



Long-term exposure to air pollution and diabetes risk in Danish Nurse Cohort study

Anne Busch Hansen^a, Line Ravnskjær^b, Steffen Loft^c, Klaus Kaae Andersen^b, Elvira Bräuner^d, Rikke Baastrup^e, Claire Yao^a, Matthias Ketzel^f, Thomas Becker^f, Ole Hertel^f, Zorana Jovanovic Andersen^a

^aCentre for Epidemiology and Screening, Department of Public Health, University of Copenhagen, Copenhagen, Denmark; ^bDanish for Cancer Research Centre, Danish Cancer Society, Copenhagen, Denmark; ^cDepartment of Public Health, University of Copenhagen, Copenhagen, Denmark; ^dResearch Centre for Prevention and Health, Capitol Region of Denmark, Rigshospitalet – Glostrup, University of Copenhagen, Glostrup, Denmark; ^eNational Institute of Public Health, University of Southern Denmark, Copenhagen, Denmark; ^fDepartment of Environmental Science, Aarhus University, Roskilde, Denmark.

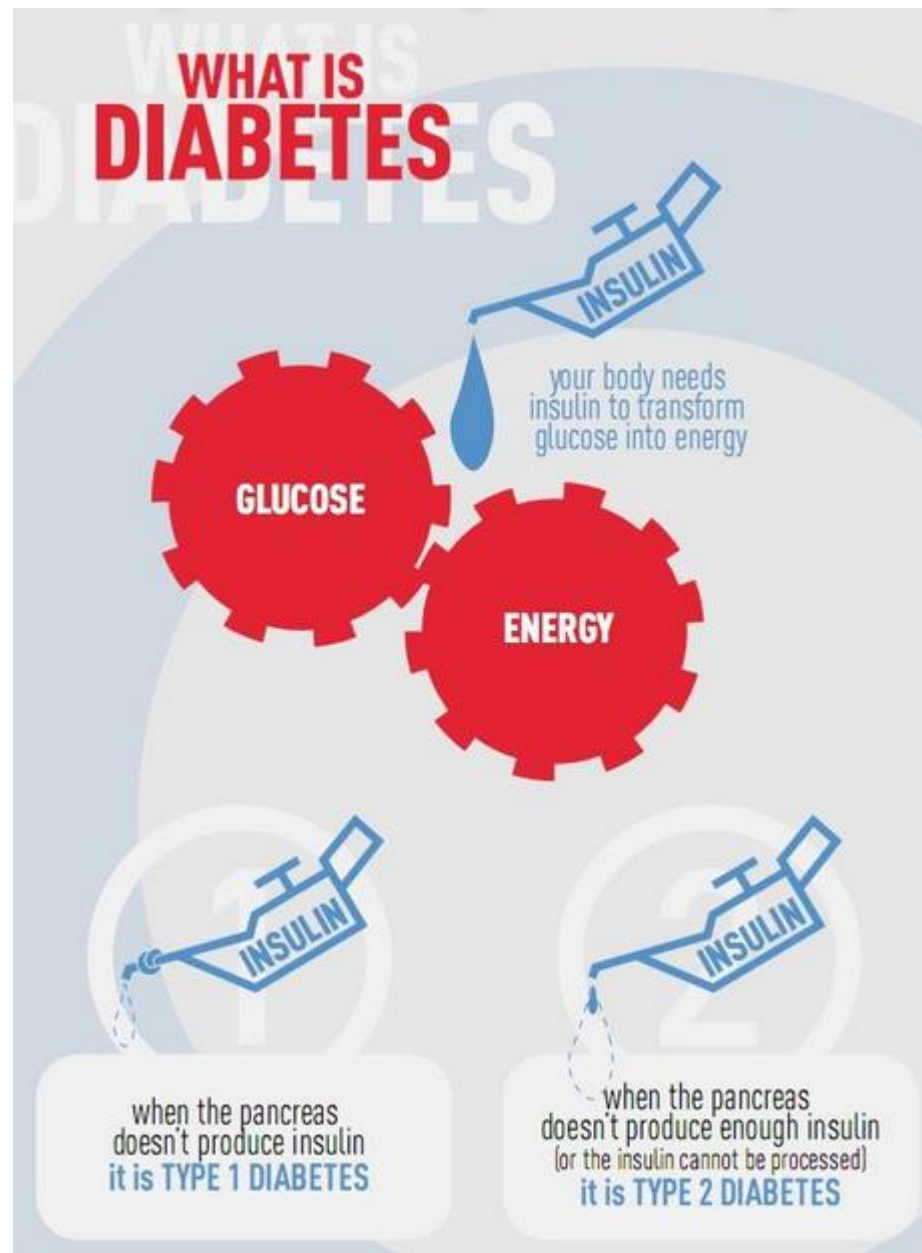
Presented by: Anne B. Hansen

Master student of Public Health Science (BSc)

Outline

- Aim
- Data
- Results
- Conclusion





Aim

- Air pollution \longrightarrow diabetes
 - PM_{2.5}
 - PM₁₀
 - NO₂
 - NO_x




Biological mechanism

Mice exposed to fine particles



Inflammation in visceral &
brown adipose tissue

Insulin resistance

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Ambient Air Pollution Exaggerates Adipose Inflammation and Insulin Resistance in a Mouse Model of Diet-Induced Obesity

