

Socioeconomic status and lung cancer incidence in men in The Netherlands: is there a role for occupational exposure?

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Abstract

Study objective – To evaluate the influence of occupational exposure to carcinogens in explaining the association between socioeconomic status and lung cancer.

Design – A prospective cohort study. Data on diet, other lifestyle factors, socio-demographic characteristics and job history were collected by means of a self administered questionnaire. Follow up for incident cancer was established by record linkage with a national pathology register and with regional cancer registries.

Setting – Population originating from 204 municipalities in The Netherlands.

Participants – These comprised 58 279 men aged 55–69 years in September 1986. After 4.3 years of follow up there were 470 microscopically confirmed incident lung cancer cases with complete data on dietary habits and job history.

Measurements and main results – Estimation of occupational exposure to asbestos, paint dust, polycyclic aromatic hydrocarbons, and welding fumes was carried out by two experts, using information on job history from the baseline questionnaire. Socioeconomic status was measured by means of highest attained level of education and two indicators based on occupation. In the initial multivariate analyses of socioeconomic status and lung cancer, adjustment was made for age, smoking habits, intake of vitamin C, beta-carotene and retinol, and history of chronic obstructive pulmonary disease or asthma. Additional adjustment for occupational exposure to the four carcinogens mentioned above did not change the inverse association between the level of education and lung cancer risk (initial model: RR highest/lowest level of education = 0.53; 95% CI 0.34, 0.82; additional model: RR highest/lowest level of education = 0.53; 95% CI 0.34, 0.84). Nor was the association between the two occupation based indicators of socioeconomic status and lung cancer risk influenced by occupational exposure to carcinogens. The effect of occupational exposure on the association between the level of education and lung cancer risk did not differ between ex-smokers and current smokers.

Conclusions – Occupational exposure to asbestos, paint dust, polycyclic aromatic

hydrocarbons, and welding fumes could not explain the inverse association between socioeconomic status and lung cancer risk. More research which explicitly addresses possible explanations for the association between socioeconomic status and lung cancer risk is needed.

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The risk of lung cancer has very often been found to be inversely related to socioeconomic status.¹ In studying this association, two possible intermediate factors arise. Firstly, smoking is the most important cause of lung cancer² which is associated with socioeconomic status.^{3,4} Secondly, occupational exposure to carcinogens is a risk factor for lung cancer^{5,6} which is also associated with socioeconomic status.⁷ Those occupational categories with a higher risk of lung cancer include metal production and processing workers, road construction workers, and chemical workers.⁸ The increased lung cancer risk within these categories is probably due to exposure to asbestos, polycyclic aromatic hydrocarbons, and metal fumes.⁸ In a previous paper we reported the findings of our research on the association between socioeconomic status and lung cancer and the role of smoking habits and other lifestyle characteristics as intermediate factors.⁹ In addition, we have examined the role of occupational exposure to carcinogens as an intermediate factor in the association between socioeconomic status and lung cancer.

In defining occupational exposure, two criteria were predetermined: the relation between exposure and lung cancer should be substantiated by means of human evidence and the occupational exposure had to occur sufficiently frequently to merit investigation by a population-based prospective study.

Four occupational exposures met these two criteria – exposure to asbestos, paint dust, polycyclic aromatic hydrocarbons (PAHs,) and welding fumes.^{10,11,12} Thus, we have studied the role of occupational exposure to carcinogens in further explaining the association between socioeconomic status and lung cancer after adjustment for relevant lifestyle characteristics like smoking and dietary habits. Furthermore, a history of chronic obstructive pulmonary disease (COPD) or asthma was also included in the analyses, because these respiratory disorders are both related to socioeconomic status¹³ as well as to the lung cancer risk.¹⁴

