

Systematic Reviews and Meta- and Pooled Analyses

Lung Cancer Risk Among Hairdressers: A Pooled Analysis of Case-Control Studies Conducted Between 1985 and 2010

Ann C. Olsson*, Yiwen Xu, Joachim Schüz, Jelle Vlaanderen, Hans Kromhout, Roel Vermeulen, Susan Peters, Isabelle Stücker, Florence Guida, Irene Brüske, Heinz-Erich Wichmann, Dario Consonni, Maria Teresa Landi, Neil Caporaso, Lap Ah Tse, Ignatius Tak-sun Yu, Jack Siemiatycki, Lesley Richardson, Dario Mirabelli, Lorenzo Richiardi, Lorenzo Simonato, Per Gustavsson, Nils Plato, Karl-Heinz Jöckel, Wolfgang Ahrens, Hermann Pohlabein, Adonina Tardón, David Zaridze, Michael W. Marcus, Andrea 't Mannelje, Neil Pearce, John McLaughlin, Paul Demers, Neonila Szeszenia-Dabrowska, Jolanta Lissowska, Peter Rudnai, Eleonora Fabianova, Rodica Stanescu Dumitru, Vladimir Bencko, Lenka Foretova, Vladimir Janout, Paolo Boffetta, Cristina Fortes, Bas Bueno-de-Mesquita, Benjamin Kendzia, Thomas Behrens, Beate Pesch, Thomas Brüning, and Kurt Straif

* Correspondence to Dr. Ann C. Olsson, Environment and Radiation Section, International Agency for Research on Cancer, 150 Cours Albert Thomas, 69372 Lyon Cedex 08, France (e-mail: olssona@iarc.fr).

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Increased lung cancer risks among hairdressers were observed in large registry-based cohort studies from Scandinavia, but these studies could not adjust for smoking. Our objective was to evaluate the lung cancer risk among hairdressers while adjusting for smoking and other confounders in a pooled database of 16 case-control studies conducted in Europe, Canada, China, and New Zealand between 1985 and 2010 (the Pooled Analysis of Case-Control Studies on the Joint Effects of Occupational Carcinogens in the Development of Lung Cancer). Lifetime occupational and smoking information was collected through interviews with 19,369 cases of lung cancer and 23,674 matched population or hospital controls. Overall, 170 cases and 167 controls had ever worked as hairdresser or barber. The odds ratios for lung cancer in women were 1.65 (95% confidence interval (CI): 1.16, 2.35) without adjustment for smoking and 1.12 (95% CI: 0.75, 1.68) with adjustment for smoking; however, women employed before 1954 also experienced an increased lung cancer risk after adjustment for smoking (odds ratio = 2.66, 95% CI: 1.09, 6.47). The odds ratios in male hairdressers/barbers were generally not elevated, except for an increased odds ratio for adenocarcinoma in long-term barbers (odds ratio = 2.20, 95% CI: 1.02, 4.77). Our results suggest that the increased lung cancer risks among hairdressers are due to their smoking behavior; single elevated risk estimates should be interpreted with caution and need replication in other studies.

case-control studies; hair bleaching agents; hair color; lung neoplasms; occupational exposure

Abbreviations: AUT-Munich, Arbeit und Technik-Munich; CI, confidence interval; LUCAS, Lungcancer i Stockholm; MORGEN, Monitoring van Risicofactoren en Gezondheid in Nederland; SYNERGY, Pooled Analysis of Case-Control Studies on the Joint Effects of Occupational Carcinogens in the Development of Lung Cancer.

The World Health Organization's International Agency for Research on Cancer classified "occupational exposures of hairdressers and barbers" as probably carcinogenic to humans (Group 2A) in 1993 and 2010 on the basis of limited evidence

for an association with bladder cancer, mainly in men (1, 2). Increases in lung cancer risk (20%–40%) have been observed in several cohort studies and a few case-control studies (3). Nevertheless, the evidence for an association between

occupation as hairdresser or barber and lung cancer is not conclusive because most of the data from cohort studies come from linkage between census data and cancer registry data, primarily in the Scandinavian countries, which provide excellent opportunities to monitor cancer risks by occupation but offer limited ability to control for tobacco smoking and other confounders (4). The case-control studies have not been convincing because of lack of power and details about type, calendar period, and duration of employment as a hairdresser (5, 6).

Hairdressers and barbers can be exposed to a wide range of chemicals, such as volatile organic chemicals (e.g., toluene, ethanol, isopropanol, ether, diaminotoluene, phenylenediamine) via hairsprays and setting lotions, as well as ammonia, ammonium persulfates, hydrogen peroxide, and organic pigments as ingredients of permanent waves, hair dyes, and hair bleaching applications (7, 8). Important changes in the composition and use of hair products have taken place over the years; many hazardous dyes have been phased out, and chlorinated solvents used as propellants in hair sprays (e.g., methylene chloride) have been replaced by less harmful organic solvents (9). Work-related skin and respiratory symptoms remain frequent, and together with musculoskeletal complaints, contribute to many hairdressers leaving their jobs within a few years of starting to work as hairdressers (10). Products used by hairdressers are used on their customers. If some of the products were carcinogenic, this would have implications for the general public. Some studies have indeed suggested an increased risk of cancer from personal use of hair dyes, but the overall evidence was not conclusive (1, 2). Occupational studies typically involve stronger exposure contrasts and might therefore provide better options to investigate a cancer risk, which in turn might also inform about carcinogenic hazards of consumer products. Pooling community-based case-control studies appears to be the best available alternative to explore why hairdressers experience an increased lung cancer risk, especially because they often work only a few years as hairdressers and because of availability of information on life-long smoking habits. The Pooled Analysis of Case-Control Studies on the Joint Effects of Occupational Carcinogens in the Development of Lung Cancer (SYNERGY) project is the largest data set of its kind with complete smoking and occupational information.

