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Cancer in the Norwegian printing industry

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Objectives The aim of this study was to investigate cancer risk among Norwegian workers in the printing industry, particularly lung and bladder cancer.

Methods Cancer incidence was investigated from 1953 through 1998 in a cohort of 10 549 male members of a trade union in the printing industry in Oslo and nearby areas. Rates from the region, were used to calculate standardized incidence ratios (SIR) separately for the skilled and unskilled workers. Smoking data from a sample of the cohort were utilized for evaluating the risk estimates of smoking-related cancers. Specific exposure data were not available.

Results Among the skilled workers, significantly elevated risks of cancer of the urinary bladder [standardized incidence ratio (SIR) 1.47, 95% confidence interval (95% CI) 1.19–1.79], liver (SIR 1.92, 95% CI 1.15–2.99), pancreas (SIR 1.46, 95% CI 1.07–1.94) and colon (SIR 1.27, 95% CI 1.05–1.55) were observed, whereas an increased risk of lung cancer in this group was confined to those born before 1910. Among the unskilled workers, there were significantly increased risks of cancer of the mouth, esophagus, stomach, larynx, lung, and all sites.

Conclusions The study showed that workers in the printing industry were at increased risk of several types of cancer. In particular the increased risk of bladder cancer among the skilled workers is suggestive of an occupational cause. However, no specific agent could be identified as an occupational carcinogen. The results did not support the hypothesis of a generally increased risk of lung cancer. The risk pattern for unskilled workers may reflect confounding by nonoccupational factors.

Key terms bladder cancer; cohort study; liver cancer; male trade-union members; occupation; social class.

Workers in the printing industry are exposed to a variety of potentially hazardous agents, including aromatic amines, inks, organic solvents, dust, noise, night work, and, previously, lead (1–7). The constituents of ink include mineral oils and pigments, in particular carbon black in black inks. Exposure circumstances in this industry are complex and heterogeneous. A Danish overview of the graphical industry reported 300 different substances, of which 26 were known or suspected carcinogens (8).

An increased risk of cancer at various sites has been reported among workers in the printing industry, but the findings are not consistent (2, 8–16). In particular an increased risk for urinary bladder cancer has been reported in several studies (11, 17–21), although no association was reported in other investigations (12, 15, 22). In addition, several studies have reported an increased risk of lung cancer (8, 17, 23–25), while other studies have failed to demonstrate any elevated risk (2, 26).

In 1996, an IARC (International Agency for Research on Cancer) working group evaluated occupational exposures in printing processes as possibly carcinogenic to humans, based on limited epidemiologic evidence for an increased risk of lung and bladder cancer (1).

The limitations of some of the available studies include small sample sizes, resulting in low statistical power, and an analysis of cancer mortality instead of cancer incidence. In the current study we investigated cancer risk in a cohort of more than 10 000 members of a union of graphical workers from Oslo, Norway. Job titles recorded in the union files enabled a reasonable subdivision of the workers. We obtained data on cancer incidence from the Cancer Registry of Norway and chose an adequate reference population that geographically corresponded to the study population.

The aim of the study was to investigate the risk of cancer, in particular the risk of lung and bladder cancer,

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among male members of the Oslo branch of the Norwegian Union of Graphical Workers.

Study population and methods

Study population

The male members of the Oslo branch of the Norwegian Union of Graphical Workers constituted our study population. The Oslo branch covered graphical establishments in the city of Oslo and, with a few exceptions, establishments in the county of Akershus. In the time period from which the study population was recruited, 95% of the workers in the printing industry in this area were members of the union (Svein Mortensen, personal communication). The study population included all workers who were union members at any time in the period January 1953 to January 1975 with the known month and year at the start of membership. The person's age at the time of entering the union had to be below 67 years. For each person, the following data were recorded: name, date of birth, country of birth, job title, month and year at the start of membership, month and year at the end of membership, any change up to 1975 in either membership status or job title, and, if relevant, the date of emigration or death. On the basis of the demographic data of 11192 subjects, personal identification numbers were added from Statistics Norway. Altogether 643 workers remained unidentified, of whom 416 either were born abroad or were registered as emigrated in the union rolls. There were no indications of emigration of the other 227 unidentified workers. The remaining 10 549 persons comprised the cohort, which has been previously studied with respect to the effect of paternal occupational exposures on offspring (27, 28).

