- 1 Socioeconomic position and outdoor nitrogen dioxide (NO₂) exposure in Western Europe: a
- 2 multi-city analysis

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ABSTRACT

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Background: Inconsistent associations between socioeconomic position (SEP) and outdoor air 74 pollution have been reported in Europe, but methodological differences prevent any direct 75 between-study comparison. 76 Objectives: Assess and compare the association between SEP and outdoor nitrogen dioxide 77 (NO₂) exposure as a marker of traffic exhaust, in 16 cities from eight Western European 78 countries. 79 80 Methods: Three SEP indicators, two defined at individual-level (education and occupation) and one at neighborhood-level (unemployment rate) were assessed in three European 81 multicenter cohorts. NO₂ annual concentration exposure was estimated at participants' 82 83 addresses with land use regression models developed within the European Study of Cohorts for Air Pollution Effects (ESCAPE; http://www.escapeproject.eu/). Pooled and city-specific 84 linear regressions were used to analyze associations between each SEP indicator and NO₂. 85 Heterogeneity across cities was assessed using the Higgins' I-squared test (I²). 86 Results: The study population included 5692 participants. Pooled analysis showed that 87 participants with lower individual-SEP were less exposed to NO₂. Conversely, participants 88 living in neighborhoods with higher unemployment rate were more exposed. City-specific 89 results exhibited strong heterogeneity (I²>76% for the three SEP indicators) resulting in 90 variation of the individual- and neighborhood-SEP patterns of NO₂ exposure across cities. 91 92 The coefficients from a model that included both individual- and neighborhood-SEP indicators were similar to the unadjusted coefficients, suggesting independent associations. 93 94 Conclusions: Our study showed for the first time using homogenized measures of outcome and exposure across 16 cities the important heterogeneity regarding the association between 95 SEP and NO₂ in Western Europe. Importantly, our results showed that individual- and 96 97 neighborhood-SEP indicators capture different aspects of the association between SEP and

exposure to air pollution, stressing the importance of considering both in air pollution health 98 effects studies. 99 100 Keywords: Europe, socioeconomic position, air pollution, environmental inequality 101 102 **ABREVIATIONS** 103 ECRHS: European Community Respiratory Health Survey 104 EGEA: French Epidemiological family-based study of the Genetics and Environment of 105 106 Asthma ESCAPE: European Study of Cohorts for Air Pollution Effects 107 LUR: land use regression 108 109 MAUP: modifiable area unit problem 110 NO₂: Nitrogen dioxide OC: occupational class 111 PM: Particulate matter 112 SAPALDIA: Swiss Cohort Study on Air Pollution and Lung and Heart Diseases in Adults 113 SEP: socioeconomic position

1. INTRODUCTION

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Environmental inequality refers to a differential distribution of environmental hazards across socioeconomic or socio-demographic groups (1). Historically, research on environmental inequality has emerged in the United States (US) following the Environmental Justice Movement (2–5). Repeatedly, US studies reported that lower socioeconomic or minority groups were more likely to be exposed to higher traffic-related air pollution exposure such as nitrogen dioxide (NO₂) or particulate matter (PM) (6). However, results from US studies cannot be extended to European countries because of very different socio-spatial characteristics, specifically in urban areas (7). For example, one of the main differences is that in general in most US cities, lower socioeconomic groups tend to live downtown when upper socioeconomic groups reside in the suburbs. In European cities, compared to US, social segregation is lower and lower socioeconomic groups rather live on the outskirts of the city (7).In Europe, a rather limited number of studies compared to US had investigated the association between socioeconomic position (SEP) and air pollution, mainly in the UK first and then in other European countries (6,8). Inconsistent results have been reported in the European literature (9). Some studies reported that populations with low SEP are more exposed to outdoor air pollution (10–14) while other studies reported an inverse association (15–18). Nonlinear association (higher exposure in middle class) (19) and no association (20) were also reported. Inconsistent results were also reported within the same country, for instance in France or Spain (20–23). However, these studies were difficult to compare with each other because they used different methodologies to assess air pollution exposure or to define SEP (6,24). Moreover, most studies relied on ecological data that can raise methodological issues such as ecological fallacy, modifiable area unit problem (MAUP) or spatial autocorrelation (19,25). Few studies used individual-level data (i.e. air pollution exposure at residential

address and individual-level SEP) or multilevel data (i.e. SEP estimated at individual- and area-level) (15,17,26–30). Recent evidence showed the importance of considering SEP at both individual and area levels because they are independently associated with health outcomes (6,10,31–33).

More generally, the association between SEP and air pollution still needs to be investigated in Europe (6,24) as SEP is one of the major potential determinants of variability in the association between air pollution and health (2,34,35).

Within the framework of the multicenter European Study of Cohorts for Air Pollution Effects (ESCAPE) (36), we had the opportunity to tackle this research gap using outdoor NO₂ annual concentrations at participants' home addresses estimated from standardized procedures across a large range of European cities (36). The main objective of the present analysis was to test the environmental justice hypothesis that people with lower SEP (defined at both individual and neighborhood level) were more exposed to traffic related air pollution exposure than people with higher SEP in Western Europe.

2. MATERIALS AND METHODS

156 2.1. Study population

This cross-sectional study included participants of three multicenter epidemiological
European cohorts that had previously collaborated together (37) and were involved in the
ESCAPE study: the French Epidemiological family-based study of the Genetics and
Environment of Asthma (EGEA2) (2003–2007) (38), and two population-based studies: the
European Community Respiratory Health Survey (ECRHSII) (1999–2002) (39) and The
Swiss Cohort Study on Air Pollution and Lung and Heart Diseases in Adults (SAPALDIA2)
(2001-2003) (40). Details on each cohort are given elsewhere (38–40) and summarized in the

supplementary materials. For the three cohorts, information on participants were collected from detailed, standardized and validated questionnaires completed by face-to-face interviews.

Initially, the ESCAPE study included a subsample of the three cohorts (n=9556 participants, Figure 1) from 20 urban areas of eight Western European countries. Of these 20 areas, we were able to recover homogenized SEP data at individual and neighborhood level for 16 (n=5692 participants: 4002, 1078 and 612 in ECRHS, EGEA and SAPALDIA respectively; Figure 1) including Norwich, Ipswich (Great Britain; GB); Antwerp (Belgium; BE); Paris, Lyon, Grenoble, Marseille (France; FR); Geneva, (Switzerland; CH); Verona, Pavia, Turin (Italy; IT); Oviedo, Galdakao, Barcelona, Albacete, Huelva (Spain; SP) (Figure S1). The areas covered by ESCAPE were of substantially different sizes (Table S1) with a range of density population from 152 to 21154 inhabitants/km² (41). Most of them could be defined as metropolitan areas (large cities with surrounding smaller suburban communities) but some areas were restricted to a single city (municipality). For purposes of clarity, we refer to these different areas as "cities".