Qinghua Sun, MD, PhD, Peibin Yue, MD, PhD, Jeffrey A. DeJulius, PhD, Carey N. Lumeng, MD, Thomas Kampfrath, MS, Michael B. Mikolaj, MD, Ying Cai, MD, Michael C. Ostrowski, PhD, Bo Lu, PhD, Sampath Parthasarathy, MBA, PhD, Robert D. Brook, MD, Susan D. Moffatt-Bruce, MD, PhD, Lung Chi Chen, PhD, and Sanjay Rajagopalan, MD
Davis Heart and Lung Research Institute (Q.S., P.Y., J.A.D., T.K., M.B.M., Y.C., S.P., S.R.), Division of Environmental Health Sciences (Q.S.), Division of Biostatistics (B.L.), Department of Molecular and Cellular Biochemistry (M.C.O.), and Division of Cardiothoracic Surgery (S.D.M.-B.), Colleges of Medicine and Public Health, Ohio State University, Columbus; Life Sciences Institute (C.N.L.) and Department of Internal Medicine (R.D.B.), University of Michigan, Ann Arbor; and Department of Environmental Medicine (L.C.C.), New York University School of Medicine, New York

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Long-term Exposure to Ambient Fine Particulate Pollution Induces Insulin Resistance and Mitochondrial Alteration in Adipose Tissue

Xiaohua Xu,* Cuiqing Liu,† Zhaobin Xu,† Kevin Tzan,* Mianhua Zhong,‡ Aixia Wang,† Morton Lippmann,‡ Lung-Chi Chen,‡ Sanjay Rajagopalan,†§ and Qinghua Sun*†§¹

**Division of Environmental Health Sciences, College of Public Health; and †Davis Heart and Lung Research Institute, College of Medicine, The Ohio State University, Columbus, Ohio 43210; ‡Department of Environmental Medicine, New York University School of Medicine, Tuxedo, New York 10012; and §Division of Cardiovascular Medicine, College of Medicine, The Ohio State University, Columbus, Ohio 43210*

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Effect of Early Particulate Air Pollution Exposure on Obesity in Mice Role of p47^{phox}

Xiaohua Xu, Zubin Yavar, Matt Verdin, Zhekang Ying, Georgeta Mihai, Thomas Kampfrath, Aixia Wang, Mianhua Zhong, Morton Lippmann, Lung-Chi Chen, Sanjay Rajagopalan, and Qinghua Sun
Division of Environmental Health Sciences (X.X., Z.Y., M.V., and Q.S.), College of Public Health, The Ohio State University, Columbus; the Davis Heart and Lung Research Institute (Z.Y., G.M., T.K., A.W., and S.R.), The Ohio State University, Columbus; the Department of Environmental Medicine (M.Z., M.L., and L.-C.C.), New York University School of Medicine, Tuxedo; and the Division of Cardiovascular Medicine (S.R. and Q.S.), College of Medicine, The Ohio State University, Columbus

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RESEARCH ARTICLE

Enhanced insulin resistance in diet-induced obese rats exposed to fine particles by instillation

Yuan-Horng Yan^{1,2}, Charles CK Chou³, Chung-Te Lee⁴, Jung-Yen Liu¹, and Tsun-Jen Cheng¹

¹Institute of Occupational Medicine and Industrial Hygiene, College of Public Health, National Taiwan University, Taipei, Taiwan, ²Department of Medical Research, Chia-Yi Christian Hospital, Chiayi, Taiwan, ³Research Center for Environmental Changes, Academia Sinica, Taipei, Taiwan, and ⁴Graduate Institute of Environmental Engineering, National Central University, Taoyuan, Taiwan

Current evidence

Research

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Risk of Incident Diabetes in Relation to Long-term Exposure to Fine Particulate Matter in Ontario, Canada

Hong Chen,^{1,2} Richard T. Burnett,³ Jeffrey C. Kwong,^{1,4,5} Paul J. Villeneuve,^{2,3} Mark S. Goldberg,^{6,7} Robert D. Brook,⁸ Aaron van Donkelaar,⁹ Michael Jerrett,¹⁰ Randall V. Martin,^{9,11} Jeffrey R. Brook,¹² and Ray Copes^{1,2}

¹Public Health Ontario, Toronto, Ontario, Canada; ²Dalla Lana School of Public Health, University of Toronto, Toronto, Ontario, Canada;

PM_{2.5} 1.11; 1.02-1.21 per 10 µg/m³

Weinmayr et al. *Environmental Health* (2015) 14:53
DOI 10.1186/s12940-015-0031-x



RESEARCH

Open Access



Long-term exposure to fine particulate matter and incidence of type 2 diabetes mellitus in a cohort study: effects of total and traffic-specific air pollution

Gudrun Weinmayr^{1,2*}, Frauke Hennig¹, Kateryna Fuks¹, Michael Nonnemacher³, Hermann Jakobs⁴, Stefan Möhlenkamp⁵, Raimund Erbe⁶, Karl-Heinz Jöckel³, Barbara Hoffmann^{1,2†}, Susanne Moebus^{3†} and on behalf of the Heinz Nixdorf Recall Investigator Group

PM_{2.5} 1.03; 0.89-1.29 per 2.3 µg/m³



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Air Pollution and Incidence of Hypertension and Diabetes in African American Women Living in Los Angeles

Patricia F. Coogan, ScD¹, Laura F. White, PhD², Michael Jerrett, PhD³, Robert D. Brook, MD⁴, Jason G. Su, PhD⁵, Edmund Seto, PhD³, Richard Burnett, PhD⁵, Julie R. Palmer, ScD¹, and Lynn Rosenberg, ScD¹

¹Stone Epidemiology Center at Boston University

²Department of Biostatistics, Boston University School of Public Health, Boston, MA

³Division of Environmental Health Sciences, School of Public Health, University of California, Berkeley, CA

⁴Division of Cardiovascular Medicine, University of Michigan Medical School, Ann Arbor, MI

⁵Healthy Environments and Consumer Safety Branch, Health Canada, Ottawa, Canada

PM_{2.5} 1.63; 0.78-3.44 per 10 µg/m³

Research

Are Particulate Matter Exposures Associated with Risk of Type 2 Diabetes?

