Bulletin of the *Transilvania* University of Braşov Series VI: Medical Sciences • Vol. 5 (54) No. 2 - 2012

LUNG CANCER AND OCCUPATION IN TWO DIFFERENT AREAS OF ROMANIA

A.P. $FILDAN^1$ D. $ALEXANDRESCU^2$

Abstract: The authors examined the relation between occupation and lung cancer in two patient cohorts from different areas of Romania: Constanta and Valcea. In 2005–2010, in Constanta and Valcea counties, 488 and 344 incident lung cancer cases were enrolled. Lifetime occupational histories (industry and job title) were coded by using standard international classifications and were translated into occupations known (list A) or suspected (list B) to be associated with lung cancer. An exposure excess of 14.6% for patients from Constanta ever employed in occupations known to be associated with lung cancer (list A) was found, with the largest contributions from the oil refinery and shipbuilding industries No overall excess was found for list B with the exception of bus and truck drivers (men) and launderers and dry cleaners (women), in both groups of patients. These results indicate that past exposure to occupational carcinogens remains an important determinant of lung cancer occurrence.

Key words: industry, lung cancer, occupations.

Introduction

Lung cancer is the most frequent neoplasm worldwide, with more than 1.6 million new cases and 1.3 million deaths in 2008 [1]. Although tobacco smoking is by large the most important cause, occupational factors play a remarkable role.

In the year 2000, it was estimated that 10% of lung cancer deaths among men (88,000 deaths) and 5% among women (14,300 deaths) worldwide were attributable to exposure to 8 occupational lung carcinogens (arsenic, asbestos, beryllium, cadmium, chromium, diesel fumes, nickel, and silica) [2–4]. In Europe, assuming attributable fractions of 7%–15% (men) and 2%–9% (women), the estimated numbers of deaths

caused by exposure to chemicals in the workplace were more than 29,300 and 3,200, respectively [2]. Prevalence of occupational exposure to carcinogens is still high: in 1990–1993, of almost 140 million workers in 15 states of the European Union, 32 million were estimated to be exposed to carcinogenic agents and about 7 million to the 8 above-mentioned carcinogens [5].

Different approaches are used to evaluate occupational exposure to carcinogens [6-8]: one makes use of lists of occupations known (list A) or suspected (list B) to be associated with lung cancer based on evaluations of carcinogenic risks by the International Agency for Research on Cancer (IARC) [9], [10]. These lists are periodically

¹ Clinical Pneumophtisiology Hospital of Constanța.

² Faculty of Medicine, *Transilvania* University Braşov.

updated and have been extensively used worldwide as a standardized tool to quantify the burden of occupational lung cancer [8], [11], [12], but there remains a need to continue to evaluate occupations and try to uncover additional jobs and occupations that may contribute to the lung cancer burden.

2. Objective

The study aimed to analyze the role of occupational exposures as a determinant of lung cancer risk.

3. Materials and Methods

The study included patients diagnosed with lung cancer in the Pneumology Departments of Constanta Clinical Pneumophtisiology Hospital and "Constantin Anastasatu" Pneumology Hospital of Râmnicu Valcea, between 2006–2010.

Extensive demographic and clinical data were collected for lung cancer cases, including morphology coded according to the International Classification of Diseases for Oncology, Third Edition [13] and categorized into major histologic subtypes based on World Health Organization/ International Association for the Study of Lung Cancer classification [14].

All subject completed a self- administered questionnaire regarding lifetime history (vears of start/stop, industry, job title) of jobs held for at least 6 months. Industries and job titles were coded following the International Standard Industrial Classification of All Economic Activities [15] and the International Standard Classification of Occupations (16). Codes were then translated into occupations known (list A) or suspected (list B) to entail a carcinogenic risk to the lung [9, 10]. The list B occupation filling station attendant, for which there are no specific codes, was identified through text search. Subjects with job titles from both lists were assigned to list A and to list B only if they had never worked in list A occupations.

We performed a statistical analysis by using the softs: GraphPad StatMate (1.01) and GraphPad Prism 4 (4.03).

