Supplemental Material

Ambient Air Pollution, Adipokines, and Glucose Homeostasis: The Framingham Heart Study

Wenyuan Li^{a,b}, Kirsten S. Dorans^{a,b,c}, Elissa H. Wilker^{a,b}, Mary B. Rice^d, Itai Kloog^e, Joel D. Schwartz^{a,f}, Petros Koutrakis^f, Brent A. Coull^g, Diane R. Gold^{f,h}, James B. Meigsⁱ, Caroline S. Fox^{j,k}, Murray A. Mittleman^{a,b}

a Department of Epidemiology, Harvard T.H. Chan School of Public Health, Boston, Massachusetts, United States b Cardiovascular Epidemiology Research Unit, Division of Cardiology, Beth Israel Deaconess Medical Center, Harvard Medical School, Boston, Massachusetts, United States c Department of Epidemiology, Tulane University School of Public Health and Tropical Medicine, New Orleans, Louisiana, United States d Division of Pulmonary, Critical Care and Sleep Medicine, Beth Israel Deaconess Medical Center, Boston, Massachusetts, United States e Department of Geography and Environmental Development, Ben-Gurion University of the Negev, Beer Sheva, Israel f Department of Environmental Health, Harvard T.H. Chan School of Public Health, Boston, Massachusetts, United States g Department of Biostatistics, Harvard T.H. Chan School of Public Health, Boston, Massachusetts, United States h Channing Division of Network Medicine, Brigham and Women's Hospital and Harvard Medical School, Boston, Massachusetts, United States i Department of Medicine, Harvard Medical School and Division of General Internal Medicine, Massachusetts General Hospital, Boston, Massachusetts, United States j National Heart, Lung, and Blood Institute's Framingham Heart Study, Framingham, Massachusetts, United States k Division of Intramural Research, National Heart, Lung, and Blood Institute, Bethesda, Maryland, United States

	Examination cycle 7	Examination cycle 1	Examination cycle 8	Examination cycle 2	Total for 2003 annual average PM _{2.5} analysis		Total for residential proximity analysis	
	(1998-2001)	(2002-2005)	(2005-2008)	(2008-2011)	No. of observations	No. of participants	No. of observations	No. of participants
Fasting Glucose	2,415	3,281	1,925	2,768	10,389	5,958	9,264	5,403
Insulin	-	2,966	1,923	2,768	7,657	5,278	6,784	4,739
HOMA-IR	-	2,966	1,923	2,768	7,657	5,278	6,784	4,739
HbA1c	-	-	1,923	2,767	4,690	4,690	4,164	4,164
Adiponectin	2,016	3,266	-	-	5,282	5,282	4,714	4,714
Resisitn	2,030	-	-	-	2,030	2,030	1,845	1,845
Leptin	-	3,266	-	-	3,266	3,266	2,881	2,881

Supplemental Table A. Sample Size of 2003 Annual Average PM_{2.5} Analysis and Distance to Major Roadways Analysis for Each Measured Biomarker.

Abbreviation: PM_{2.5}, fine particulate matter; HOMA-IR, homeostasis model assessment of insulin resistance; HbA1c, hemoglobin A1c.

Air Pollutants	No. of	Moon (SD)	Interquartile	Spearman Correlation Coefficients			
All Follutants	Observations	Mean (SD)	Range	BC	SO 4 ²⁻	NO _x	O ₃
PM _{2.5} (µg/m ³)	6,990	9.7 (5.9)	5.7	0.72	0.82	0.43	0.01
BC ($\mu g/m^3$)	6,983	0.8 (0.4)	0.5		0.54	0.58	-0.26
$SO_4^{2-} (\mu g/m^3)$	6,145	2.9 (2.4)	2.1			0.31	0.13
NO _x (ppb)	6,719	36.5 (20.2)	19.3				-0.54
O ₃ (ppb)	6,997	23.7 (10.9)	14.4				

Supplemental Table B. Characteristics and Spearman Correlations Between 1-Day Moving Averages of Pollutants.

Abbreviation: SD, standard deviation; $PM_{2.5}$, fine particulate matter; BC, black carbon; SO_4^{2-} , sulfates; NO_x , nitrogen oxides; O_3 , ozone.