Our objective was to study the potential association between employment as a hairdresser and increased lung cancer risk in SYNERGY, while adjusting for tobacco smoking. We stratified analyses by sex, type of hairdresser, calendar period of employment, and lung cancer histology. We also compared smoking habits between hairdressers and nonhairdressers.

MATERIALS AND METHODS

Study population and data collection

The SYNERGY project includes data from 16 case-control studies on lung cancer conducted in 13 European countries, Canada, New Zealand, and China between 1985 and 2010. The Lungcancer i Stockholm (LUCAS) Study, a study of lung cancer in France, and a case-control study of male lung cancer, occupational exposures, and smoking in Hong Kong were restricted to men, and the lung cancer study in Paris included

only former and current smokers. The International Agency for Research on Cancer Multicenter Case-Control Study of Occupation, Environment, and Lung Cancer in Central and Eastern Europe (INCO-COPERNICUS) is a multicenter study in Central and Eastern Europe and includes participants from the Liverpool Lung Project in the United Kingdom (11). The 7 countries in that study are participating as separate studies in SYNERGY. Monitoring van Risicofactoren en Gezondheid in Nederland (MORGEN) is a case-control study nested in the prospective European Prospective Investigation Into Cancer and Nutrition (EPIC) cohort in the Netherlands, and the study participants filled in a questionnaire at recruitment (12). Besides MORGEN, all studies have provided data on lifetime smoking habits and complete occupational history. Cases were recruited from hospitals or cancer registries, and the case definition varied slightly across the original studies. In most studies, cases were eligible if they: 1) were younger than 75 years; 2) had been a resident of the study area for at least 1 year, and 3) had a final diagnosis of lung cancer confirmed by histology or cytology. Controls were recruited from the general population (81%) or hospitals (19%) and were individually matched or frequency matched to cases by sex and age (± 3 years). Information was collected predominantly by interviews with the subjects themselves, though next-of-kin respondents were accepted in LUCAS (Sweden), Investigations Cancers Respiratoires et Environnement (ICARE; France), a case-control study of environmental causes of lung cancer in Montreal, Canada, the study in Hong Kong (China), and Occupational Cancer in New Zealand (OCANZ; New Zealand) if subjects were unavailable (9.1% of cases, 6.6% of controls). In most studies, face-to-face interviews (87% of study population) were conducted; however, in LUCAS and MORGEN, questionnaires were sent via mail, and in the study in Hong Kong, the study in Montreal, a lung cancer case-control study in Toronto, Canada, and OCANZ, parts of the study populations were interviewed via telephone. More information about the SYNERGY project is available at <http://synergy.iarc.fr/> and in previously published papers (13–15). The subtype of lung cancer was classified according to World Health Organization guidelines by pathologists associated with the participating hospitals. Reference pathology was performed for the German cases (16). For the original studies, ethics approval was obtained in accordance with legislation in each country. In addition, for the pooling project, ethics approval was obtained from the International Agency for Research on Cancer Ethics Committee, and therefore the project has been conducted in accordance with the ethics standards in the 1964 Declaration of Helsinki and subsequent amendments.

Identification of hairdressers and barbers

The occupational data was coded or recoded to the International Standard Classification of Occupations issued by International Labour Office in 1968 (17). We studied all hairdressers (International Standard Classification of Occupations: 5-70.20 and 5-70.30), as well as women's hairdressers (International Standard Classification of Occupations: 5-70.20) and barbers (International Standard Classification of Occupations: 5-70.30) separately. Women's hairdressers cut

