



Occupational variation in the risk of female breast cancer in the Nordic countries

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Abstract

Objective This study aimed to determine occupational variations in the incidence of breast cancer in the population-based cohort of Nordic Occupational Cancer Study (NOCCA).

Methods The study included long-term follow-up data from almost 7.5 million Nordic women. Participants were assigned to one of the 54 occupational categories based on census records at the ages of 30–64 years. Sixty-two thousand cases of breast cancer were identified through record linkages between nationwide cancer registries in Finland, Sweden, Norway, Denmark, and Iceland, followed up between 1961 and 2005. Country-specific standardized incidence ratios (SIRs) with 95% confidence intervals were estimated.

Results Overall, the highest risk elevations were seen among military personnel (SIR 1.58, 95% CI 1.03–2.32), dentists (SIR 1.43, 95% CI 1.31–1.56), and physicians (SIR 1.35, 95% CI 1.26–1.46). The lowest risks were observed among gardeners (SIR 0.76, 95% CI 0.74–0.78), farmers (SIR 0.80, 95% CI 0.78–0.82), and woodworkers (SIR 0.75, 95% CI 0.70–0.81). Welders, tobacco workers, and painters had higher SIRs for breast cancer diagnosed at age < 50. A reduced risk was observed among forestry workers, welders, and fishery workers for breast cancers diagnosed both before and after age 50. The SIRs for breast cancer did not vary substantially by histology. A significantly increased risk of breast cancer was observed among laboratory workers in the latest calendar period (1991–2005) compared with earlier periods (1976–1990 and 1961–1975). Occupations such as farming, forestry, driving, and gardening had low SIRs during all periods.

Conclusions The study suggests that the risk of breast cancer varies by occupation. Heterogeneity is also observed in some occupational categories according to age (before or after 50), histology, and calendar period.

Keywords Breast cancer · Nordic · Occupational exposure · Risk factors

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Introduction

Breast cancer is the most frequent cancer diagnosis among women globally, and the second largest cause of death among women in developed countries [1]. In the Nordic countries, the age-specific incidence of breast cancer has more than doubled during the last 65 years [2].

The most consistently reported risk factors for breast cancer are reproductive and hormonal factors including exogenous and endogenous hormones, genetic factors, lifestyle factors including alcohol consumption and obesity, exposure to ionizing radiation, and exposure to some chemicals [3–5]. However, the strength and direction of the association between these risk factors varies depending on the age at diagnosis, owing to differences in hormonal levels by menopausal status [6, 7]. Also, the association between endogenous and exogenous hormones and the risk of breast cancer varies by histology: lobular breast cancer tends to be more sensitive to endogenous and exogenous hormones than ductal breast cancer [8, 9]. A large proportion of breast cancers cannot be explained by established risk factors [10]. Recently, several studies have reported on potential environmental and occupational exposures as potential risk factors for breast cancer, although these associations are controversial [11–13]. The strongest suggestion in relation to occupational and environmental risk factors is seen for night shift work [14, 15] and for exposure to ionizing radiation, as well as suggestive evidence related to endocrine-disrupting chemicals in the textile, rubber, and plastics industries [16, 17]. Some of the effects of modern working life, such as increasing sedentary work or the demand for longer education and career planning, may lead women to postpone childbirth and thus may influence their breast cancer risk. Migration studies also suggest the influence of women's environmental exposures on the risk of breast cancer [18].

There is a significant gap in the understanding of work-related exposures and breast cancer risk. To our knowledge, most published studies are based on small populations, and only a few have taken age at diagnosis or histology into account [19, 20]. Globally, the workforce consists of increasing numbers of women, and it is crucial to identify the risk factors for breast cancer in the work environment. In this article, we present results from a large cohort study with 45 years of follow-up, aiming to describe occupational variations in breast cancer incidence in the Nordic countries. This might aid the search for specific risk factors in this insufficiently explored area.

Materials and methods

The NOCCA study

The Nordic Occupational Cancer (NOCCA) Study [21] is a large population-based cohort study consisting of

14.9 million individuals (7,454,847 women) from all five Nordic countries (Denmark, Finland, Iceland, Norway, and Sweden) who participated in one or more population censuses in 1960, 1970, 1980/1981, or 1990. The unique personal identity codes assigned to all residents in Nordic countries were used to link data from the census to nationwide registers of cancer, death, and emigration. Personal identity codes have been systematically assigned to individuals in Finland since 1967, Sweden since 1947, Denmark since 1968, Iceland since 1953, and Norway since 1964. The NOCCA Study was approved by relevant permission authorities according to the rules of each participating country.

The census questionnaire included questions related to individuals' economic activity, occupation, and industry. In Finland, Norway, and Sweden, occupations were coded according to national adaptations of the Nordic Occupational Classification, a Nordic version of the International Standard Classification of Occupations (ISCO). In Iceland, occupations were coded according to a national adaptation of ISCO. In Denmark, occupations were coded according to a special national nomenclature, with distinctions among self-employed persons, family workers, salaried employees, skilled workers, and unskilled workers. The basis of the coding of occupations was free-text information, provided by individuals at the time of the census, on education, occupation, industry, and employer's name and address. For the present study, the original national occupation codes were converted into a common classification with 54 occupational categories, one of which was economically inactive persons.