	Fasting plasma glucose	HBA1c	Adiponectin	Resistin
_	% Difference (95% CI)	% Difference (95% CI)	% Difference (95% CI)	% Difference (95% CI)
Median Age ^b				
\leq Median age	-0.11 (-0.35, 0.14)	-0.13 (-0.35, 0.10)	-1.00 (-3.19, 1.25)	-1.18 (-3.98, 1.69)
> Median age	-0.05 (-0.31, 0.22)	0.04 (-0.19, 0.27)	0.55 (-1.86, 3.01)	-3.13 (-5.89, -0.29)
Sex ^c				
Women	-0.16 (-0.41, 0.10)	-0.04 (-0.26, 0.17)	-1.33 (-3.48, 0.87)	-3.08 (-5.69, -0.41)
Men	0.02 (-0.27, 0.31)	-0.06 (-0.31, 0.18)	1.01 (-1.44, 3.52)	-0.93 (-3.93, 2.16)
Educational Attainment ^d				
\leq High school	0.30 (-0.19, 0.78)	0.15 (-0.28, 0.59)	0.62 (-3.30, 4.69)	-3.05 (-7.16, 1.25)
\geq College	-0.15 (-0.36, 0.07)	-0.09 (-0.27, 0.09)	-0.41 (-2.25, 1.46)	-2.06 (-4.40, 0.34)
Obesity Status ^e				
$BMI > 30 \text{ kg/m}^2$	-0.12 (-0.34, 0.11)	-0.03 (-0.22, 0.16)	-0.33 (-2.23, 1.59)	-1.61 (-3.98, 0.81)
BMI $\leq 30 \text{ kg/m}^2$	0.04 (-0.32, 0.39)	-0.11 (-0.41, 0.19)	-0.18 (-3.46, 3.22)	-3.55 (-7.16, 0.20)
Prediabetes Status ^f				
No Prediabetes	-0.04 (-0.20, 0.11)	0.00 (-0.19, 0.19)	-0.96 (-2.86, 0.97)	-2.18 (-4.75, 0.46)
Prediabetes	0.04 (-0.20, 0.27)	-0.10 (-0.38, 0.18)	1.10 (-2.07, 4.38)	-1.90 (-4.96, 1.26)
	Insulin	HOMA-IR	Leptin	
-	% Difference (95% CI)	% Difference (95% CI)	% Difference (95% CI)	-
Median Age ^b				-
\leq Median age	-0.14 (-1.64, 1.39)	-0.18 (-1.79, 1.45)	-0.45 (-3.33, 2.52)	
> Median age	-1.37 (-2.90, 0.19)	-1.47 (-3.10, 0.19)	0.80 (-2.14, 3.83)	
Sex ^c				
Women	-0.59 (-2.10, 0.94)	-0.73 (-2.34, 0.91)	-0.58 (-3.32, 2.23)	
Men	-0.92 (-2.61, 0.80)	-0.91 (-2.72, 0.93)	1.18 (-1.91, 4.37)	
Educational Attainment ^d				
\leq High school	0.77 (-2.15, 3.78)	1.07 (-2.06, 4.30)	0.44 (-4.92, 6.11)	
\geq College	-1.06 (-2.30, 0.20)	-1.21 (-2.53, 0.14)	0.01 (-2.28, 2.35)	
Obesity Status ^e				
$BMI > 30 \text{ kg/m}^2$	-0.76 (-2.07, 0.58)	-0.81 (-2.22, 0.61)	0.08 (-2.29, 2.51)	
BMI $\leq 30 \text{ kg/m}^2$	-0.60 (-2.67, 1.53)	-0.72 (-2.94, 1.54)	0.55 (-3.82, 5.12)	
Prediabetes Status ^f				
No Prediabetes	-0.39 (-1.69, 0.92)	-0.44 (-1.78, 0.92)	0.14 (-2.19, 2.53)	
Prediabetes	-1.25 (-3.20, 0.74)	-1.21 (-3.22, 0.86)	0.54 (-4.16, 5.46)	

Supplemental Table C. Associations of 2003 Annual Average Fine Particulate Matter (PM_{2.5}) With Adipokines And Biomarkers of Glucose Homeostasis, **by Median Age, Sex, Educational Attainment, Obesity and Prediabetes Status** ^a.

Abbreviation: PM_{2.5}, fine particulate matter; HBA1c, hemoglobin A1c; HOMA-IR, homeostasis model assessment of insulin resistance; CI, confidence intervals.

a) Results were scaled to equivalent to 1.5 μ g/m³ higher in PM_{2.5} concentrations.

b) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin. Median age for fasting glucose analyses was 50 years old, for insulin and HOMA-IR analyses was 47 years old, for HBA1c analysis was 53 years old, for adiponectin analysis was 46 years old, for leptin 40 years old, and for resistin was 59 years old.