Table 1. Description of the Studies Included in the SYNERGY Project, 1985–2010

| First Author, Year (Reference No.) | Study (Short Names) | Country | Data Collection Period | Cases | | Controls | | Source of Controls | Data Source | Interviewee |
|---------------------------------------|------------------------|----------------|------------------------------|--------|---------------------|----------|---------------------|-----------------------|----------------|-------------|
| | | | | No. | Response Rate, % | No. | Response Rate, % | | | |
| Bruske-Hohlfeld, 2000 (23) | AUT-Munich | Germany | 1990–1995 | 3,180 | 77 | 3,249 | 41 | P | I | S |
| Jöckel, 1998 (24) | HdA | Germany | 1988–1993 | 1,004 | 69 | 1,004 | 68 | P | I | S |
| Consonni, 2010 (25) | EAGLE | Italy | 2002–2005 | 1,943 | 87 | 2,116 | 72 | P | I | S |
| Richiardi, 2004 (26) | TURIN/VENETO | Italy | 1990–1994 | 1,132 | 79 | 1,553 | 80 | P | I | S |
| Fortes, 2003 (27) | ROME | Italy | 1993–1996 | 347 | 74 | 365 | 63 | H | I | S |
| Stücker, 2002 (28) | LUCA | France | 1989–1992 | 309 | 98 | 302 | 98 | H | I | S |
| Kazma, 2012 (29) | PARIS | France | 1988–1992 | 173 | 95 | 234 | 95 | H | I | S |
| Guida, 2011 (30) | ICARE | France | 2001–2007 | 2,926 | 87 | 3,555 | 81 | P | I | S and NOK |
| Lopez-Cima, 2007 (31) | CAPUA | Spain | 2000–2010 | 875 | 91 | 838 | 96 | H | I | S |
| Riboli, 1997 (12) | MORGEN ^a | Netherlands | 1993–1997 | 71 | N/A | 202 | N/A | P | Q | S |
| Scelo, 2004 (32) | INCO | Czech Republic | 1999–2002 | 304 | 94 | 453 | 80 | H | I | S |
| Scelo, 2004 (32) | INCO | Hungary | 1998–2001 | 402 | 90 | 315 | 100 | H | I | S |
| Scelo, 2004 (32) | INCO | Poland | 1998–2002 | 800 | 88 | 841 | 88 | P and H | I | S |
| Scelo, 2004 (32) | INCO | Slovakia | 1998–2002 | 346 | 90 | 285 | 84 | H | I | S |
| Scelo, 2004 (32) | INCO | Romania | 1998–2002 | 181 | 90 | 228 | 99 | H | I | S |
| Scelo, 2004 (32) | INCO | Russia | 1998–2001 | 600 | 96 | 580 | 90 | H | I | S |
| Scelo, 2004 (32) | INCO-LLP | United Kingdom | 1998–2005 | 442 | 78 | 918 | 84 | P | I | S |
| Gustavsson, 2000 (33) | LUCAS | Sweden | 1985–1990 | 1,042 | 87 | 2,356 | 85 | P | Q | S and NOK |
| Corbin, 2011 (34) | OCANZ | New Zealand | 2003–2009 | 457 | 53 | 792 | 48 | P | I and T | S and NOK |
| Ramanakumar, 2007 (35) | MONTREAL | Canada | 1996–2002 | 1,203 | 85 | 1,509 | 69 | P | I and T | S and NOK |
| Brenner, 2012 (36) | TORONTO | Canada | 1997–2002 | 425 | 62 | 910 | 71 | P and H | I and T | S |
| Tse, 2012 (37) | HONG KONG | China | 2003–2007 | 1,207 | 96 | 1,069 | 48 | P | I and T | S and NOK |
| Overall | | | 1985–2010 | 19,369 | 82 | 23,674 | 67 | | | |

Abbreviations: AUT-Munich, Arbeit und Technik–Munich; CAPUA, Cancer de Pulmon en Asturias; EAGLE, Environment and Genetics in Lung Cancer Etiology; H, control subjects enrolled from hospitals; HdA, Humanisierung des Arbeitslebens; HONG KONG, male lung cancer, occupational exposures, and smoking—a case-control study in Hong Kong; I, face-to-face interview; ICARE, Investigations Cancers Respiratoires et Environnement; INCO, International Agency for Research on Cancer Multicenter Case-Control Study of Occupation, Environment, and Lung Cancer in Central and Eastern Europe; LLP, Liverpool Lung Project; LUCA, study of lung cancer in France; LUCAS, Lungcancer i Stockholm; MONTREAL, Montreal case-control study of environmental causes of lung cancer; MORGEN, Monitoring van Risicofactoren en Gezondheid in Nederland; N/A, not applicable; NOK, next-of-kin, for example, husband or wife of the study participant; OCANZ, Occupational Cancer in New Zealand; P, control subjects enrolled from the general population; PARIS, lung cancer study in Paris; Q, self-administered questionnaire; ROME, Rome lung cancer case-control study; S, study participant; T, over-the-phone interview; TORONTO, Toronto lung cancer (case-control) study; TURIN/VENETO, population-based case-control study of lung cancer in the city of Turin and in the Eastern part of Veneto Region.

^a Nested case-control study.

Table 2. General Characteristics of Hairdressers/Barbers and Nonhairdressers/Nonbarbers in the SYNERGY Project, 1985–2010

