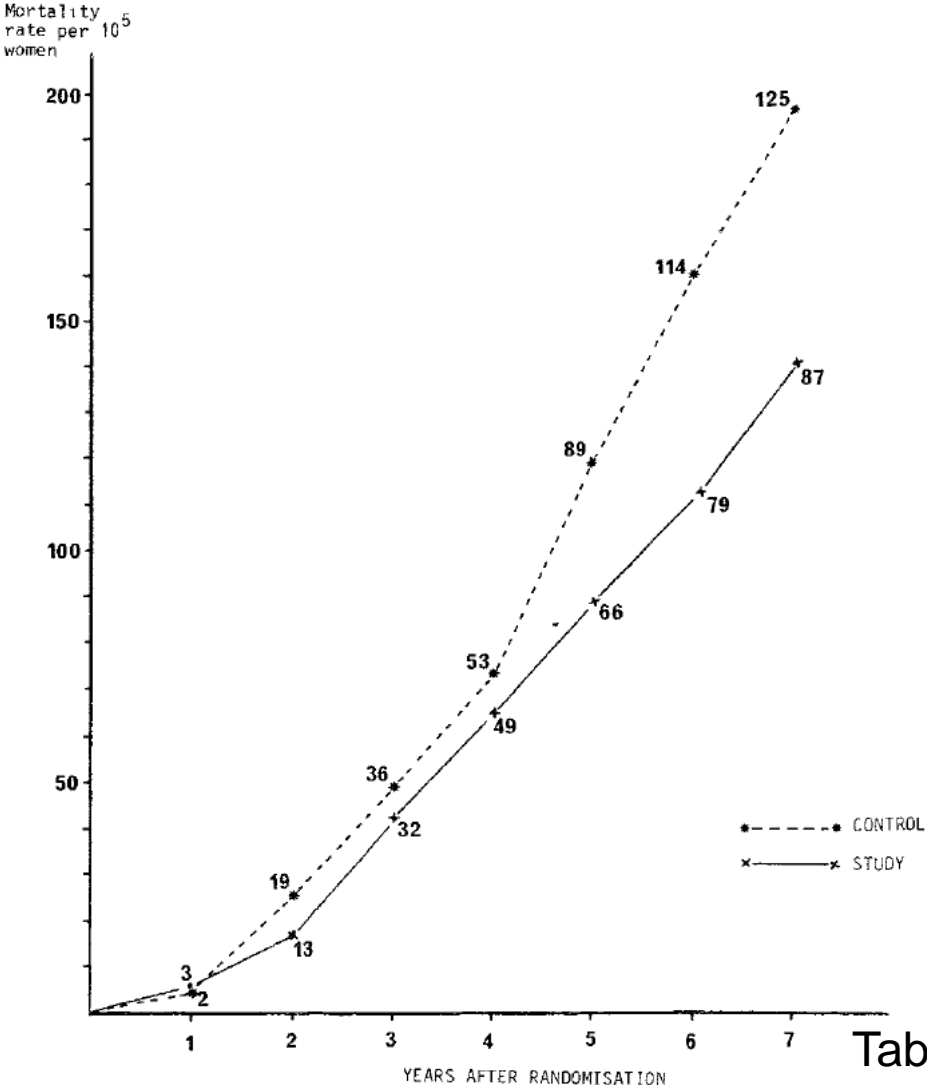
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# Controversies in Mammography Screening and Screening between 40 and 50 years

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# Cumulative mortality rates women aged 40 -74 at entry