Robin C. Puett,¹ Jaime E. Hart,^{2,3} Joel Schwartz,^{2,3,4} Frank B. Hu,^{2,3,5} Angela D. Liese,⁶ and Francine Laden^{2,3,4}

Research

Traffic-Related Air Pollution and Incident Type 2 Diabetes: Results from the SALIA Cohort Study

Ursula Krämer,^{1*} Christian Herder,^{2*} Dorothea Sugiri,¹ Klaus Strassburger,³ Tamara Schikowski,¹ Ulrich Ranft,¹ and Wolfgang Rathmann³

¹Institut für Umweltmedizinische Forschung (IUF), Leibniz Center at Heinrich Heine University Düsseldorf, Düsseldorf, Germany;

²Institute for Clinical Diabetology, and ³Institute of Biometrics and Epidemiology, German Diabetes Center, Leibniz Center for Diabetes Research at Heinrich Heine University Düsseldorf, Düsseldorf, Germany

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Long-term air pollution exposure and diabetes in a population-based Swiss cohort

Ikenna C. Eze^{a,b}, Emmanuel Schaffner^{a,b}, Evelyn Fischer^{a,b}, Tamara Schikowski^{a,b,c}, Martin Adam^{a,b}, Medea Imboden^{a,b}, Ming Tsai^{a,b}, David Carballo^d, Arnold von Eckardstein^e, Nino Künzli^{a,b}, Christian Schindler^{a,b}, Nicole Probst-Hensch^{a,b,*}



Epidemiology/Health Services Research

ORIGINAL ARTICLE

Diabetes Incidence and Long-Term Exposure to Air Pollution

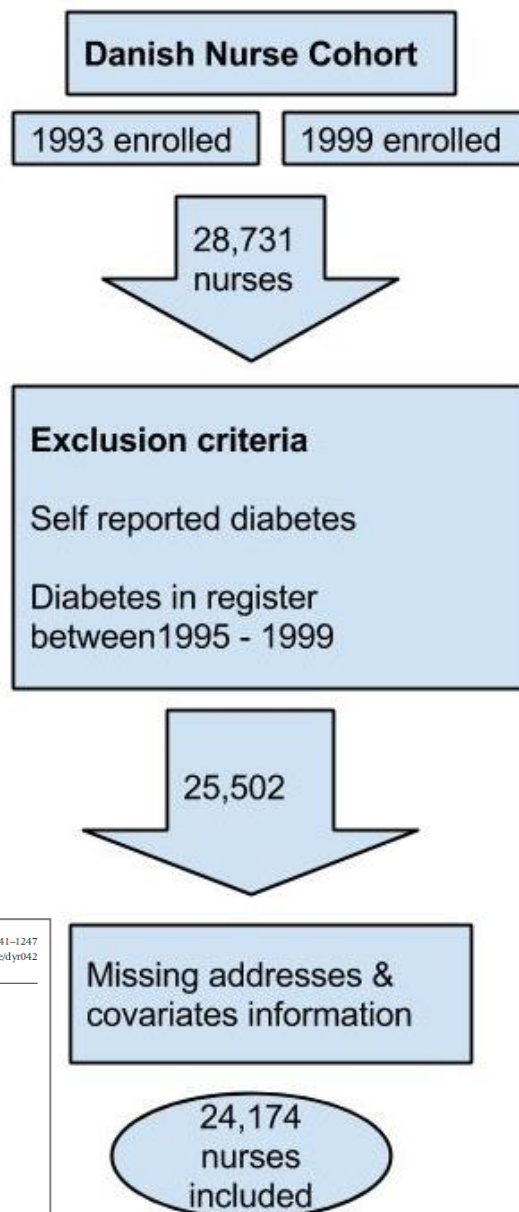
A cohort study

ZORANA J. ANDERSEN, PHD¹
ØLE RAASCHOU-NIELSEN, PHD¹
MATTHIAS KETZEL, PHD²
STEEN S. JENSEN, PHD²
MARTIN HVIDBERG, BS²

STEFFEN LOFT, DMSC³
ANNE TJØNNELAND, DMSC¹
KIM OVERVAD, PHD^{4,5}
METTE SØRENSEN, PHD¹

diabetes correlated with the release of toxicants into the air (3), whereas diabetic people appeared more vulnerable than nondiabetic people to cardiovascular health effects associated with exposure to

Data



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COHORT PROFILE

Cohort Profile: The Danish nurse cohort

Yrsa A Hundrup,^{1*} Mette K Simonsen,² Torben Jørgensen¹ and Erik B Obel¹

¹Research Centre for Prevention and Health, Glostrup University Hospital, Nordre Ringvej, Glostrup, Denmark and
²The Finsen Centre, Rigshospitalet, Blegdamsvej, Copenhagen Ø, Denmark

*Corresponding author. Research Centre for Prevention and Health, Glostrup University Hospital, Nordre Ringvej 57, DK-2600 Glostrup, Denmark. E-mail: yrsand01@glo.regionh.dk

Accepted 15 February 2011

How did the cohort come about?