| Characteristics | Hairdressers | | | | Nonhairdressers | | | |
|--|--------------------------|------|--------------------------|------|--------------------------|------|--------------------------|------|
| | Cases | | Controls | | Cases | | Controls | |
| | No. | % | No. | % | No. | % | No. | % |
| Men | | | | | | | | |
| Total | 100 | | 107 | | 15,095 | | 18,109 | |
| Age, years | 64.7 (8.3) ^a | | 64.0 (8.9) ^a | | 62.7 (9.0) ^a | | 62.2 (9.5) ^a | |
| ≤40 | 1 | 1.0 | 1 | 0.9 | 200 | 1.3 | 398 | 2.2 |
| 41–50 | 4 | 4.0 | 7 | 6.5 | 1,362 | 9.0 | 1,857 | 10.2 |
| 51–60 | 19 | 19.0 | 26 | 24.3 | 4,106 | 27.2 | 4,698 | 25.9 |
| 61–70 | 52 | 52.0 | 46 | 43.0 | 6,206 | 41.1 | 7,373 | 40.7 |
| 71–80 | 23 | 23.0 | 25 | 23.4 | 3,165 | 21.0 | 3,736 | 20.6 |
| >80 | 1 | 1.0 | 2 | 1.9 | 56 | 0.4 | 47 | 0.3 |
| Smoking status | | | | | | | | |
| Never | 4 | 4.0 | 24 | 22.4 | 464 | 3.1 | 4,707 | 26.0 |
| Former | 28 | 28.0 | 49 | 45.8 | 4,887 | 32.4 | 7,925 | 43.8 |
| Current | 68 | 68.0 | 34 | 31.8 | 9,742 | 64.5 | 5,473 | 30.2 |
| Pack-years of cigarette smoking ^b | 48.8 (31.0) ^a | | 32.0 (26.5) ^a | | 43.8 (28.2) ^a | | 27.4 (24.3) ^a | |
| Time since quitting smoking | 5.0 (9.5) ^a | | 11.1 (12.9) ^a | | 5.2 (9.1) ^a | | 11.7 (13.6) ^a | |
| Current smoker | 68 | 68.0 | 34 | 31.8 | 9,742 | 64.5 | 5,473 | 30.2 |
| Former smoker | | | | | | | | |
| 2–7 years | 10 | 10.0 | 10 | 9.4 | 1,641 | 10.9 | 1,170 | 6.5 |
| 8–15 years | 7 | 7.0 | 9 | 8.4 | 1,470 | 9.7 | 1,809 | 10.0 |
| 16–25 years | 6 | 6.0 | 17 | 15.9 | 1,091 | 7.2 | 2,220 | 12.3 |
| ≥26 years | 5 | 5.0 | 12 | 11.2 | 609 | 4.0 | 2,441 | 13.5 |
| Never smoker | 4 | 4.0 | 24 | 22.4 | 464 | 3.1 | 4,707 | 26.0 |
| List A | | | | | | | | |
| Ever | 9 | 9.0 | 4 | 3.7 | 2,130 | 14.1 | 1,663 | 9.2 |

Table continues

and dress primarily women's hair or can serve mixed customers, including men and children. Barbers cut and dress the hair of men and shave or trim their beards. Thirteen participants had worked as both women's hairdressers and barbers and therefore contributed to both subanalyses.

Statistical analysis

Differences in mean lifetime smoking consumption (log pack-years) between hairdressers and nonhairdressers among ever-smoking control subjects were evaluated with the *t* test. The Pearson χ^2 test was used to compare the distributions in hairdressers versus nonhairdressers with regard to smoking status and ever having been employed in a job with known lung cancer risk. Odds ratios and 95% confidence intervals of lung cancer risk were estimated with unconditional logistic regression. We stratified analyses by sex, types of hairdressers, and calendar period of employment. We divided the period into 2 segments and chose the cutpoint at the median start of first employment as hairdresser/barber in our study population (1954) to ensure a sufficient number of persons in each

category. With regard to histology, we first included all lung cancers (including all cell types, unknown cell type, and mixed cell types). We thereafter looked at each of the major cell types separately, using the same controls and excluding all cases with another or unknown or mixed cell type. Duration of employment was studied in categories, with tertiles based on the distribution of employment duration of all hairdressers in the control population. Subjects who had never worked as a hairdresser or barber comprised the reference group. *P* for trend was calculated with the maximum likelihood estimates based on the categorical variables. Odds ratios were adjusted for potential confounders in a stepwise manner: Odds ratio 1 was adjusted for log(age) and study (22 study centers). Odds ratio 2 was further adjusted for smoking in pack-years (log(cigarette pack-years + 1)) as a continuous variable and time since quitting smoking all types of tobacco as a categorical variable (current smokers; former smokers who stopped smoking 2–7 years, 8–15 years, 16–25 years, or ≥26 years before interview or diagnosis; or never smokers). Odds ratio 3 was additionally adjusted for ever employment in a job with known lung cancer risk (list A), as a proxy for exposure

Table 2. Continued

| Characteristics | Hairdressers | | | | Nonhairdressers | | | |
|--|--------------------------|------|--------------------------|------|--------------------------|------|--------------------------|------|
| | Cases | | Controls | | Cases | | Controls | |
| | No. | % | No. | % | No. | % | No. | % |
| Women | | | | | | | | |
| Total | 70 | | 60 | | 3,585 | | 5,000 | |
| Age, years | 56.7 (11.2) ^a | | 55.1 (10.5) ^a | | 60.6 (10.1) ^a | | 60.3 (11.2) ^a | |
| ≤40 | 7 | 10.0 | 3 | 5.0 | 99 | 2.8 | 266 | 5.3 |
| 41–50 | 15 | 21.4 | 19 | 31.7 | 543 | 15.2 | 699 | 14.0 |
| 51–60 | 19 | 27.1 | 20 | 33.3 | 1,043 | 29.1 | 1,319 | 26.4 |
| 61–70 | 24 | 34.3 | 15 | 25.0 | 1,227 | 34.2 | 1,733 | 34.7 |
| 71–80 | 5 | 7.1 | 2 | 3.3 | 653 | 18.2 | 969 | 19.4 |
| >80 | | | 1 | 1.7 | 20 | 0.6 | 14 | 0.3 |
| Smoking status | | | | | | | | |
| Never | 8 | 11.4 | 21 | 35.0 | 961 | 26.8 | 2,997 | 59.9 |
| Former | 14 | 20.0 | 18 | 30.0 | 680 | 19.0 | 1,019 | 20.4 |
| Current | 48 | 68.6 | 21 | 35.0 | 1,944 | 54.2 | 979 | 19.6 |
| Pack-years of cigarette smoking ^b | 33.9 (22.4) ^a | | 22.1 (22.4) ^a | | 34.5 (23.0) ^a | | 19.3 (18.3) ^a | |
| Time since quitting smoking | 2.3 (4.8) ^a | | 7.2 (10.9) ^a | | 3.4 (7.7) ^a | | 6.9 (11.7) ^a | |
| Current smoker | 48 | 68.6 | 21 | 35.0 | 1,944 | 54.2 | 979 | 19.6 |
| Former smoker | | | | | | | | |
| 2–7 years | 6 | 8.6 | 3 | 5.0 | 246 | 6.9 | 194 | 3.9 |
| 8–15 years | 5 | 7.1 | 5 | 8.3 | 202 | 5.6 | 231 | 4.6 |
| 16–25 years | 3 | 4.3 | 5 | 8.3 | 143 | 4.0 | 281 | 5.6 |
| ≥26 years | | | 3 | 5.0 | 80 | 2.2 | 277 | 5.5 |
| Never | 8 | 11.4 | 21 | 35.0 | 961 | 26.8 | 2,997 | 59.9 |
| List A | | | | | | | | |
| Ever | 3 | 4.3 | 1 | 1.7 | 86 | 2.4 | 58 | 1.2 |