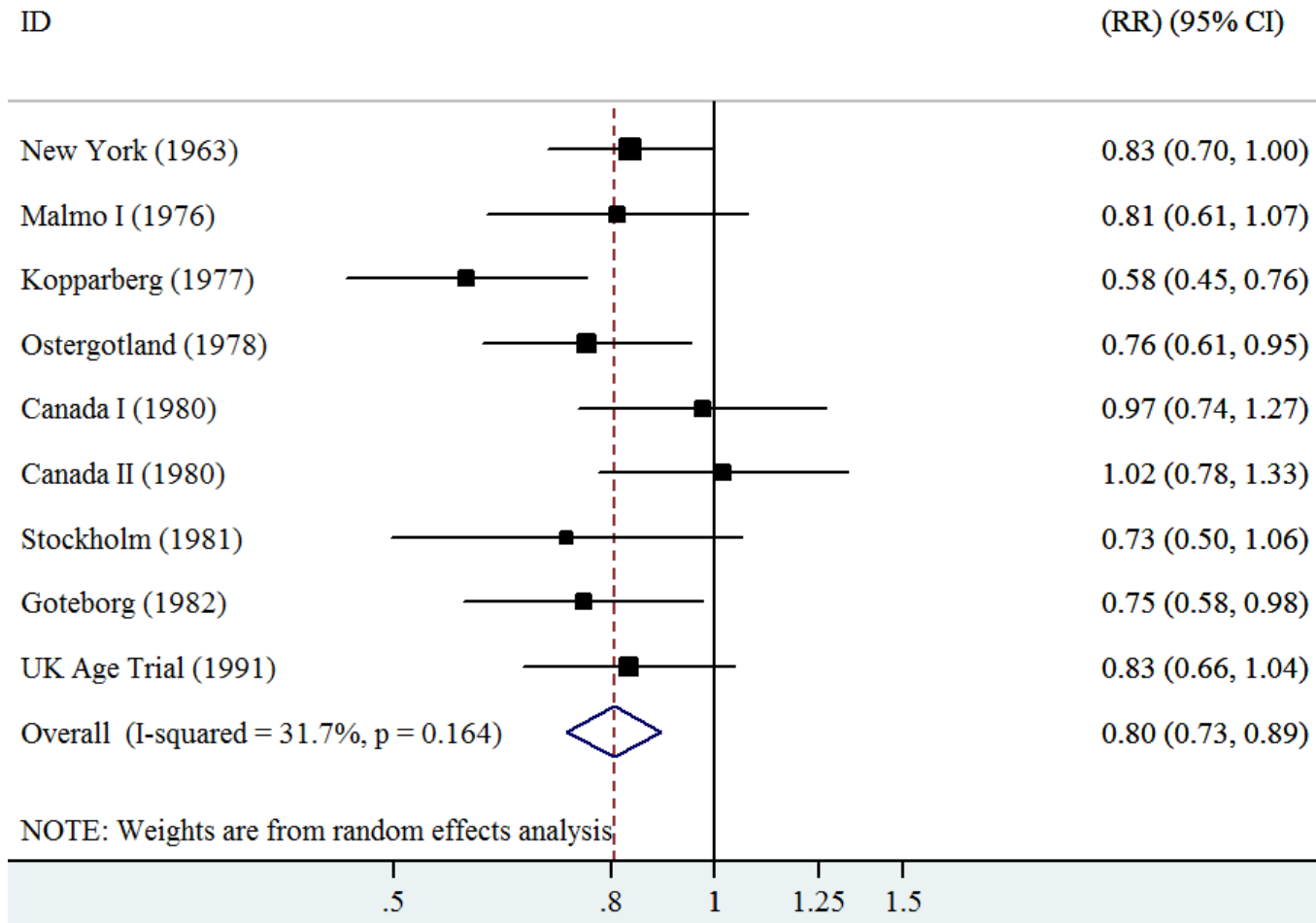
# Early service screening programs in Europe

- Sweden, Västmanland 1986
- Netherlands 1988
- Great Britain 1988
- Finland 1989
- Sweden, Stockholm 1989

# Criticism

- Validity of studies questioned by Nordic Cochrane foundation
  - Olsen and Gøtzsche, Lancet 2001
- Criticism refuted by WHO:
  - Screening reduces mortality in participants by 30 %. IARC Working Group, in : Vaino and Bianchini eds. Lyon 2002