Statistical analysis

The individuals in the NOCCA Study were followed up from 1 January of the year following the first available census, provided that on that day the individual was 30–64 years of age. The follow-up ended at emigration, at death, or on 31 December 2003 (Norway and Denmark), 2004 (Iceland), or 2005 (Finland and Sweden), whichever came first. The information on emigration dates and death dates was obtained from the population registries. Information on breast cancer diagnoses from the Nordic cancer registries was grouped into ductal and lobular subgroups based on national topography and morphology coding systems.

The standardized incidence ratio (SIR) was used to estimate the relative risk of breast cancer incidence for each occupational category, the reference rates being the breast cancer incidence rates for the entire national female study populations. The SIR was counted as the ratio of observed and expected numbers of cases. For each country, the observed numbers of cancer cases and person years were stratified into five-year age categories and five-year calendar periods. The expected number of cases for each

country was estimated by multiplying the incidence rates by the respective numbers of person years at risk in the NOCCA Study by country. Where there were fewer than 100 observed cases, the exact 95% confidence interval (CI) for the SIR was defined assuming a Poisson distribution of observed cancer. When the number of observed cases was ≥ 100 , the CI was calculated based on the normal approximation to Poisson distribution. The aggregate SIR for all Nordic countries was calculated as the ratio of the total number of observed cases to the total number of expected cases in the five countries. The analyses were performed using Stata 14.

We performed analysis stratified by age group at diagnosis (< 50 and 50+ years) as a surrogate variable for pre- and postmenopausal breast cancers, given that the large majority of women in the Nordic countries below the age of 50 years are premenopausal [22, 23]. Stratified analysis was also performed for breast cancer by histological subtype (ductal and lobular breast cancer) and calendar periods (1961–1975, 1976–1990, and 1991–2005). The ratio of the SIRs of ductal and lobular breast cancer (SIR_{DC}/SIR_{LC}) and the 95% CIs were estimated using R statistical software [24].

Results

Altogether 373,361 cases of breast cancer were reported during the follow-up between 1961 and 2005. The SIRs for breast cancer are presented overall and based on age at diagnosis (< 50 years and 50+ years), histology (ductal and lobular), and calendar periods (1961–1975, 1976–1990, and 1991–2005).

Table 1 presents the SIRs for all types of breast cancer combined, for subtypes of ductal and lobular breast cancer, and the ratio of SIRs of ductal and lobular breast cancer subtypes for different occupational categories. Overall, the highest SIR was observed for military personnel (SIR 1.58, 95% CI 1.03–2.32), followed by dentists (SIR 1.43, 95% CI 1.31–1.56), and physicians (SIR 1.35, 95% CI 1.26–1.46). The lowest SIRs of breast cancer were seen among gardeners (SIR 0.76, 95% CI 0.74–0.78), farmers (SIR 0.80, 95% CI 0.78–0.82), woodworkers (SIR 0.75, 95% CI 0.70–0.81), and building caretakers (SIR 0.86, 95% CI 0.84–0.87).

A similar pattern of elevated SIRs for both lobular and ductal carcinomas was observed among journalists, nurses, dentists, physicians, and administrators. Engine operators and woodworkers had the lowest risk of lobular breast cancer, with SIRs of 0.43 (95% CI 0.24–0.72) and 0.53 (95% CI 0.39–0.71), respectively. For ductal breast cancer, gardeners had the lowest risk (SIR 0.74, 95% CI 0.71–0.77), followed by farmers and woodworkers. Significantly increased SIR_{DC}/SIR_{LC} was observed among transport workers, woodworkers, building caretakers, and engine operators. The highest

SIR_{DC}/SIR_{LC} ratio was observed among transport workers (SIR 1.49, 95% CI 1.00–2.30). Decreased SIR_{DC}/SIR_{LC} was observed among occupational groups such as journalists, physicians, administrators, teachers, and nurses. The country-specific SIRs and 95% CIs for different occupations are shown in Table 2. For the occupations with increased and decreased overall SIRs, country-specific SIRs across the five Nordic countries were mostly consistent.

The risk of breast cancer was significantly increased in women aged < 50 or 50+ years among highly educated occupational groups such as dentists, physicians, journalists, administrators, and technical workers. Welders, tobacco workers, and painters showed higher SIRs for women aged < 50. A reduced risk of breast cancer was observed for forestry workers, welders, and fishery workers for both age groups (Fig. 1).