2.2. NO₂ exposure assessment

We considered nitrogen dioxide (NO₂) as a marker of near-road traffic-related air pollution (42). The major sources of NO2 are motorized road traffic, industry, shipping and heating (41). In the framework of ESCAPE, a single harmonized exposure assessment protocol has been developed to estimate the NO₂ annual concentrations. A common protocol described in detail in Beelen et al. was used to ensure high standardization of all procedures (i.e. measurement and estimation model) across the study areas (36). Briefly, in each city covered, two-week integrated NO₂ measurements at approximately 40 urban sites were made in three different seasons over a one-year period between 2008 and 2011. City-specific land use

regression (LUR) models (see supplementary materials) were developed to explain the spatial variation of NO_2 using a variety of geographical data including traffic, population and land use variables. The model explained variances (R^2) of the LUR models ranged from 55% in Huelva to 92% in Pavia, 10 out of the 16 cities have a R^2 above 75% (36). These LUR models were used to assign estimates of NO_2 annual average concentrations at each participant's geocoded residential address. Back-extrapolated estimates were also derived because ESCAPE measurement campaigns took place after the health surveys for the three cohorts (43). Correlations between back-extrapolated and non-back-extrapolated concentrations were high (Pearson correlation coefficient=0.95) so we only considered the non-back-extrapolated data in the present analysis.

- 2.3. Markers of socioeconomic position
- We indexed SEP defined at two different levels:
- 202 2.3.1. Individual-level SEP
 - We characterized individual-level SEP based on educational level and occupational class. For the three cohorts, educational level corresponded to the age at completion of full-time education. We categorized the continuous educational variable into country-specific tertiles (high, medium and low). Occupational class was based on the longest job held between baseline and follow-up (in average 10–12 years), and categorized in five classes according to the International Standard Classification of Occupation (ISCO-1988) (44): Manager and Professional (Occupational Class-I); Technician & associate (OC-II); Other non-manual (OC-III); Skilled, semi-skilled and unskilled manual (OC-IV) and "not in labor force".
- 2.3.2. Neighborhood-level SEP

To characterize the socioeconomic residential environment of the participants, we used the neighborhood unemployment rate (i.e. proportion of unemployed persons of the labor force). The neighborhood level corresponded to the smallest geographical level unit (with a population size ranging from 169 to 2000 inhabitants) with census-based data available in the different countries (see Table S2 for neighborhood specific characteristics). We obtained the unemployment rate variable from 2001 national censuses (except for France: 2008 and Switzerland: 2006). As the magnitude of the unemployment rate varied across European countries, we standardized it using country-specific z-scores to take this variability into account.

- 2.4. Strategy of analysis
- *2.4.1. Main analyses*
- The strategy of analysis aimed to test the hypothesis that the NO₂ annual concentration (dependent variable) differs according to the individual- and neighborhood- SEP of the participants (explanatory variables).

We performed analyses considering first the pooled dataset and then each city separately, due to the heterogeneity of the associations between SEP and air pollution among the cities (assessed with the Higgins' I-squared test (I^2) (45)) We ran several multilevel linear regression models (Table S3) with neighborhood random effects (plus city random effects for the pooled dataset) including one individual SEP indicator (education or occupation) mutually adjusted for neighborhood unemployment rate. In the supplementary materials, we present the results for the single-level linear regression models that ignore the nested structure of the observations.

We transformed NO_2 using a natural log transformation to obtain a normally distributed variable. For ease of interpretation, we converted the regression coefficients (β s) into percent change (and 95% Confidence Interval (CI)) per one unit increase in the explanatory factor using the formula [exp(β)-1]*100 (a 95% CI which does not include zero indicates the presence of significant differences). The considered unit for unemployment rate was 1 standard deviation (SD). For the individual-level SEP variables, we considered each subgroup and tested the statistical differences of the coefficients against the highest group (thus reference group were high educational level and OC-I for occupational class). We deliberately did not show results for participants who were not in the labor force as this class was too heterogeneous to draw any kind of conclusion (i.e. housepersons, unemployed, not working because of poor health, full-time student and retired). This category was excluded to assess the trend across the occupational groups.

2.4.2. Additional analyses

We ran a sensitivity analysis using logistic regression models considering high vs. low exposure (high exposure was defined as an exposure above the 75th percentile of the distribution for each city). All models were adjusted for cohort, age and sex. We checked for potential interactions between SEP and sex, SEP and age and between individual- and neighborhood-level SEP (supplementary materials). Analyses were conducted using R statistical software (Version 3.0.3) and SAS 9.3.

As pointed out above some "cities" included in this analysis had a wide geographic coverage. For example, the city labelled "Paris" (FR) covered actually the metropolitan area of Paris-Region (*i.e.* 12,000 km²). Therefore, we ran a sensitivity analysis by examining more in detail this area: instead of considering participants of Paris in only one area, we considered three distinctive areas (i.e. City of Paris, the inner-suburbs and the outer-suburbs) defined by particular sociodemographic and geographic situations that could influence the association

between SEP and air pollution. The methods and results are presented in detail in the supplementary materials and discussed in the main article.

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3. RESULTS

3.1. Study population characteristics

The study population (Table 1a) was composed of 48% males, with a mean age (±standard deviation; \pm SD) of 44 (\pm 11) years. Regarding the NO₂ distribution, we found substantial variability between cities with a mean ranging from 21 (\pm 5) (Pavia; IT) to 57 (\pm 14) µg m⁻³ (Barcelona; ES). Substantial variability was also found within cities. The average range for NO₂ (difference between the highest and the lowest annual average) within each area was 50.3 µg m⁻³. The largest variation for NO₂ was found in the two largest cities Paris (FR) (85.0) and Barcelona (SP) (92.8). Regarding the socioeconomic characteristics of the population (Table 1b), participants completed their education on average at age 20 (± 4) years. The proportion of manual workers ranged from 6% (Paris; FR) to 38% (Galdakao; SP) and was generally higher in the Spanish cities. On average, participants with lower educational attainment were employed in less skilled occupations (p-value for trend <0.001) (Table S4). The neighborhood unemployment rate varied from 3% (Pavia; IT) to 22% (Huelva; SP). Participants with lower educational attainment or less skilled occupations were more likely to live in neighborhoods with higher unemployment rate. However, the associations did not reach the level of significance in 7 and 6 out of the 16 cities for education and occupation respectively (Tables S5a-S5b).