Linkage

The cohort was linked with population and emigration registers at Statistics Norway to identify deceased and emigrated members. It was further linked with the Cancer Register to identify incident cases of cancer. Dates of emigration, death, or cancer diagnosis, as well as topographical and morphological codes of cancer, based on the seventh revision of the International Classification of Diseases (ICD-7) (29), were added. Linkage was performed using the unique personal identification number assigned to all people resident in Norway.

Follow-up

The date of the start of follow-up was 1 January 1953, the date of beginning membership, or the date of the

person's 14th birthday, whichever came last; the date of the end of follow-up was the date of death, the date of emigration, or 1 January 1999, whichever came first. Loss to follow-up occurred on emigration. Persons who were lost to follow-up contributed person-time and cases until date of emigration.

Exposure

Potentially toxic agents occurring in the printing industry include organic solvents, mineral oils, pigments, resins, lead, and paper dust. Workers in this industry have been exposed to these factors through both inhalation and dermal contact.

Common exposures in traditional letterpress techniques include lead dust and fumes, benzene, toluene, xylene, carbon tetrachloride, and ink mist (1, 7). In the 1970s, the traditional letterpress gradually disappeared from the scene and was substituted by offset techniques, while rotogravure techniques, for publication purposes, in Norway were used until the end of the century. Solvent exposures in offset establishments included white spirits, methylene chloride, isopropanol, 1,1,1-trichloroethane, and ethanol. In rotogravure establishments, toluene represented the major solvent exposure, but other solvents were also present, including trichloroethylene (3). To our knowledge, benzene, as such, has not been used in the Norwegian printing industry, but there may have been exposure to it as a contaminant of toluene.

Statistical analysis

For each cancer site, the standardized incidence ratio (SIR), the ratio between observed and expected numbers of new cancer cases, was computed along with 95% confidence intervals (95% CI). For standardized incidence ratios based on less than 100 observed cases, exact Poisson 95% confidence intervals were used (30). For standardized incidence ratios based on 100 observed cases or more, the 95% confidence intervals were based on the approximation of the standard error of the log SIR. A result was regarded as statistically significant if the 95% confidence interval did not include 1. The expected numbers were computed for 5-year observation periods as age-specific rates with 5-year age groups. The population of the city of Oslo and the county of Akershus, the area in which the members worked and lived for most of their lives, served as the reference population. The statistical package Epicure was used to compute the person-years, expected cases, and standardized incidence ratios (31).

The statistical analysis was performed separately for the skilled and unskilled workers. Four persons had held both a skilled and an unskilled job; their person-years

were assigned to both groups. Table 1 reports selected jobs for the skilled and unskilled workers with corresponding tasks. Table 2 summarizes the mean and median year of birth, the start of membership, and the duration of membership of the cohort members.

For lung and bladder cancer and for the cancer sites with significantly increased risk among the skilled workers, the additional analysis of the standardized incidence ratio stratified on attained age, year of birth, and year of first membership was performed. In an additional analysis of alcohol-related cancers, the following cancer sites were included: tongue, mouth, pharynx, esophagus, liver, larynx.

An analysis of the risk of cirrhosis was performed by computing the standardized mortality ratio (SMR), the ratio of observed to expected deaths, with data from Statistics Norway, which has information on cause-specific mortality in Norway from 1960 to 1993. The follow-up period, accordingly, for this analysis was 1960–1993.

Smoking

We obtained information on tobacco smoking for 311 cohort members who were involved in a survey on obstructive lung disease performed in Oslo in 1972 (32). We compared the age-adjusted prevalence of smoking

among the cohort members with that of the males in the entire survey population ($N=8128$). Overall the age-adjusted proportion of ever smokers was 82% (95% CI 78–86%) among the printing industry workers versus 72% in the total survey population. The proportions of ever smokers among the skilled and unskilled workers were 81.5% and 83.4%, respectively. Among the trade union members born before 1935, the proportion of ever smokers was 83% versus 81% in the total survey population, whereas among the workers born after 1935 the proportion of ever smokers was 82% versus 61% in the total survey population.