The SIRs of breast cancer for different occupational categories were also calculated for three 15-year time periods (Figs. 2, 3). Journalists had the highest risk of breast cancer in 1961–1975 (SIR 1.76, 95% CI 1.29–2.34), falling to 1.28 (95% CI 1.15–1.42) in 1991–2005. Significantly, an increased risk of breast cancer was observed for laboratory workers only in the latest period (1991–2005), while there was no excess in the periods before that (1976–1990 and 1961–1975). Farmers, forestry workers, drivers, and gardeners had low risks during all time periods.

Discussion

The present study explored the association between breast cancer and occupation among women in the Nordic countries. The occupational groups with an overall increased risk of breast cancer were healthcare professionals, religious workers, artistic workers, journalists, women in administrative and clerical work, sales agents, transport workers, print workers, military personnel, postal workers, hairdressers, and public safety workers. The SIR for lobular cancer was comparatively higher than for ductal cancer in most of the occupations that showed an increased risk, while some occupational groups such as domestic assistants, building caretakers, chimney sweeps, and hairdressers had a significant elevation for lobular cancer only. Most of the SIR_{DC}/SIR_{LC} was insignificant, showing that the risk of breast cancer in those occupations did not differ strongly by histology. Increased SIR_{DC}/SIR_{LC} was observed among transport workers, woodworkers, building caretakers, and engine operators, while a decreased SIR_{DC}/SIR_{LC} was observed among groups such as journalists, physicians, administrators, teachers, and nurses.

A wide range of white-collar jobs in our study—including administrative work, clerical work, teaching, health work, and technical work—were associated with an

Table 1 Observed number (Obs), standardized incidence ratio (SIR) with 95% confidence intervals (95% CI) for all breast in selected occupational categories in five Nordic countries in the period 1961–2005, by histology and ratio of SIR of ductal and lobular breast cancer (SIR_{DC}/SIR_{LC}) with its 95% CI

Abbreviations	Occupational categories			All cancer			Ductal cancer			Lobular cancer			SIR _{DC} /SIR _{LC}		
	Obs	SIR	95% CI	Obs	SIR	95% CI	Obs	SIR	95% CI	Obs	SIR	95% CI	Ratio	95% CI	
Mi	26	1.58	1.03–2.32	13	1.41	0.75–2.42	1	0.59	0.01–3.30	2.39	0.36–102				
De	538	1.43	1.31–1.56	233	1.39	1.22–1.58	48	1.66	1.22–2.20	0.84	0.61–1.72				
Jo	550	1.36	1.24–1.47	271	1.30	1.15–1.47	67	1.79	1.39–2.28	0.72	0.56–0.97				
Ph	702	1.35	1.26–1.46	316	1.25	1.11–1.39	79	1.71	1.35–2.13	0.73	0.57–0.94				
Ad	3,164	1.29	1.24–1.33	1,297	1.28	1.21–1.35	256	1.51	1.33–1.71	0.84	0.74–0.97				
Ar	1,119	1.26	1.18–1.33	539	1.27	1.17–1.39	106	1.45	1.18–1.75	0.88	0.71–1.10				
TeW	2,312	1.24	1.19–1.29	1,096	1.25	1.18–1.33	208	1.27	1.1–1.450	0.99	0.85–1.15				
Te	15,311	1.23	1.20–1.25	6,731	1.22	1.19–1.25	1,321	1.36	1.29–1.44	0.89	0.84–0.95				
La	1,188	1.20	1.13–1.27	606	1.19	1.10–1.29	102	1.14	0.93–1.39	1.04	0.84–1.30				
Cl	43,043	1.20	1.18–1.21	18,380	1.18	1.16–1.20	3,280	1.23	1.19–1.27	0.96	0.93–1.00				
Re	5,560	1.19	1.16–1.22	2,974	1.17	1.13–1.22	571	1.23	1.13–1.33	0.95	0.87–1.04				
Nu	7,679	1.18	1.15–1.21	3,392	1.19	1.15–1.23	654	1.32	1.22–1.43	0.90	0.83–0.98				
Tr	398	1.15	1.04–1.27	243	1.31	1.15–1.49	27	0.88	0.58–1.28	1.49	1.00–2.30				
OHW	5,248	1.15	1.12–1.18	2,526	1.14	1.09–1.18	448	1.22	1.11–1.33	0.94	0.85–1.04				
Pr	1,208	1.15	1.08–1.21	513	1.10	1.01–1.20	82	1.06	0.84–1.32	1.04	0.82–1.33				
Pu	317	1.11	0.99–1.24	166	1.08	0.92–1.25	30	1.20	0.81–1.71	0.90	0.61–1.37				
Sa	4,152	1.10	1.06–1.13	1,987	1.11	1.07–1.17	336	1.17	1.04–1.30	0.96	0.85–1.08				
Po	5,191	1.08	1.05–1.11	2,328	1.05	1.01–1.09	399	1.11	1.01–1.23	0.94	0.85–1.05				
Ha	1,983	1.06	1.01–1.10	804	1.05	0.98–1.13	143	1.09	0.92–1.28	0.96	0.81–1.16				
To	222	1.01	0.88–1.15	52	1.22	0.91–1.60	7	1.15	0.46–2.37	1.06	0.48–2.77				
ShL	840	1.01	0.94–1.08	342	0.98	0.88–1.09	50	0.90	0.67–1.18	1.10	0.81–1.50				
Sho	20,177	1.01	0.99–1.02	7,712	1.00	0.97–1.02	1,102	0.91	0.85–0.96	1.10	1.03–1.17				
Wa	4,071	0.99	0.96–1.02	1,767	0.99	0.94–1.04	245	0.88	0.77–0.99	1.13	0.99–1.30				
Tex	9,336	0.99	0.97–1.01	3,498	1.01	0.97–1.04	499	0.92	0.84–1.00	1.10	1.00–1.20				
Be	195	0.99	0.85–1.14	40	0.91	0.65–1.23	10	1.35	0.65–2.49	0.67	0.33–1.50				
Co	3,693	0.98	0.94–1.01	1,671	0.95	0.91–1.00	252	0.95	0.83–1.07	1.00	0.88–1.16				
AsN	8,058	0.95	0.93–0.97	3,467	0.95	0.92–0.98	610	0.92	0.85–1.00	1.03	0.95–1.13				
OW	6,816	0.95	0.93–0.97	2,910	0.93	0.89–0.96	449	0.99	0.90–1.09	0.94	0.85–1.04				
Pac	2,872	0.95	0.92–0.99	1,428	0.97	0.92–1.02	211	0.90	0.78–1.03	1.07	0.93–1.24				
El	1,343	0.94	0.89–0.99	518	1.00	0.92–1.09	72	0.78	0.61–0.99	1.28	1.00–1.66				
Min	24	0.93	0.60–1.39	18	1.38	0.82–2.18	2	0.90	0.11–3.23	1.54	0.37–13.66				