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3.2. Pooled results

Pooled results are shown in Table 2. In the model taking into account only clustering within cities, low educational level and manual occupations were associated with a lower NO₂

exposure (Percent difference (95% CI) Low vs. high educational level= -6.9% (-9.1; -4.7); OC-IV vs. OC-I=-5.6% (-8.2; -3.0)). Conversely, higher neighborhood unemployment rate was associated with higher NO₂ exposure (7.3% (6.2; 8.5) per 1 SD increase in the unemployment rate). The introduction of individual- and neighborhood-SEP in the same model did not substantially alter effect estimates (Low vs. High educational level= -8.7% (-10.8; -6.5) and 7.8% (6.7; 8.9) per 1 SD increase in the unemployment rate). Accounting for both city and neighborhood clustering decreased the effect size of both the individual- and neighborhood-SEP. Associations remained significant for educational level and the unemployment rate.

3.3. City-specific results

In the city-specific analyses using standard linear regression models (Table S4), associations with NO $_2$ were highly heterogeneous for all SEP indicators (I 2 >76%, p<0.001). Using multilevel linear regression models, individual-SEP was weakly or not associated with NO $_2$ exposure for most cities (14 out of 16 cities). For educational level (Table 3a), significant associations were only found in Lyon (FR) (Low vs. High =-3.6 (-12.3; -5.9)) and Verona (IT) (-16.1 (-26.5; -4.3)). For occupational class (Table 3b), significant associations were found for the middle class in Paris (FR) (OC-III vs. OC-I= -3.3 (-6.4; -0.1) and Oviedo (-8.7 (-15.7; -1.2)). Living in a neighborhood with higher unemployment rate was associated with higher NO $_2$ exposure (regardless of the individual-SEP marker included in the model) in 11 out of 16 cities. In Oviedo (ES) and Barcelona (ES) an inverse association was observed.

3.4. Additional analyses

Results from the logistic regression models (high vs. low exposure) were consistent with the linear regression ones for the educational level (Table S6a) as well for occupational class (Table S6b).

In Paris-Region (FR), when considering participants in three distinctive areas (i.e. city of Paris, inner suburbs and outer suburbs; supplementary materials), participants with lower educational level or occupational class were less exposed to air pollution (not significant) but those living in neighborhood with higher unemployment rate were more exposed. These

results are consistent with those observed when considering participants in one area.

We investigated, in three European cohorts, whether SEP evaluated at both individual- and

4. DISCUSSION

neighborhood-level was associated with traffic related air pollution exposure across sixteen Western European cities. The pooled analyses masked important heterogeneity across the cities showing that city appeared to be the major predictor of the association between SEP and NO₂ exposure.

The associations between individual-SEP and NO₂ were generally weak and inconsistent across the cities. This is in accordance with those of the three studies that used a comparable approach to ours (17,20,46). Education and occupation showed the same pattern with NO₂ in the pooled data and in most cities, in the city specific analyses, showing that both indicators measured the same concept (47,48). The associations between neighborhood-SEP and NO₂ were in the opposite direction (higher exposure in lower neighborhood-SEP) compared to the individual-SEP variables, both in the pooled data and in most cities in the city-specific models. This has also been observed in other studies in Europe (30) and in Montreal, Canada (49).

One possible explanation for the difference in direction is that the neighborhood-SEP is capturing aspects beyond the SEP of the population living in that area, such as how industrialized the neighborhood may be. Moreover, NO2 variability was relatively small across the individual-SEP groups, and after adjusting for neighborhood-SEP there was little evidence of potential confounding by individual-SEP. Place of residence is strongly patterned by social position and outdoor air pollution is spatially located within cities, therefore the degree to which air pollution is socially patterned is likely to occur more at area-level as well (33).Accounting for both city and neighborhood clustering using a two level random intercept model drastically decreased the size effects of the associations for both individual- and area-SEP markers compared to the single level linear regression model (Table S7). This has been observed in other studies (30,35,50) showing the importance to accounting for clustering in analyses including spatially nested data. With the multilevel approach the effect of unemployment rate remained in all cities but the effect of the individual-SEP decreased and even became null for several cities showing that variability was mainly explained by the city first then by the neighborhoods and for a smaller part by the individual-SEP. We looked at some socioeconomic variables at city level (e.g. population density, gross domestic product, etc.) to try to explain the heterogeneity of the association between SEP and NO₂ among the cities using a meta-regression. However, none of the tested variables explained this heterogeneity (not shown). To the best of our knowledge this is the first study including a large sample of cities geographically representative of Western Europe, with important within- and between-area variability of air pollution exposure. We used NO₂ as a traffic-related pollutant known to have a great intra-urban variability and thus was the most appropriate to study socioeconomic differences at individual-level (10,41,51). The NO₂ annual concentrations have been

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estimated at participant's residential address with a single harmonized exposure assessment protocol across the cities. The measurement time of NO₂ does not overlap with the questionnaire data from the cohorts. However, we assume that spatial contrasts in outdoor NO₂ pollution were stable over time; an assumption supported from observations in different settings in European countries (52,53). We used homogenized SEP indicators at both individual- and neighborhood-level. Recent evidence showed the importance of accounting SEP at both levels because they were independently associated with health outcomes (32– 34,46,54,55) but this had rarely been investigated with air pollution exposure (10,28,29). We used an area-based indicator defined at the smallest geographical unit available in each country to avoid MAUP as recommended (49,56–58). Our study has some limitations. Due to data confidentiality, we did not have access to participants' geographical coordinates for the present analysis and we were not able to analyze their spatial distribution. We applied an aspatial multilevel model to take into account the clustering of the participants within neighborhoods (46,59) but the proportion of neighborhoods containing only one participant was relatively high in some cities (60). This highlights a common problem in studies that were not originally designed to study area-level determinants. We compared a large number of European cities, but the sample in some cities was quite small and could explain the absence of associations and large confidence intervals. The different areas were also of different sizes and with different population density. However, the additional analysis performed for the Paris-Region suggested that the results were not sensitive to this aspect. We considered he unemployment rate, the sole indicator of neighborhood SEP uniformly available for most of the cities with ESCAPE NO₂ estimates. This single indicator does not fully describe participants' neighborhood-SEP (33) but has been used in other studies that compared different countries regarding air pollution (61) and has been associated with

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adverse health outcomes neighborhood level (61–64). We performed additional analyses with country-specific deprivation indices that were available at neighborhood level only for 12 out of the 16 cities (65–68) and we found consistent results compared to the ones with the neighborhood unemployment rate (Table S8).

Finally, we did not have information on other type of exposures such as occupational and indoor exposures or time-activity patterns (69) which could contribute to create or reinforce

5. CONCLUSIONS

environmental inequalities.

Unequal distribution to air pollution exposure according to SEP groups is complex in European cities and no general pattern exists across cities, but rather inequalities need to be specifically assessed in each city. Importantly, our results highlighted the importance of taking into account both individual- and neighborhood-SEP in order to fully describe and understand the complexity of current patterns of social inequalities relating to air pollution.