c) Models were adjusted for centered age, (centered age)², body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

d) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

e) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

f) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

_	Fasting plasma glucose	HBA1c	Adiponectin	Resistin
	% Difference (95% CI)	% Difference (95% CI)	% Difference (95% CI)	% Difference (95% CI)
Median Age ^b				
\leq Median age	0.21 (-0.09, 0.51)	0.06 (-0.22, 0.34)	0.66 (-2.07, 3.47)	0.80 (-2.39, 4.09)
> Median age	0.35 (0.03, 0.67)	0.00 (-0.28, 0.27)	-2.15 (-4.86, 0.64)	3.57 (0.37, 6.87)
Sex ^c				
Women	0.40 (0.09, 0.71)	0.06 (-0.21, 0.33)	0.82 (-1.84, 3.55)	2.20 (-0.82, 5.31)
Men	0.14 (-0.19, 0.48)	-0.01 (-0.30, 0.28)	-2.53 (-5.32, 0.34)	2.20 (-1.21, 5.72)
Educational Attainment ^d				
\leq High school	0.16 (-0.30, 0.62)	0.31 (-0.11, 0.72)	1.22 (-2.67, 5.27)	2.43 (-1.32, 6.33)
\geq College	0.32 (0.06, 0.58)	-0.05 (-0.28, 0.17)	-1.11 (-3.36, 1.19)	1.95 (-0.97, 4.95)
Obesity Status ^e				
BMI $\leq 30 \text{ kg/m}^2$	0.17 (-0.09, 0.44)	-0.11 (-0.35, 0.12)	-0.23 (-2.50, 2.10)	3.28 (0.55, 6.09)
BMI $>30 \text{ kg/m}^2$	0.55 (0.14, 0.96)	0.37 (0.01, 0.73)	-2.18 (-5.86, 1.65)	-0.10 (-4.12, 4.09)
Prediabetes Status ^f				
No Prediabetes	0.15 (-0.04, 0.33)	-0.02 (-0.26, 0.22)	0.20 (-2.09, 2.53)	1.91 (-0.99, 4.88)
Prediabetes	0.15 (-0.12, 0.42)	0.01 (-0.31, 0.33)	-3.06 (-6.62, 0.64)	2.65 (-0.97, 6.41)
	Insulin	HOMA-IR	Leptin	
-	% Difference (95% CI)	% Difference (95% CI)	% Difference (95% CI)	-
Median Age ^b				-
\leq Median age	0.93 (-0.89, 2.78)	1.12 (-0.83, 3.10)	2.82 (-0.66, 6.43)	
> Median age	0.25 (-1.63, 2.16)	0.54 (-1.47, 2.59)	-0.27 (-4.01, 3.61)	
Sex ^c				
Women	0.81 (-1.04, 2.70)	1.18 (-0.81, 3.21)	0.87 (-2.61, 4.46)	
Men	0.37 (-1.61, 2.39)	0.46 (-1.65, 2.63)	2.15 (-1.60, 6.03)	
Educational Attainment ^d				
\leq High school	0.25 (-2.63, 3.21)	0.50 (-2.58, 3.68)	-0.76 (-6.51, 5.33)	
\geq College	0.79 (-0.74, 2.35)	1.04 (-0.61, 2.71)	2.02 (-0.85, 4.98)	
Obesity Status ^e				
BMI $\leq 30 \text{ kg/m}^2$	0.40 (-1.18, 2.00)	0.56 (-1.13, 2.28)	2.96 (-0.03, 6.04)	
BMI $>30 \text{ kg/m}^2$	1.13 (-1.37, 3.69)	1.58 (-1.11, 4.33)	-3.10 (-8.01, 2.06)	
Prediabetes Status ^f				
No Prediabetes	-0.23 (-1.79, 1.35)	-0.15 (-1.76, 1.49)	2.61 (-0.30, 5.60)	
Prediabetes	1.61 (-0.67, 3.93)	1.71 (-0.65, 4.12)	-3.05 (-8.36, 2.57)	

Supplemental Table D. Associations of Residential Proximity to Major Roadways With Adipokines And Biomarkers of Glucose Homeostasis, **by Median Age, Sex, Educational Attainment, Obesity and Prediabetes Status**^a.