^a Values expressed as mean and standard deviation.

^b Among ever smokers only.

to occupational lung carcinogens. List A is a list of occupations and industries known to present an excess risk of lung cancer, which were identified by Ahrens and Merletti in 1998 (18) and updated by Mirabelli et al. in 2001 (19). Stability of the results was assessed by restricting the analyses to never smokers and by exploring potential heterogeneity with I^2 measuring the variation in risk estimates attributable to heterogeneity between studies, countries, size of the study ($\pm 1,500$ participants), year of the study (end of data collection ± 1995), and type of controls (population, hospital, or mixed). Statistical analyses were conducted in SAS, version 9.2 (SAS Institute, Inc., Cary, North Carolina) and STATA, version 12.1 (StataCorp LP, College Station, Texas). A P value of ≤ 0.05 was considered statistically significant.

RESULTS

Table 1 describes the studies included in the SYNERGY project. Study participants who did not provide complete

data for calculation of smoking pack-years (519 cases and 398 controls) were excluded. Thus, 18,850 cases and 23,276 controls were included in these analyses.

Table 2 shows the characteristics of hairdressers/barbers and nonhairdressers/nonbarbers by sex. Comparison of hairdressers versus nonhairdressers with a focus on ever-smoking control subjects showed that the mean cumulative smoking consumption (log pack-years) was not statistically different among men ($P = 0.21$) or women ($P = 0.80$). However, the frequency distribution across never, former, and current smokers was significantly different between female hairdressers and nonhairdressers ($P < 0.001$ in controls) but not between male hairdressers/barbers and nonhairdressers/nonbarbers ($P = 0.70$ in controls). The proportion of participants having worked in a job with known lung cancer risk (list A) was $< 5\%$ among women overall, and no significant difference was observed between hairdressers and nonhairdressers ($P = 0.72$ in controls). Among men, hairdressers/barbers had less often been employed in a list A job than had nonhairdressers/nonbarbers

($P=0.05$ in controls). Adjustment for list A in the analyses (odds ratio 3) did not influence the results, so only odds ratio 1 and odds ratio 2 are displayed in Table 3.

Lung cancer risk among hairdressers

The proportion of men having worked as women's hairdressers was 0.20% in cases ($n=30$) and 0.22% in controls ($n=40$), and the proportion of men having worked as barbers was 0.51% in cases ($n=77$) and 0.40% in controls ($n=73$) (Table 3). We observed no significant increased risk of lung cancer among male hairdressers, either before or after adjustment for smoking. Among barbers, we observed a nonsignificant trend of increasing odds ratios for lung cancer with longer duration of employment, with odds ratios ranging from 0.83 (95% confidence interval (CI): 0.43, 1.61) to 1.62 (95% CI: 0.88, 2.98), with $P=0.32$.

Employment as a hairdresser/barber with regard to time of first employment did not reveal a different risk pattern: Odds ratio 2 was 0.92 (95% CI: 0.52, 1.63) before 1954 and 0.95 (95% CI: 0.65, 1.38) in or later than 1954. With regard to histology, long-term barbers (>26 years of employment) had an increased odds ratio for adenocarcinoma, on the basis of 12 cases (odds ratio 2 = 2.20, 95% CI: 1.02, 4.77).

A meta-analysis by study resulted in an overall odds ratio 2 of 0.93 (95% CI: 0.67, 1.28), with an I^2 of 0% and $P=0.90$. Further sensitivity analyses among men revealed no significant heterogeneity in risk estimates with regard to country, calendar period of data collection, size of study, or type of control group, and all overall odds ratios remained <1 (data not shown). In never smokers, odds ratio 1 for male hairdressers/barbers was 1.60 (95% CI: 0.53, 4.82), on the basis of 4 cases.

