

PREMATURE DEATHS AND INCIDENCE OF CARDIO-VASCULAR DISEASES RESULTS OF THE EP-PARTICLES STUDY



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CONFLICT OF INTEREST
None declared



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FOUNDING

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The
Ella Roberta
Foundation

“A life cut short. On 15th February 2013, just three weeks after her ninth birthday, Ella died of a fatal asthma attack contributed to by exposure to excessive air pollution.”

CORONER'S RECOMMENDATIONS

1. National air pollution limits should be in line with WHO guidelines.
2. Raise awareness of the dangers of air pollution amongst the general public.
3. Ensure health professionals are aware of the dangers of air pollution, and that they tell their patients about it.

EYES

Because there is a high flow of blood in the eyes, they are especially sensitive to small pollution particles like those found in PM2.5. Conditions such as dry eye syndrome, retinopathy, glaucoma, and cataracts have been connected to high air pollution exposure.



BRAIN

Air pollution exposure has been linked to a variety of neurological and cognitive impacts, including memory impairment, learning disabilities, anxiety, depression, schizophrenia, ADHD, and neurological conditions including dementia, Alzheimer's disease, Parkinson's disease, and stroke. Studies have even linked precise air pollution decreases to lowered dementia risk.



LUNGS

A slew of respiratory impacts are attributed to dirty air, from respiratory inflammation to asthma development to chronic loss of pulmonary function. Because most air pollutants are breathed in, the respiratory system is often the place where air pollution-related disease is most readily observed.



HEART

Cardiovascular disease and death are closely linked to air pollution, with outcomes of heart disease, heart failure, cardiac arrest, and arrhythmias. Some studies have even shown a stronger correlation between cardiovascular damage and death after air pollution exposure than observed with respiratory diseases.



STOMACH

Studies have demonstrated a link between poor air quality and gastrointestinal diseases, including inflammatory bowel disease (IBD), irritable bowel syndrome (IBS), and appendicitis, as the inhalation of air pollution is associated with changes to the gut microbiota. Long-term exposure to high concentrations of NO2 and PM has been linked to the early onset of Crohn's Disease.



LIVER

Animal studies find that long-term exposure to ambient air pollution is associated with metabolic-associated fatty liver disease, a disease that affects a quarter of the global population. The disease can progress to liver cancer and liver-related death.



BONES

A 2020 study found that ambient air pollution exposure is linked to lower bone mass. Other studies have found that air pollution may act as a risk factor for osteoporosis and be linked to higher rates of hospitalization for bone fractures, though research is limited.



REPRODUCTION

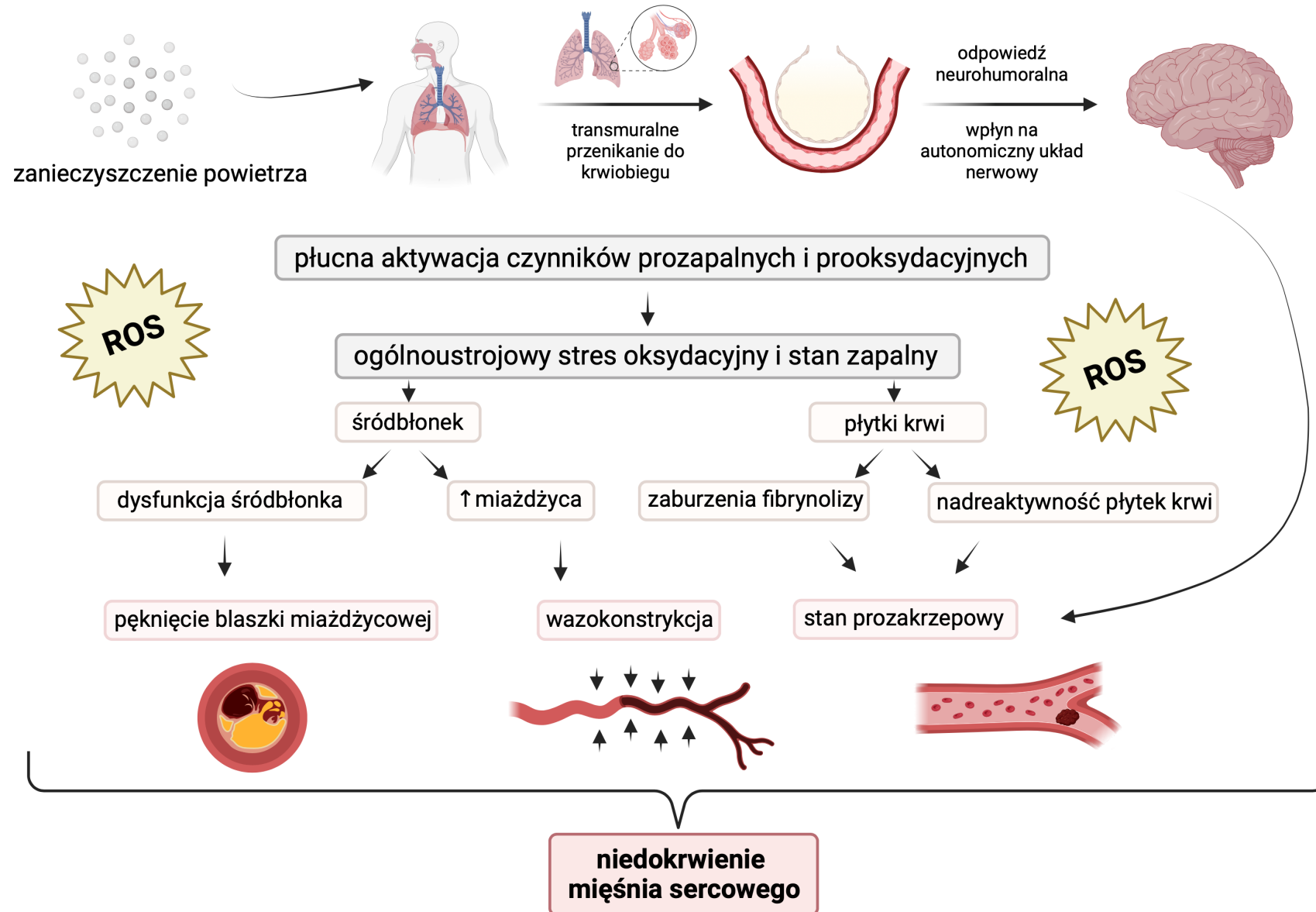
Though the mechanisms behind it are not yet understood, exposure to higher levels of air pollution has been associated with lower levels of fertility and more difficulty in conceiving, including in those undergoing in vitro fertilization, as well as in a variety of sperm quality parameters. Further research is needed to investigate how exactly air pollution acts on the reproductive system.



Zanieczyszczenie powietrza powoduje:

ponad **8 milionów** dodatkowych **zgonów** rocznie na całym świecie

globalną średnią utratę oczekiwanej długości życia wynoszącą **2,9 roku**



Grafika 1. Potencjalne patomechanizmy wpływu zanieczyszczeń powietrza

Exposure to air pollution—a trigger for myocardial infarction? A nine-year study in Białystok—the capital of the Green Lungs of Poland (BIA-ACS registry)