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405 REFERENCES

- 406 1. Gabriele Bolte et al. Environmental Health Inequalities in Europe. Copenhagen; 2012.
- 407 2. O'Neill MS, Jerrett M, Kawachi I, Levy JI, Cohen AJ, Gouveia N, et al. Health, 408 Wealth, and Air Pollution: Advancing Theory and Methods. Environmental Health
- 409 Perspectives. 2003 Sep 2;111(16):1861–70.
- 410 3. Morello-Frosch R, Zuk M, Jerrett M, Shamasunder B, Kyle AD. Understanding The
 411 Cumulative Impacts Of Inequalities In Environmental Health: Implications For Policy.
 412 Health Affairs. 2011 May 1;30(5):879–87.
- 4. Evans GW, Kantrowitz E. Socioeconomic status and health: the potential role of environmental risk exposure. Annual review of public health. 2002 May;23(1):303–31.
- Bowen W. An Analytical Review of Environmental Justice Research: What Do We Really Know? Environmental Management. 2002 Jan 11;29(1):3–15.
- Hajat A, Hsia C, O'Neill MS. Socioeconomic Disparities and Air Pollution Exposure: a
 Global Review. Current Environmental Health Reports. 2015 Dec 18;2(4):440–50.
- Musterd S. Social and Ethnic Segregation in Europe: Levels, Causes, and Effects.
 Journal of Urban Affairs. 2005 Aug;27(3):331–48.
- Pye S, Skinner I, Energy a E a, Meyer-ohlendorf N, Leipprand A. Addressing the social dimensions of environmental policy. 2008;(July):1–9.
- Deguen S, Zmirou-Navier D. Social inequalities resulting from health risks related to
 ambient air quality--A European review. The European Journal of Public Health. 2010
 Feb 1;20(1):27–35.
- 10. Chaix B, Gustafsson S, Jerrett M, Kristersson H, Lithman T, Boalt A, et al. Children's exposure to nitrogen dioxide in Sweden: investigating environmental injustice in an egalitarian country. Journal of epidemiology and community health. 2006 Mar 1;60(3):234–41.
- Helsinki: microenvironment, behavioral and sociodemographic factors. Journal of Exposure Analysis and Environmental Epidemiology. 2001 Jun;11(3):216–23.
- 433 12. Schikowski T, Sugiri D, Reimann V, Pesch B, Ranft U, Krämer U. Contribution of 434 smoking and air pollution exposure in urban areas to social differences in respiratory 435 health. BMC Public Health. 2008 Jan;8(1):179.
- Wheeler BW, Ben-Shlomo Y. Environmental equity, air quality, socioeconomic status, and respiratory health: a linkage analysis of routine data from the Health Survey for England. Journal of epidemiology and community health. 2005 Nov 1;59(11):948–54.
- Hand JS, Jones AP, Bateman IJ, Lovett AA, Fallon PJ. Modelling environmental equity: Access to air quality in Birmingham, England. Environment and Planning A. 2002;34(4):695–716.
- Forastiere F, Stafoggia M, Tasco C, Picciotto S, Agabiti N, Cesaroni G, et al.
 Socioeconomic status, particulate air pollution, and daily mortality: Differential
 exposure or differential susceptibility. American Journal of Industrial Medicine. 2007
 Mar;50(3):208–16.
- Nafstad P, Håheim LL, Wisløff T, Gram F, Oftedal B, Holme I, et al. Urban air
 pollution and mortality in a cohort of Norwegian men. Environmental health

- perspectives. 2004;112(5):610–5.
- 449 17. Fernandez-Somoano A, Tardon A. Socioeconomic status and exposure to outdoor NO2 450 and benzene in the Asturias INMA birth cohort, Spain. Journal of Epidemiology & 451 Community Health. 2014 Jan 1;68(1):29–36.
- Wheeler BW. Health-related environmental indices and environmental equity in England and Wales. Environment and Planning A. 2004;36(5):803–22.
- Havard S, Deguen S, Zmirou-Navier D, Schillinger C, Bard D. Traffic-Related Air
 Pollution and Socioeconomic Status. Epidemiology. 2009 Mar;20(2):223–30.
- Vrijheid M, Martinez D, Aguilera I, Ballester F, Basterrechea M, Esplugues A, et al.
 Socioeconomic status and exposure to multiple environmental pollutants during
 pregnancy: evidence for environmental inequity? Journal of Epidemiology &
 Community Health. 2012 Feb 1;66(2):106–13.
- Padilla CM, Kihal-Talantikite W, Vieira VM, Rossello P, Nir G Le, Zmirou-Navier D, et al. Air quality and social deprivation in four French metropolitan areas—A localized spatio-temporal environmental inequality analysis. Environmental Research. Elsevier; 2014 Oct 5;134:315–24.
- Fernández-Somoano A, Hoek G, Tardon A. Relationship between area-level socioeconomic characteristics and outdoor NO2 concentrations in rural and urban areas of northern Spain. BMC Public Health. 2013 Jan 25;13(1):71.
- Morelli X, Rieux C, Cyrys J, Forsberg B, Slama R. Air pollution, health and social deprivation: A fine-scale risk assessment. Environmental Research. 2016;147:59–70.
- 469 24. Miao Q, Chen D, Buzzelli M, Aronson KJ. Environmental Equity Research: Review
 470 With Focus on Outdoor Air Pollution Research Methods and Analytic Tools. Archives
 471 of Environmental & Occupational Health. 2015 Jan 2;70(1):47–55.
- Jerrett M, Finkelstein M. Geographies of risk in studies linking chronic air pollution exposure to health outcomes. Journal of toxicology and environmental health Part A. 2005;68(13–14):1207–42.
- Llop S, Ballester F, Estarlich M, Iñiguez C, Ramón R, Gonzalez M, et al. Social factors associated with nitrogen dioxide (NO2) exposure during pregnancy: The INMA-Valencia project in Spain. Social Science & Medicine. 2011 Mar;72(6):890–8.
- Chaix B, Leyland AH, Sabel CE, Chauvin P, Råstam L, Kristersson H, et al. Spatial clustering of mental disorders and associated characteristics of the neighbourhood context in Malmö, Sweden, in 2001. Journal of epidemiology and community health.
 2006 May 1;60(5):427–35.
- 482 28. Naess O, Piro FN, Nafstad P, Smith GD, Leyland AH. Air pollution, social deprivation, and mortality: a multilevel cohort study. Epidemiology (Cambridge, Mass). 2007;18(6):686–94.
- Cesaroni G, Badaloni C, Romano V, Donato E, Perucci C a, Forastiere F.
 Socioeconomic position and health status of people who live near busy roads: the
 Rome Longitudinal Study (RoLS). Environmental Health. 2010 Jan;9(1):41.
- 488 30. Goodman A, Wilkinson P, Stafford M, Tonne C. Characterising socio-economic 489 inequalities in exposure to air pollution: a comparison of socio-economic markers and 490 scales of measurement. Health & place. Elsevier; 2011 May;17(3):767–74.
- 491 31. Bell ML, O'Neill MS, Cifuentes LA, Braga ALF, Green C, Nweke A, et al. Challenges and recommendations for the study of socioeconomic factors and air pollution health