From the 1970s to the mid-1990s, ~20–30% of Danish women between 55 and 65 years of age used hormone replacement therapy (HRT).^{1,2} It was primarily used

In 1993, a questionnaire was mailed to 23 170 female Danish nurses >44 years of age. Followed by two reminders, a total of 19 898 (86%) nurses replied with a completed questionnaire. The cohort was rein-



Danish Diabetes Register

All the participants were merged with diabetes status from the Danish Diabetes Register.

We categorized the cohort members as diabetic if they fulfilled either of the following criteria:

- 1) was hospitalized with diabetes as discharge diagnosis (ICD-10 code E10-14, DH36.0, DO24) in the National Patient Register
- 2) used chiropody as a diabetic patient
- 3) made a second purchase of insulin or oral anti-diabetic drugs within 6 months, registered in Danish National Prescription Registry.

Type 2 diabetes

Scandinavian Journal of Public Health, 2011; 39(Suppl 7): 58–61

DESCRIPTION OF DANISH REGISTERS

The National Diabetes Register

BENDIX CARSTENSEN^{1,2}, JETTE KOLDING KRISTENSEN³, MORTEN MUNK MARCUSSEN⁴ & KNUT BORCH-JOHNSEN⁵

¹Steno Diabetes Center, Gentofte, Denmark, ²Department of Biostatistics, University of Copenhagen, Copenhagen, Denmark, ³Department of General Medical Practice, Aarhus University, Aarhus, Denmark, ⁴National Board of Health, Copenhagen, Denmark, and ⁵Research Center for Quality in Health Care, Institute of Public Health, University of Southern Denmark, Odense, Denmark



Method - analyses

Cox Proportional hazards regression model with age as underlying time, adjusted for:

Model 1: Adjusted for age

Model 2: Adjusted for age, calendar time, smoking status, smoked gram of tobacco, level of physical activity, consumed gram of alcohol per week, avoidance of fatty meat, eat fruit or vegetables, employment status and marital status.

Model 3: Model 2 + hypertension, MI (myocardial infarct)

Model 4: Model 3 + BMI



Method - Air Pollution exposure

AirGIS dispersion model, sum of:

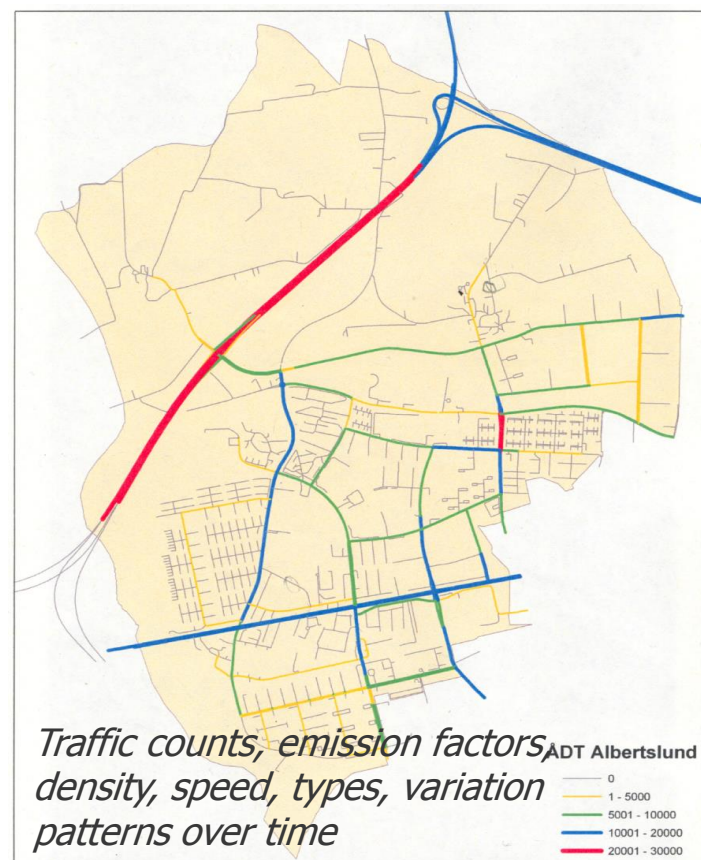
- 1) **regional** background, 2) **urban** background, &
- 3) **street level** contribution