Abbreviation: PM_{2.5}, fine particulate matter; HBA1c, hemoglobin A1c; HOMA-IR, homeostasis model assessment of insulin resistance; CI, confidence intervals.

a) Analysis was restricted to participants who lived within 1,000 m from major roadways. Results were scaled to comparing participants who lived 64 m from major roadways to those who lived 413 m away.

b) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin. Median age for fasting glucose analyses was 50 years old, for insulin and HOMA-IR analyses was 47 years old, for HBA1c analysis was 53 years old, for adiponectin analysis was 46 years old, for leptin 40 years old, and for resistin was 59 years old.

c) Models were adjusted for centered age, (centered age)², body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

d) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.
e) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.
f) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.
f) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

Condition	Emm	Fasting plasma glucose	HBA1c	Adiponectin	Resistin
Condition	Exposure	% Difference (95% CI)	% Difference (95% CI)	% Difference (95% CI)	% Difference (95% CI)
1 Minimally adjusted 8	PM _{2.5} °	-0.17 (-0.37, 0.02)	-0.08 (-0.24, 0.08)	-0.18 (-1.82, 1.49)	-1.24 (-3.21, 0.77)
1. Minimally adjusted ^a	Distance ^d	0.39 (0.16, 0.63)	0.09 (-0.11, 0.29)	-1.61 (-3.60, 0.43)	2.22 (-0.09, 4.58)
2. PM _{2.5} 2003-2005 average ^b	PM _{2.5} ^c	-0.06 (-0.24, 0.12)	-0.03 (-0.18, 0.12)	0.69 (-0.85, 2.26)	-1.78 (-3.61, 0.09)
3. Exclude observations with 2003 annual $PM_{2.5} > 12 \ \mu g/m^{3 b}$	$PM_{2.5}$ °	-0.02 (-0.28, 0.23)	-0.09 (-0.30, 0.12)	0.38 (-1.81, 2.61)	-2.09 (-4.55, 0.42)
4. Include those who lived >1,000 m from the nearest major roadway ^b	Distance ^d	0.18 (-0.02, 0.39)	-0.05 (-0.22, 0.13)	-0.56 (-2.31, 1.21)	1.78 (-0.32, 3.92)
5. Restrict to participant with both	PM _{2.5} °	-0.03 (-0.26, 0.20)	0.00 (-0.19, 0.19)	-0.61 (-2.57, 1.39)	-1.70 (-4.05, 0.71)
PM _{2.5} and proximity data ^b	Distance ^d	0.28 (0.05, 0.51)	0.03 (-0.17, 0.22)	-0.73 (-2.68, 1.26)	2.20 (-0.09, 4.54)
1 V		× , , ,	· · · · ·	· · · · ·	· · · · ·
	Eurocum	Insulin	HOMA-IR	Leptin	
Condition	Exposure			× · · /	
Condition	Exposure PM _{2.5} ^c	Insulin	HOMA-IR	Leptin	
		Insulin % Difference (95% CI)	HOMA-IR % Difference (95% CI)	Leptin % Difference (95% CI)	
Condition	PM _{2.5} ^c	Insulin % Difference (95% CI) -1.36 (-2.68, -0.03)	HOMA-IR % Difference (95% CI) -1.55 (-2.97, -0.12)	Leptin % Difference (95% CI) -0.83 (-3.87, 2.30)	
Condition 1. Minimally adjusted ^a	PM _{2.5} ^c Distance ^d	Insulin % Difference (95% CI) -1.36 (-2.68, -0.03) 1.45 (-0.18, 3.11)	HOMA-IR % Difference (95% CI) -1.55 (-2.97, -0.12) 1.81 (0.05, 3.59)	Leptin % Difference (95% CI) -0.83 (-3.87, 2.30) 5.25 (1.17, 9.51)	
Condition 1. Minimally adjusted ^a 2. PM _{2.5} 2003-2005 average ^b 3. Exclude observations with	PM _{2.5} ^c Distance ^d PM _{2.5} ^c	Insulin % Difference (95% CI) -1.36 (-2.68, -0.03) 1.45 (-0.18, 3.11) -0.52 (-1.58, 0.55)	HOMA-IR % Difference (95% CI) -1.55 (-2.97, -0.12) 1.81 (0.05, 3.59) -0.60 (-1.73, 0.54)	Leptin % Difference (95% CI) -0.83 (-3.87, 2.30) 5.25 (1.17, 9.51) -0.01 (-1.95, 1.98)	
Condition 1. Minimally adjusted ^a 2. PM _{2.5} 2003-2005 average ^b 3. Exclude observations with 2003 annual PM _{2.5} > 12 µg/m ^{3 b} 4. Include those who lived >1,000 m	PM _{2.5} ^c Distance ^d PM _{2.5} ^c PM _{2.5} ^c	Insulin % Difference (95% CI) -1.36 (-2.68, -0.03) 1.45 (-0.18, 3.11) -0.52 (-1.58, 0.55) -0.53 (-2.03, 1.00)	HOMA-IR % Difference (95% CI) -1.55 (-2.97, -0.12) 1.81 (0.05, 3.59) -0.60 (-1.73, 0.54) -0.57 (-2.18, 1.06)	Leptin % Difference (95% CI) -0.83 (-3.87, 2.30) 5.25 (1.17, 9.51) -0.01 (-1.95, 1.98) 1.12 (-1.75, 4.07)	

Supplemental Table E. Sensitivity Analyses for Associations of 2003 Annual Average Fine Particulate Matter (PM_{2.5}) and Residential Proximity to Major Roadways With Adipokines And Biomarkers of Glucose Homeostasis.