The proportions of women who had ever been employed as hairdressers among cases and controls were 1.89% ($n=69$) and 1.17% ($n=59$), respectively. Only 1 female case and 1 female control had worked as barbers, so women's hairdressers and barbers were not analyzed separately. A significant increased lung cancer risk among hairdressers was observed (odds ratio 1 = 1.65, 95% CI: 1.16, 2.35), which was reduced and no longer statistically significant when adjusted for smoking (odds ratio 2 = 1.12, 95% CI: 0.75, 1.68). The highest odds ratio was observed among those who had worked <8 years as hairdressers. No trend in relation to duration was observed, with $P=0.71$. The risk changed with time period of employment; women who had been employed as hairdressers before 1954 experienced an increased lung cancer risk before and after adjustment for smoking (odds ratio 1 = 3.01, 95% CI: 1.38, 5.59; odds ratio 2 = 2.66, 95% CI: 1.09, 6.47), whereas women employed as hairdressers in or later than 1954 did not (odds ratio 1 = 1.41, 95% CI: 0.94, 2.12; odds ratio 2 = 0.89, 95% CI: 0.56, 1.40). Table 4 shows lung cancer risk by calendar period, duration of employment, and lung cancer histology. Female hairdressers first employed before 1954 experienced increased risk of all major lung cancer types, and the strongest association was observed for adenocarcinoma (odds ratio 2 = 3.10, 95% CI: 1.14, 8.43). Across all lung cancer types, the elevated risks were restricted to the short-term hairdressers (<8 years).

A meta-analysis by study resulted in an overall odds ratio 2 of 1.13 (95% CI: 0.74, 1.73), with an I^2 of 0% and $P=0.84$. The odds ratio 2 for the Arbeit und Technik-Munich (AUT-

Munich) study alone was 3.25 (95% CI: 1.03, 10.23); when AUT-Munich was excluded, the overall odds ratio 2 decreased to 1.07 (95% CI: 0.80, 1.43). Exclusion of AUT-Munich from the analysis of female hairdressers employed before 1954 resulted in an odds ratio 2 of 2.72 (95% CI: 0.93, 8.02) and in an odds ratio 2 of 3.91 (95% CI: 1.22, 12.50) for adenocarcinoma alone. With regard to the calendar period of data collection in the different studies (taking 1995 as cutpoint), we observed an odds ratio 2 of 1.84 (95% CI: 0.85, 3.98) for women enrolled in the earlier studies and an odds ratio 2 of 1.00 (95% CI: 0.63, 1.59) for the more recent studies, with an I^2 of 42% ($P=0.19$). When comparing the risk estimates for the different sources of control subjects, we observed an odds ratio 2 of 1.26 (95% CI: 0.82, 1.95) for population-based case-control studies and odds ratios <1 for hospital-based and mixed case-control studies, with an I^2 of 0% ($P=0.64$). No heterogeneity was observed with regard to country or study size (data not shown). In never smokers, female hairdressers experienced an odds ratio 1 of 1.33 (95% CI: 0.57, 3.08), on the basis of 8 cases.

DISCUSSION

We investigated hairdressers' and barbers' lung cancer risks compared with nonhairdressers/nonbarbers in the SYNERGY population. Our results show the importance of adjusting for the major risk factor of a certain cancer when investigating a presumably less pronounced and less prevalent other potential risk factor and how this can be done efficiently in a pooled data set of independently conducted studies. In our case, an association reported from a large linkage study could no longer be observed.

The odds ratio for women overall (including both time periods) was significantly elevated before adjustment for smoking but not afterward. Female hairdressers were more often smokers than were nonhairdressers, whereas no significant difference in smoking habits was seen between male hairdressers/barbers and nonhairdressers/nonbarbers in this study. In subgroup analyses, our results revealed an increased risk among women with first employment before 1954 and who had worked <8 years as hairdressers but not in women with first employment after 1954 or who had worked long term as hairdressers, and our results revealed no increased risk among men, except for an increased odds ratio for adenocarcinoma in long-term barbers (>26 years).

Strengths of this study include: 1) The study size was large, as is necessary to study a relatively rare occupation in the general population and to stratify the results by sex and type of hairdresser. 2) Most of the original studies were initiated to study occupational risk factors and therefore collected detailed lifetime work histories. 3) Most interviews were conducted face to face with the study participants. 4) We could adjust for smoking, the most important risk factor for lung cancer. Smoking was a confounder, particularly in women. Our adjustment for list A jobs did not reveal a confounding effect, probably because so few hairdressers/barbers worked in list A jobs that overall are more common in men.

All case-control studies can be affected by some degree of recall bias. However, studies in SYNERGY recorded occupational histories but did not solicit direct information

Table 3. Lung Cancer Risk Associated With Hairdressing and Duration of Employment Among 18,850 Cases and 23,276 Controls in the SYNERGY Project, 1985–2010