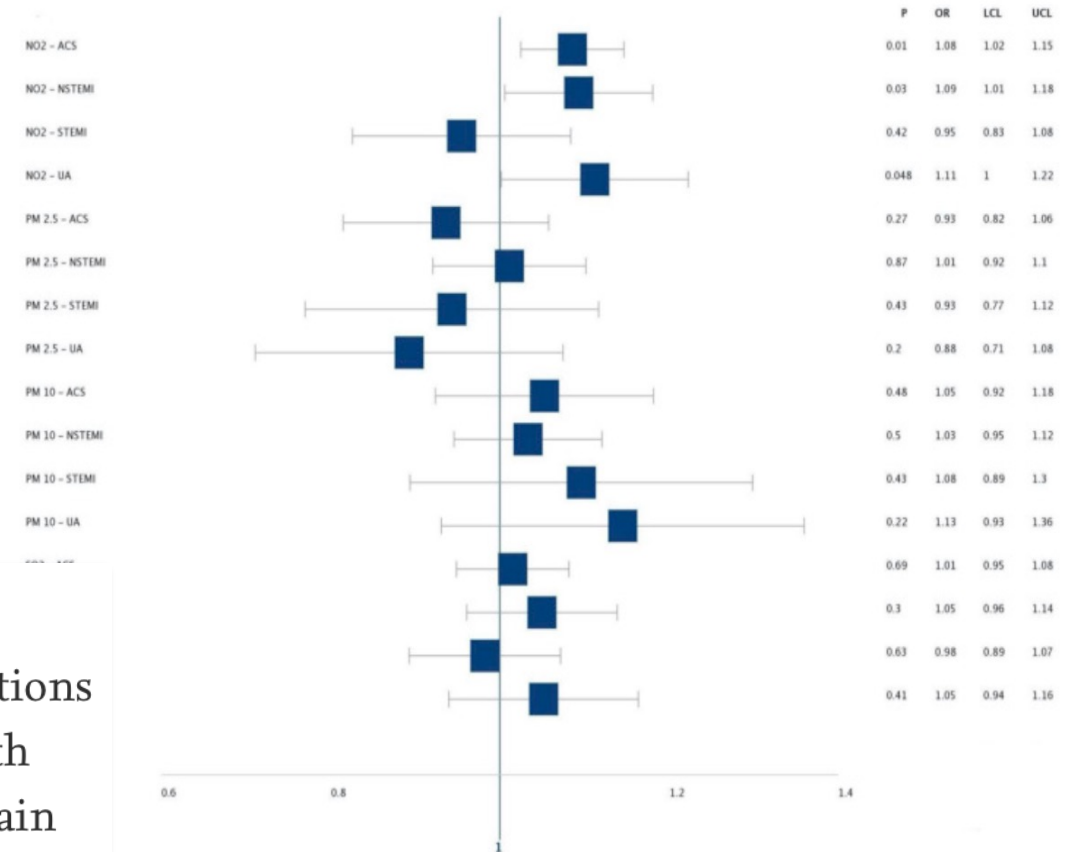
Łukasz Kuźma ^a ✉, Szymon Pogorzelski ^a, Krzysztof Struniawski ^a, Hanna Bachórzewska-Gajewska ^a,
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Conclusion

The study showed that the effects of air pollution and weather conditions on the number of ACS hospitalizations are also observed in cities with moderately polluted or good air quality. NO₂ was identified as the main air pollutant affecting the incidence of ACS.



Gender Differences in Association between Air Pollution and Daily Mortality in the Capital of the Green Lungs of Poland—Population-Based Study with 2,953,000 Person-Years of Follow-Up



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Variables	RR	Lower 95% CI for RR	Upper 95% CI for RR	p
NO ₂ µg/m ³ * + Meteorological parameters	0.99	0.96	1.04	0.96
SO ₂ µg/m ³ * + Meteorological parameters	1.10	1.04	1.17	0.002
PM _{2.5} µg/m ³ ** + Meteorological parameters	1.03	0.99	1.06	0.13
PM ₁₀ µg/m ³ ** + Meteorological parameters	0.99	0.96	1.03	0.99
Temp. °C ***	1.01	0.99	1.04	0.29

Variables	RR	Lower 95% CI for RR	Upper 95% CI for RR	p
NO ₂ µg/m ³ * + Meteorological parameters	1.02	0.97	1.07	0.40
SO ₂ µg/m ³ * + Meteorological parameters	1.02	1.00	1.04	0.08
PM _{2.5} µg/m ³ ** + Meteorological parameters	1.07	1.02	1.12	0.01
PM ₁₀ µg/m ³ ** + Meteorological parameters	0.95	0.90	1.01	0.60
Temp. °C ***	1.02	0.98	1.07	0.25

Variables	RR	Lower 95% CI for RR	Upper 95% CI for RR	p
NO ₂ µg/m ³ * + Meteorological parameters	0.98	0.94	1.03	0.31
SO ₂ µg/m ³ * + Meteorological parameters	1.05	1.01	1.10	0.009
PM _{2.5} µg/m ³ ** + Meteorological parameters	0.99	0.96	1.03	0.73
PM ₁₀ µg/m ³ ** + Meteorological parameters	1.03	0.98	1.07	0.17
Temp. °C ***	1.04	1.02	1.07	0.003

Variables	RR	Lower 95% CI for RR	Upper 95% CI for RR	p
NO ₂ µg/m ³ * + Meteorological parameters	0.99	0.93	1.05	0.69
SO ₂ µg/m ³ * + Meteorological parameters	1.02	0.99	1.05	0.19
PM _{2.5} µg/m ³ ** + Meteorological parameters	1.00	0.95	1.05	0.86
PM ₁₀ µg/m ³ ** + Meteorological parameters	1.03	0.97	1.09	0.36
Temp. °C ***	1.08	1.04	1.13	<0.001

Conclusions

1. Air quality and atmospheric conditions had an impact on the mortality of Białystok residents.
2. The main air pollutant that influenced the mortality rate was SO₂, and there were no gender differences in the impact of this pollutant. In the male population, an increased exposure to PM_{2.5} concentration was associated with significantly higher cardiovascular mortality.

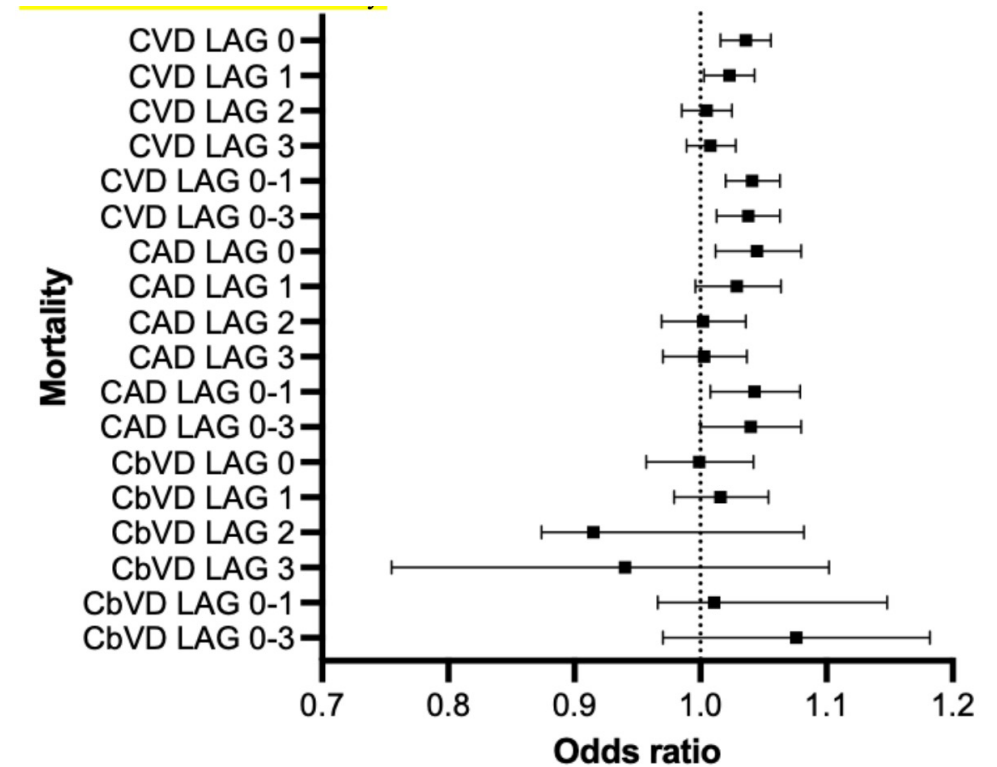
Association between air pollution and case-specific mortality in north-eastern part of Poland. Case-crossover study with 4,500,000 person-years of follow-up (PL-PARTICLES study)

Objectives: To assess the short-term impact of air pollution on cardiovascular (CVD)-, coronary artery-related (CAD)-, and cerebrovascular-related (CbVD) mortality.

Patients and methods: The analysis with 4,500,000 person-years of follow-up with a time-stratified case-crossover design was performed.