Abbreviation: PM_{2.5}, fine particulate matter; HBA1c, hemoglobin A1c; HOMA-IR, homeostasis model assessment of insulin resistance; CI, confidence intervals.

a) Models were adjusted for centered age, (centered age)², sex, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

b) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

c) Results were scaled to equivalent to 1.5 μ g/m³ higher in PM_{2.5} concentrations.

d) Analysis was restricted to participants who lived within 1,000 m from major roadways. Results were scaled to comparing participants who lived $64 \text{ m} (25^{\text{th}} \text{ percentile})$ from major roadways to those who lived $413 \text{ m} (75^{\text{th}} \text{ percentile})$ away.

Supplemental Table F. Associations of the 2003 Annual Average Fine Particulate Matter (PM_{2.5}) Concentrations and Residential Proximity to Major Roadways With Adipokines and Biomarkers of Glucose Homeostasis Among Participants Who Lived Within 50 km From the Central Site ^a.

	Fasting Plasn	na Glucose	HBA	.1c	Adipor	nectin	Resi	stin
	% Difference	95% CI	% Difference	95% CI	% Difference	95% CI	% Difference	95% CI
2003 annual PM _{2.5} ^{b,c}	-0.16	-0.53, 0.21	-0.18	-0.50, 0.13	0.20	-2.86, 3.34	-1.69	-5.32, 2.08
2003 annual $PM_{2.5} + \Delta PM_{2.5}$ ^{c,d}	-0.14	-0.51, 0.24	-0.21	-0.53, 0.12	0.79	-2.31, 3.99	-1.20	-4.90, 2.65
Living closer to major roadways e,f	0.08	-0.21, 0.36	0.02	-0.22, 0.26	-0.45	-2.81, 1.97	2.02	-0.65, 4.77
Distance categories ^c								
<50 m	0.45	-0.21, 1.12	0.03	-0.52, 0.58	-1.31	-6.66, 4.35	2.73	-3.25, 9.09
50-<100 m	0.25	-0.58, 1.09	0.12	-0.59, 0.83	0.91	-5.98, 8.31	-3.72	-11.07, 4.24
100-<200 m	0.04	-0.66, 0.74	0.56	-0.04, 1.15	0.24	-5.47, 6.29	5.23	-1.38, 12.29
200-<400 m	0.26	-0.38, 0.91	0.12	-0.42, 0.65	-0.04	-5.30, 5.52	-2.66	-8.23, 3.25
400-<1,000 m	0 (RI	EF)	0 (RI	EF)	0 (R	EF)	0 (R	EF)
	Insu	lin	HOM	A-IR	Lep	tin	_	
	% Difference	95% CI	% Difference	95% CI	% Difference	95% CI	_	
2003 annual $PM_{2.5}$ ^{b,c}	-0.34	-2.51, 1.88	-0.49	-2.81, 1.88	0.60	-3.31, 4.68		
2003 annual $PM_{2.5} + \Delta PM_{2.5}^{c,d}$	-0.53	-2.71, 1.70	-0.68	-3.01, 1.70	0.25	-3.69, 4.34		
Living closer to major roadways ^{e,f}	0.42	-1.28, 2.16	0.45	-1.37, 2.31	3.92	0.61, 7.35		
Distance categories ^c								
<50 m	1.59	-2.37, 5.72	1.76	-2.49, 6.19	6.14	-1.67, 14.58		
50-<100 m	-3.20	-7.94, 1.77	-3.25	-8.30, 2.08	2.18	-7.00, 12.27		
100-<200 m	-0.26	-4.40, 4.05	-0.29	-4.70, 4.33	-4.91	-12.11, 2.88		
200-<400 m	0.45	-3.39, 4.43	0.41	-3.69, 4.68	-6.30	-12.97, 0.88		
400-<1,000 m	0 (RI	EF)	0 (RI	EF)	0 (R	EF)		

Abbreviation: CI, confidence interval; PM_{2.5}, fine particulate matter; HBA1c, hemoglobin A1c; HOMA-IR, homeostasis model assessment of insulin resistance.

a) Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract

population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

b) Sample size was: 7,007 for fasting plasma glucose; 3,069 for HbA1c; 3,599 for adiponectin; 1,479 for resistin; 4,996 for fasting insulin and HOMA-IR; and 2,129 for leptin.

c) Results were scaled to equivalent to 1.5 μ g/m³ higher in PM_{2.5} concentrations.

d) $\Delta PM_{2.5} = (1 \text{-day moving average of } PM_{2.5} \text{ measured at central site}) - (model-based 2003 annual average } PM_{2.5}).$

e) Analysis was further restricted to participants who lived within 1,000 m from major roadways.

f) Results were scaled to comparing participants who lived 64 m (25th percentile) from major roadways to those who lived 413 m (75th percentile) away.