Results

During the follow-up, 318 443 person-years of observation were accumulated, 4240 persons died, and 1652 incident cancer cases were registered.

The standardized incidence ratios for selected cancer sites are presented in table 3. Among the skilled workers, there was no excess risk for all cancer sites (SIR 1.03, 95% CI 0.97–1.10), whereas among the unskilled workers there was a very slight excess (SIR 1.08, 95% CI 1.00–1.17).

The risk of bladder cancer was significantly elevated among the skilled workers on the basis of 98 observed versus 67 expected cases (SIR 1.47). Seventeen of the cases occurred before the age of 55 years (SIR 1.81, 95% CI 1.05–2.90) (table 4). Among the unskilled workers there was no excess risk.

No significant excess of lung cancer was found among the skilled workers. Among the skilled workers born before 1910, however, there was a significantly increased risk of lung cancer based on 74 cases versus 50 expected (SIR 1.48, 95% CI 1.16–1.85) (table 4). The lung cancer risk was significantly elevated among the unskilled workers on the basis of 130 observed versus 88 expected cases (SIR 1.49, 95% CI 1.25–1.76). A two-fold risk was observed among the unskilled workers born in 1935 or later on the basis of 16 cases versus 8 expected (SIR 1.99, 95% CI 1.14–3.24) (not shown in table). Furthermore, the unskilled workers, but not the skilled workers, experienced an excess of cancers of the oral cavity, esophagus, stomach, and larynx.

There was an excess risk for colon cancer among the skilled workers on the basis of 101 observed cases versus 79 expected (SIR 1.27, 95% CI 1.05–1.55). Among the unskilled workers there was a significantly reduced risk for this cancer.

Among the skilled workers, there was an excess of liver cancer on the basis of 19 observed versus 10 expected cases (SIR 1.92), and, among the unskilled workers, there was a nonsignificant excess (SIR 1.62). In the

Table 1. Selected job titles and their respective typical worktasks.

Job titles	Worktasks
Skilled workers	
Compositors	Set type by hand or with machines
Printers	Set, operate, clean, and oil printing machines, clean printing plates, check print quality
Reproduction photographers	Photograph originals for reproduction to printing plates, develop film
Etchers	Mount film for copying to metal plates, etch clichés, print tests of clichés
Bookbinders	Bind books and periodicals, operate book-binding machines, and related work
Unskilled workers	
Printer's assistants	Lay on paper, operate printing machines, clean printing plates
Assistants	Carry rolls of paper, remove paper, clean, any odd jobs
Bookbinder's assistants	Operate folding machines, other tasks in the finishing department

Table 2. Mean and median duration of membership, year of birth, first year of membership and follow-up time.

	Skilled workers			Unskilled workers		
	Median	Mean	SD	Median	Mean	SD
Duration of membership (years)	12.2	17.6	12.2	3.7	9.0	11.6
Year of birth	1934	1928	19.7	1931	1928	17.1
Start of membership (year)	1957	1951	18.4	1961	1958	12.2
Follow-up time (years)	32.6	30.6	11.8	30.4	29.3	10.9

Table 3. Observed number of selected cancer cases (O), standardized incidence ratios (SIR), and 95% confidence intervals (95% CI) for the skilled and unskilled workers.