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Methods

THE COHORT STUDY

The Netherlands cohort study (NLCS) on diet, other lifestyle factors, sociodemographic characteristics, job history, and cancer risk started in 1986.¹⁵ The cohort included 58 279 men aged 55–69 years at the beginning of the study, originating from 204 municipalities in The Netherlands. Data on diet, various lifestyle variables, job history, and other risk factors for cancer were collected by means of a self administered questionnaire. The case-cohort approach was used for reasons of efficiency for data processing and analysis. In a case-cohort approach, cases are derived from the entire cohort, while the person-years at risk are estimated from a random sample of 1688 subjects (subcohort). The subcohort can also be used for studying tumours other than lung cancer.¹⁵ Since the baseline exposure measurement, the subcohort has been followed up biennially for vital status information. Between September 1986 and December 1990 (4.3 years of follow up) no subcohort members were lost to follow up. Follow up for incident cancer has been established by record linkage with a national pathology register (PALGA) and with all regional cancer registries in The Netherlands.¹⁶ The current analysis is restricted to microscopically confirmed lung cancer incidence after 4.3 years of follow up. In this period completeness of follow up of the cohort through linkage with the cancer registries and PALGA was estimated to be at least 96%.¹⁷ After excluding subjects with self reported prevalent cancer other than skin cancer, 677 lung cancer cases were detected. Prevalent cancer cases other than skin cancer were also excluded from the subcohort ($n = 58$). Furthermore, people with incomplete information about job history or dietary habits were excluded, leaving 1245 subcohort members and 470 lung cancer cases available for analysis.

SOCIOECONOMIC STATUS

Socioeconomic status was measured by means of highest attained level of education and by means of occupational history, two of the recommended measures for socioeconomic status.¹⁸ Educational level was classified as primary school, lower vocational school, junior high school, senior high school, higher vocational school, university, and other education. Information about occupational history was coded according to the job coding system of the Central Bureau of Statistics (CBS) frequently used in The Netherlands.¹⁹ For the present analyses, these CBS codes were aggregated according to occupational sector and required training (EGP) and according to social standing (U&S). The EGP coding scheme is a reconstruction of the scheme developed by Erikson, Goldthorpe, and Portocarero,²⁰ which is still comparable with the original list.²¹ The U&S score is based on an ordering of occupational titles according to social standing and is also comparable with international classifications.²² Other factors relevant to the association between socioeconomic status and

lung cancer risk that were measured in the baseline questionnaire are age, smoking habits, intake of vitamin C, beta-carotene and retinol, and history of COPD or asthma. For the construction of the smoking variables the following information is used: smoking status (never/ex/current), total years of smoking, and the number of cigarettes, cigars, and/or pipes smoked per day. For the multivariate analyses smoking status and pack-years of cigarette smoking for current and ex-smokers were used. Mean individual intakes of vitamin C, beta carotene, and retinol per day were computed by using information about the usual consumption of food and beverages from a 150 item food frequency questionnaire²³ and information about nutrient contents from the Dutch food composition table of 1986.²⁴

ASSESSMENT OF OCCUPATIONAL EXPOSURE TO CARCINOGENS

For the exposure assessment by experts, information from the baseline questionnaire of the NLCS was used. This information can be best described as a self administered job history, containing data on job title, name of company, type of company, time period, and information about what was being produced at the department. The construction of the coding system is described in detail elsewhere (Van Loon *et al* unpublished data).

Briefly, an exposure estimate was assigned to each job by an occupational hygienist (IJK) and an occupational epidemiologist (GMHS), based on information about job title, name of the company, type of company, and time period. Each job was classified into one of four categories: no exposure, possible exposure (<30% probability), probable exposure (30–90% probability), and nearly certain exposure (>90% probability). Furthermore, a weight was assigned to each exposure category as follows: no exposure – weight 0, possible exposure – weight 0.15, probable exposure – weight 0.6, and nearly certain exposure – weight 0.95. Next, the cumulative index of exposure was calculated by multiplying the weight given to each exposure category by the number of years exposed. Subsequently, for each person, all exposures were summed up for the four carcinogens separately.