Table 1 (continued)

Abbreviations	Occupational categories	All cancer			Ductal cancer			Lobular cancer			SIR _{DC} /SIR _{LC}	
		Obs	SIR	95% CI	Obs	SIR	95% CI	Obs	SIR	95% CI	Ratio	95% CI
Pa	Painters	132	0.93	0.78–1.10	84	1.07	0.85–1.33	12	0.91	0.47–1.59	1.18	0.64–2.37
Do	Domestic assistants	9,907	0.93	0.91–0.95	3,881	0.92	0.89–0.95	544	0.80	0.74–0.87	1.15	1.05–1.26
Me	Mechanics	1714	0.92	0.88–0.97	631	0.91	0.84–0.99	105	0.82	0.67–1.00	1.11	0.90–1.38
Gl	Glassmakers, etc	1,805	0.90	0.86–0.95	742	0.89	0.83–0.96	109	0.78	0.64–0.94	1.14	0.93–1.41
La	Launderers	1,754	0.89	0.85–0.94	523	0.94	0.86–1.03	57	0.69	0.52–0.89	1.37	1.04–1.83
Ch	Chemical process workers	725	0.89	0.83–0.96	259	0.94	0.83–1.06	36	0.81	0.57–1.12	1.16	0.81–1.69
Oc	“Other construction workers”	200	0.87	0.76–1.00	136	0.89	0.75–1.06	16	0.70	0.40–1.14	1.27	0.76–2.28
Bu	Building caretakers	14,938	0.86	0.84–0.87	5,825	0.85	0.83–0.87	771	0.76	0.70–0.81	1.13	1.05–1.22
Foo	Food workers	3,076	0.85	0.82–0.89	1,177	0.86	0.81–0.91	146	0.76	0.64–0.89	1.13	0.95–1.35
Sm	Smelting workers	189	0.84	0.72–0.97	89	0.90	0.72–1.11	15	0.84	0.47–1.39	1.07	0.61–1.99
En	Engine operators	255	0.83	0.73–0.94	153	0.84	0.71–0.98	14	0.43	0.24–0.72	1.94	1.12–3.63
Dr	Drivers	567	0.81	0.74–0.88	228	0.80	0.70–0.91	45	0.87	0.63–1.16	0.92	0.67–1.30
Far	Farmers	6,445	0.80	0.78–0.82	2,618	0.80	0.77–0.84	317	0.78	0.70–0.87	1.03	0.92–1.16
Ga	Gardeners	7,300	0.76	0.74–0.78	3,984	0.74	0.72–0.77	549	0.68	0.63–0.74	1.09	1.00–1.19
For	Forestry workers	85	0.75	0.60–0.93	35	0.62	0.44–0.87	5	0.56	0.18–1.30	1.12	0.44–3.66
We	Welders	74	0.75	0.59–0.95	36	0.65	0.46–0.90	12	1.21	0.62–2.11	0.54	0.28–1.14
Wo	Wood workers	725	0.75	0.70–0.81	424	0.80	0.73–0.88	46	0.53	0.39–0.71	1.51	1.11–2.09
Fi	Fishermen	44	0.69	0.50–0.93	21	0.65	0.40–1.00	4	0.94	0.25–2.40	0.70	0.24–2.80
All	All categories	373,153	1.00	0.99–1.01	134,905	1.00	0.99–1.01	21,496	1.00	0.99–1.01	1.00	0.99–1.00