Site ^a	Skilled (N= 6553) ^b			Unskilled (N=4 000) ^c			Total (N=10 549) ^{d, e}		
	O	SIR	95%CI	O	SIR	95% CI	O	SIR	95% CI
Lip (140)	2	0.35	0.04–1.27	2	0.58	0.07–2.11	4	0.44	0.12–1.13
Tongue (141)	9	1.52	0.69–2.88	4	1.10	0.30–2.81	13	1.36	0.72–2.32
Mouth (143–144)	10	1.21	0.58–2.22	11	2.12	1.06–3.79	21	1.56	0.96–2.38
Pharynx (145–146)	5	0.48	0.16–1.12	5	0.79	0.26–1.85	10	0.60	0.29–1.10
Esophagus (150)	19	1.14	0.69–1.79	20	2.08	1.27–3.22	39	1.49	1.06–2.03
Stomach (151)	56	0.83	0.63–1.08	53	1.41	1.06–1.85	109	1.04	0.86–1.25
Colon (153)	101	1.27	1.05–1.55	29	0.59	0.40–0.85	130	1.01	0.85–1.21
Rectum (154)	46	0.93	0.68–1.24	39	1.26	0.90–1.72	85	1.06	0.85–1.31
Liver (155) ^f	19	1.92	1.15–2.99	10	1.62	0.78–2.98	29	1.80	1.21–2.59
Pancreas (157)	47	1.46	1.07–1.94	27	1.37	0.90–1.99	74	1.42	1.12–1.79
Nose (160)	2	0.77	0.09–2.77	2	1.25	0.15–4.52	4	0.95	0.26–2.43
Larynx (161)	9	0.65	0.30–1.24	18	2.03	1.20–3.21	27	1.19	0.78–1.73
Lung (162)	150	1.12	0.95–1.31	130	1.49	1.25–1.76	280	1.26	1.12–1.42
Prostate (177)	160	0.92	0.78–1.07	103	0.98	0.81–1.19	263	0.94	0.83–1.06
Testis (178)	9	0.60	0.27–1.14	6	0.71	0.26–1.55	15	0.64	0.36–1.06
Kidney (180)	44	1.17	0.85–1.57	29	1.20	0.81–1.73	73	1.18	0.93–1.49
Bladder (181)	98	1.47	1.19–1.79	42	0.97	0.70–1.31	140	1.27	1.08–1.50
Melanoma (190)	39	0.91	0.65–1.25	16	0.60	0.34–0.97	55	0.79	0.60–1.03
Other skin (191)	33	1.09	0.75–1.53	14	0.74	0.40–1.23	47	0.95	0.70–1.27
Brain (193)	23	0.81	0.52–1.22	11	0.63	0.31–1.13	34	0.74	0.51–1.04
Lymphoma (206)	29	0.83	0.55–1.19	20	0.93	0.57–1.43	49	0.87	0.64–1.15
Leukemia (207) ^g	32	0.85	0.58–1.20	18	0.79	0.47–1.25	50	0.83	0.62–1.09
Other sites	61	0.90	0.69–1.15	40	0.97	0.69–1.32	101	0.92	0.76–1.12
All sites (140–207)	1003	1.03	0.97–1.10	649	1.08	1.00–1.17	1652	1.05	1.00–1.10

^a Code of the International Classification, 7th revision, in parentheses.^b 201 283 person-years.^c 117 177 person-years.^d 318 443 person-years.^e Four persons contributed person-years to both groups of workers.^f Including intrahepatic bile ducts.^g Leukemia and other hematopoietic neoplasms.**Table 4.** Observed number of cancer cases (O), standardized incidence ratios (SIR), and 95% confidence intervals (95% CI) for selected cancer sites among the skilled workers, stratified by year of birth, first year of membership, duration of membership in the trade union (with 20 years of lag) and age at time of diagnosis.