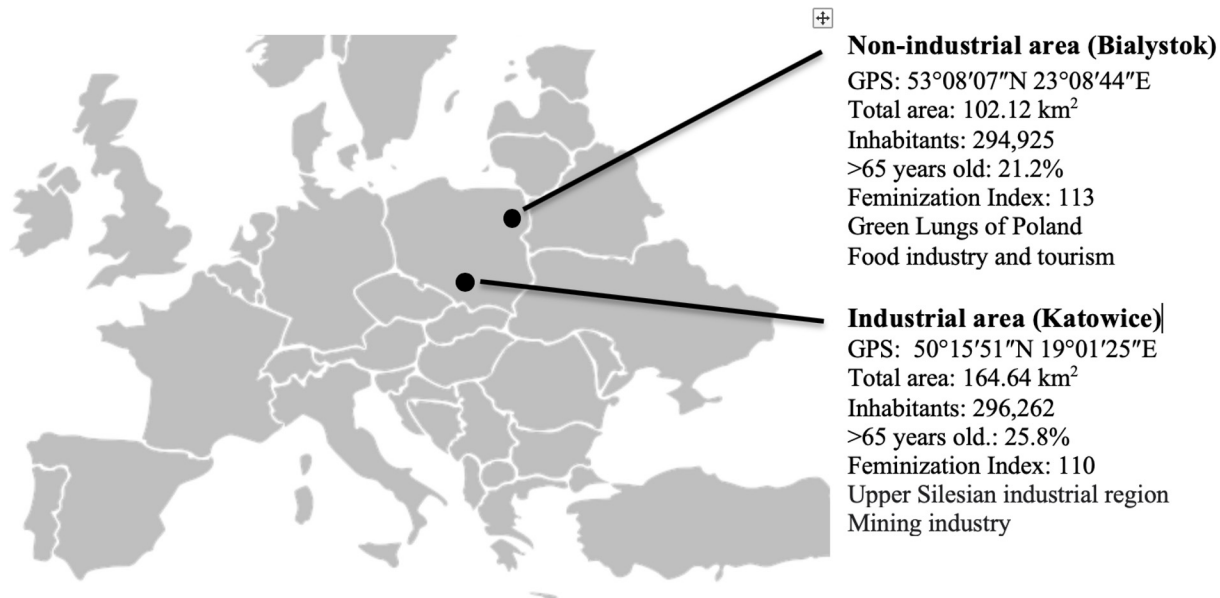
Results: The interquartile range (IQR) increase in $PM_{2.5}$ (OR 1.036, 95%CI 1.016–1.056, $P < 0.001$) and PM_{10} concentration (OR 1.034, 95%CI 1.015–1.053, $P < 0.001$) was associated with increased CVD mortality on lag 0 and this effect persisted on following days. The effects of PMs were more expressed in association with CAD-related mortality (OR for $PM_{2.5}$ = 1.045, 95%CI 1.012–1.080, $P = 0.008$), (OR for PM_{10} = 1.044, 95%CI 1.010–1.078, $P = 0.011$). Additionally, IQR increase in NO_2 concentration was associated with increased CAD-related mortality at lag 0-1 (OR = 1.055, 95%CI 1.004–1.108, $P = 0.032$).

Conclusions: The impact of PMs on CVD mortality is also observed in moderately polluted areas. This adverse health effect was more apparent in CAD mortality. Differences in effect size and seasonality may depend on the source of air pollution.



Impact of short-term air pollution exposure on acute coronary syndrome in two cohorts of industrial and non-industrial areas: A time series regression with 6,000,000 person-years of follow-up (ACS - Air Pollution Study)

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6,000,000 person - years

Data on hospitalization for ACS from 2008 to 2017 was obtained and extracted from the National Health Fund reports. We used data from patients registered as residents in the city of Białystok – non industrial area and Katowice city - industrial area. The limit of the daily mean value for PM_{2.5} according to the WHO guidelines was exceeded on 45.2% days in industrial area and on 24.9% days in non-industrial area. The daily WHO upper limit for PM₁₀ was exceeded on 27.6% days in industrial area and on 9.1% days in non-industrial area. The WHO limit for SO₂ was exceeded sporadically in non-industrial area (0.4%) and on 18.4% days in industrial area.

Exposure to Air Pollution and Renal Function - An Underestimated Threat?

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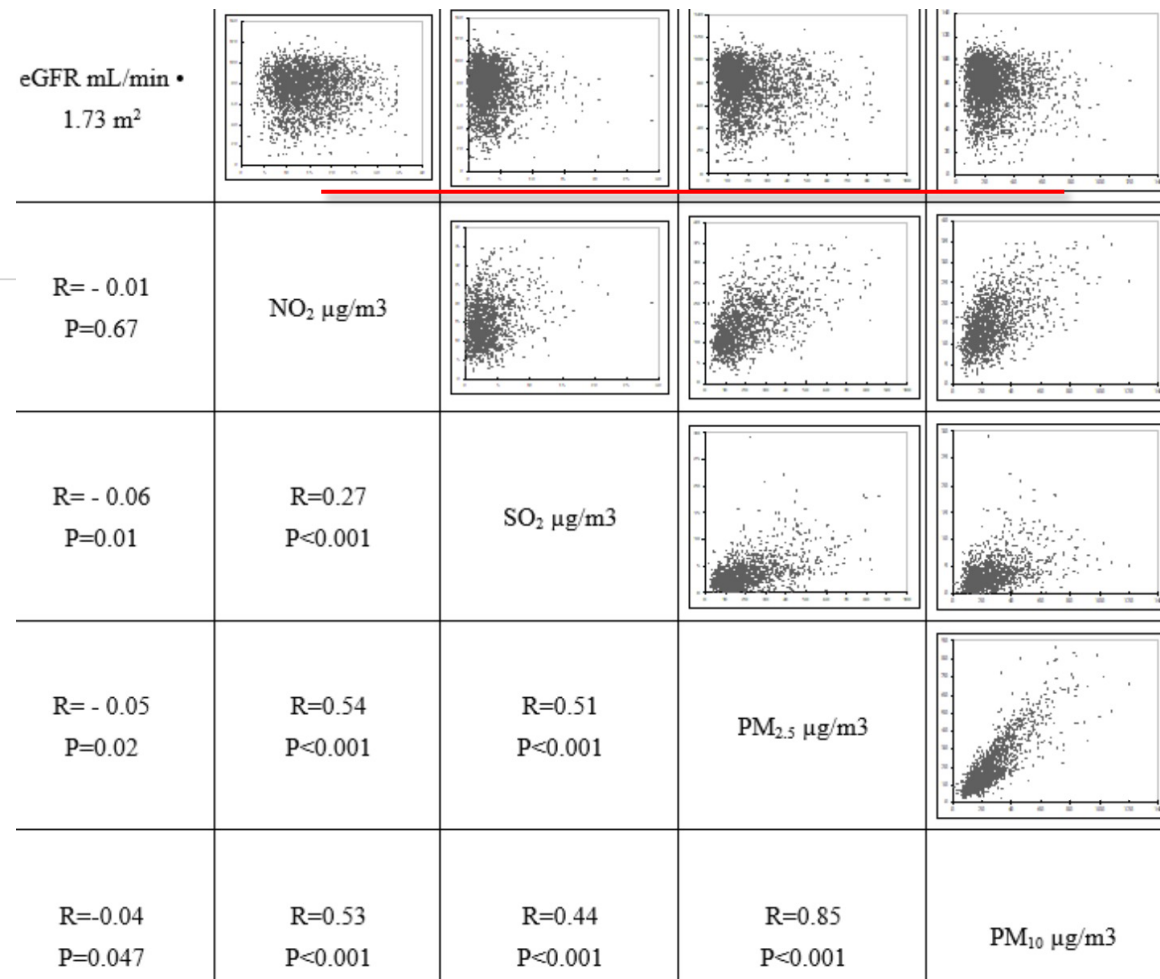
Medical University of Białystok