| Employment | Cases | | Controls | | OR1 | 95% CI | OR2 | 95% CI |
|-----------------------------------|--------|------|----------|------|------|------------|------|------------|
| | No. | % | No. | % | | | | |
| Men | | | | | | | | |
| All hairdressers ^a | | | | | | | | |
| Never | 15,095 | 99.3 | 18,019 | 99.4 | 1.00 | | 1.00 | |
| Ever | 100 | 0.7 | 107 | 0.6 | 1.04 | 0.79, 1.37 | 0.91 | 0.66, 1.25 |
| <8 years | 34 | 0.2 | 33 | 0.2 | 1.14 | 0.70, 1.85 | 0.82 | 0.48, 1.41 |
| 8–26 years | 26 | 0.2 | 34 | 0.2 | 0.83 | 0.50, 1.39 | 0.68 | 0.38, 1.22 |
| >26 years | 40 | 0.3 | 40 | 0.2 | 1.14 | 0.73, 1.78 | 1.26 | 0.74, 2.12 |
| Women's hairdressers ^b | | | | | | | | |
| Never | 15,095 | 99.8 | 18,109 | 99.8 | 1.00 | | 1.00 | |
| Ever | 30 | 0.2 | 40 | 0.2 | 0.84 | 0.52, 1.35 | 0.69 | 0.40, 1.19 |
| <8 years | 18 | 0.1 | 14 | 0.1 | 1.37 | 0.68, 2.78 | 1.12 | 0.50, 2.52 |
| 8–26 years | 19 | 0.1 | 13 | 0.1 | 0.77 | 0.33, 1.81 | 0.61 | 0.24, 1.60 |
| >26 years | 3 | 0.0 | 13 | 0.1 | 0.29 | 0.08, 1.01 | 0.25 | 0.06, 0.98 |
| Barber hairdressers ^c | | | | | | | | |
| Never | 15,095 | 99.5 | 18,109 | 99.6 | 1.00 | | 1.00 | |
| Ever | 77 | 0.5 | 73 | 0.4 | 1.17 | 0.84, 1.61 | 1.09 | 0.76, 1.59 |
| <8 years | 21 | 0.1 | 22 | 0.1 | 1.06 | 0.58, 1.94 | 0.83 | 0.43, 1.61 |
| 8–26 years | 23 | 0.2 | 24 | 0.1 | 1.05 | 0.59, 1.87 | 0.91 | 0.47, 1.74 |
| >26 years | 33 | 0.2 | 27 | 0.2 | 1.35 | 0.81, 2.27 | 1.62 | 0.88, 2.98 |
| Women | | | | | | | | |
| All hairdressers ^a | | | | | | | | |
| Never | 3,585 | 98.1 | 5,000 | 98.8 | 1.00 | | 1.00 | |
| Ever | 70 | 1.9 | 60 | 1.2 | 1.65 | 1.16, 2.35 | 1.12 | 0.75, 1.68 |
| <8 years | 37 | 1.0 | 26 | 0.5 | 2.07 | 1.25, 3.46 | 1.28 | 0.72, 2.29 |
| 8–26 years | 15 | 0.4 | 20 | 0.4 | 1.00 | 0.51, 1.97 | 0.93 | 0.42, 2.02 |
| >26 years | 18 | 0.5 | 13 | 0.3 | 1.96 | 0.95, 4.03 | 1.10 | 0.48, 2.51 |

Abbreviations: CI, confidence interval; ISCO, International Standard Classification of Occupations 1968; OR1, odds ratio adjusted for age and study; OR2, odds ratio additionally adjusted for cigarette pack-years and time since quitting smoking.

^a ISCO: 5-70.20 and 5-70.30.

^b ISCO: 5-70.20.

^c ISCO: 5-70.30.

on the use of specific chemicals, which is especially prone to positive recall bias, and no emphasis had been put on employment as barber or hairdresser. Furthermore, no special alert was present in the general population on a possible cancer risk linked to these occupations, which could have induced cancer patients to report them more frequently than controls. Next-of-kin were interviewed instead of the index subject in a few studies, but these represented a small proportion (<10%) of the cases. Recall bias in the smoking history could have resulted in residual confounding when adjustment was made for smoking; however, the smoking-adjusted increased risks in women are unlikely to be due to residual confounding by smoking. Low response rates among control subjects in some studies might have resulted in selection bias if hairdressers were more likely than other control subjects to be nonrespondents. AUT-Munich, with a response rate of 41%

in controls, was the only single study with a significantly elevated odds ratio in female hairdressers. In the pooled data set and in *Humanisierung des Arbeitslebens* (HdA), another German study, the lung cancer risk was not increased.

A limitation of the present study was the lack of information on determinants of exposure—for example, to which specific agents these groups of hairdressers were exposed and whether protective devices had been used. The SYNERGY data were collected between 1985 and 2010, and the time of working as a hairdresser covered the period from around the 1930s onward. Both the types and the quantities of products used by hairdressers have changed substantially during this time span, and some exposures might have increased in the late 1940s before they were reduced or changed again in the mid-1980s. Among such exposures, worth mentioning are the use of talc (talc products before 1973 could have been contaminated

Table 4. Lung Cancer Risk Associated With Hairdressing and Duration of Employment by Start of First Employment in Women Overall (n = 8,585) and in Women by Major Lung Cancer Cell Types in the SYNERGY Project, 1985–2010