Occupations with < 10 observed cases in total were excluded

Table 2 Observed number (Obs) and standardized incidence ratio (SIR) with 95% confidence intervals (CI) for all breast cancer in selected occupational categories in five Nordic countries in the period 1961–2005, by country

Occupational categories	Denmark			Finland			Iceland			Norway			Sweden		
	Obs	SIR	95% CI	Obs	SIR	95% CI	Obs	SIR	95% CI	Obs	SIR	95% CI	Obs	SIR	95% CI
Military personnel	5	2.14	0.70–5.01	10	1.52	0.73–2.80	0	–	0–0	3	2.00	0.41–5.84	8	1.32	0.57–2.60
Dentists	102	1.78	1.45–2.16	184	1.57	1.35–1.82	1	1.31	0.03–7.27	53	1.26	0.95–1.65	198	1.24	1.08–1.43
Journalists	34	1.45	1.01–2.03	181	1.29	1.11–1.49	5	1.63	0.53–3.81	53	1.36	1.02–1.78	277	1.38	1.23–1.56
Physicians	108	1.64	1.35–1.99	211	1.41	1.23–1.62	1	0.57	0.01–3.20	66	1.23	0.95–1.56	316	1.28	1.14–1.43
Administrators	781	1.37	1.28–1.47	689	1.40	1.30–1.51	8	1.05	0.45–2.07	563	1.26	1.16–1.37	1,123	1.19	1.12–1.26
Artistic workers	128	1.34	1.12–1.60	300	1.30	1.16–1.46	10	0.90	0.43–1.66	166	1.22	1.04–1.42	515	1.24	1.13–1.35
Technical workers, etc	247	1.22	1.08–1.39	771	1.32	1.23–1.41	8	0.94	0.41–1.85	156	1.36	1.15–1.59	1,130	1.18	1.11–1.25
Teachers	1,967	1.28	1.23–1.34	3,467	1.35	1.31–1.40	109	1.16	0.95–1.40	2,231	1.16	1.11–1.21	7,537	1.18	1.15–1.21
Laboratory assistants	191	1.17	1.01–1.35	358	1.18	1.06–1.31	15	1.25	0.70–2.06	167	1.29	1.10–1.50	457	1.20	1.09–1.31
Clerical workers	7,651	1.25	1.22–1.28	10,490	1.20	1.18–1.23	343	1.24	1.11–1.38	6,903	1.21	1.18–1.24	17,656	1.17	1.15–1.19
Religious workers etc	230	1.16	1.02–1.32	1,918	1.19	1.14–1.25	20	1.47	0.90–2.26	570	1.21	1.11–1.32	2,822	1.18	1.14–1.23
Nurses	1,321	1.15	1.09–1.22	2,024	1.31	1.25–1.37	60	1.17	0.89–1.51	1,153	1.13	1.07–1.20	3,121	1.14	1.10–1.18
Transport workers	2	0.51	0.06–1.83	217	1.30	1.14–1.49	6	1.45	0.53–3.17	34	1.22	0.84–1.70	139	0.96	0.81–1.14
“Other health workers”	835	1.29	1.20–1.38	1,367	1.18	1.11–1.24	19	1.21	0.73–1.89	1,008	1.05	0.98–1.12	2,019	1.13	1.08–1.18
Printers	156	1.20	1.02–1.40	386	1.13	1.02–1.24	6	0.74	0.27–1.61	192	1.37	1.18–1.58	468	1.08	0.99–1.19
Sales agents	63	1.23	0.94–1.57	1,171	1.07	1.01–1.13	32	1.21	0.83–1.71	813	1.16	1.08–1.24	2,073	1.09	1.04–1.13
Postal workers	207	1.03	0.89–1.18	1,204	1.05	0.99–1.11	44	0.84	0.61–1.12	1,170	1.11	1.04–1.17	2,566	1.09	1.05–1.13
Domestic assistants	1,313	0.90	0.85–0.95	1,346	0.92	0.87–0.97	19	1.09	0.66–1.71	2,003	0.93	0.89–0.98	5,226	0.93	0.91–0.96
Mechanics	213	0.83	0.72–0.94	306	0.85	0.75–0.95	2	1.87	0.23–6.77	152	1.05	0.89–1.23	1,041	0.95	0.90–1.01
Glassmakers, etc	206	0.82	0.71–0.94	510	0.84	0.77–0.92	4	0.74	0.20–1.88	260	0.99	0.87–1.12	825	0.94	0.88–1.01
Launderers	423	0.90	0.82–0.99	250	0.88	0.77–0.99	16	0.84	0.48–1.37	257	0.79	0.70–0.89	808	0.93	0.87–1.01
Chemical process workers	186	0.93	0.80–1.07	146	0.90	0.76–1.06	2	0.62	0.07–2.24	108	0.88	0.73–1.07	283	0.88	0.78–0.98
Building caretakers	3,678	0.87	0.84–0.90	3,291	0.83	0.80–0.86	102	0.76	0.62–0.92	3,404	0.86	0.83–0.89	4,463	0.87	0.84–0.90
Food workers	799	0.82	0.76–0.88	689	0.85	0.78–0.91	164	0.84	0.72–0.98	628	0.85	0.79–0.92	796	0.90	0.84–0.97
Smelting workers	17	0.68	0.40–1.09	55	0.77	0.58–1.00	1	4.23	0.11–23.5	16	1.03	0.59–1.67	100	0.88	0.72–1.07
Engine operators	4	1.40	0.38–3.57	157	0.86	0.73–1.00	0	0.00	0.0–13.02	11	1.01	0.50–1.80	83	0.75	0.60–0.93
Drivers	77	0.75	0.59–0.94	74	0.69	0.55–0.87	1	0.43	0.01–2.42	84	0.88	0.70–1.09	331	0.84	0.75–0.93
Farmers	1,993	0.75	0.71–0.78	1,489	0.79	0.75–0.83	90	0.82	0.66–1.01	2,176	0.82	0.79–0.86	697	0.89	0.82–0.96
Gardeners	283	0.83	0.73–0.93	4,230	0.74	0.72–0.76	4	1.07	0.29–2.74	1,184	0.75	0.71–0.80	1,599	0.83	0.79–0.87
Forestry workers	4	1.09	0.30–2.79	22	0.73	0.46–1.11	0	0.00	0.00–6.23	19	0.62	0.37–0.97	40	0.84	0.60–1.14
Welders	0	–	0–0	22	0.65	0.41–0.99	0	0.00	0.0–30.40	10	0.76	0.36–1.40	42	0.82	0.59–1.11
Wood workers	89	0.78	0.62–0.96	439	0.73	0.66–0.80	1	0.60	0.02–3.37	58	1.07	0.81–1.38	138	0.71	0.60–0.84
Fishermen	8	0.57	0.25–1.12	7	0.47	0.19–0.97	7	1.49	0.60–3.07	17	0.74	0.43–1.19	5	0.72	0.23–1.67