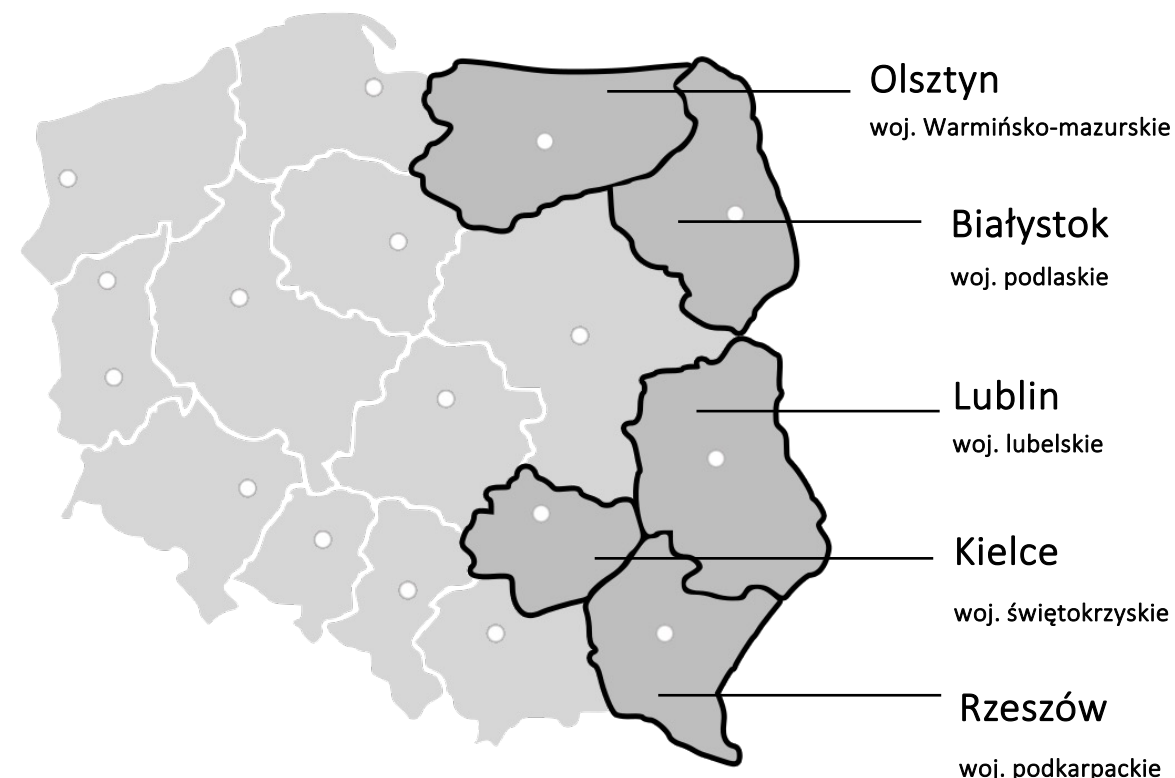
Sławomir Dobrzycki

Medical University of Białystok

Results: 3,554 patients were included into the final analysis. The median age was 66 (IQR 15) and men were in the majority (53.2%, N=1891). Chronic kidney disease (CKD) was diagnosed in 21.5% (N=764). The long-term increase in annual average concentration of PM_{2.5} (OR for IQR increase=1.07; 95% CI 1.01 – 1.15, P=0.037) and NO₂ (OR for IQR increase=1.05; 95% CI 1.01 – 1.10, P=0.047) resulted in an increased number of patients with CKD. In short-term observation the IQR increase in weekly PM_{2.5} concentration was associated with a 2% reduction in eGFR (OR=0.98, 95%CI 0.97 – 0.99, P=0.03)

Conclusions: The effects of air pollution on renal function were observed. Long- and short-term exposure to elevated air pollution levels was associated with a decrease in eGFR. The main pollutant affecting the kidneys was PM_{2.5}.





Zgony N=831 246

Zgony **CVD** N=**377 344** (45,5%)

Hospitalizacje CVD N=2 141 213

139 697 przypadków **ACS** (45,2% STEMI)

Analiza statystyczna
na poziomie gmin dla wzrostu o $10 \mu\text{g}/\text{m}^3$ stężeń zanieczyszczeń



analiza czasowo-przestrzenna

I etap

Oszacowanie wpływu zanieczyszczenia powietrza w poszczególnych powiatów przy użyciu uogólnionych modeli addytywnych z regresją Poissona. W analizie uwzględniono wpływ dnia tygodnia, świąt państwowych, okresów epidemii grypy i SARS-CoV-2 oraz naturalne splajny z 6 stopniami swobody (df) dla 0-1-dniowej średniej ruchomej temperatury i 0-6 średnich ruchomych i 3 df dla wilgotności względnej i ciśnienia atmosferycznego, tendencje sezonowe, naturalny splajn z 6 df na rok),



II etap

Metaanaliza z zastosowaniem efektów losowych
Heterogeniczność między gminami określono ilościowo za pomocą statystyki I².

EP-PARTICLES COHORT

Observation years: 2011- 2020

Main outcomes

3,141,213 hospitalizations due to CVD

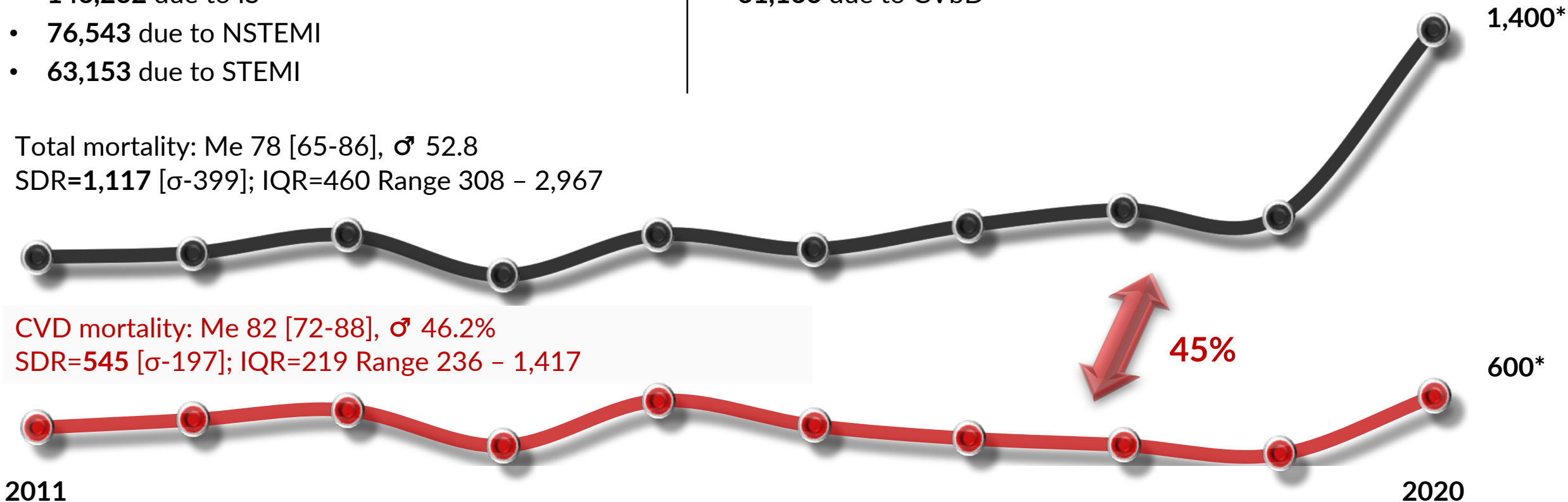
- 822,144 due to HF
- 252,566 due to AF
- 146,262 due to IS
- 76,543 due to NSTEMI
- 63,153 due to STEMI

831,246 recorded deaths

- 377,344 due to CVD
- 83,031 due to ACS
- 61,166 due to CVbD

Total mortality: Me 78 [65-86], ♂ 52.8
SDR=1,117 [σ -399]; IQR=460 Range 308 - 2,967

CVD mortality: Me 82 [72-88], ♂ 46.2%
SDR=545 [σ -197]; IQR=219 Range 236 - 1,417



*Age standardized death rate /100,000 population/year – based on European Standard Population (ESP) structure

EP-PARTICLES COHORT



Observation years: 2011- 2020

Aggregated at NUTS3

	GDP	Mean age	Fem index	BMI >25	Tobacco use	Alcohol use*	HT**	DM**	ESRD**
Poland	50,050	42.1	108	53%	29.1%	74.7%	315	86.4	74.1

Lubelskie	31,000	39.8	107	45.9%	27%	71%	342	87.2	74.2
Podkarpackie	33,417	41.6	104	52%	23.7%	71.4%	314	82.5	73.1
Podlaskie	32,547	42.3	110	44%	31%	77%	291	89	82.4
Świętokrzyskie	33,229	42.5	105	54.8%	25%	71%	323	90.6	83.3
Warmińsko-mazurskie	29,312	41.8	104.2	46.2%	28%	74%	296	78.9	57.5
	P=0.48	P<0.001	P<0.001	P<0.001	P<0.001	P<0.001	P<0.001	P<0.001	P<0.001

* more than one time per week

**per /1,000 population

Combined data from National Health Found Reports/Central Statistical Office

Abbreviations: ESRD, end-stage renal disease; DM, diabetes mellitus; HT, hypertension