Input for AirGIS model

- Street/building geometry
- Street network and traffic data
- Meteorology



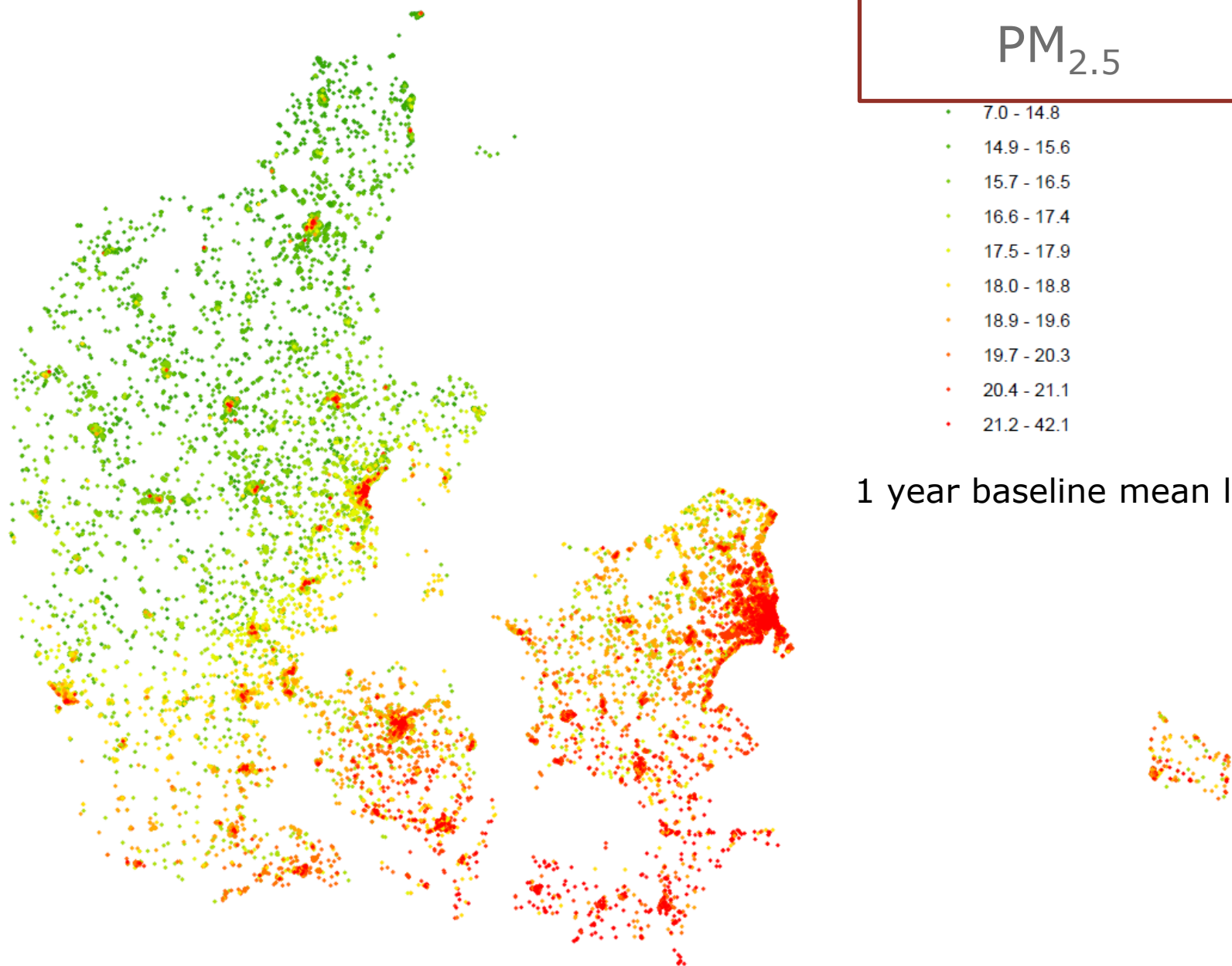
*GIS Maps building height,
street width, open sector*



Method - Air Pollution exposure

AirGIS collected annual concentration, in a 5 year time window, since 1990 to 2013, at each address for each participants.