Only occupations with significantly increased and decreased SIR for all countries combined are shown

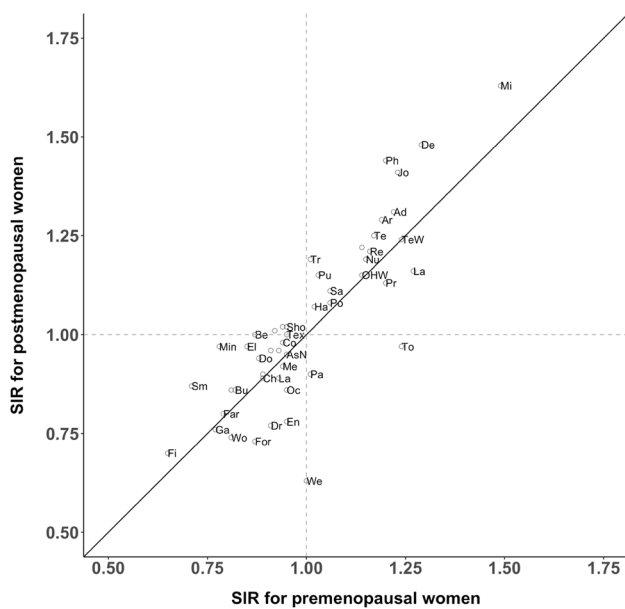


Fig. 1 Correlation of premenopausal and postmenopausal breast cancer SIRs for different occupational categories. Occupations with < 10 cases in total were excluded. Abbreviations for the occupations are explained in Table 1

increased risk of breast cancer. Our results are consistent with other findings from Nordic [25] and other countries [20, 26, 27]. It is difficult to interpret the association of these occupations with an increased risk of breast cancer, since we do not have information on the specific exposures related to each occupation. However, one possible link

for the high SIRs observed in these occupations, which require higher levels of education, might be such women’s postponement of first childbirth and their birthing of fewer children in their lifetime. Early age at first childbirth and an increased number of children are established protective factors in relation to breast cancer [28–30]. These factors could not be adjusted for in our study; however, one large Swedish cohort study showed that, even after adjustment for an extensive set of reproductive factors including breastfeeding, hormonal replacement therapy, and lifestyle factors, the risk for white-collar workers was still high [25]. This and other studies suggest that estrogen-related pathways explain only a small proportion of the elevated breast cancer risk among professional women, and social stress pathways might potentially be more important [31, 32]. Chronic stress at work results in systematically elevated levels of cortisol [31]. Normal levels of cortisol contribute to the lobuloalveolar development of mammary glands, and also aid the process of lactation [32]. However, a prolonged elevated level of cortisol production is likely to expose the mammary cells to an adverse environment, as well as contributing to breast tumorigenesis by altering the generation or activity of estrogen [33, 34].