	Colon			Liver ^a			Pancreas			Lung			Bladder		
	O	SIR	95% CI	O	SIR	95% CI	O	SIR	95% CI	O	SIR	95% CI	O	SIR	95% CI
Year of birth															
<1910	45	1.40	1.02–1.87	8	1.72	0.74–3.40	21	1.34	0.83–2.05	74	1.48	1.16–1.85	34	1.42	0.98–1.98
1910–1929	47	1.52	1.11–2.02	8	2.28	0.98–4.49	16	1.38	0.79–2.25	53	0.95	0.71–1.24	41	1.44	1.03–1.96
≥1930	9	0.56	0.26–1.06	3	1.70	0.35–4.97	10	1.97	0.95–3.63	23	0.81	0.52–1.22	23	1.59	1.01–2.38
Starting Year															
<1925	27	1.51	1.00–2.20	5	1.99	0.65–4.65	11	1.27	0.63–2.26	28	1.12	0.74–1.61	16	1.35	0.77–2.19
1925–1934	18	1.28	0.76–2.02	1	0.50	0.01–2.77	5	0.76	0.25–1.77	34	1.35	0.94–1.89	18	1.42	0.84–2.25
1935–1944	13	1.09	0.58–1.87	6	3.96	1.45–8.61	8	1.73	0.75–3.42	28	1.35	0.89–1.95	9	0.84	0.38–1.59
1945–1954	27	1.36	0.89–1.97	1	0.48	0.01–2.67	13	1.80	0.96–3.07	39	1.05	0.75–1.44	33	1.89	1.30–2.65
≥1955	16	1.03	0.59–1.67	6	3.35	1.23–7.29	10	1.94	0.93–3.57	21	0.80	0.50–1.23	22	1.55	0.97–2.35
Age (years)															
<55	11	0.99	0.49–1.77	2	1.63	0.20–5.88	7	1.67	0.67–3.45	17	0.84	0.49–1.35	17	1.81	1.05–2.90
55–74	58	1.29	0.98–1.67	11	1.73	0.86–3.09	32	1.59	1.09–2.24	107	1.19	0.98–1.43	59	1.50	1.14–1.93
≥75	32	1.37	0.94–1.93	6	2.58	0.95–5.61	8	1.00	0.43–1.98	26	1.09	0.71–1.59	22	1.21	0.76–1.84

^a Including intrahepatic bile ducts.

analysis stratified by starting year, the largest excess occurred after 1934 (table 4).

The risk of pancreatic cancer was significantly elevated among the skilled workers (SIR 1.46, 95% CI

1.07–1.94) and was of similar magnitude among the unskilled workers (SIR 1.37, 95% CI 0.90–1.99). The skilled workers employed after 1934 had the highest risks (table 4).

In a pooled analysis of alcohol-related cancers, there was a significantly increased risk of these cancers among the unskilled workers (SIR 1.71, 95% CI 1.33–2.17) that was not present among the skilled workers (SIR 1.09, 95% CI 0.85–1.38). There was no increased risk of cirrhosis among the skilled workers on the basis of 16 observed cases versus 24 expected (SMR 0.66, 95% CI 0.37–1.06). For the unskilled workers there was a nonsignificantly elevated risk based on 23 cases versus 16 expected (SMR 1.43, 95% CI 0.90–2.14).

Discussion

Among the skilled workers, an excess of bladder cancer was observed. In the same occupational group statistically significant excesses were observed also for cancer of the liver, pancreas, and colon. The increase in lung cancer incidence among the skilled workers was restricted to cohort members born before 1910. Among the unskilled workers significant excesses were observed for cancer of the oral cavity, esophagus, stomach, larynx, lung, and all sites.

The advantages of our study include its large size with more than 300 000 person-years of observation and more than 1500 expected cancer cases. High-quality cancer incidence data from the Cancer Registry of Norway provide accurateness in the outcome measures. The use of union rolls as a base for a cohort as in our study may weaken external validity, but, in our study, this problem would have been limited due to a high percentage of organized labor. A major limitation of the study was the lack of detailed information on occupational exposure. This problem may have reduced our ability to detect potential associations between occupation and cancer risk and to identify specific carcinogens. Another limitation of the study was the inability to directly control for potential confounders.

Potential confounders for some of the aforementioned associations are social class, tobacco smoking, and alcohol drinking. Strong negative social gradients have been observed for stomach cancer in several studies (33, 34). Negative gradients have also been reported for cancer of the mouth, pharynx, esophagus, liver, larynx, and lung, while a positive gradient has been found for cancer of the colon and melanoma. Therefore, the pattern of results among unskilled workers in our study resembles a pattern associated with low socioeconomic status. The skilled workers, on the other hand, compared with the unskilled workers, had lower risk for some cancers associated with a negative social gradient (stomach and laryngeal cancer) and a higher risk of cancers with a positive gradient (colon cancer and melanoma). The results draw attention to the importance of

controlling for social class, even in studies within a single industry.