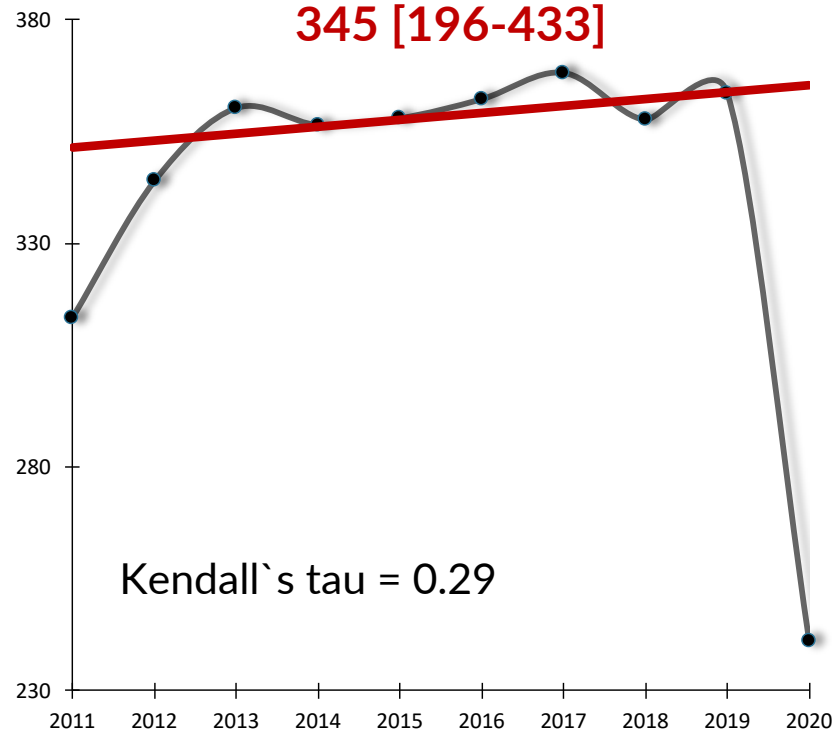
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Time trends (aggregated at LAU 2)

Atrial fibrillation

345 [196-433]

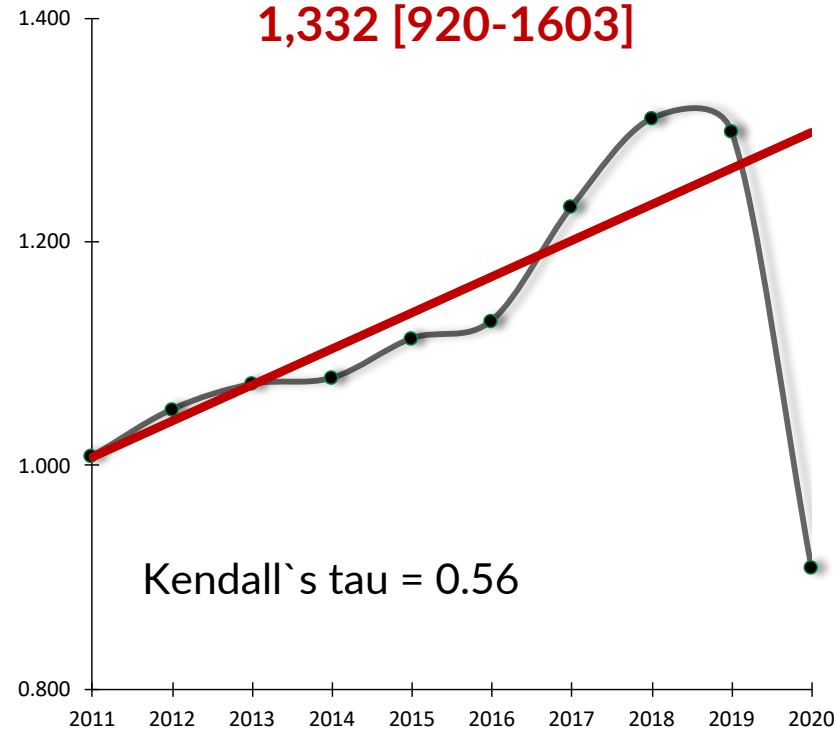


Kendall`s tau = 0.29

Sen`s slope:
1.54 (-2.33 - 5.07)
P=0.29

Heart Failure

1,332 [920-1603]

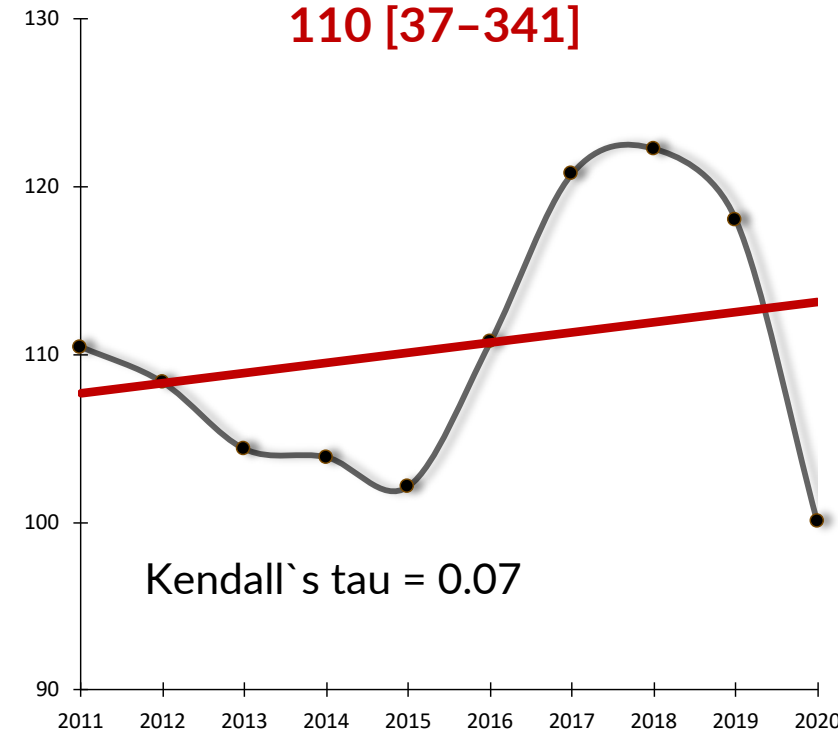


Kendall`s tau = 0.56

Sen`s slope:
0.03 (0.01 - 0.04)
P=0.03

NSTEMI

110 [37-341]