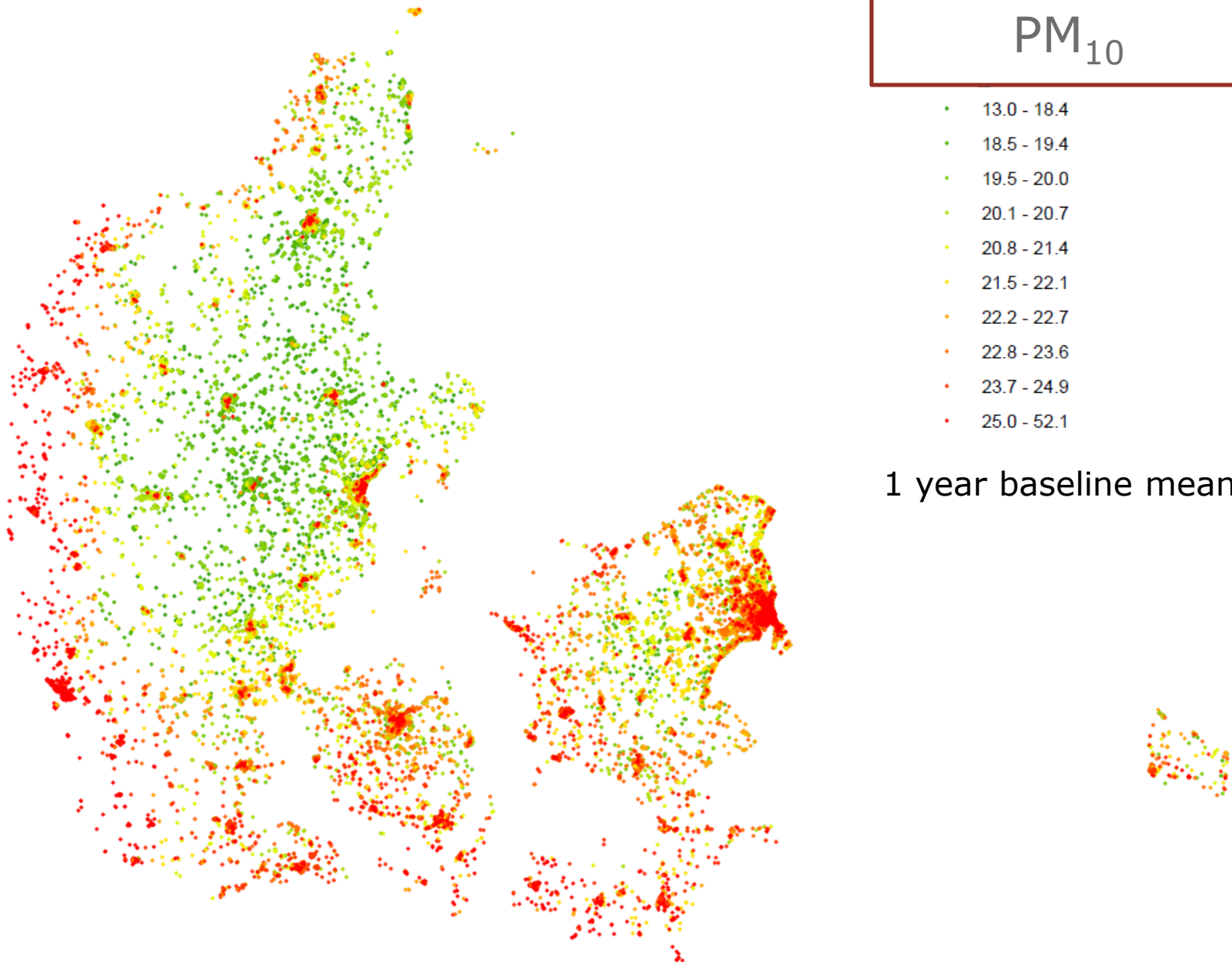




PM_{10}

- 13.0 - 18.4
- 18.5 - 19.4
- 19.5 - 20.0
- 20.1 - 20.7
- 20.8 - 21.4
- 21.5 - 22.1
- 22.2 - 22.7
- 22.8 - 23.6
- 23.7 - 24.9
- 25.0 - 52.1