- 493 effects. Environmental Science and Policy. 2005;8(5):525–33.
- 494 32. Stafford M. Neighbourhood deprivation and health: does it affect us all equally? International Journal of Epidemiology. 2003 Jun 1;32(>3):357–66.
- 496 33. Diez Roux A-V. Neighborhoods and health: where are we and were do we go from here? Revue d'Épidémiologie et de Santé Publique. 2007 Feb;55(1):13–21.
- Bell ML, O'Neill MS, Cifuentes L a., Braga ALF, Green C, Nweke A, et al.
 Challenges and recommendations for the study of socioeconomic factors and air
 pollution health effects. Environmental Science & Policy. 2005 Oct;8(5):525–33.
- Jerrett M, Burnett RT, Willis A, Krewski D, Goldberg MS, DeLuca P, et al. Spatial analysis of the air pollution-mortality relationship in the context of ecologic confounders. Journal of toxicology and environmental health Part A. 2011;66(16–19):1735–77.
- 36. Beelen R, Hoek G, Vienneau D, Eeftens M, Dimakopoulou K, Pedeli X, et al.
 Development of NO2 and NOx land use regression models for estimating air pollution exposure in 36 study areas in Europe The ESCAPE project. Atmospheric Environment. 2013 Jun;72(2):10–23.
- 509 37. Boudier A, Curjuric I, Basagaña X, Hazgui H, Anto JM, Bousquet J, et al. Ten-year follow-up of cluster-based asthma phenotypes in adults a pooled analysis of three cohorts. American Journal of Respiratory and Critical Care Medicine.
 512 2013;188(5):550–60.
- 513 38. Siroux V, Boudier A, Bousquet J, Bresson J-L, Cracowski J-L, Ferran J, et al.
 514 Phenotypic determinants of uncontrolled asthma. Journal of Allergy and Clinical

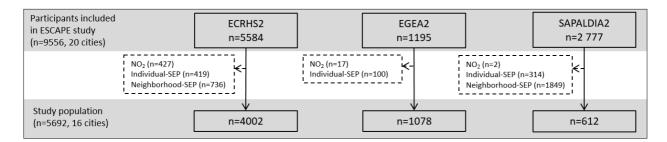
515 Immunology. 2009 Oct;124(4):681–687.e3.

- Jarvis D, ECRHS. The European Community Respiratory Health Survey II. European
 Respiratory Journal. 2002 Nov 1;20(5):1071–9.
- 518 40. Ackermann-Liebrich U, Kuna-Dibbert B, Probst-Hensch NM, Schindler C, Dietrich DF, Stutz EZ, et al. Follow-up of the Swiss Cohort Study on Air Pollution and Lung Diseases in Adults (SAPALDIA 2) 1991-2003: Methods and characterization of participants. Sozial- und Praventivmedizin. 2005;50(4):245–63.
- Cyrys J, Eeftens M, Heinrich J, Ampe C, Armengaud A, Beelen R, et al. Variation of
 NO2 and NOx concentrations between and within 36 European study areas: Results
 from the ESCAPE study. Atmospheric Environment. 2012;62:374–90.
- 42. WHO Regional Office for Europe. Health effects of transport-related air pollution.
 Krzyzanowski M, editor. 2005.
- 527 43. Beelen R, Raaschou-Nielsen O, Stafoggia M, Andersen ZJ, Weinmayr G, Hoffmann B, et al. Effects of long-term exposure to air pollution on natural-cause mortality: an analysis of 22 European cohorts within the multicentre ESCAPE project. Lancet. 2014 530 Mar 1;383(9919):785–95.
- 531 44. International Standard Classification of Occupations, Revised edition ISCO-88.
 532 Geneva, Switzerland: International Labour Office; 1991.
- Higgins JPT, Thompson SG, Deeks JJ, Altman DG. Measuring inconsistency in metaanalyses. BMJ (Clinical research ed). 2003 Sep 6;327(7414):557–60.
- Hajat A, Diez-Roux A V., Adar SD, Auchincloss AH, Lovasi GS, O'Neill MS, et al.
 Air Pollution and Individual and Neighborhood Socioeconomic Status: Evidence from
 the Multi-Ethnic Study of Atherosclerosis (MESA). Environmental Health

- Ferspectives. 2013 Sep 27;121(11):1325–33.
- 539 47. Galobardes B. Diet and socioeconomic position: does the use of different indicators matter? International Journal of Epidemiology. 2001 Apr 1;30(2):334–40.
- 541 48. Stronks K, van de Mheen H, van den Bos J, Mackenbach J. The interrelationship 542 between income, health and employment status. International Journal of Epidemiology. 543 1997 Jun 1;26(3):592–600.
- 544 49. Crouse DL, Ross N a, Goldberg MS. Double burden of deprivation and high 545 concentrations of ambient air pollution at the neighbourhood scale in Montreal, 546 Canada. Social Science & Medicine. Elsevier Ltd; 2009 Sep;69(6):971–81.
- 547 50. Havard S, Deguen S, Bodin J, Louis K, Laurent O, Bard D. A small-area index of 548 socioeconomic deprivation to capture health inequalities in France. Social Science & 549 Medicine. 2008;67(12):2007–16.
- Jerrett M, Arain A, Kanaroglou P, Beckerman B, Potoglou D, Sahsuvaroglu T, et al. A
 review and evaluation of intraurban air pollution exposure models. Journal of Exposure
 Analysis and Environmental Epidemiology. 2005 Mar 4;15(2):185–204.
- 553 52. Eeftens M, Beelen R, Fischer P, Brunekreef B, Meliefste K, Hoek G. Stability of measured and modelled spatial contrasts in NO2 over time. Occupational and environmental medicine. 2011;68(10):765–70.
- 53. Beevers SD, Westmoreland E, de Jong MC, Williams ML, Carslaw DC. Trends in
 NOx and NO2 emissions from road traffic in Great Britain. Atmospheric Environment.
 2012 Jul;54(2):107–16.
- 559 54. Chaix B, Leal C, Evans D. Neighborhood-level Confounding in Epidemiologic Studies. Epidemiology. 2010 Jan;21(1):124–7.
- 561 55. Krieger N, Waterman PD, Gryparis A, Coull B a. Black carbon exposure more strongly
 562 associated with census tract poverty compared to household income among US black,
 563 white, and Latino working class adults in Boston, MA (2003–2010). Environmental
 564 Pollution. 2014 Jul;190:36–42.
- 565 56. Diez Roux A V. Commentary: Estimating and understanding area health effects. International Journal of Epidemiology. 2005 Mar 31;34(2):284–5.
- 567 57. Maantay J. Mapping environmental injustices: pitfalls and potential of geographic information systems in assessing environmental health and equity. Environmental health Perspectives. 2002 Apr;110 Suppl(Supplement 2):161–71.
- 570 58. Mujahid MS, Diez Roux A V, Morenoff JD, Raghunathan T. Assessing the 571 measurement properties of neighborhood scales: from psychometrics to ecometrics. 572 American journal of epidemiology. 2007 Apr 15;165(8):858–67.
- 573 59. Havard S, Reich BJ, Bean K, Chaix B. Social inequalities in residential exposure to road traffic noise: an environmental justice analysis based on the RECORD Cohort Study. Occupational and environmental medicine. 2011 May;68(5):366–74.
- Bell B, Morgan G, Kromrey J, Ferron J. The impact of small cluster size on multilevel
 models: a Monte Carlo examination of two-level models with binary and continuous
 predictors. JSM Proceedings, Section on Survey Research Methods. 2010;4057–67.
- 579 61. Samoli E, Peng R, Ramsay T, Pipikou M, Touloumi G, Dominici F, et al. Acute effects 580 of ambient particulate matter on mortality in Europe and North America: Results from 581 the APHENA study. Environmental Health Perspectives. 2008;116(11):1480–6.