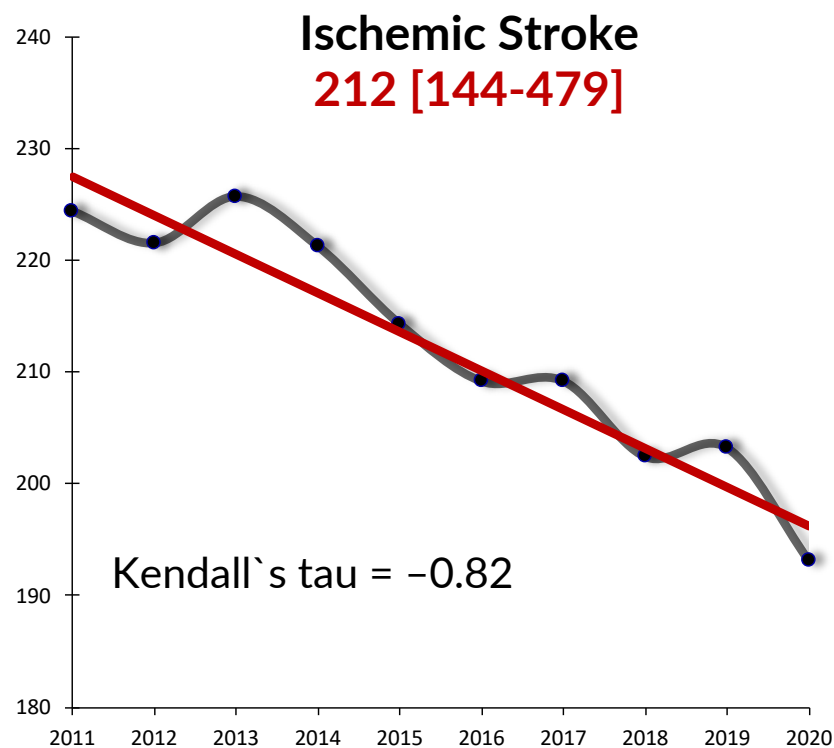
Kendall`s tau = 0.07

Sen`s slope:
0.6 (-2.06 - 2.49)
P=0.86

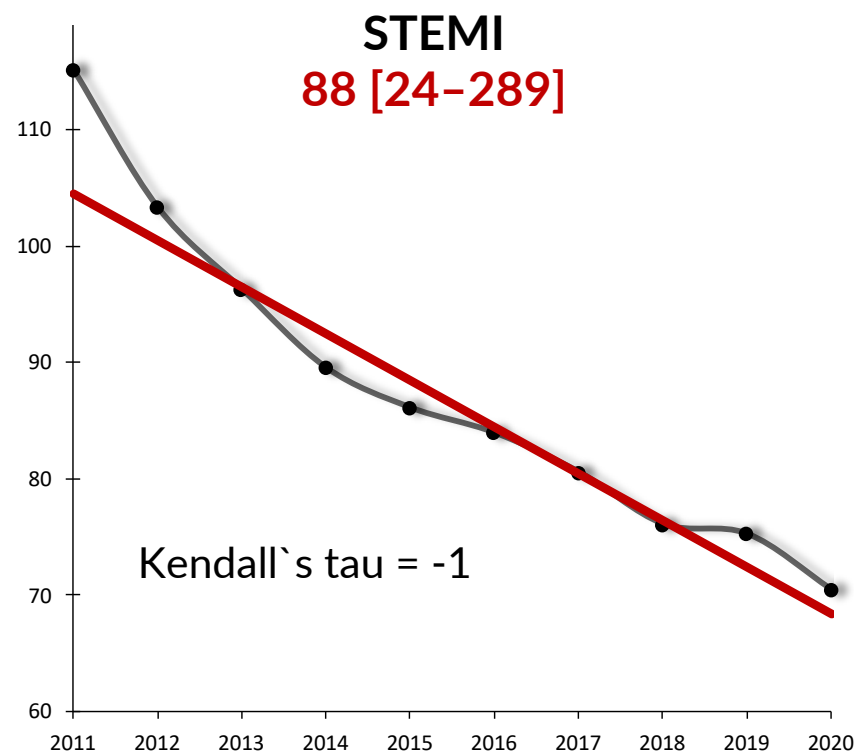
*Age standardized rate /100,000 population/year - based on European Standard Population (ESP) structure

EP-PARTICLES COHORT

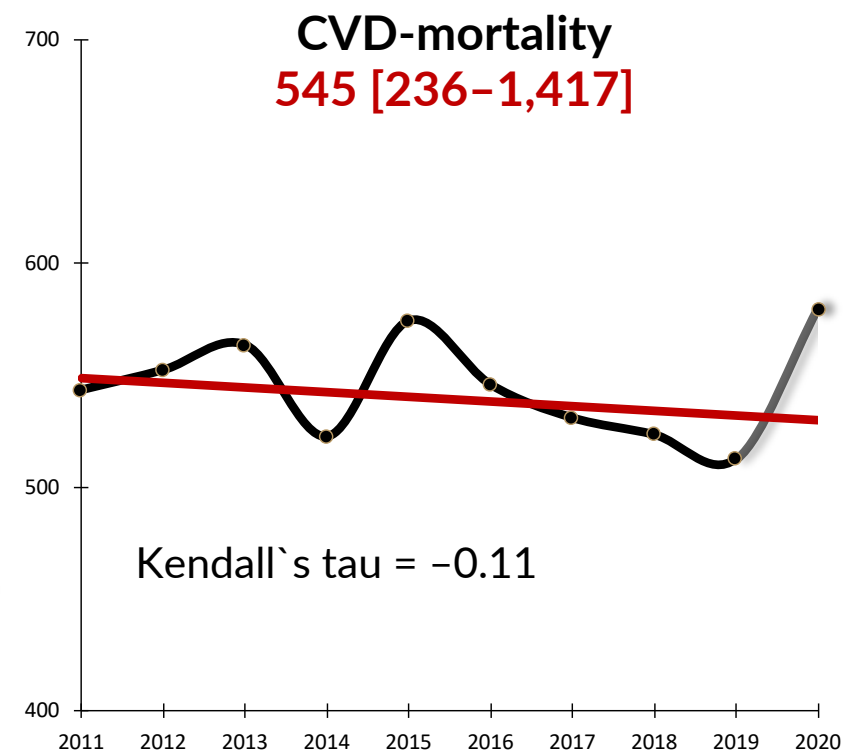
Time trends (aggregated at LAU 2)



Sen`s slope:
-3.47 (-4.65 - -2.62)
P<0.001



Sen`s slope:
-4.01 (-5.08 - -3.33)
P<0.001


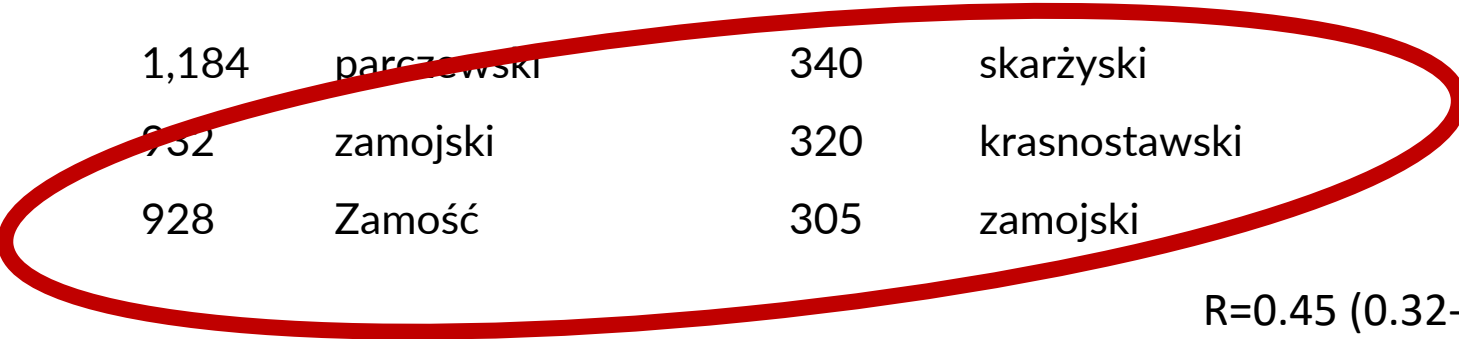



Sen`s slope:
-2.09 (-8.46 - 5.45)
P=0.73

*Age standardized rate /100,000 population/year - based on European Standard Population (ESP) structure

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Geographical variation (aggregated at LAU 1)

	Heart Failure*		Atrial Fibrillation		Ischemic Stroke		
Max 	3,326	bieszczadzki	1,184	parczewski	340	skarżyski	
	3,114	włodawski	952	zamojski	320	krasnostawski	
	2,748	parczewski	928	Zamość	305	zamojski	
Me	Age 75 [63-82] ♂ 51.4% Skewness 1.24 Kurtosis 3.54		Age 70 [63-79] ♂ 47.2% Skewness 1.42 Kurtosis 2.43		Age 76 [66-83] ♂ 48% Skewness 0.96 Kurtosis 1.98		R=0.45 (0.32-0.62)
Min 	478	rycki	108	bieszczadzki	123	Suwałki	
	439	ostrowiecki	63	kolbuszowski	105	leski	
	404	piski	60	kazimierski	65	kazimierski	

*Age standardized rate /100,000 population/year – based on European Standard Population (ESP) structure