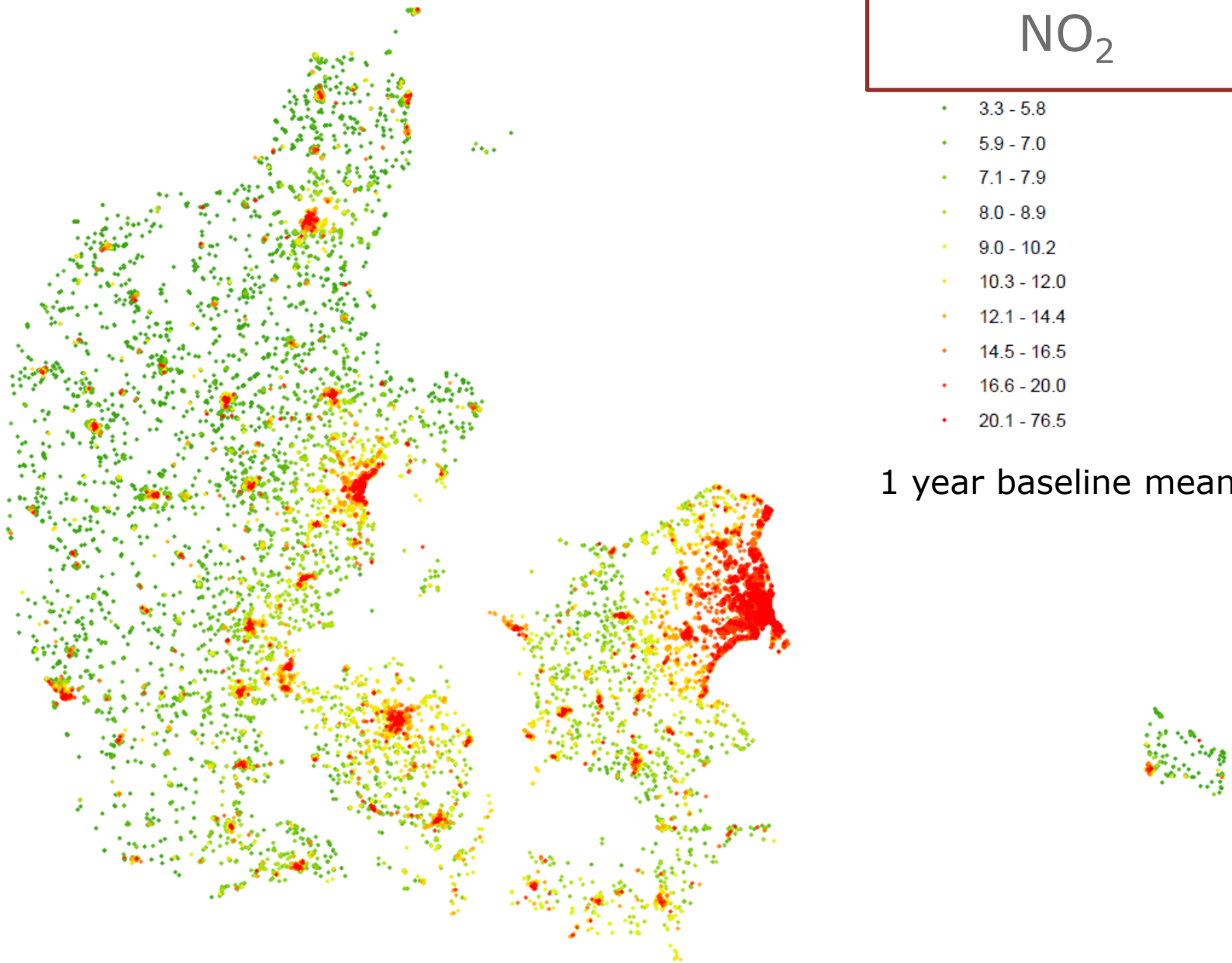
1 year baseline mean level

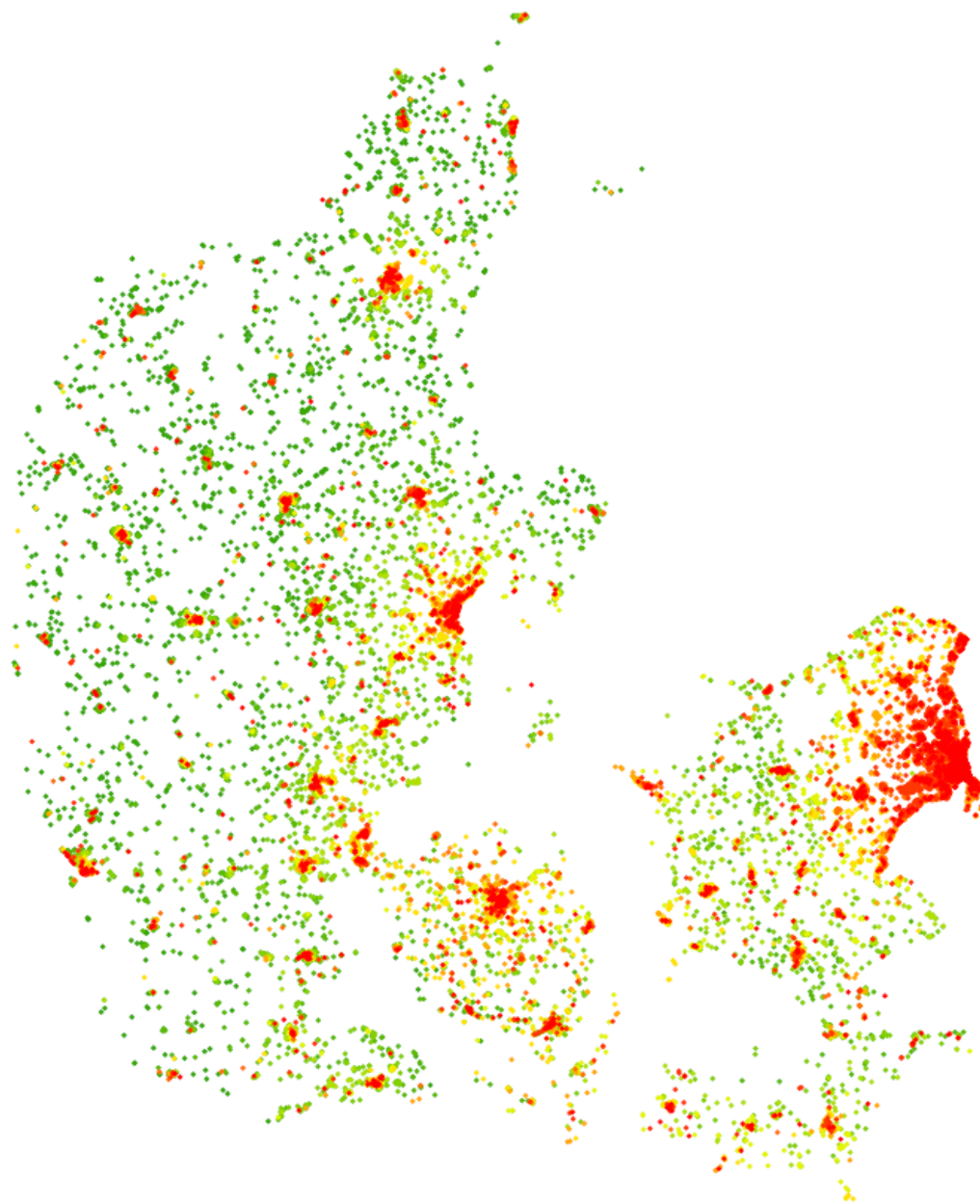


NO_2

- 3.3 - 5.8
- 5.9 - 7.0
- 7.1 - 7.9
- 8.0 - 8.9
- 9.0 - 10.2
- 10.3 - 12.0
- 12.1 - 14.4
- 14.5 - 16.5
- 16.6 - 20.0
- 20.1 - 76.5

1 year baseline mean level



 NO_x

- 3.5 - 6.3
- 6.4 - 7.6
- 7.7 - 8.8
- 8.9 - 10.0
- 10.1 - 11.8
- 11.9 - 14.6
- 14.7 - 17.7
- 17.8 - 20.5
- 20.6 - 33.2
- 33.3 - 389.1

1 year baseline mean level

Results

Follow-up from: 1995 or 1999 until 2013

Mean follow up time: 15.3 years, 370,367 person-years

Incident diabetes cases: 3 new cases per 1,000 person-years

Baseline mean age: 54.0

The nurses who developed diabetes:

↑higher BMI, smoked more, consumed more fat and had higher rate of hypertension and MI at baseline

↓had lower physical activity level, less fruit and vegetables intake, consumed less alcohol

	Total N = 24,174	Diabetes N = 1,137	Non diabetic N=23,037
Exposure measures, baseline values			
PM _{2.5} (µg/m ³), mean(SD)	18.1 (2.8)	18.7 (2.8)	18.1 (2.8)
PM ₁₀ (µg/m ³), mean(SD)	21.7 (2.9)	22.3 (2.9)	21.7 (2.9)
NO ₂ (µg/m ³), mean(SD)	12.5 (7.9)	13.4 (8.7)	12.5 (7.9)
NO _x (µg/m ³), mean(SD)	18.4 (22.7)	19.9 (23.2)	18.3 (22.6)

Results – cont.