DATA ANALYSIS

The prevalence of exposure to the four carcinogens was compared between the case and subcohort groups and between socioeconomic status categories. To study the association between socioeconomic status and lung cancer risk and the role of possible intermediate factors, data were analysed according to the case-cohort approach,^{25,26} using the *GLIM* statistical package.²⁷ In the multivariate analyses, rate ratios and 95% confidence intervals for lung cancer were computed for the different socioeconomic status indicators, after adjustment for age, smoking (never/ex/current and pack years of past and current smokers), intake of beta-carotene, vitamin C and retinol (all as

Table 1 Occupational exposure to carcinogens: number (%) of persons who were ever exposed*

	Asbestos	Paint dust	PAHs	Welding fumes
Subcohort (n = 1245)	109 (8.8)	15 (1.2)	66 (5.3)	134 (10.8)
Cases (n = 470)	66 (14.0)	17 (3.6)	31 (6.6)	59 (12.6)

* Only respondents with complete dietary data and job history.
PAHs = polycyclic aromatic hydrocarbons

Table 2 Association between three different indicators of socioeconomic status (SES) and occupational exposure to carcinogens: number (%) of people in the subcohort who were ever exposed*

SES indicator	Asbestos	Paint dust	PAHs	Welding fumes
Highest level of education*				
Primary school	15 (5.1)	7 (2.4)	12 (4.1)	19 (6.5)
Lower vocational	49 (18.8)	6 (2.3)	29 (11.2)	65 (25.0)
Junior high school	20 (5.8)	2 (0.6)	14 (4.0)	23 (6.6)
Senior high school	2 (1.9)	0 (0)	1 (1.0)	4 (3.8)
Higher vocational/university	23 (9.6)	0 (0)	10 (4.2)	23 (9.6)
EGP score†: last occupation				
Blue collar	55 (12.1)	12 (2.6)	38 (8.4)	82 (18.0)
Lower white collar	12 (6.7)	2 (1.1)	8 (4.5)	14 (7.8)
Upper white collar	36 (9.0)	1 (0.2)	17 (4.2)	36 (9.0)
Other	6 (2.8)	0 (0)	3 (1.4)	2 (0.9)
U&S score‡: last occupation				
1 (low)	19 (8.1)	12 (5.1)	10 (4.3)	28 (11.9)
2	40 (12.7)	1 (0.3)	28 (8.9)	48 (15.2)
3	16 (4.6)	2 (0.6)	13 (3.7)	26 (7.4)
4	14 (7.3)	0 (0)	6 (3.1)	16 (8.4)
5 (high)	20 (13.1)	0 (0)	9 (5.9)	16 (10.5)

* Only respondents with complete dietary data and job history (n = 1245). † EGP score: an ordering based on occupational sector and required training.
‡ U&S score: an ordering based on social standing.
PAHs = polycyclic aromatic hydrocarbons.

Table 3 Rate ratio for lung cancer according to three different socioeconomic status (SES) indicators in multivariate analysis*

SES indicator	No of cases in cohort	Person years in subcohort	RR†	95% CI	RR‡	95% CI
Highest level of education						
Primary school	137	1214	1†		1†	
Lower vocational	121	1097	1.28	0.89,1.85	1.27	0.87,1.85
Junior high school	121	1442	0.94	0.65,1.36	0.97	0.67,1.41
Senior high school	31	437	0.70	0.40,1.21	0.73	0.42,1.28
Higher vocational/university	57	992	0.53	0.34,0.82	0.53	0.34,0.84
Test for trend ² (p value)				19.38 (<0.01)		16.34 (<0.01)
EGP score¶: last profession						
Blue collar	198	1884	1†		1†	
Lower white collar	55	752	0.76	0.49,1.18	0.82	0.52,1.28
Upper white collar	144	1668	0.91	0.66,1.25	0.95	0.68,1.31
Other**	73	891	0.75	0.51,1.10	0.81	0.54,1.20
Test for trend χ^2 (p value)				0.83 (0.36)		0.25 (0.62)
U&S score‡: last profession						
1 (low)	102	980	1†		1†	
2	129	1317	1.22	0.82,1.80	1.23	0.82,1.86
3	120	1462	0.89	0.60,1.32	0.97	0.65,1.46
4	61	796	0.85	0.53,1.36	0.91	0.56,1.48
5 (high)	58	639	1.15	0.72,1.83	1.18	0.73,1.92
Test for trend χ^2 (p value)				0.33 (0.56)		0.05 (0.83)