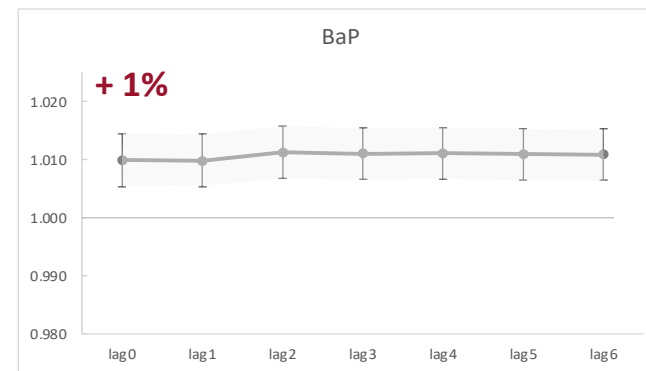
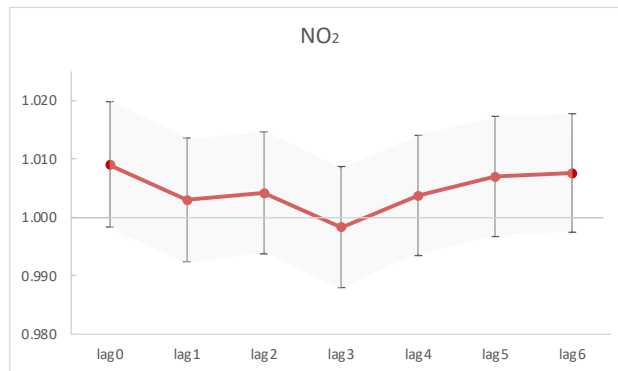
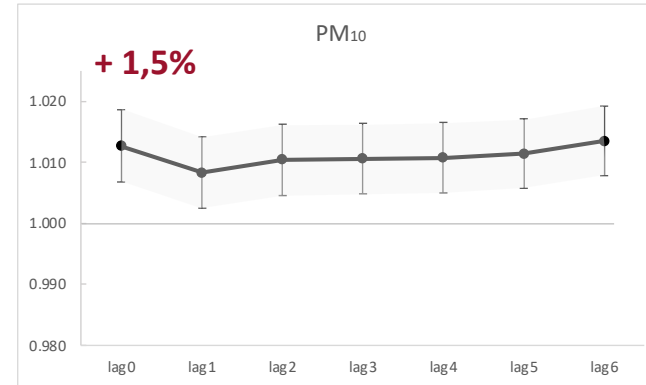
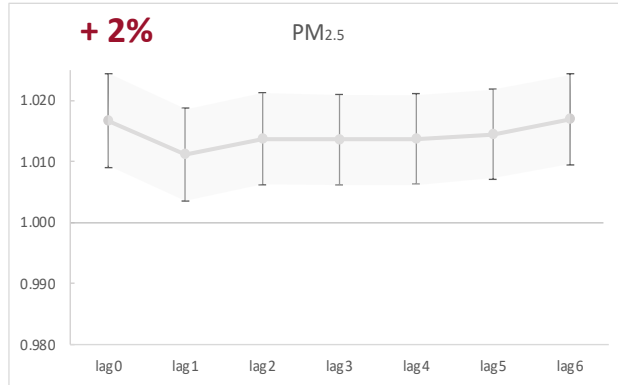
EP-PARTICLES COHORT

Geographical dispersion (aggregated at LAU 1)

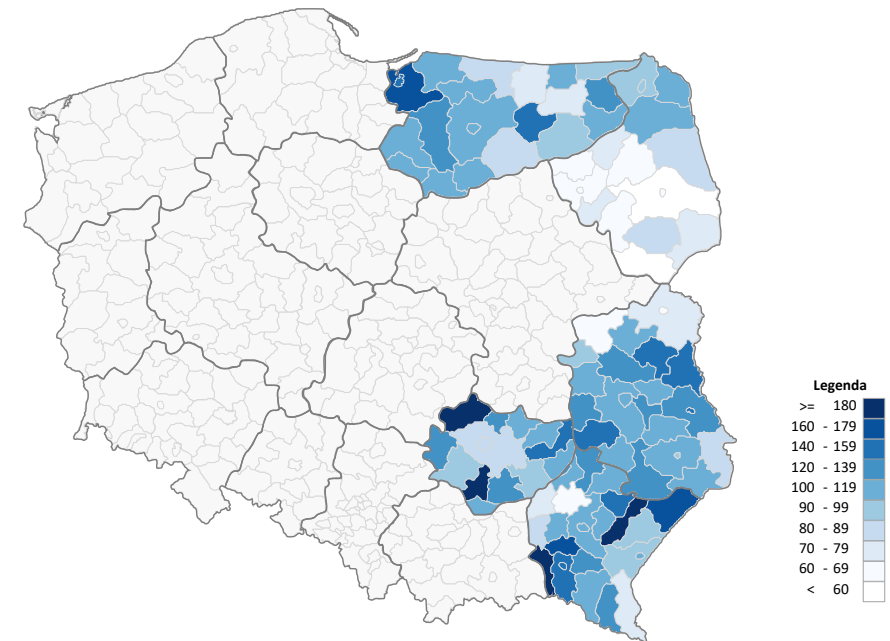
R=0.35 (0.15-0.52)

	STEMI*		NSTEMI		CVD mortality	
Max ↑	174	sandomierski	209	pińczowski	1,148	Chełm
	154	skarżyski	200	jasielski	1,141	Zamość
	153	jędrzejowski	191	konecki	1,121	Krosno
Me	Age 67 [59-77] ♂ 64% Skewness 1.43 Kurtosis 4.58		Age 70 [62-80] ♂ 60% Skewness 0.98 Kurtosis 2.39		Age 82 [72-88] ♂ 46.2% Skewness 0.96 Kurtosis 1.98	
	58	łukowski	60	białostocki	188	suwalski
	52	dębicki	53	Białystok	178	przemyski
Min ↓	37	grajewski	50	siemiatycki	174	chełmski

*Age standardized rate /100,000 population/year – based on European Standard Population (ESP) structure

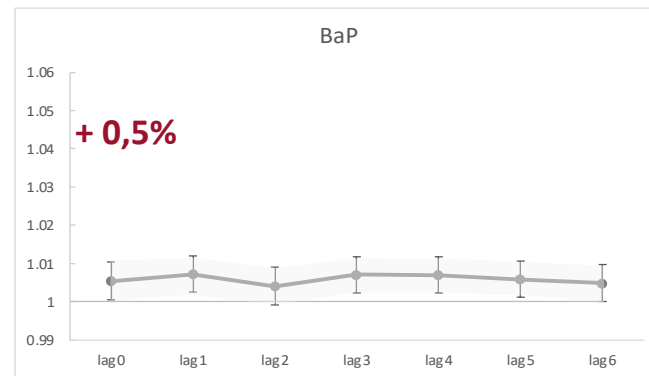
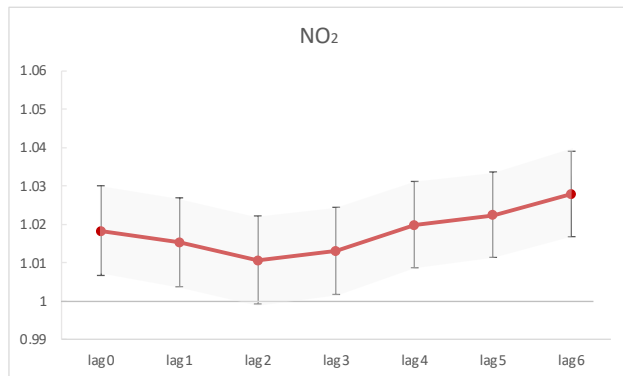
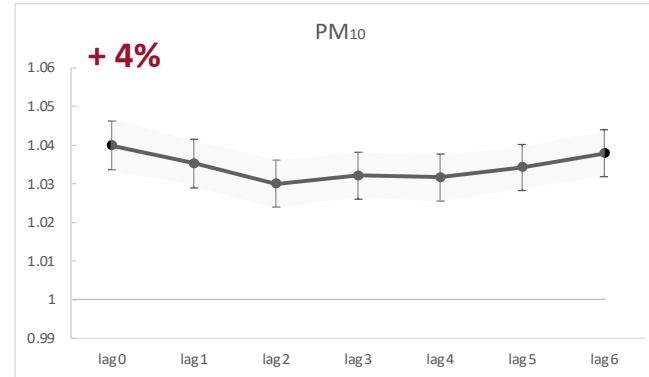
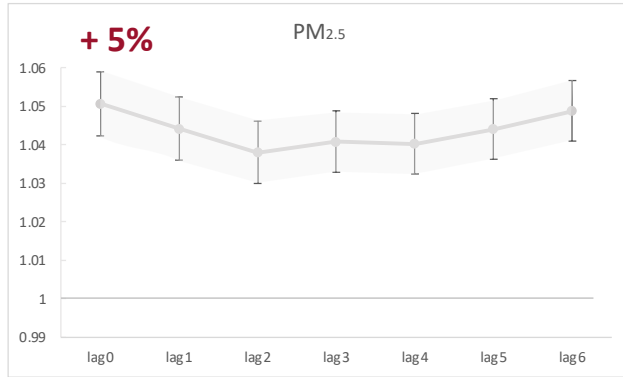