# Meta-analysis of 11 randomized trials: risk reduction by invitation to screening 20%



# Observational Studies

Type	N	Breast cancer mortality reduction	Comment
Trend analysis	12	1-9%/year pre vs postscreening: 28% - 36%	Least reliable
Incidence based mortality	7	Invitation 25% Participation 38%	Meta-analysis
Case control	7	Invitation 31% Participation 48%	Meta-analysis

# Overdiagnosis

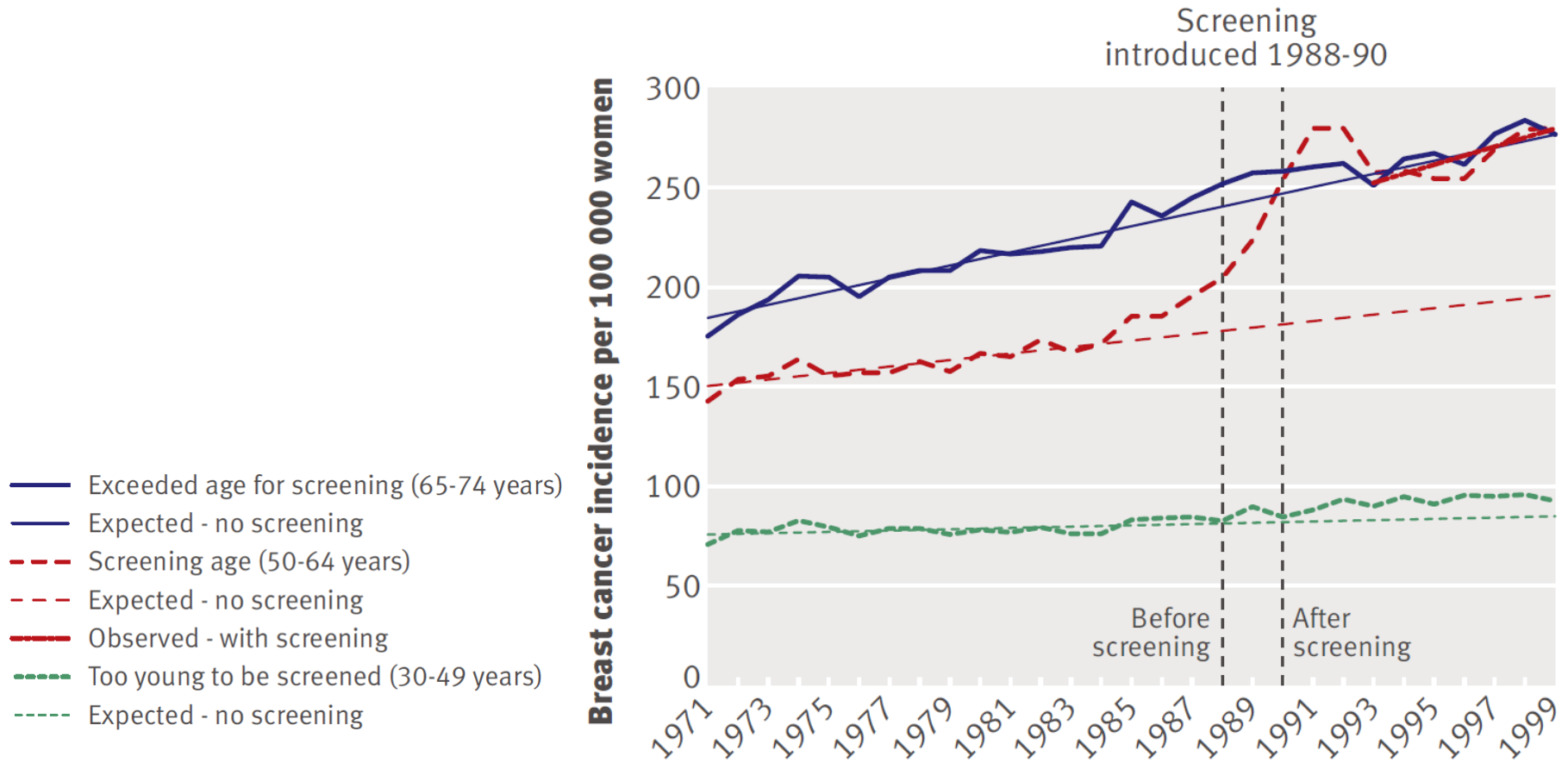
- Carcinoma are detected, which would not have become symptomatic during the remaining lifetime of the screened person
- These carcinoma are treated as all other carcinomas, since they cannot be discriminated from other carcinoma.
- This leads to more therapies in a screened population with associated morbidity and cost