4. Results

Of the 822 lung cancer cases registered in Pneumology Departments, during the 5year period, there were 478 patients from Constanta and 344 from Ramnicu Valcea. The main characteristics of lung cancer patients are given in Table 1.

Table 1

	All (%)	Constanta (CT)		Valcea (VL)		P value
	822	No=478	%	No=344	%	
Age, years (mean±SD)	63.03±10.05	62.22±10.05		65.26±9.76		< 0.05
Gender (sex ratio)		4.3/1 (388/90)		5.5/1 (291/53)		
Smoking	822					
Never	72 (8.8)	44	9.2	28	8.1	ns
Former (quit >6 months ago)	175 (21.2)	98	20.5	77	22.4	ns
Current	575 (69.9)	336	70.3	239	69.5	ns
Cigarette pack-years (mean±SD)	61.5±10.84	63.5±10.02		59.07±11.34		< 0.05
Histology	723 (88)					
Squamous	291 (40.2)	168	39.8	121	40.2	ns
Adenocarcinoma	207 (28.6)	124	29.3	85	28.2	ns
SCLC	104 (14.4)	63	14.9	44	14.6	ns
Large cell carcinoma	45 (6.2)	30	7.1	19	6.3	ns
Others	76(10.5)	37	8.7	32	10.6	ns

Selected Characteristics of Lung Cancer Patients

28

The patients from CT developed the disease at an earlier age than patients from VL (62.22 versus 65.26 years; P < 0.05). Less than 10% of patients were never smokers (9.2% CT group, 8.1% VL group). Almost a fifth of the patients (175/822 cases) were former smokers. In both groups, about 70% of patients were current smokers but the patients from CT group had smoked greater numbers of cigarettes (63.5 pack-years versus 59.07 pack-years, P < 0.05).

The majority of lung cancers were squamous cell carcinomas (41.5%) in both groups.

Among all patients, 70 cases (14.64%) from CT group and 27 cases (7.84%) from VL group, had ever worked in list A occupations (P<0.05) (Table 2). The most frequent exposures in CT patients were in oil refinery (15 cases) and shipbuilding (14 cases). For VL group of patients, the exposures in constructions (9 cases) and chemical industry (9 cases) were more frequents.

7	7~	L	1	~	2	
L	a	D	l	e	4	

Industry	CT group	of patients	VL group of patients		
	No	%	No	%	
Agriculture (using arsenical insecticides)	2	0.41	3	0.87	
Construction and construction materials	13	2.71	9	2.61	
Metals (iron and steel, basic)	4	0.83	1	0.29	
Chemical industry	5	1.04	9	2.61	
Oil refinery	15	3.13	0	0	
Mining and quarring	0	0	4	1.16	
Oil and gas extraction	12	2.51	0	0	
Schipbuilding	14	2.92	0	0	
Others (construction, automobile painters)	5	1.04	1	0.29	
Total	70	14.64	27	7.84	
<i>P</i> value			< 0.05		

The number and	nercentage of	natients accora	ling to l	ist A o	ccupations
The number and	percentage of	patients accord	$m_{\Sigma} i \circ i$		companions

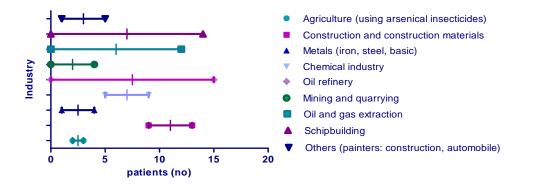


Fig. 1. Exposure to list A occupations

In total, 114 cases (13.86%): 56 cases (11.71%) from CT group of patients and 48 cases (13.95%) from VL group of patients had been working in occupations

in list B, with no statistical differences between the two groups (Table 3). We found a marked elevated exposure for bus and truck drivers in both groups (16 cases in CT group and 14 cases in VL group of patients). For women, we found increased level of exposure for launderers and dry

cleaners (5 cases in both groups of patients).