- van Lenthe FJ, Borrell LN, Costa G, Diez Roux A V, Kauppinen TM, Marinacci C, et al. Neighbourhood unemployment and all cause mortality: a comparison of six countries. Journal of epidemiology and community health. 2005 Mar;59(3):231–7.
- 585 63. Bosma H, Van De Mheen HD, Borsboom GJJM, Mackenbach JP. Neighborhood 586 socioeconomic status and all-cause mortality. American Journal of Epidemiology. 587 2001;153(4):363–71.
- Payne JN, Coy J, Milner PC, Patterson S. Are deprivation indicators a proxy for morbidity? A comparison of the prevalence of arthritis, depression, dyspepsia, obesity and respiratory symptoms with unemployment rates and Jarman scores. Journal of public health medicine. 1993 Jun;15(2):161–70.
- 592 65. Pornet C, Delpierre C, Dejardin O, Grosclaude P, Launay L, Guittet L, et al.
 593 Construction of an adaptable European transnational ecological deprivation index: the
 594 French version. Journal of Epidemiology & Community Health. 2012 Nov
 595 1;66(11):982–9.
- 596 66. Carstairs V, Morris R. Deprivation: explaining differences in mortality between Scotland and England and Wales. BMJ (Clinical research ed). 1989;299(6704):886–9.
- 598 67. Alguacil Gómez J, Camacho Gutiérrez J, Hernández Ajá A. La vulnerabilidad urbana 599 en España. Identificación y evolución de los barrios vulnerables. Empiria Revista de 600 metodología de ciencias sociales. 2013 Dec 18;(27):73.
- 601 68. Caranci N, Biggeri A, Grisotto L, Pacelli B, Spadea T, Costa G. [The Italian deprivation index at census block level: definition, description and association with general mortality]. Epidemiologia e prevenzione. 2010;34(4):167–76.
- 604 69. Schweizer C, Edwards RD, Bayer-Oglesby L, Gauderman WJ, Ilacqua V, Juhani 605 Jantunen M, et al. Indoor time-microenvironment-activity patterns in seven regions of 606 Europe. Journal of Exposure Science and Environmental Epidemiology. 2007 607 Mar;17(2):170-81.

Figure 1: Flow chart of the study population



611 Dotted frame: missing data

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612 ESCAPE: European Study of Cohorts for Air Pollution Effects

613 ECRHS: European Community Respiratory Health Survey (1999-2002)

614 EGEA: Epidemiological study on Genetics and Environment of Asthma (2003-2007)

SAPALDIA: Swiss Cohort Study on Air Pollution and Lung and Heart Diseases in Adults (2001-2003)

Table 1a: Characteristics of the population (by city and data pooled)

City	Country	n	Sex	Age	NO ₂ (μg*m ⁻³)	
			Men, %	mean ±sd	mean ±sd	Q1 – Q3
Norwich ^a	UK	242	43.0	43.6 ±6.5	25.6 ±5.7	22.8 - 28.7
Ipswich ^a	UK	338	42.3	42.4 ± 6.8	24.2 ± 4.0	22.7 - 26.0
Antwerp ^a	Belgium	500	49.9	42.7 ± 6.9	39.4 ± 9.0	32.7 - 45.6
Paris ^{a b}	France	785	48.3	41.7 ± 12.9	36.4 ± 13.4	27.4 - 42.6
Lyon a	France	210	46.7	48.4 ± 15.3	28.7 ± 13.5	16.9 - 40.6
Grenoble a b	France	690	52.9	44.9 ± 13.4	27.5 ± 8.2	20.8 - 32.9
Marseille b	France	119	43.7	49.2 ± 15.8	26.1 ± 8.2	21.4 - 31.1
Geneva c	Switzerland	612	49.4	52.1 ± 11.3	26.5 ± 7.0	21.1 - 31.3
Verona a	Italy	179	44.1	42.6 ± 7.1	30.7 ± 13.8	22.6 - 40.2
Pavia ^a	Italy	188	53.7	44.2 ± 6.6	20.5 ± 4.8	17.6 - 21.8
Turin ^a	Italy	170	46.6	42.9 ± 7.0	54.9 ± 10.1	49.2 - 61.9
Oviedo ^a	Spain	315	49.8	42.9 ± 7.1	36.6 ± 12.5	29.3 - 43.9
Galdakao ^a	Spain	408	48.5	40.7 ± 7.3	23.9 ± 6.6	18.6 - 28.3
Barcelona a	Spain	284	44.4	41.9 ± 7.1	57.4 ± 14.1	49.6 - 62.4
Albacete ^a	Spain	419	46.8	40.8 ± 7.3	28.6 ± 14.8	19.5 - 38.1
Huelva ^a	Spain	233	50.2	41.1 ± 7.2	25.2 ± 6.4	20.6 - 29.8
Pooled data		5692	48.2	43.9 ±10.6	31.8 ±13.6	22.4 - 38.6

Cities are sorted from north to south.