Supplemental Table G. Associations of the 2003 Annual Average Fine Particulate Matter (PM_{2.5}) Concentrations and Residential Proximity to Major Roadways With Adipokines and Biomarkers of Glucose Homeostasis ^a, Restricting to Participants With Two Measurements.

	Fasting Glucose	Insulin	HOMA-IR
	% Difference (95% CI)	% Difference (95% CI)	% Difference (95% CI)
2003 annual PM _{2.5} ^{b,c} , µg/m ³	-0.17 (-0.38, 0.05)	-0.99 (-2.39, 0.42)	-1.14 (-2.63, 0.37)
Living closer to major roadways d,e	0.22 (-0.03, 0.47)	-0.10 (-1.80, 1.63)	0.07 (-1.75, 1.92)
Distance Category ^d , m			
<50	0.65 (0.07, 1.23)	-1.02 (-4.86, 2.98)	-0.59 (-4.71, 3.72)
50-<100	0.11 (-0.61, 0.83)	-2.96 (-7.53, 1.83)	-2.95 (-7.83, 2.19)
100-<200	0.06 (-0.54, 0.67)	-2.33 (-6.29, 1.79)	-2.58 (-6.79, 1.83)
200-<400	0.47 (-0.09, 1.02)	-1.99 (-5.66, 1.83)	-1.89 (-5.82, 2.20)
400-<1,000	0 (REF)	0 (REF)	0 (REF)

Abbreviation: CI, confidence interval; $PM_{2.5}$, fine particulate matter; HOMA-IR, homeostasis model assessment of insulin resistance. **a)** Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, and sine and cosine season. An exam identifier was added for fasting glucose, insulin, HOMA-IR, HBA1c, and adiponectin.

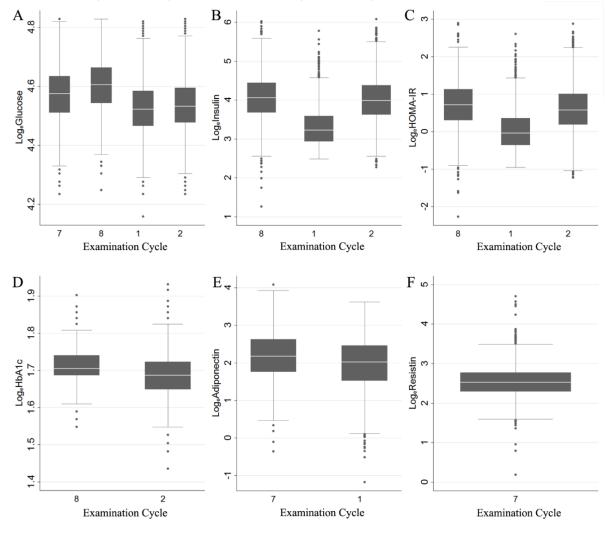
b) Analyses were restricted to participants with two measurements: 4,431 for fasting plasma glucose; and 2,379 for fasting insulin and HOMA-IR.

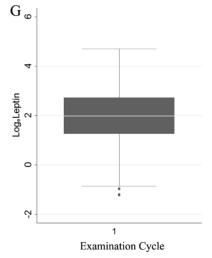
c) Results were scaled to equivalent to $1.5 \,\mu g/m^3$ higher in PM_{2.5} concentrations.

d) Analysis was further restricted to participants who lived within 1,000 m from major roadways.