- Main results of estimated risk of diabetes from pollutants
- Increase per unit

Pollutant	Unit (µg/m ³)	Model 1 ^a HR (95% CI)	Model 2 ^b HR (95% CI)	Model 3 ^c HR (95% CI)	Model 4 ^d HR (95% CI)
PM_{2.5}	3.1	1.14 (1.04-1.24)	1.11 (1.02-1.22)	1.11 (1.01-1.22)	1.11 (1.02-1.22)
PM₁₀	2.8	1.08 (1.00-1.16)	1.06 (0.99-1.14)	1.06 (0.99-1.14)	1.06 (0.98-1.14)
NO₂	7.5	1.06 (1.00-1.12)	1.03 (0.97-1.10)	1.03 (0.97-1.10)	1.05 (0.99-1.12)
NO_x	10.2	1.02 (0.99-1.05)	1.01 (0.98-1.04)	1.01 (0.98-1.04)	1.01 (0.98-1.05)
PM_{2.5}	10	1.50 (1.13-2.00)	1.41 (1.06-1.88)	1.39 (1.04-1.86)	1.41 (1.05-1.88)
PM₁₀	10	1.32 (1.02-1.72)	1.24 (0.95-1.62)	1.24 (0.95-1.62)	1.22 (0.93-1.59)

^a Adjusted for age at entry

^b Included covariates in Model 1^a + calendar time, smoking status, smoked gram of tobacco, level of physical activity, consumed gram of alcohol per week, avoidance of fatty meat, eat fruit or vegetables, employment status and marital status.

^c Included covariates in Model 2^b + hypertension, MI (myocardial infarct)

^d Included covariates in Model 3^c + body mass index.



Results – cont.

Table - Modification of associations^a between diabetes incidence and PM_{2.5} per interquartile range by baseline characteristics and co-morbid conditions among 24,174 participants in Danish Nurse cohort.

Covariates	Covariate level	n	IR	HR (95%CI)	p ^b
Smoking status	Never	358	2.7	1.24 (1.09-1.42)	0.01
	Previous	455	3.5	1.19 (1.03-1.36)	
	Current	324	2.8	0.97 (0.86-1.10)	
BMI	Underweight (BMI < 25)	13	1.5	0.86 (0.43-1.75)	0.37
	Normal	470	1.7	1.08 (0.96-1.22)	
	Overweight (25 ≤ BMI < 30)	433	5.3	1.09 (0.96-1.24)	
	Obese (BMI ≥ 30)	221	12.4	1.25 (1.06-1.47)	
MI	Yes	19	9.3	1.32 (0.86-2.02)	0.43
	No	1,118	3.0	1.11 (1.01-1.21)	
Hypertension	Yes	313	7.5	1.15 (1.00-1.33)	0.55
	No	824	2.5	1.10 (0.99-1.21)	
Urbanity	Urban	197	3.5	1.10 (0.92-1.31)	0.33
	Rural	446	2.9	1.16 (1.01-1.33)	
	Provincial	494	3.0	1.02 (0.88-1.18)	

^aFully adjusted for age, calendar time, smoking status, smoked gram of tobacco, level of physical activity, consumed gram of alcohol per week, avoidance of fatty meat, eat fruit or vegetables, hypertension, MI (myocardial infarct), employment status, marital status and BMI (body mass index).

^bFrom likelihood ratio test for interaction.

IR, incidence; HR, hazard ratio; CI, confidence interval



Strengths and Limitations

Strengths

- Big cohort, well defined covariate information
- Diabetes register, objective measure of incidence
- High response rate with high accuracy in answers

Limitations

- Misclassification in modeled concentrations since these are only proxies of personal exposure
- Missing information on indoor exposures (gas cooking, passive tobacco smoke, air conditioning ventilation habits), air pollution at work, commuting habits & personal activity patterns
- Type 2 diabetes status
- Family history of diabetes unknown



Conclusion



- 7.5 $\mu\text{g}/\text{m}^3$ NO_2 increases the risk of diabetes with 5% insignificantly (1.05; 0.99-1.12)

Author	Hazard ratio in fully adjusted model, incl. HP, MI & BMI
Andersen et al., 2012	HR = 1.04 (1.01-1.07) NO_2 per 4.9 $\mu\text{g}/\text{m}^3$

- Long-term exposure to air pollution increases risk of diabetes, fine particles are most relevant
- 3.1 $\mu\text{g}/\text{m}^3$ $\text{PM}_{2.5}$ increases the risk of diabetes with 11% significantly



Declaration

No conflict-of-interest

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Anne Busch Hansen^a, Line Ravnskjær^b, Steffen Loft^c, Klaus Kaae Andersen^b, Elvira Bräuner^d, Rikke Baastrup^e, Claire Yao^a, Matthias Ketzelf^f, Thomas Becker^f, Ole Hertelf^f, Zorana Jovanovic Andersen^a

^aCenter for Epidemiology and Screening, Department of Public Health, University of Copenhagen, Copenhagen, Denmark; ^bDanish for Cancer Research Center, Danish Cancer Society, Copenhagen, Denmark; ^cDepartment of Public Health, University of Copenhagen, Copenhagen, Denmark; ^dResearch Center for Prevention and Health, Capitol Region of Denmark, Rigshospitalet – Glostrup, University of Copenhagen, Glostrup, Denmark; ^eNational Institute of Public Health, University of Southern Denmark, Copenhagen, Denmark; ^fDepartment of Environmental Science, Aarhus University, Roskilde, Denmark.



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E-mail: hansen.anneb@gmail.com

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