Smoking is related to cancer of the lung, bladder, and several other organs (35). In our study there was a considerably higher proportion of smokers among both the skilled and unskilled workers than in the total survey population. However, among those born before 1935, the group of people contributing the largest proportion of cancer cases, the difference was small. The skilled workers in this cohort experienced an excess of pancreatic cancer and bladder cancer but not of lung cancer, except for the workers in the oldest cohorts. It therefore seems likely that factors other than smoking accounted for the excess of pancreatic cancer and bladder cancer among the skilled workers. The difference in risk for lung cancer and other smoking-related cancers between the skilled and unskilled workers may be due to differences in smoking habits for the two groups, although available data suggest a similar prevalence of smokers. It can be speculated that after 1972 skilled workers, to a greater extent than unskilled workers, had given up smoking, following the pattern of differences between social classes. As the excess of lung cancer in our study was almost solely restricted to the unskilled workers, except for in the old cohorts, and the proportion of ever smokers was higher than in the general population, the increased risk of lung cancer observed in this study must be interpreted with caution with regard to occupational causes.

The alcohol drinking habits of the cohort were unknown. Cancer of the larynx, oral cavity, pharynx, esophagus, and liver are associated with alcohol consumption (36). Among the unskilled workers, there was a significantly elevated risk of all alcohol-related cancers, except for cancer of the pharynx. On the other hand, except for liver cancer, there was no excess of alcohol-related cancers among the skilled workers. This finding, along with the absence of any increased cirrhosis mortality, indicates that excess alcohol consumption in general did not occur in this group of workers.

In general, cancer risk among unskilled workers seems to be confined to cancer sites expected to be influenced by confounders like social class, tobacco smoking, and alcohol drinking. These factors would not be expected to explain the excess cancer risk among skilled workers, and occupational exposure may therefore be a more likely cause in the latter group.

Several case-control studies have shown excess risks of bladder cancer in this industry, although few have been statistically significant. In the current study the excess of bladder cancer was only present for skilled workers. A study from England and Wales based on death certificates of men less than 50 years of age reported odds ratios for bladder cancer of 1.1 for exposure (high or low) to printing inks and 5.0 for high

exposure (17). In a study from Italy, men ever employed in the printing industry had an odds ratio of 1.7 for bladder cancer, adjusted for smoking habits (4). A study from Yorkshire in the United Kingdom reported an elevated odds ratio of 3.1 (95% CI 1.4–6.8) for this cancer for printers (21). A Swedish study presenting risk estimates adjusted for smoking and year of birth reported a relative risk of 2.0 (95% CI 0.8–4.9) for urothelial cancer for exposure to carbon black, and odds ratios of 3.2 (95% CI 0.4–27.1), 0.5 (95% CI 0.1–2.1), and 3.6 (95% CI 0.8–12.1) for estimated low, moderate, and high exposure to printing inks, respectively (37). Among cohort and register linkage studies, several incidence studies have shown increased risks of 1.3–1.4 for bladder cancer (8, 20, 23, 38), and also some mortality studies have shown excess risks of this magnitude (2, 11) although other mortality studies have failed to detect any increased risk (12, 14, 39).

It is recognized that exposure to aromatic amines may result in an increased risk of bladder cancer, but, even though there has been use of such chemicals in the printing industry, it is not clear to what extent they may have contributed to the increased risks in the current and other studies. Other occupations and industries with mixed exposures, including exposures to pigments, have shown an increased risk for bladder cancer (4, 18, 22, 40–50).

An increased risk of lung cancer in the printing industry has been reported in several studies, and ink mist has been suggested as an etiologic agent for lung cancer (24). In our study there was no overall increase among skilled workers. The excess of lung cancer among the oldest cohorts, together with an absence of excess among younger cohorts, may indicate a shift in occupational exposures over time, but the result may also be due to chance or uncontrolled confounding by smoking.