Table 3

Industry	CT group of patients		VL group of patients	
	No	%	No	%
Machine and equipments industry	4	0.83	1	0.29
Leather (tanners, processors)	3	0.62	1	0.29
Motor vehicle manufacturing and repair	8	1.67	5	1.45
Trade: filling station attendants	6	1.25	4	1.16
Manufacture of fabricated wood products	2	0.41	4	1.16
Rubber and plastisc industry	3	0.62	7	2.03
Printing (machine-rooms workers, printing pressmen)	2	0.41	0	0
Transport (bus, truck drivers, railroad workers)	16	3.34	14	4.07
Glas (art glas, container and pressed ware)	4	0.83	5	1.45
Food (butchers and meat workers)	5	1.04	5	1.45
Others (launderers, dry cleaners, joiners)	3	0.62	2	0.58
Total	56	11.71	48	13.95
<i>P</i> value		0.0885 CI 95% -1067 to 1207		

The number and percentage of patients according to list B occupations

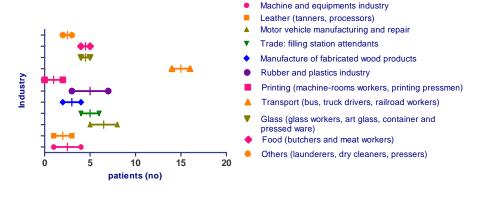


Fig. 2. Exposure to list B occupations

5. Discussion

Lung cancer incidence in Romania (1) and, particularly, in Constanta county is rising dramatically for several decades. Although tobacco smoking is by large the most important cause, occupational factors play a remarkable role.

This study was performed in 2005–2010 in Constanta, eastern Romania—one of the most populated, 724000 inhabitants, 65%

living in urban areas, economically relevant, and industrialized region in Romania, high lung cancer incidence and mortality rates (higher than national rates) and in Vâlcea– southwestern Romania, 412000 inhabitants, 55% living in rural areas, lower incidence and mortality rates than national values, in order to analyze the role of occupational exposures as a determinant of lung cancer risk. We found an exposure excess of 14.6% for patients from Constanta ever employed in occupations known (list A) to be associated with lung cancer, with the largest contributions from the oil refinery and shipbuilding industries. For patients from Valcea, the frequency of exposure was lower (7.8%, P<0.05), with higher contributions from constructions and chemical industries. These findings are similar with results reported in other studies [17, 18].

Statistical analysis showed differences regarding the exposure according with occupation list A (P<0.05).

Of the occupations suspected (list B) to be associated with lung cancer, we found a suggestive increases for bus, truck drivers and railroad workers (men) in both groups of patients, with no statistical differences regarding the frequency of exposure (P=0.0885 CI 95% -1067 to 1207). The findings for individual occupations in list B are suggestive because the number of exposed subjects was great. The increased risk for bus and truck drivers deserves mention because it was based on a substantial number of exposed workers (16 cases in Constanta and 14 cases in Vâlcea group of patients). For women, we found a moderate risk increase for launderers and dry cleaners, a finding reported in other studies [17].

This study confirmed the important role of past occupational exposures as a determinant of lung cancer risk at the beginning of the new century. The low exposure level for list A occupations among women was expected, given that exposure to most occupational lung carcinogens occurred in workplaces in which women constituted a minority.

In conclusion, the findings of this study confirm the need for continuous monitoring and improved control of workrelated exposures, both for prevention and workers compensation purposes. Future occupational health studies should improve their ability to address interindividual variability in response to the lower exposures in work settings [19, 20, 21].

6. Conclusions

The frequency of exposure for patients from Constanța ever employed in occupations known to be associated with lung cancer (list A) was higher than the level of exposure for Vâlcea group of patients. Over representation of oil refinery and shipbuilding industry exposure was observed in the group of patients from Constanța.

Of the occupations suspected to be associated with lung cancer (list B), the higher exposure was found for bus, truck drives and railroad workers, in both groups of patients.