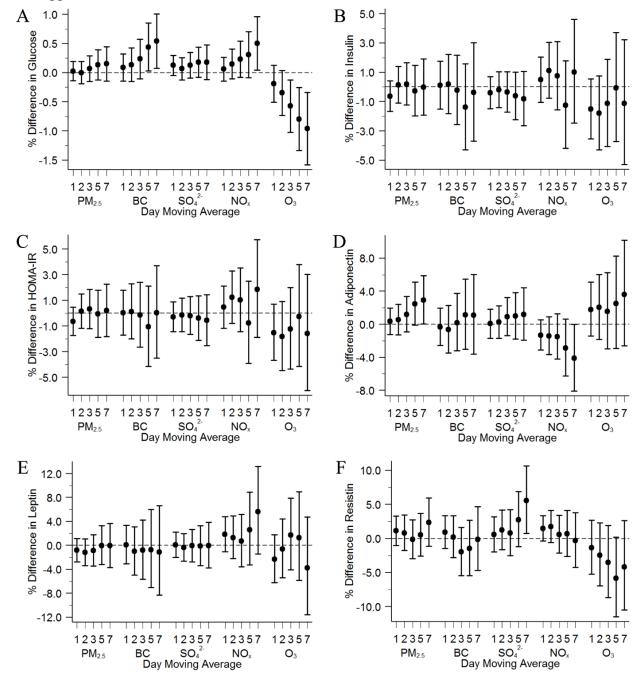
e) Results were scaled to comparing participants who lived 64 m (25th percentile) from major roadways to those who lived 413 m (75th percentile) away.

Supplemental Figure A. Boxplots of A) Fasting Glucose, B) Insulin, C) HOMA-IR, D) HbA1c, E) Adiponectin, F) Resistin, and G) Leptin Among Participants From the Framingham Offspring Cohort Examination 7 (1998-2001), Examination 8 (2005-2008), Third Generation Cohort Examination 1 (2002-2005), or Examination 2 (2008-2011).

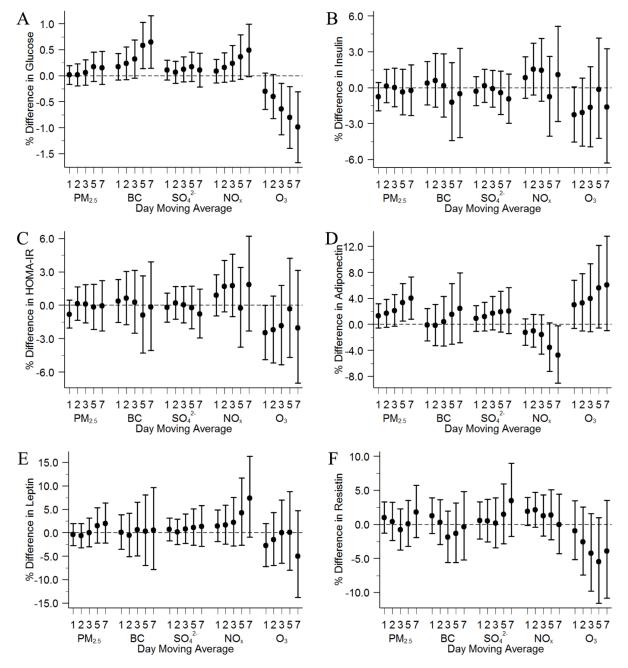




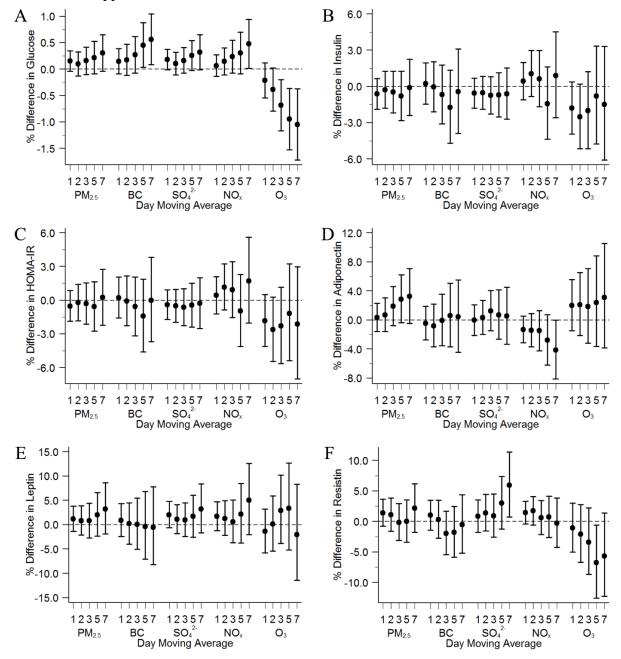
Supplemental Figure B. Associations of **1-**, **2-**, **3-**, **5-**, **and 7-Day Moving Averages** of Air Pollutants With A) Fasting Glucose, B) Insulin, C) HOMA-IR, D) Adiponectin, E) Leptin, and F) Resistin. Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, sine and cosine season, day of week, temperature, and relative humidity. An exam identifier was added for glucose, insulin, HOMA-IR, and adiponectin. Results were scaled to 5 μ g/m³ for fine particulate matter (PM_{2.5}), 0.5 μ g/m³ for black carbon (BC), 2 μ g/m³ for sulfate (SO₄²⁻), 20 ppb for nitrogen oxides (NO_x), and 15 ppb for ozone (O₃). Error bars indicate the 95% confidence intervals.