* Only respondents with complete dietary data and job history.

† Reference category.

‡ Adjusted for age, smoking behaviour (never/ex/current and packyears), intake of beta carotene, vitamin C and retinol, history of chronic obstructive pulmonary disease (COPD), and asthma.

§ For age, smoking behaviour (never/ex/current and packyears), intake of beta carotene, vitamin C, retinol, history of COPD and asthma and exposure to asbestos, paint dust, PAHs and welding fumes.

¶ EGP score: an ordering based on occupational sector and required training.

** Excluded from test for trend.

‡‡ U&S score: an ordering based on social standing.

continuous variables), and history of COPD or asthma and after additional adjustment for lifetime exposure to asbestos, paint dust, PAHs, and welding fumes (also as continuous variables). Finally, the role of occupational exposure to carcinogens in explaining the association between socioeconomic status and lung cancer was studied more extensively by conducting a multivariate analysis in the different smoking categories.

Results

The prevalence of occupational exposure to the relevant carcinogens is presented in table 1. A higher proportion of cases was exposed to asbestos, paint dust, PAHs, or welding fumes. The distribution of socioeconomic status indicators and covariates in the case and subcohort groups after 3.3 years of follow up is presented elsewhere.⁹ Briefly, cases were on average older than members of the subcohort and current smoking was more prevalent in the case group. Cases had a lower educational level and were more often employed in blue collar occupations than members of the subcohort. Finally, the prevalence of COPD or asthma was higher among cases (13%) than in with subcohort members (10%).

The prevalence of occupational exposure within each socioeconomic status category in the subcohort is presented in table 2. Occupational exposure to carcinogens was most prevalent among men with lower vocational schooling. The prevalence of occupational exposure to asbestos or welding fumes was also relatively high among men with higher vocational schooling/university education. According to the occupation-based socioeconomic status indicators the prevalence of exposure to asbestos, paint dust, PAHs, or welding fumes was higher among men whose most recent occupation was a blue collar- or a low social standing occupation. The prevalence of exposure to asbestos, PAHs, and welding fumes was also high among men within the highest social standing category.

Table 3 shows the results of the multivariate analyses without and with adjustment for lifetime occupational exposure to asbestos, paint dust, PAHs, and welding fumes. Initially, adjustment was made for age, smoking habits, intake of beta-carotene, vitamin C and retinol, and history of COPD or asthma. The significant inverse association between the highest level of education and lung cancer risk (RR highest/lowest level of education = 0.53; 95% CI 0.34,0.82, trend p<0.01) did not change after additional adjustment for exposure to the four carcinogens (RR highest/lowest level of education = 0.53; 95% CI 0.34,0.84, trend p<0.01). The association between the EGP score and lung cancer risk (RR lower white collar/blue collar = 0.76, 95% CI 0.49–1.18) was slightly changed after additional adjustment (RR lower white collar/blue collar = 0.82; 95% CI 0.52,1.28). There was no association found between the U&S score and lung cancer risk.