| Employment as Hairdressers ^a | Controls | | All Lung Cancer | | | Adenocarcinoma | | | Squamous Cell Carcinoma | | | Small Cell Carcinoma | | |
|---|----------|-------|-----------------|------|-------------|----------------|-------|-------------|-------------------------|-------|--------------|----------------------|-------|--------------|
| | No. | No. | No. | OR2 | 95% CI | No. | OR2 | 95% CI | No. | OR2 | 95% CI | No. | OR2 | 95% CI |
| Never | 5,000 | 3,585 | 1,547 | 1.00 | | 736 | 1.00 | | 573 | 1.00 | | | | |
| Ever (≤1953) | 9 | 22 | 10 | 2.66 | 1.09, 6.47 | 10 | 3.10 | 1.14, 8.43 | 7 | 1.80 | 0.48, 6.73 | 3 | 2.51 | 0.44, 14.33 |
| <8 years | 2 | 15 | 6 | 12.9 | 2.67, 62.44 | 6 | 14.66 | 2.75, 78.02 | 5 | 14.09 | 1.63, 121.62 | 2 | 50.99 | 4.63, 561.67 |
| 8–26 years | 5 | 4 | 1 | 0.67 | 0.13, 3.37 | 1 | 0.50 | 0.05, 5.45 | 2 | 0.56 | 0.07, 4.85 | 1 | 0.78 | 0.03, 21.38 |
| >26 years | 2 | 3 | 3 | 0.56 | 0.09, 3.48 | 3 | 1.46 | 0.24, 9.07 | 0 | | | 0 | | |
| Ever (>1953) | 50 | 48 | 16 | 0.89 | 0.56, 1.40 | 16 | 0.73 | 0.39, 1.35 | 9 | 1.08 | 0.47, 2.45 | 13 | 1.26 | 0.60, 2.65 |
| <8 years | 24 | 22 | 8 | 0.67 | 0.35, 1.30 | 8 | 0.65 | 0.28, 1.54 | 3 | 0.49 | 0.13, 1.91 | 6 | 1.12 | 0.40, 3.17 |
| 8–26 years | 15 | 11 | 6 | 1.02 | 0.42, 2.50 | 6 | 1.48 | 0.53, 4.12 | 1 | 0.91 | 0.11, 7.85 | 3 | 1.82 | 0.35, 9.37 |
| >26 years | 11 | 15 | 2 | 1.28 | 0.51, 3.17 | 2 | 0.30 | 0.06, 1.46 | 5 | 2.92 | 0.80, 10.65 | 4 | 1.23 | 0.32, 4.77 |

Abbreviations: CI, confidence interval; ISCO, International Standard Classification of Occupations 1968; OR2, odds ratio adjusted for age, study, cigarette pack-years, and time since quitting smoking.

^a ISCO 5-70.20 and 5-70.30.

by asbestos (20)), propellants, coloring agents, and passive smoking.

We observed an increased risk of lung cancer among female hairdressers first employed before 1954, but it was restricted to women working <8 years in this job, and we observed a somewhat higher odds ratio for the earlier studies than for the more recent studies, which could indicate that hairdressers previously experienced an increased risk of lung cancer. This result should, however, be interpreted with caution because the number of hairdressers employed before 1954 was low (22 cases, 9 controls), resulting in wide confidence intervals, and risks were increased only in women with the shortest duration of employment. Also, this hypothesis does not get support from cohort studies. A Finnish study linked census data with cancer registry data for 1970–1987 and found 13 lung cancer cases among female hairdressers, resulting in a standardized incidence ratio of 1.72 (95% CI: 0.92, 2.94). The standardized incidence ratio for lung cancer was below unity for 1970–1981 and was significantly elevated in the later period, 1982–1987 (standardized incidence ratio = 2.92, 95% CI: 1.46, 5.22) (21). A similar study in Sweden investigated hairdressers in 1960–1990 and found an increased risk of lung cancer in both male and female hairdressers, with standardized incidence ratios of 1.38 (95% CI: 1.16, 1.68) and 1.35 (95% CI: 1.15, 1.58), respectively. When they analyzed the earlier period (before 1960) separately, the risk was similar for men (standardized incidence ratio = 1.41, 95% CI: 1.18, 1.68) and somewhat lower (standardized incidence ratio = 1.22, 95% CI: 1.00, 1.47) for female hairdressers (22). However, those results are not directly comparable with the present study, because employment as barbers/hairdressers is often abandoned early and census information refers to the current job, so it is not likely that the employment periods in these cohorts correspond to the employment periods in SYNERGY.

We observed an association in men between long-term employment as a barber and adenocarcinoma, but not squamous cell carcinoma or small cell carcinoma. Interestingly, in the Nordic Occupational Cancer Study (NOCCA), a similar observation was made in the combined Nordic population followed up during 1961–2005. Among male hairdressers, the standardized incidence ratio for all lung cancer types combined was 1.22 (95% CI: 1.12, 1.33), but it was 1.33 (95% CI: 1.10, 1.60) for adenocarcinoma alone. The standardized incidence ratio for all lung cancer types combined in women hairdressers was 1.30 (95% CI: 1.19, 1.42), and it was 1.38 (95% CI: 1.19, 1.61) for adenocarcinoma (4).

Our results in never smokers (slightly elevated nonsignificant odds ratios) were limited by their small numbers.

In summary, our observed association with adenocarcinoma in both men and women is supported by results from the Nordic Occupational Cancer Study, whereas our other findings are not directly comparable with results in the previous literature. Although the statistical power to detect an increased risk was limited in the subanalyses, and our extensive stratification leading to multiple comparisons could have resulted in high variation of the risk estimates and spurious findings by chance, it was important to conduct these analyses because this is a rare opportunity to study hairdressers in a population-based study with detailed smoking information covering such a long calendar period.

Our results suggest that the increased lung cancer risk among female hairdressers is due to smoking behavior among this occupational group and is not directly related to occupational exposure. Single elevated risks among the many subgroup analyses should be interpreted with caution unless replicated in other studies.

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