# Problems with estimation of overdiagnosis

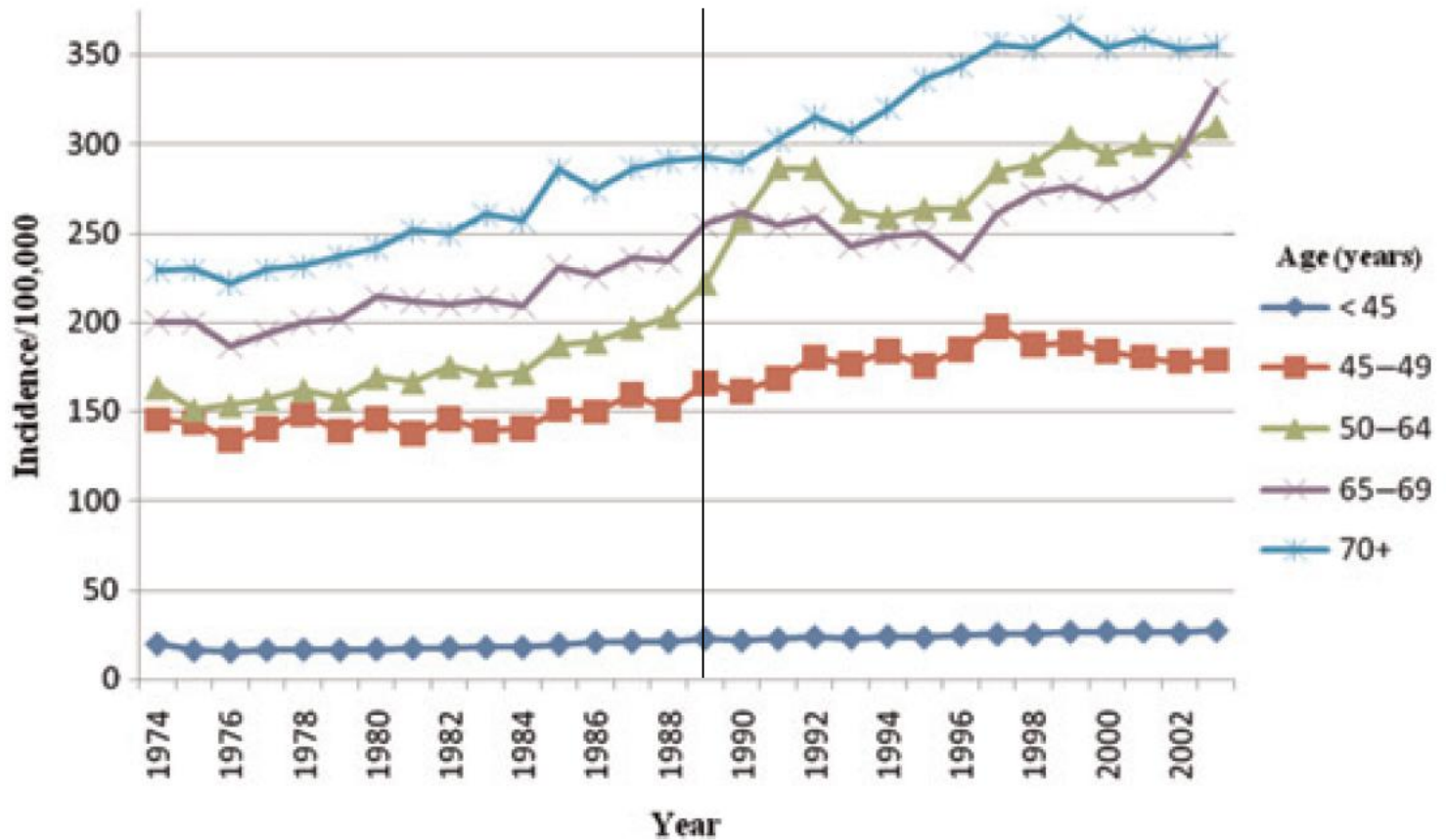
- Changing background incidence in screened and control group
- Lead time bias