In a previous analysis of the NLCS data, the inverse association between the highest level of education and lung cancer risk was only found among current smokers.⁹ Therefore, we have studied the effect of occupational exposure to carcinogens on the association between socioeconomic status and lung cancer within the different smoking categories (table 4). Due to the small number of lung cancer cases with complete data in the non-smoking group (n = 3), the analysis could not be carried out for the non-smokers only. We found significant inverse associations between highest level of education

Table 4 Rate ratio for lung cancer according to highest level of education, by category of smoking in multivariate analysis*, without and with adjustment for occupational exposures

	RR†	95% CI	RR‡	95% CI
Ex-smokers				
Highest level of education				
Primary school	1§		1§	
Lower vocational	1.12	0.61,2.07	1.24	0.59,2.60
Junior high school	1.02	0.57,1.84	1.13	0.58,2.18
Senior high school	1.12	0.48,2.58	1.25	0.52,30.3
Higher voc./university	0.37	0.17,0.82	0.41	0.18,0.94
Test for trend χ^2 (p value)	7.08	(<0.01)	5.41	(0.02)
Current smokers				
Highest level of education				
Primary school	1§		1§	
Lower vocational	1.49	0.92,2.41	1.48	0.90,2.43
Junior high school	0.86	0.54,1.37	0.91	0.57,1.48
Senior high school	0.58	0.29,1.18	0.62	0.30,1.25
Higher voc./university	0.60	0.35,1.03	0.63	0.36,1.09
Test for trend χ^2 (p value)	12.65	(<0.01)	9.26	(<0.01)

*Only respondents with complete dietary data and job history.

†Adjusted for age, packyears, dietary intake of beta carotene, vitamin C and retinol, history of COPD or asthma.

‡Adjusted for age, packyears, dietary intake of beta-carotene, vitamin C and retinol, history of COPD or asthma and occupational exposure to asbestos, paint dust, PAHs and welding fumes.

§Reference category.

and lung cancer risk among ex-smokers and among current smokers after adjustment for pack-years, dietary intake of beta-carotene, vitamin C and retinol, and history of COPD or asthma. However, among ex-smokers, only the rate ratio for men with higher vocational school or university education was below one. Additional adjustment for occupational exposure to asbestos, paint dust, PAHs, and welding fumes showed only marginal changes, both among ex-smokers and current smokers.

Discussion

In a previous paper we reported an inverse association between socioeconomic status and lung cancer in the NLCS, after adjustment for age and lifestyle variables, after 3.3 years of follow up.⁹ In addition, we have studied the influence of occupational exposure to carcinogens on the association between socioeconomic status and lung cancer after 4.3 years of follow up. The statistically significant inverse association between the level of education and lung cancer risk did not change after additional adjustment for occupational exposure to asbestos, paint dust, PAHs, and welding fumes. The non-significant inverse association between the EGP score and lung cancer risk changed marginally after additional adjustment for occupational exposure to carcinogens. No association was found between social standing and lung cancer risk. The effect of occupational exposure on the association between the level of education and lung cancer risk was not different between ex-smokers and current smokers.

Earlier results from the NLCS showed statistically significant positive associations between lung cancer risk and occupational exposure to asbestos or paint dust (Van Loon *et al.*, unpublished data). Occupations with a high probability of exposure to asbestos in the NLCS were blacksmith, motor mechanic, pipefitter, welder, marine engineer, and barge-man. Most of the people working in these occupations had lower vocational school as their highest attained level of education. This

might be an explanation for the higher lung cancer risk among men with lower vocational schooling. However, the proportion of men with higher vocational school or university education as the highest attained level of education who were exposed to asbestos was also relatively high – although most were in the lowest exposure tertiles (not presented). This indicates that there is no straightforward association between the level of education and occupational exposure to asbestos. Consequently, this could be explain why adjustment for occupational exposure to asbestos did not change the risk estimates either when occupational exposure was considered as a continuous variable or when categorical variables were used (not presented).