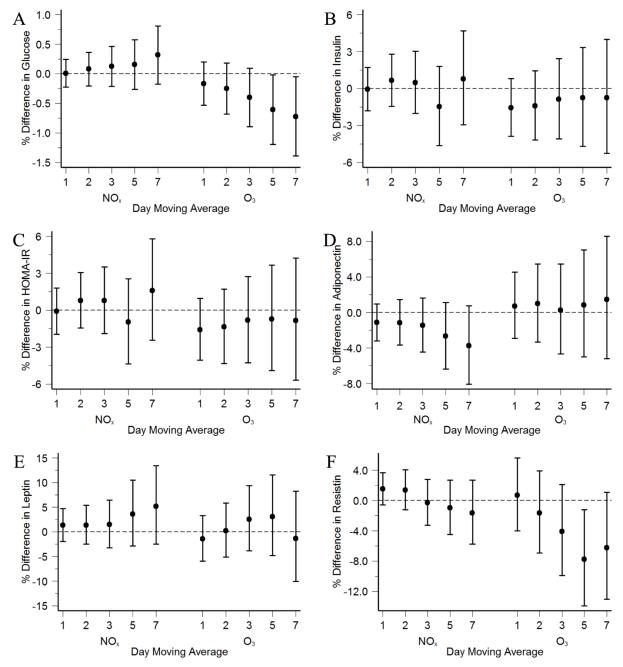
Supplemental Figure C. Associations of 1- to 7-Day Moving Averages of Air Pollutants With A) Fasting Glucose, B) Insulin, C) HOMA-IR, D) Adiponectin, E) Leptin, and F) Resistin Among Participants **Who Lived Within 40 Km From the Central Site.** Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, sine and cosine season, day of week, temperature, and relative humidity. An exam identifier was added for glucose, insulin, HOMA-IR, and adiponectin. Results were scaled to $5 \,\mu g/m^3$ for fine particulate matter (PM_{2.5}), 0.5 $\mu g/m^3$ for black carbon (BC), $2 \,\mu g/m^3$ for sulfate (SO₄²⁻), 20 ppb for NO_x, and 15 ppb for O₃. Error bars indicate the 95% confidence intervals.



Supplemental Figure D. Associations of 1- to 7-Day Moving Averages of Air Pollutants With A) Fasting Glucose, B) Insulin, C) HOMA-IR, D) Adiponectin, E) Leptin, and F) Resistin. **Observations with fine particulate matter (PM2.5)** > **35 µg/m³ in any of the 7 days prior to examination visit were excluded.** Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, sine and cosine season, day of week, temperature, and relative humidity. An exam identifier was added for glucose, insulin, HOMA-IR, and adiponectin. Results were scaled to 5 µg/m³ for fine particulate matter (PM_{2.5}), 0.5 µg/m³ for black carbon (BC), 2 µg/m³ for sulfate (SO₄²⁻), 20 ppb for NO_x, and 15 ppb for O₃. Error bars indicate the 95% confidence intervals.



Supplemental Figure E. Associations of 1- to 7-Day Moving Averages of Air Pollutants With A) Fasting Glucose, B) Insulin, C) HOMA-IR, D) Adiponectin, E) Leptin, and F) Resistin. Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, sine and cosine season, day of week, temperature, and relative humidity. An exam identifier was added for glucose, insulin, HOMA-IR, and adiponectin. Results were scaled to 20 ppb for NO_x and 15 ppb for O₃. Error bars indicate the 95% confidence intervals. **The same day moving averages of both pollutants were included in the same model.**



Supplemental Figure F. Associations of **1-**, **3-**, **and 7-Day Moving Averages** of Air Pollutants With A) Fasting Glucose, B) Insulin, and C) HOMA-IR, **Restricting to Participants With Two Measurements**. Models were adjusted for centered age, (centered age)², sex, body mass index, smoking status, pack years, alcohol intake, educational attainment, physical activity, census tract median household income, census tract median value of owner occupied housing units, census tract population density, usual occupation, date of examination visit, sine and cosine season, day of week, temperature, and relative humidity. An exam identifier was added for glucose, insulin, HOMA-IR, and adiponectin. Results were scaled to $5 \,\mu g/m^3$ for fine particulate matter (PM_{2.5}), $0.5 \,\mu g/m^3$ for black carbon (BC), $2 \,\mu g/m^3$ for sulfate (SO₄²⁻), 20 ppb for nitrogen oxides (NO_x), and 15 ppb for ozone (O₃). Error bars indicate the 95% confidence intervals.