A recent study [32] suggested that working women might have experienced more interpersonal stress in the 1970s compared with recent years due to societal norms. We also observed differences in risk among white-collar occupations in different study calendar periods, with lower risks in the most recent calendar periods compared with earlier years. It is conceivable that white-collar workers might be subjected

Fig. 2 Breast cancer standardized incidence ratio (SIR), by occupational categories with increased overall risk and time period (1961 through 2005) among women in five Nordic countries. *p* value of linear trend for each occupational category is shown

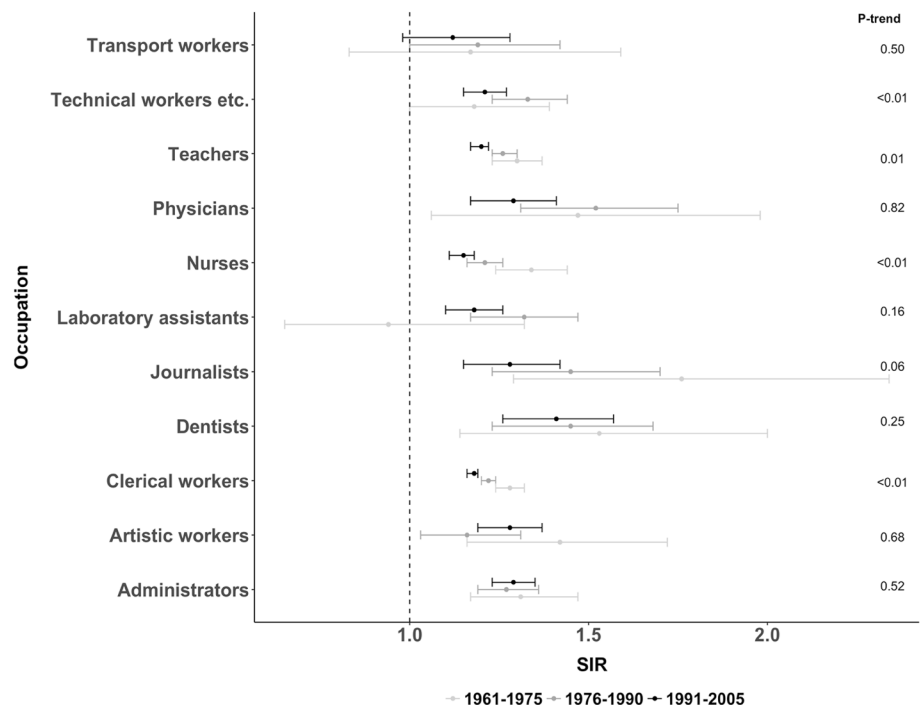




Fig. 3 Breast cancer standardized incidence ratio (SIR), by occupational categories with decreased overall risk and time period (1961 through 2005) among women in 5 Nordic countries. *p* value of linear trend for each occupational category is shown

to a higher level of stress due to higher work demands, and might thus be at higher risk of breast cancer. However, the few studies conducted on the association between work-related stress and breast cancer have been limited to the nursing profession only, and have shown no indication of any such risk association [35, 36]. We also noted that the SIR for lobular cancer compared with ductal breast cancer (SIR_{DC}/SIR_{LC}) was significantly higher among many white-collar workers, which is conceivably because lobular cancer is hormone sensitive; a consistently stronger association between age at first childbirth and exogenous hormones has been shown for lobular breast cancer compared with ductal breast cancer [37–41].

Furthermore, the increase in breast cancer risk among white-collar workers could be related to such workers' higher educational status and income. Studies show that more educated women and those with higher incomes are more likely to participate in breast cancer screening [42–45]. In the Nordic countries, the incidence of breast cancer increased markedly after the introduction of mammographic screening, and it remains elevated among women who continue to be screened compared with women who do not get screening [46, 47]. Compared with the early 1980s, there has been a steep increase in breast cancer incidence in recent years, with a peak at the age of 65 years. In addition, the incidence of breast cancer has increased in recent years among premenopausal women over the age of 40, which

might indicate self-initiated opportunistic mammographic screening in these age groups [2].

The International Agency for Research on Cancer (IARC) has classified shift work that involves circadian disruption as probably carcinogenic to humans, based on sufficient evidence in experimental animals but limited evidence in humans [13]. Exposure to light during night-time leads to the suppression of the production of the pineal hormone melatonin, thus influencing a rise in estrogen production and leading to the possibility of breast cancer development [48]. A study among Finnish women reported the decreased risk of breast cancer by degree of visual impairment which suggest towards the positive association between visible light at night and breast cancer risk via alteration in melatonin hormone [49]. Some recent systematic reviews and meta-analyses [50, 51] have suggested that shift work increases the risk of breast cancer by 48%; however, one meta-analysis has indicated insufficient evidence of, or minimal or no effect on, the risk of breast cancer [52]. A cohort study among Swedish twins suggested that night work was associated with breast cancer risk, but only after long-term exposure [53]. Similarly, a study among Finnish cabin attendants did not show any statistically significant association with sleep rhythm disruption [54]. Studies on night shift work among nurses have shown a positive association with breast cancer [55, 56]. An increased risk was also observed for military women in previous studies [57, 58]. Nurses and military workers in our study showed an increased risk of breast cancer in pre- and postmenopausal age groups and during each time period, but these findings are inconclusive in the absence of information regarding the frequency and duration of night shifts worked or adjustment for other possible risk factors.