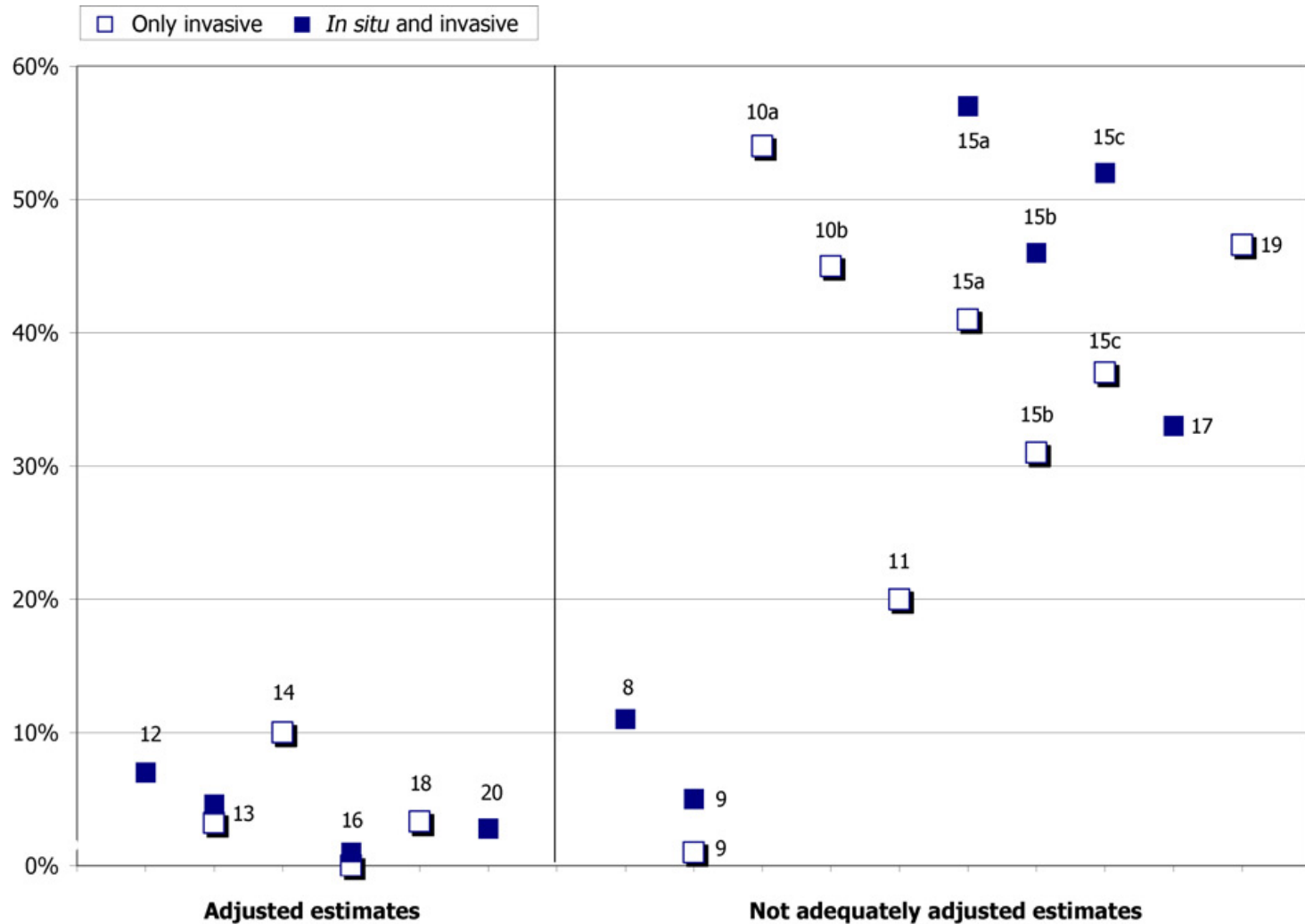
# Overdiagnosis in UK equals 53%



# Overdiagnosis in UK equals 4 – 7 %

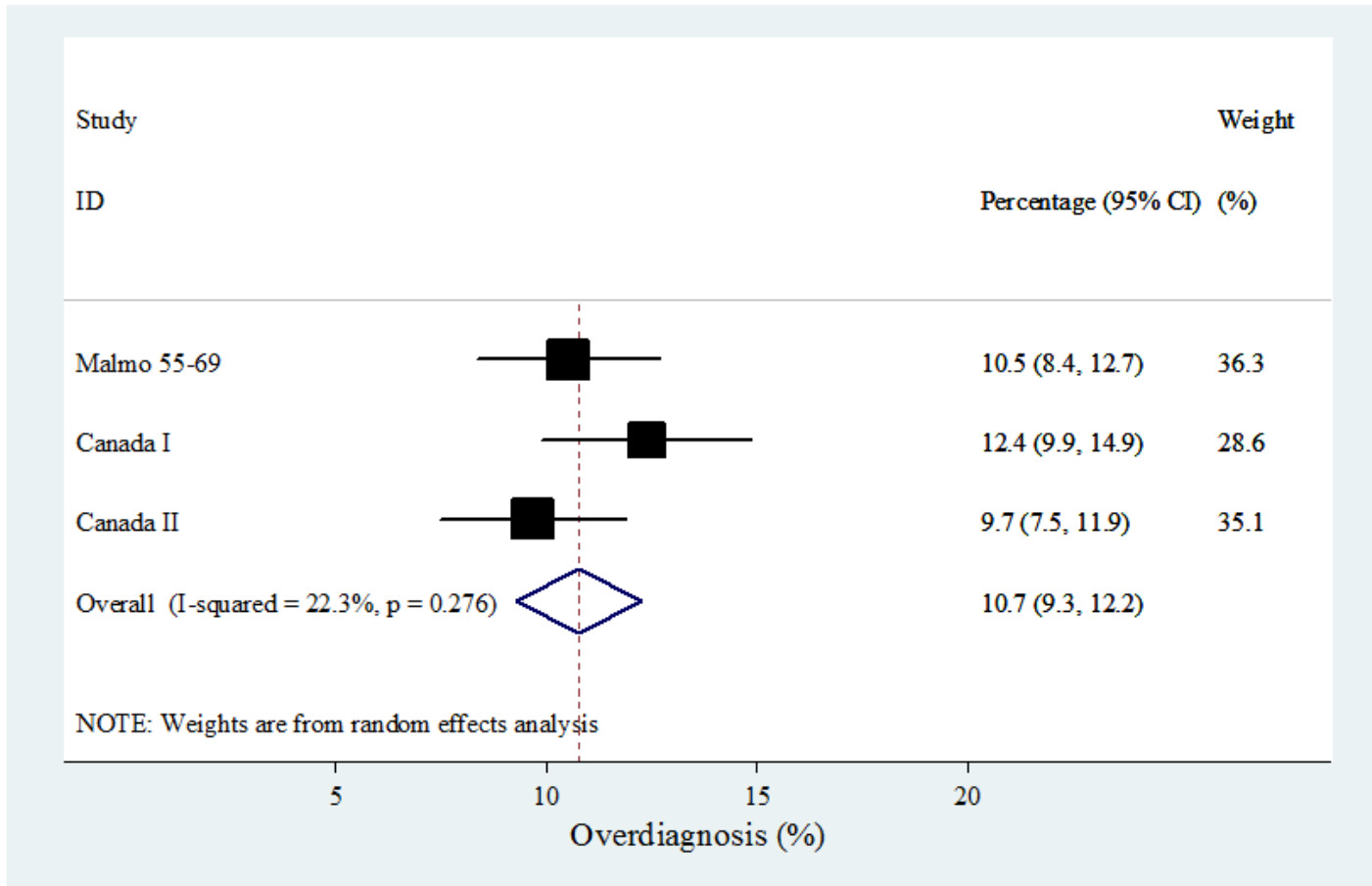