An increased risk of colon cancer has been reported in cohort and record linkage studies in the printing industry (8, 11, 16, 48, 51), and, in a case-control study, colon cancer was reported with a nearly twofold increase among workers in the printing industry (52). Mineral oil is a candidate as a causative agent as it is one of the constituents of ink, and there is some evidence of an increased risk of cancer of the gastrointestinal tract among workers exposed to mineral oils (53, 54). On the other hand, colon cancer risk has been associated with a positive social gradient, and the excess among skilled workers in our study may, in part, be attributed to an effect of social class.

There was a marked excess of cancer of the liver among the skilled workers. Cancer of the liver is associated with alcohol consumption (36). As neither the risk of the other alcohol-related cancers combined nor the risk of cirrhosis was increased among the skilled workers, it seems unlikely that alcohol consumption can

explain the excess. There has been, and still is, a heavy use of organic solvents in the printing industry. Epidemiologic studies have reported excesses of cancer of the liver among persons exposed to organic solvents (55–57). Such exposures may have contributed to the excess of liver cancer in our study. Elevated risks of liver cancer among workers in the printing industry have been reported in some studies from Nordic countries (8, 15, 38), while few other studies do so, although two studies from the United States (11, 16) reported excesses of liver cancer. Differences in exposures between study populations could account for the discrepancy in the risk estimates. However, an excess of liver cancer due to an occupational cause may more easily be detected in a study conducted in a low incidence population using incidence data, as is the case in our study.

No major risk factor except tobacco has been established for pancreatic cancer, a further cancer site with an increased incidence among skilled workers. There is no evidence of a consistent social differential for pancreatic cancer, nor is there evidence of a direct carcinogenic effect of alcohol drinking. An excess of pancreatic cancer was reported among printers with more than 10 years of employment in a hospital-based case-control study (58), and a population-based case-control study reported a moderate, statistically significant increase in pancreatic cancer risk in the printing industry (59), but most studies in the printing industry report no excess of this cancer. Pancreatic cancer has been reported with elevated risks also in other mixed exposure circumstances including those for chemical workers, rubber workers, and paper and pulp workers (60). Organic solvents have been associated with pancreatic cancer risk (61, 62), as has dyes (60). Although our results are suggestive of an association between pancreatic cancer risk and employment in the printing industry, they can also be due to chance.

Although the excess cancer risk among the unskilled workers could have been due to confounding, an effect of occupational exposures should not be ruled out, as, in some occupational settings, the latest arrived unskilled worker gets the worktasks with the highest exposures. In particular, the unskilled workers experienced an excess of cancers of the liver, pancreas, and kidney, neoplasms potentially associated with organic solvent exposure.

The printing industry has changed greatly in Norway over the past 20 years, including digitalization of the prepress processes. The findings in our study may, therefore, not be relevant to the printing industry today. However, the exposure experienced by this study population may still occur today in less developed countries, and artists in many countries are still using some of the traditional techniques.

The standardized incidence ratio for all cancer sites was close to unity. This finding may indicate that an

appropriate reference population was chosen, using rates from Oslo and Akershus as the basis for the expected cases. The printing industry is traditionally an urban industry. Approximately 60% of the members of the Norwegian Union of Graphical Workers are members of its Oslo branch, while one quarter of the general population of Norway lives in the same area. In studies of this industry, on a national level, problems with the correct choice of a reference population may arise because the geographic settlement pattern of the whole population may differ from that of the workers in the industry. In our study the use of national rates would have yielded (eg, for skilled workers) a standardized incidence ratio of 3.5 for liver cancer (compared with 1.9 in table 3).

Our study strongly suggests that printing industry workers are at increased risk of several types of cancer. The risk pattern among the unskilled workers did not allow any firm conclusion of an occupational etiology, as some of the cancers are greatly influenced by nonoccupational factors like alcohol drinking and tobacco smoking. Among the skilled workers, the risk pattern differed, and some of the excess cancer risk was probably due to occupational exposure.

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