Participants were from ^aECRHS, ^bEGEA, ^cSAPALDIA; Paris: ECRHS n=386, EGEA n=399, Grenoble: ECRHS n=350, EGEA n=340. 620

Table1b: Socioeconomic characteristics of the population (by city and data pooled) 621

City	n		Neighborhood-level SEP					
		Age at end of school	Occupational Class, %					Unemployment rate*
		mean ±SD	Managers and Professionals (OC-I)	Technicians & Associate Professionals (OC-II)	Other non-manuals (OC-III)	Manuals (OC-IV)	Not in labor force	mean ±SD (min-max)
Norwich ^a	242	17.6 ± 3.1	25.6	19.4	27.3	24.0	3.7	11.1 ±7.2 (2.1-34.1)
Ipswich ^a	338	17.1 ± 2.6	22.5	16.6	30.8	22.2	8.0	$10.4 \pm 6.6 (2.4-32.0)$
Antwerp ^a	500	20.2 ± 3.1	33.0	18.6	31.0	16.8	0.7	$8.2 \pm 5.9 (0.8 - 31.2)$
Paris ^{a b}	785	21.3 ± 3.6	41.7	23.6	18.5	6.2	10.1	$10.6 \pm 4.0 (3.0 - 28.0)$
Lyon ^a	210	19.5 ± 3.7	20.5	24.8	26.2	21.0	7.6	$9.1 \pm 3.8 (3.4-25.1)$
Grenoble ab	690	20.8 ± 3.8	37.5	20.1	17.4	13.9	11.0	$9.8 \pm 4.5 (3.4 - 31.3)$
Marseille b	119	20.6 ± 3.4	46.2	20.2	14.3	9.3	10.1	$12.1 \pm 5.5 (4.9-35.0)$
Geneva c	612	20.5 ± 4.3	32.4	20.4	24.8	11.4	11.0	$4.3 \pm 1.4 (0.7-9.1)$
Verona a	179	19.0 ± 4.7	25.8	13.7	29.0	23.7	7.9	$4.5 \pm 3.0 (1.0 - 15.4)$
Pavia ^a	188	18.7 ± 4.6	25.8	13.7	29.0	23.7	7.9	$3.4 \pm 2.5 (0.7 - 14.3)$
Turin ^a	170	19.5 ± 5.2	21.6	13.1	36.4	22.1	6.8	$7.4 \pm 4.1 (1.4 - 21.7)$
Oviedo ^a	315	19.3 ± 4.6	26.7	10.8	29.2	28.6	4.8	$14.0 \pm 3.0 (7.5 - 33.3)$
Galdakao ^a	408	18.2 ± 4.1	17.9	8.6	25.3	37.7	10.5	$10.7 \pm 3.5 (3.1-21.9)$
Barcelona a	284	18.8 ± 4.9	28.9	14.4	29.6	21.1	6.0	$10.9 \pm 3.3 (4.1-26.4)$
Albacete ^a	419	17.7 ± 4.9	17.0	10.0	29.4	33.2	10.5	$14.6 \pm 5.3 (7.7-60.4)$
Huelva ^a	233	18.0 ± 4.6	17.6	9.4	27.9	30.5	14.6	$21.8 \pm 6.7 (10.7-41.4)$
Pooled data	5692	19.5 ±4.3	29.1	17.0	25.6	19.6	8.7	10.0 ±6.0 (0.7-60.4)

⁶²² Cities are sorted from north to south

⁶²³ SD=standard deviation

Participants were from ^a ECRHS, ^bEGEA, ^cSAPALDIA; Paris: ECRHS n=386, EGEA n=399, Grenoble: ECRHS n=350, EGEA n=340 624

OC= Occupational class. Not in labor force participants (in italics) included unemployed, retired, housepersons and students 625 626

^{*} The neighborhood unemployment rate has been assigned individually to participants using their residential addresses.

Table 2: Pooled results for the association between NO₂ concentration (µg*m-3) and SEP markers (n=5692) in percent change (95%CI)

			Multilevel model with city at level*			Multilevel model with neighborhood (level 2) and city (level 3) [†]			
		n	Adjusted for Mutually adjusted for individual		Adjusted for individual factors	Mutually adjusted for individual and neighborhood SEP			
Individual-level SEP									
Educational level	High (ref)	1917	-	-		-	-		
	Medium	2001	-4.5 (-6.6; -2.3)	-5.1 (-7.1; -3.0)		-1.3 (-2.7; -0.2)	-1.3 (-2.7; 0.2)		
	Low	1774	-6.9 (-9.1; -4.7)	-8.7 (-10.8; -6.5)		-1.7 (-3.2; -0.1)	-1.8 (-3.3; -0.2)		
p-value for trend [‡]			< 0.0001	< 0.0001		0.04	0.03		
Occupational class	OC-I (ref)	1657	_		-	_		-	
•	OC-II	967	-2.6 (-5.3; 0.2)		-2.7 (-5.4;0.01)	1.0 (-0.8; 2.9)		1.0 (-0.8; 2.9)	
	OC-III	1457	-1.0 (-3.5; 1.6)		-2.0 (-4.1; 0.5)	-0.6 (-2.3;1.0)		-0.7 (-2.3; 1.0)	
	OC-IV	1118	-5.6 (-8.2; -3.0)		-7.9 (-10.4; -5.3)	-0.6 (-2.5;1.2)		-0.8 (-2.6; 1.1)	
p-value for trend [‡]			0.001		< 0.0001	0.03		0.03	
Neighborhood-level SEI									
Unemployment rate§		5692	7.3 (6.2; 8.5)	$7.8 (6.7; 8.9)^{\P}$	7.7 (6.6; 8.8)#	3.33 (0.71; 6.01)	$3.2 (1.5; 5.0)^{\P}$	3.3 (1.5; 5.1)#	

^{*} A multilevel model was performed with city at level-2 (random intercept for city level).

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[†] A multilevel model was performed with neighborhood at level-2 and city at level-3 (random intercept for city and neighborhood levels).

^{\$\}frac{1}{2}\$ The unemployment rate has been transformed in z-score, the change in NO2 is showed for 1 standard deviation.

[¶] Mutually adjusted for educational level and neighborhood unemployment rate.

[#] Mutually adjusted for occupational class and neighborhood unemployment rate.

All models are adjusted for cohort, age and sex.

Results are expressed in percent change in NO₂ (µg*m-3) concentration adjusted for cohort, age, sex. Negative value means a decrease in NO₂ (in percent) compared to the

reference class for categorical variable and for 1SD increase for the continuous variable; p-value for trend were calculated by introducing the categorical variables in continuous.

Occupational class (OC): OC-I: Managers and Professionals, OC-II: Technician and associate professionals, OC-III: other non-manuals, OC-IV: skilled, semi-skilled and unskilled manuals.