# Overdiagnosis in mammographic screening for breast cancer in Europe: observational studies



# Overdiagnosis in mammographic screening: randomized studies

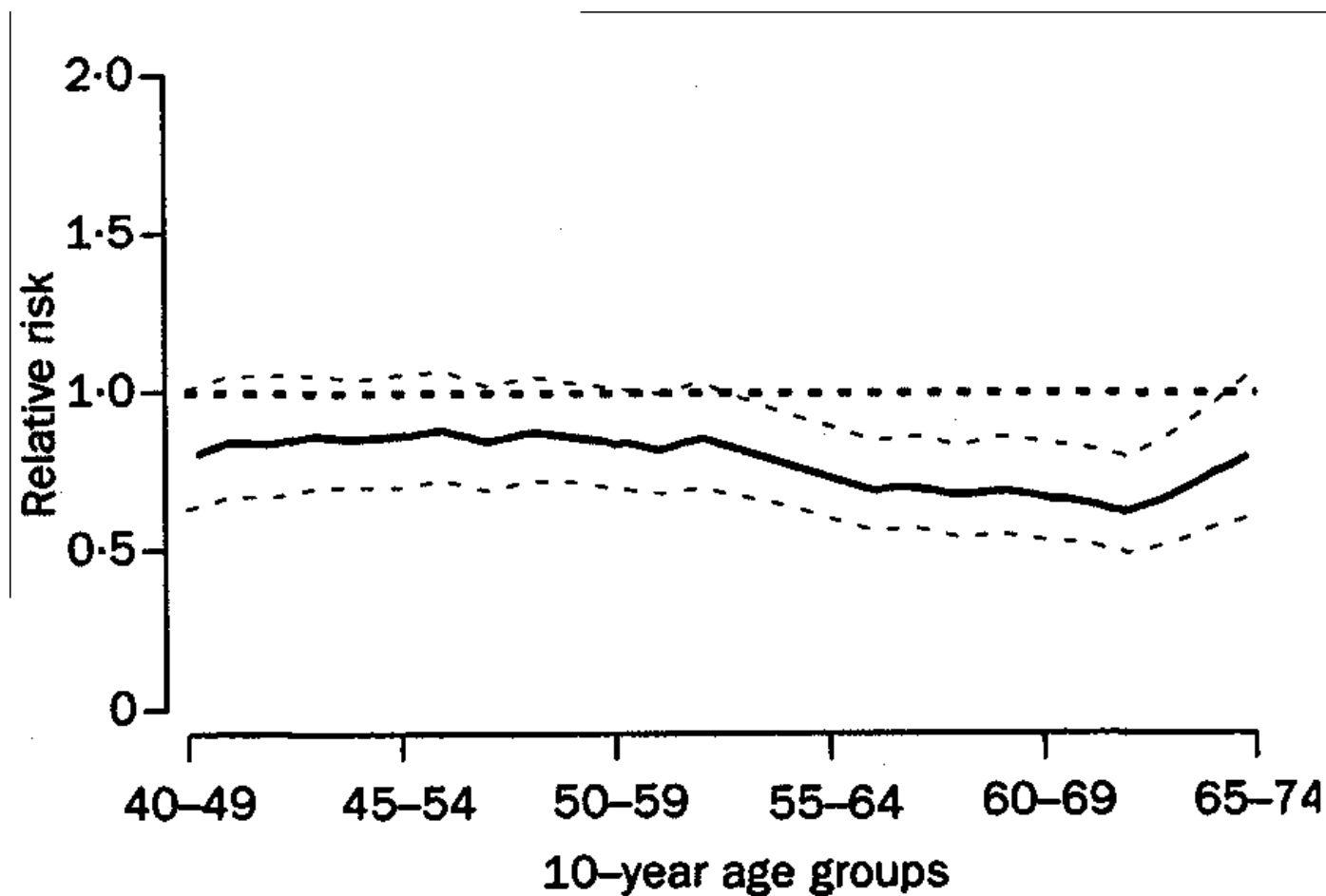
## proportion of cancers over long-term follow-up