The proportion of men who were ever exposed to asbestos, PAHs, or welding fumes and whose last occupation was a white collar profession or a high social standing occupation was rather high. In about half of the instances the probability of exposure to asbestos in the white collar or high social standing profession was actually assessed to be greater than zero (marine engineer, management of metal industry, or metallurgist). In the remainder, people whose last occupation was a white collar profession or a high social standing occupation were exposed to asbestos, PAHs, or welding fumes during an earlier episode of their job history, in a blue collar profession or a low social standing occupation. This explains the finding that occupational exposure to asbestos, PAHs, or welding fumes was not restricted to men whose last occupation was a blue collar profession or a low social standing occupation. Exposure to paint dust is mainly found among house painters, which appears to be a 'lifelong' occupation in the NLCS. Therefore, exposure to paint dust is only found among blue collar workers and within low social standing jobs. However, because the proportion of house painters in the cohort is small, this will not explain differences in lung cancer risks among socioeconomic status categories.

We found only one study that evaluated the association between social class and lung cancer after adjustment for occupational exposure and tobacco.⁷ The main goal of that study was to assess the proportion of male lung cancer cases due to occupational exposure. The authors reported that social class (based on the last occupation) was only a minor determinant of lung cancer when occupation and smoking habits were controlled for (RR high social class / low social class=0.8). They did not present information about the association between social class and lung cancer without adjustment for occupational exposure.

The NLCS has been performed in a large sample of the general population aged 55–69 years at baseline. The follow up period of 4.3 years resulted in 470 male lung cancer cases with complete dietary data and job history. The follow up of person-years was 100% complete and the completeness of cancer follow up was also very high, indicating that selection bias due to loss to follow up is unlikely. Besides, information bias due to random mis-

classification may have influenced the results. Socioeconomic status is operationalised as the highest attained level of education, EGP score (functional level), and social standing (U&S score) – the last two both based on the last occupation. In individuals, the highest level of education is stable over time and therefore it will avoid the risk of reverse causation, but this stability can mask important changes in individual circumstances after education has been completed. The occupation based socioeconomic status indicators reflect the more recent situation. However, it is not clear whether knowledge of the socioeconomic status at an older age or younger age is more relevant in studying the association between socioeconomic status and lung cancer risk.

The lifetime occupational exposure to carcinogens was calculated by multiplying the duration of exposure by the likelihood of exposure. The accuracy of the job histories did not allow an estimation of the actual exposure concentrations that were experienced in the past. This may lead to misclassification of exposure. However, since the associations between occupational exposure and lung cancer risk found in the NLCS are comparable with findings from other studies (van Loon *et al.*, unpublished data), it is likely that the exposure assessment reflects the actual exposure to carcinogens to a reasonable extent.

Misclassification of smoking habits may lead to residual confounding. We have measured smoking habits very carefully and included smoking status (never/ex/current smokers) and pack-years of cigarette smoking into the model. Therefore, both duration and amount were taken into account. Moreover, we studied the effect of occupational exposure to carcinogens on the association between socioeconomic status and lung cancer among smokers and ex-smokers separately, with comparable results. Consequently, it is not likely that residual confounding of smoking will explain the finding that adjustment for occupational exposure to carcinogens does not influence the association between socioeconomic status and lung cancer risk.

Although more articles have been published on the association between socioeconomic status and lung cancer risk, we did not find papers explicitly addressing possible explanations for these associations. Studies on socioeconomic health differences in The Netherlands point to the role of work related factors like the ability to regulate working conditions,²⁸ psychosocial factors,²⁹ and differences in social participation and possibilities to control one's fate.³⁰ Since these factors are associated with cancer,^{31 32} they may also influence the association between socioeconomic status and lung cancer. However, it is not possible to investigate these factors in the NLCS, because there were no items about psychosocial characteristics included in the baseline questionnaire. Also, air pollution^{33 34} may play a part in explaining the association between socioeconomic status and lung cancer. For the same reason mentioned above, it is not possible to investigate this in the NLCS.

In conclusion, after adjustment for smoking, dietary habits, and history of lung disease there was still an inverse association between socioeconomic status and lung cancer risk after 4.3 years of follow up. Occupational exposure to asbestos, paint dust, PAHs, and welding fumes could not explain this association.

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