References

- Jemal, A., Siegel, R., Xu, J., Ward, E.: *Cancer statistics 2010.* In: Cancer J Clin 2010; 60:277-300.
- Driscoll, T., Nelson, D.I., Steenland, K., et al.: *The global burden of disease due to occupational carcinogens*. In: Am J Ind Med 2005;48(6):419-431.
- Fingerhut, M., Nelson, D.I., Driscoll, T., et al.: *The contribution of* occupational risks to the global burden of disease: summary and next steps. In: Med Lav 2006; 97(2):313-321.
- Nelson, D.I., Concha-Barrientos, M., Driscoll, T., et al.: *The global burden* of selected occupational diseases and injury risks: methodology and summary. In: Am J Ind Med 2005; 48(6):400-418.
- 5. Kauppinen, T., Toikkanen, J., Pedersen, D., et al.: *Occupational exposure to carcinogens in the European Union*. In: Occup Environ Med 2000;57(1):10-18.

- 6. Mannetje, A., Kromhout, H.: *The use* of occupation and industry classifications in general population studies. In: Int J Epidemiol 2003; 32(3):419-428.
- Teschke, K., Olshan, A.F., Daniels, J.L., et al.: Occupational exposure assessment in case-control studies: opportunities for improvement. In: Occup Environ Med 2002;59(9):575-593.
- Siemiatycki, J., Richardson, L., Boffetta, P.: Occupation. In: Schottenfeld D, Fraumeni JF Jr, (eds.) Cancer Epidemiology and Prevention. 3rd ed. New York, NY: Oxford University Press; 2006. p. 322-354.
- IARC Monographs on the Evaluation of Carcinogenic Risks to Humans. Lyon, France: International Agency for Research on Cancer; (http://monographs.iarc.fr/ENG/Classi fication/index.php). (Accessed October 15, 2010).
- Simonato, L., Saracci, R.: International Labor Organization (ILO), *Encyclopedia of Occupational Safety* and Health. Geneva, Switzerland: ILO; 1983. Cancer, occupational; p. 369-375.
- 11. De Matteis, S., Consonni, D., Bertazzi, P.A.: *Exposure to occupational carcinogens and lung cancer risk. Evolution of epidemiological estimates of attributable fraction.* In: Acta Biomed 2008; 79:34-42.
- Siemiatycki, J., Richardson, L., Straif, K., et al.: *Listing occupational carcinogens*. In: Environ Health Perspect 2004;112(15):1447-1459.
- Fritz, A.G., Percy, C., Jack, A., et al.: International Classification of Diseases for Oncology. 3rd ed. Geneva, Switzerland: World Health Organization; 2000.

- Brambilla, E., Travis, W., Colby, T., Corrin, B., Shimosato, Y.: *The new World Health Organization classification of lung tumors*. In: Eur J Respir 2001 Dec;18(6):1059-1068.
- International Standard Industrial Classification of All Economic Activities (ISIC). United Nations Publications ST/STAT/M.4/Rev.2/Add.1, Sales No.: E.71.XVII.8. New York, NY: Publishing Service, United Nations; 1971.
- International Labour Office. International Standard Classification of Occupations. Geneva, Switzerland: International Labour Office; 1968.
- Richiardi, L., Boffetta, P., Simonato, L., et al.: Occupational risk factors for lung cancer in men and women: a population-based case-control study in Italy. In: Cancer Causes Control 2004;15(3):285-294.
- Pohlabeln, H., Boffetta, P., Ahrens, W., et al.: Occupational risks for lung cancer among nonsmokers. In: Epidemiology 2000;11(5):532-538.
- Alexandrescu, D.: Integrated positronemission tomography and the staging of nodular non-small-cell lung cancer. In: Bulletin of the Transilvania University of Braşov, vol I (50), series VI Medical Science, 2008, ISSN 2065-2224, 157-162.
- Man, M., Alexandrescu, D.: *Pulmonary cancer among nonsmokersrisk factors*. In: Bulletin of the Transilvania University of Braşov, vol I (50), series VI Medical Science, 2008, ISSN 2065-2224, 157-162.
- Alexandrescu, D., Man, M.: Estimarea probabilității de malignitate a nodulilor pulmonari solitari (NPS) cu ajutorul modelului Swensen. In: Clujul Medical, vol LXXXI 2008, nr.1, ISSN 1222-2119, 96-99.