# Balance of mortality reduction and overdiagnosis

	Per 1000 participants	Participants per event
Breast cancer diagnosis	71	14
Mortality reduction	8	120
Overdiagnosis	4	250

# Age dependent mortality reduction



# Coverage of new breast cancers by age groups

on the basis of new cases in Germany before the screening program started

50 – 69 years	47 %
50 – 74 years	57 %
45 – 74 years	65 %

## Significant randomized studies of screening from 40 to 50 years

Study	Breast cancer mortality	95% CI	Citation
Gothenburg	44%	32-98%	Bjurstam et al, 1997, Cancer 80:2091
Malmö	64%	45-89%	Andersson u. Janzon, 1997, JNCI Monogr: 63



# Meta-analysis

- Several meta-analyses of the randomized studies have confirmed reduced breast cancer mortality from mammography screening of women 40 to 50 years

# Nijmegen case-control study

Age at index-invitation	Cases	Referents	Odds ratio (95% CI)
	Screened (unscreened)	Screened (unscreened)	
40–49	50 (26)	596 (154)	0.50 (0.30–0.82)
50–59	69 (39)	350 (107)	0.54 (0.35–0.85)
60–69	53 (35)	107 (46)	0.65 (0.38–1.13)

## Reduction of breast cancer mortality from service screening of women from 40 to 50 years in Sweden

Age	Reference	%	95% CI
40 - 44	invited	17	0 - 30
40 - 44	screened	18	0 - 33
45 - 49	invited	32	22 - 41
45 - 49	screened	37	25 - 46

Hellquist et al, 2011

# Radiation risk of mammography

breast cancer mortality per 100000 women

age	induced	avoided
50 - 74	$1,6 \times 2 = 3,2$	1121
40 - 74	$3,7 \times 2 = 7,4$	1302
40 - 49	$2,1 \times 2 = 4,2$	181

nach: de Gelder et al, (2011) B J Cancer 104:1214

# Studies in Great Britain

- Randomized extension of lower and upper age range from 50 - 70 to 47 - 73 years
- Pilot study with 60000 women successfully closed
- Recruitment will be finished in 2016

Moser et al (2011) J Med Screen 18:96

# Summary and Conclusion

- Recent research has confirmed the mortality reducing effect of breast cancer screening with mammography
- Overdiagnosis with following overtreatment is the most important hazard, but is in an acceptable range if analyzed properly

# Summary and Outlook

- Recent results have confirmed the influence of mammography screening on breast cancer mortality in women between 40 and 50 years of age
- The relative mortality reduction seems to be similar to age 50 and 70 years
- A lowering of the entry age of organized screening programs should be considered