Table 3a: Percent change (95%CI) in NO_2 concentration (μg^*m -3) in association to educational level mutually adjusted for neighborhood unemployment rate (n=5692)

City	n	Educa	Neighborhood Unemployment rate [*]		
		Medium	Low	P-value	
				for trend	
Norwich	242	-0.9 (-5.7; 4.3)	-1.1 (-7.7; 6.0)	0.71	9.4 (5.1; 13.8)
Ipswich	338	2.0 (-0.6; 4.7)	0.5 (-2.8; 3.8)	0.69	4.9 (1.0; 8.9)
Antwerp	500	0.6 (-2.2; 3.4)	1.2 (-1.9; 4.3)	0.45	14.9 (11.8; 18.2)
Paris	785	0.1 (-2.6; 2.9)	-0.3 (-3.1; 2.6)	0.84	13.7 (9.7; 17.8)
Lyon	210	-9.4 (-17.0; -0.9)	-3.6 (-12.3; -5.9)	0.58	12.6 (2.2; 24.0)
Grenoble	690	0.5 (-2.1; 3.0)	0.8 (-1.9; 3.7)	0.56	9.3 (5.1; 13.7)
Marseille	119	-1.9 (-10.4; 7.3)	-7.1 (-16.1; 2.9)	0.13	12.1 (7.1; 17.4)
Geneva	612	-2.0 (-4.5; 0.6)	-1.8 (-4.4; 0.9)	0.18	9.5 (4.7; 14.6)
Verona	179	-0.9 (-15.8; 16.8)	-16.1 (-26.5; -4.3)	0.01	14.0 (3.6; 25.3)
Pavia	188	0.1 (-4.2; 4.6)	-1.4 (-5.4; 2.6)	0.48	2.6 (-1.0; 6.4)
Turin	170	2.8 (-5.9; 12.3)	5.9 (-3.9; 16.6)	0.22	2.3 (-1.4; 6.1)
Oviedo	315	-0.4 (-7.2; 7.0)	-5.0 (-12.3; 3.0)	0.25	-14.1 (-23.6; -3.3)
Galdakao	408	-1.3 (-5.1; 2.8)	-3.3 (-7.8; 1.5)	0.18	21.8 (14.1; 30.1)
Barcelona	284	3.3 (-2.7; 9.7)	3.7 (-3.3; 11.2)	0.28	-7.7 (-12.7; -2.4)
Albacete	419	-10.3 (-21.1; 1.9)	-8.4 (-18.4; 2.9)	0.11	-7.9 (-17.5; 2.9)
Huelva	233	-1.0 (-6.1; 4.3)	-2.6 (-8.5; 3.6)	0.39	1.9 (-2.3; 6.4)

Cities are sorted from north to south.

A multilevel linear regression model (PROC MIXED) was performed with neighborhood at level-2 (random intercept for neighborhood level); adjusted for cohort, age and sex.

Results are expressed in percent change in NO_2 (μg^*m -3) concentration. Negative value means a decrease in NO_2 (in percent) compared to the reference class for the categorical variable; p-value for trend were calculated by introducing the categorical variables in continuous. The unemployment rate has been transformed in z-score, the change in NO_2 is showed for 1 standard deviation.

Table 3b: Percent change (95%CI) in NO₂ concentration (μ g*m-3) in association to occupational class mutually adjusted for neighborhood unemployment rate (n=5692)

City	n		Neighborhood Unemployment rate*			
		OC-II	OC-III	OC-IV	P-value for trend	
Norwich	242	-0.1 (-6.1; 6.2)	0.1 (-6.1; 6.7)	4.9 (-1.5; 11.8)	0.45	9.7 (5.3; 14.3)
Ipswich	338	2.3 (-1.2; 5.8)	1.6 (-1.4; 4.7)	0.6 (-2.5; 3.7)	0.99	5.0 (1.2; 9.1)
Antwerp	500	0.9 (-2.5; 4.4)	1.6 (-1.4; 4.6)	-1.7 (-5.0; 1.7)	0.63	15.1 (11. 9; 8.3)
Paris	785	-2.3 (-5.0; 0.6)	-3.3 (-6.4; -0.01)	-4.8 (-9.5; 0.1)	0.03	13.7 (9.7; 17.8)
Lyon	210	3.2 (-5.7; 12.9)	-3.9 (-12.5; 5.5)	-2.1 (-11.7; 8.6)	0.78	13.0 (2.5; 24.6)
Grenoble	690	1.8 (-1.1; 4.8)	1.1 (-2.1; 4.3)	3.1 (-0.4; 6.7)	0.20	9.1 (4.9; 13.5)
Marseille	119	-8.6 (-16.6; 0.1)	-6.9 (-15.2; 2.2)	-4.8 (-15.8; 7.7)	0.07	12.1 (7.0; 17.3)
Geneva	612	1.7 (-1.3; 4.8)	-1.0 (-3.7; 1.9)	-0.7 (-4.1; 2.8)	0.72	9.3 (4.4; 14.3)
Verona	179	1.9 (-20.8; 31.0)	-2.7 (-18.3; 15.8)	-12.9 (-28.1; 5.4)	0.07	13.3 (2.9;4.7)
Pavia	188	-2.6 (-8.2; 3.4)	-3.7 (-7.8; 0.7)	-2.5 (-7.6; 2.8)	0.17	2.7 (-0.9; 6.4)
Turin	170	9.5 (-3.6; 24.4)	9.6 (-0.6; 20.8)	11.7 (-0.1; 25.0)	0.07	2.3 (-1.3; 6.1)
Oviedo	315	0.8 (-9.5; 12.3)	-8.7 (-15.7; -1.2)	-5.9 (-13.2; 2.1)	0.07	-13.7 (-23.6; -2.8)
Galdakao	408	3.9 (-3.1; 11.4)	3.6 (-1.6; 9.0)	3.3 (-1.8; 8.6)	0.67	21.4 (13.6; 29.6)
Barcelona	284	3.4 (-4.8; 12.2)	3.4 (-2.8; 10.1)	4.1 (-2.6; 11.2)	0.16	-7.7 (-12.7; -2.5)
Albacete	419	-3.7 (-18.2; 13.5)	-6.1 (-18.2; 7.8)	-4.6 (-16.5; 9.1)	0.34	-8.3 (-18.0; 2.6)
Huelva	233	8.5 (-0.1; 17.9)	4.1 (-2.1; 10.8)	6.8 (0.1; 13.8)	0.15	1.0 (-3.2; 5.3)

Cities are sorted from north to south.

 A multilevel linear regression model (PROC MIXED) was performed with neighborhood at level-2 (random intercept for neighborhood level); adjusted for cohort, age and sex. Results are expressed in percent change in NO_2 (μg^*m -3) concentration. Negative value means a decrease in NO_2 (in percent) compared to the reference class for the categorical variable; p-value for trend were calculated by introducing the categorical variables in continuous. The unemployment rate has been transformed in z-score, the change in NO_2 is showed for 1 standard deviation.

Occupational class (OC): OC-I: Managers and Professionals (ref), OC-II: Technicians and associate professionals, OC-III: other non-manuals, OC-IV: skilled, semi-skilled and unskilled manuals. P-value for trend were calculated by introducing the categorical variables in continuous.