## Milk intake and risk of mortality and fractures in women and men: cohort studies

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## Abstract

Objective To examine whether high milk consumption is associated with mortality and fractures in women and men.

Design Cohort studies.

Setting Three counties in central Sweden.

Participants Two large Swedish cohorts, one with 61 433 women (39-74 years at baseline 1987-90) and one with 45 339 men (45-79 years at baseline 1997), were administered food frequency questionnaires. The women responded to a second food frequency questionnaire in 1997.

Main outcome measure Multivariable survival models were applied to determine the association between milk consumption and time to mortality or fracture.

Results During a mean follow-up of 20.1 years, 15 541 women died and 17 252 had a fracture, of whom 4259 had a hip fracture. In the male cohort with a mean follow-up of 11.2 years, 10 112 men died and 5066 had a fracture, with 1166 hip fracture cases. In women the adjusted mortality hazard ratio for three or more glasses of milk a day compared with less than one glass a day was 1.93 (95% confidence interval 1.80 to 2.06). For every glass of milk, the adjusted hazard ratio of all cause mortality was 1.15 (1.13 to 1.17) in women and 1.03 (1.01 to 1.04) in men. For every glass of milk in women no reduction was observed in fracture risk with higher milk consumption for any fracture (1.02, 1.00 to 1.04) or for hip fracture (1.09, 1.05 to 1.13). The corresponding adjusted hazard ratios in men were 1.01 (0.99 to 1.03) and 1.03 (0.99 to 1.07). In subsamples of two additional cohorts, one in males and one in females, a positive association was seen between milk intake and both urine 8-iso-PGF2a (a biomarker of oxidative stress) and serum interleukin 6 (a main inflammatory biomarker).

Conclusions High milk intake was associated with higher mortality in one cohort of women and in another cohort of men, and with higher fracture incidence in women. Given the observational study designs with the inherent possibility of residual confounding and reverse causation phenomena, a cautious interpretation of the results is recommended.

Table 2

Milk consumption and time to death and fracture in Swedish Mammography Cohort\* and Cohort of Swedish Mer Categories of daily milk intake ⟨1 glass (⟨200 g/d) 1-2 glasses (200-399 g/d) 2-3 glasses (400-599 g/d) ≥3 glasses (≥600 g/d) Swedish Mammography Cohort Total mortality: 5422 No of deaths 229648 80961 1 231 818 Person years of follow-up 444 724 476 485 Rate/1000 person years† .20 (2.06 to 2.35) 1.18 (1.16 to 1.20) .00 (reference) 1.83 (1.75 to 1.91) 1.30 (1.25 to 1.35) Age adjusted HR (95% C 1.00 (reference) 1.21 (1.16 to 1.25) 1.60 (1.53 to 1.68) 1.93 (1.80 to 2.06) Multivariable adjusted HR (95% C) Cardiovascular mortality: No of deaths 80 961 Person years of follow-up 444 724 229 648 476 485 Rate/1000 person years\* 2.19 (1.96 to 2.46) 1.20 (1.16 to 1.23) 1.26 (1.18 to 1.34) 1.85 (1.71 to 1.99) 1.00 (reference) Age adjusted HR (95% CI) 1.59 (1.47 to 1.73) Multivariable adjusted HR (95 1.90 (1.69 to 2.14) 1.15 (1.12 to 1.19) Cancer mortality: No of deaths 80 961 231 818 Person years of follow-up 476 485 444 724 229 648 Rate/1000 person years† 1,24 (1.12 to 1.37) 1,16 (1.04 to 1.29) 1.55 (1.33 to 1.80) 1.09 (1.05 to 1.13) 1.00 (reference) .11 (1.02 to 1.19) Age adjusted HR (95% CI) Multivariable adjusted HR (95 1.00 (reference .07 (0.99 to 1.15 1.44 (1.23 to 1.69) 11.07 (1.02 to 1.11 Any fracture: No of fractures Person years of follow-up Rate/1000 person years† 210 610 1.00 (reference 16 (1.08 to 1.24) 1.02 (1.00 to 1.00 Age adjusted HR (95% 1.16 (1.11 to 1.21) 1.07 (1.04 to 1.11) 1.00 (reference) 1.16 (1.08 to 1.25) 1,02 (1,00 to 1,04 Multivariable adjusted HR (95% CI Hip fracture: No of fractures Person years of follow-up 225 969 Rate/1000 person years† .69 (1.55 to 1.85 1.76 (1.54 to 2.02) 1.00 (reference) 1.24 (1.16 to 1.33) 1.11 (1.08 to 1.15 Age adjusted HR (95% CI) ‡ 1.00 (reference) 1.55 (1.41 to 1.69 1.60 (1.39 to 1.84) Multivariable adjusted HR (95% C 1.19 (1.11 to 1.28) 1.09 (1.05 to 1.1 Cohort of Swedish Men Total mortality:

Person years of follow-up	221 381	127 248		103 049		82 415	534094
Rate/1000 person years†	18.2		18.3		19.6		18.9
Age adjusted HR (95% CI)		1.00 (0.95 to 1.05)		1.07 (1.02 to 1.13)		1.13 (1.07 to 1.19)	1.03 (1.02 to 1.05)
Multivariable adjusted HR (95% CI)‡	1.00 (reference)	0.99 (0.94 to 1.05)		1.05 (1.00 to 1.11)		1.10 (1.03 to 1.17)	1.03 (1.01 to 1.04)
Cardiovascular mortality:							
No of deaths	1468		1161		1098	841	4568
Person years of follow-up	221 381	127 248			103049		534 094
Rate/1000 person years†	7.9		8.4		9	9,6	8.6
Age adjusted HR (95% CI)	1.00 (reference)	1.06 (0.98 to 1.15)		1.14 (1.05 to 1.23)		1.21 (1.11 to 1.32)	1.05 (1.03 to 1.07)
Multivariable adjusted HR (95% CI)‡	1.00 (reference)	1,04 (0.96 to 1.12)		1.10 (1.01 to 1.19)		1.16 (1.06 to 1.27)	1.05 (1.03 to 1.07)
Cancer mortality:							
No of deaths	1077		704		616	484	2881
Person years of follow-up	221 381	127 248		103 049		82 415	534 094
Rate/1000 person years†	5.5		5.2		5.4	5.6	
Age adjusted HR (95% CI)		0.95 (0.87 to 1.05)		0.97 (0.88 to 1.07)		1.03 (0.92 to 1.14)	1.00 (0.98 to 1.03)
Multivariable adjusted HR (95% CI)‡	1.00 (reference)	0.97 (0.88 to 1.07)		0.97 (0.87 to 1.07)		1.01 (0.90 to 1.13)	0.99 (0.97 to 1.02)
Any fracture:							
No of fractures	2095		1326		1095		
Person years of follow-up	211 554	121 162		98 290		78 <b>35</b> 5	509 361
Rate/1000 person years†	10.5		10.7		10.5		
Age adjusted HR (95% CI)	1.00 (reference)	1.02 (0.95 to 1.10)		1.01 (0.94 to 1.09)		1.03 (0.96 to 1.12)	1.01 (1.00 to 1.03)
Multivariable adjusted HR (95% CI)‡	1.00 (reference)	1.02 (0.96 to 1.10)		1,01 (0.93 to 1.08)		1.03 (0.94 to 1.11)	1.01 (0.99 to 1.03)
Hip fracture:							
No of fractures	439		309		318		
Person years of follow-up	219 925	126 241		102 126		81 755	530 047
Rate/1000 person yearsf	2,4		2.3		2.7	2.3	
Age adjusted HR (95% CI)		0.95 (0.82 to 1.10)		1.12 (0.96 to 1.29)		0.97 (0.82 to 1.15)	1.02 (0.98 to 1.06)
Multivariable adjusted HR (95% CI)‡	1.00 (reference)	0.95 (0.82 to 1.11)		1.13 (0.97 to 1.31)		1.01 (0.85 to 1.20)	1,03 (0.99 to 1.07)

HR=hazard ratio

\*Had access to repeat exposure information; exposures and covariates were treated as cumulative averages