In our study, other health professionals apart from nurses also showed an elevated risk of breast cancer, including dentists, laboratory assistants, and physicians. Health workers might be exposed to ionizing radiation, which the IARC states is carcinogenic to the human breast, with sufficient evidence and shown in other studies [59–61]. In a Finnish study that assessed the association between cumulative radiation exposure and cancer incidence among physicians by systematically monitoring radiation exposure, there was a significantly increased risk of breast cancer among exposed physicians compared with the non-exposed group [62]. Laboratory workers are likely to be exposed to carcinogenic chemicals and organic solvents that have been shown to be associated with an increased risk of breast cancer [63, 64]. In our study, the increased risk among laboratory workers in recent years might reflect advances in medical laboratory technologies and the increased exposure of these workers to ionizing radiation and other potentially carcinogenic chemical compounds.

While ionizing radiation has an established association with breast cancer, the IARC has classified exposure to non-ionizing radiation—specifically, exposure to electromagnetic fields (EMF)—as having possible but inadequate evidence for a breast cancer risk. Some studies have identified an increased risk of breast cancer associated with electrical work [65, 66] or electromagnetic radiation [67]; however, we did not observe any such association among electrical workers in any time period for the Nordic countries.

The IARC has stated that there is inadequate evidence for the carcinogenicity of printing inks in humans or of the possibility that printing processes might be carcinogenic to human beings. In our study, printing and publishing workers had an increased risk of breast cancer in both age groups (<50 years and \geq 50 years). A study by Lynge et al. [68] based on Danish data, which overlaps with the NOCCA data, suggested a significantly higher SIR of breast cancer (SIR 1.35, 95% CI 1.08–1.66) among women working in the printing and publishing industry. In agreement with our Nordic study, a case–control study of breast cancer among premenopausal women based on the British Columbia cancer registry showed an elevated risk of breast cancer among younger women working in printing [65].

Some previous studies have suggested an elevated risk of breast cancer among agricultural populations [65, 69] that might be related to exposure to pesticides during farming; for example, not wearing protective equipment during chemical use increases the risk of breast cancer. Chemical compounds such as organochlorines and dichlorodiphenyltrichloroethane, which are frequently used as pest and insect control measures, are reportedly carcinogenic to the human breast [70, 71]. Studies have recently suggested that these estrogenic environmental compounds, often referred to as ‘xenoestrogens,’ are formed by the interaction of many chemicals in the environment, and are stored in breast tissue and interfere with endogenous estrogen. Xenoestrogen has been linked to the neoplastic transformation of the human breast [72, 73]. However, in the current study’s long-term follow-up of the Nordic population, we did not find any elevated risk of breast cancer among farmers or gardeners, which is similar to findings from some other recent studies [20, 74]. On the contrary, women in farming had a deficit risk of breast cancer. With reference to a Finnish survey among working-age cancer cases, there is increased parity among agricultural occupations compared with academic occupations, which might possibly be one explanation for the protective effect observed among females in these occupations in the Nordic population [75].

Women in sedentary occupations have been reported to be at increased risk of breast cancer compared with women in non-sedentary occupations [76–78]. In this study, we could not precisely categorize or define sedentary occupations.

However, the protective effect on breast cancer among farmers, gardeners, woodworkers, and forestry workers might tend to suggest that an active lifestyle is associated with a decreased risk [79]. On the other hand, sedentary work environments, such as those of administrative workers, had an increased risk.

The strength of this study is its large study population and its long-term follow-up of individuals. Further, the high quality and completeness of the Nordic countries’ nationwide population registers, and the record linkage system via unique personal identity codes, ensures accurate follow-up measurements and accurate assessments of breast cancer cases [80].

Our study also has limitations. There were some heterogeneous occupational categories, and it was not always possible to evaluate the actual occupational risk associated with specific exposures within such categories. Information on each individual’s occupation was obtained only at the first census in which the individual participated, and might possibly have varied across the individual’s lifetime. Another major limitation was that there was a lack of data regarding potential confounding by well-known risk factors for breast cancer, and hence we could not adjust for these factors.

In conclusion, this study suggests that breast cancer risk is heterogeneously distributed among different occupations in the Nordic population. The study also suggests that breast cancer risks based on occupation differ markedly by histology in some occupational categories. It was not within the scope of our study to separate the roles of strictly occupation-related exposures from the effects of individual and lifestyle factors, but evidently there would be a need for further studies to precisely explain the variations in breast cancer risk by occupation.

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