Chorobowość hospitalizowana NSTEMI

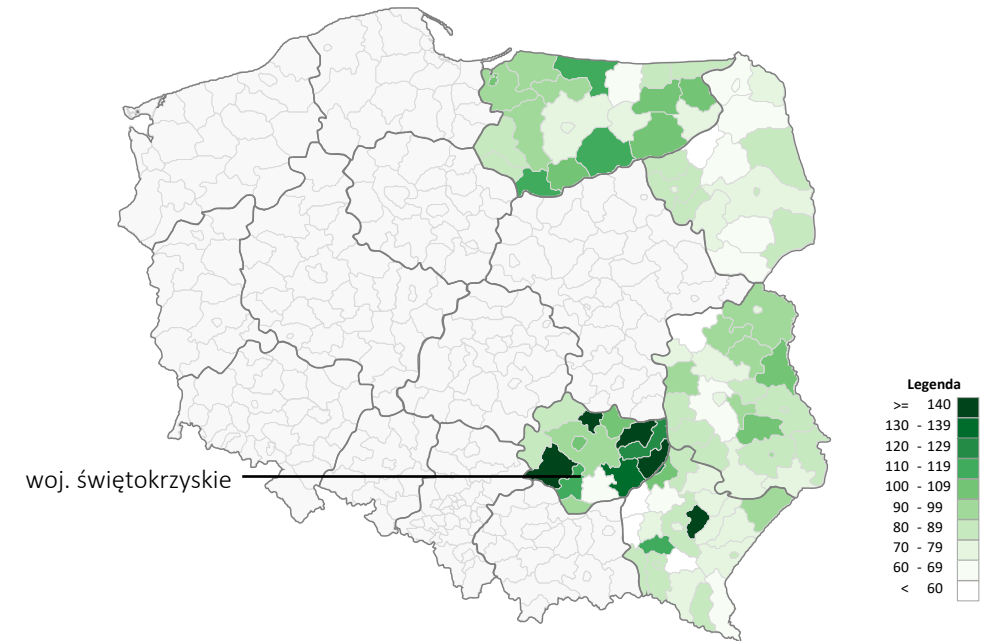


Standaryzowana wg wieku i płci liczba przypadków/100 tys./rok
(Standardowe populacje europejska – EP27)

Rycina 1. Zwiększone hospitalizacje NSTEMI związane ze wzrostem o 10 µg/m³ stężeń zanieczyszczeń

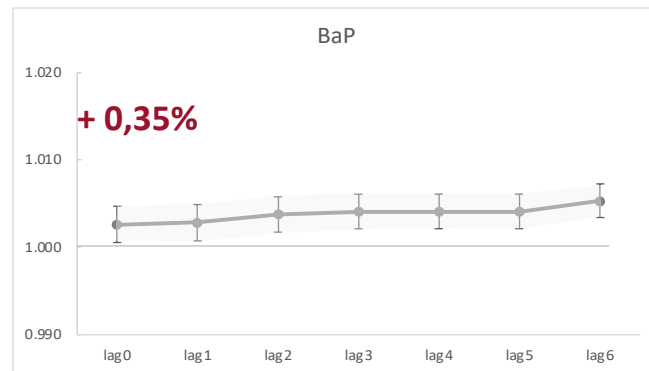
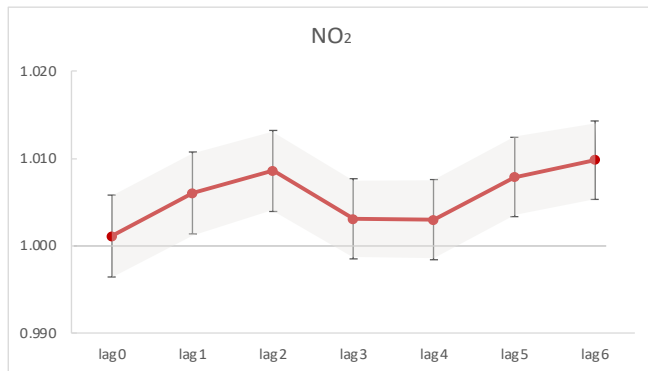
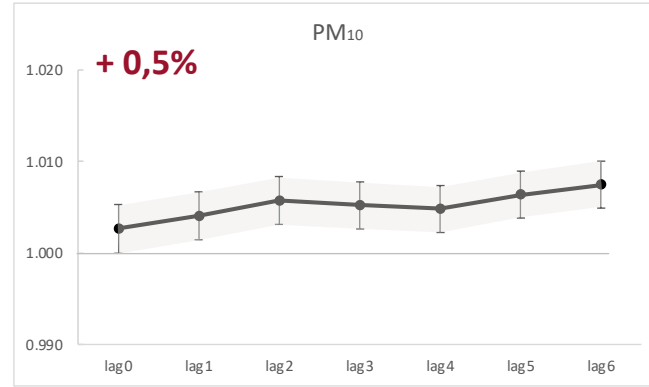
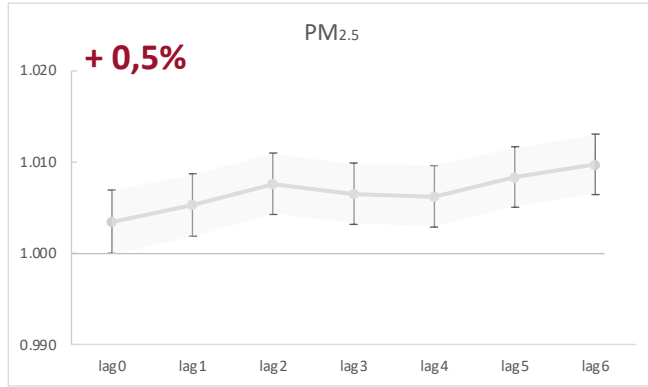


Chorobowość hospitalizowana STEMI

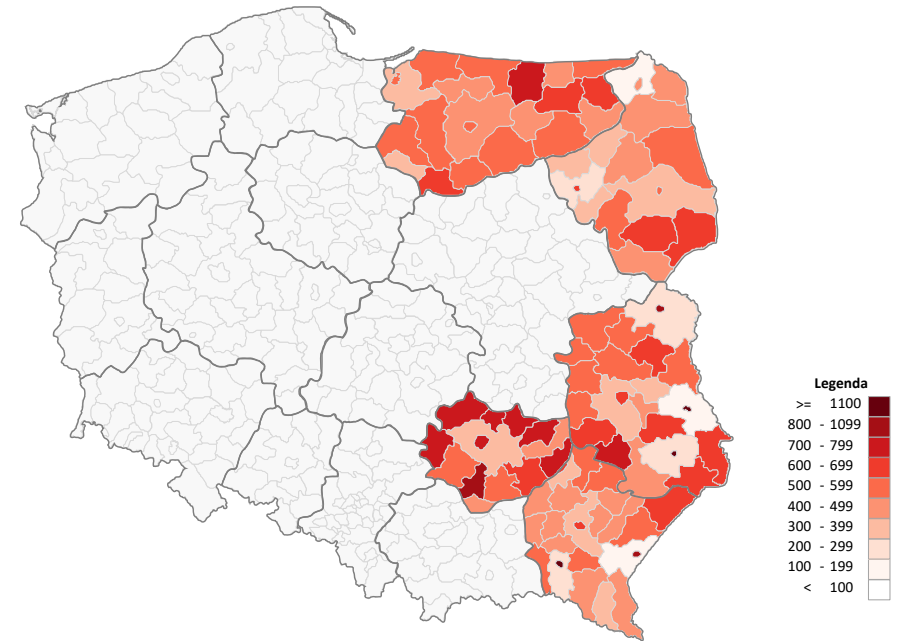


Standaryzowana wg wieku i płci liczba przypadków/100 tys./rok
(Standardowe populacje europejska – EP27)

Rycina 2. Zwiększone hospitalizacje STEMI związane ze wzrostem o 10 µg/m³ stężeń zanieczyszczeń



Śmiertelność sercowo-naczyniowa



Standaryzowana wg wieku i płci liczba zgonów/100 tyś/rok
(Standardowe populacje europejska – EP27)

Zmiany występowania incydentów sercowo-naczyniowych związanych z miesięcznym wzrostem stężenia:

STEMI (RR=**1.214**, 95%CI 1.002 - 1.009), P=0.005

NSTEMI (OR=**1.017**, 95%CI 1.009 - 1.026), P<0.001

CVD mortality (OR=**1.057** 95%CI 1.048 - 1.067), P<0.001

Overall (OR=**1.036**, 95%CI 1.028 - 1.053), P<0.001

na wzrost o IQR PM_{2.5} µg/m³

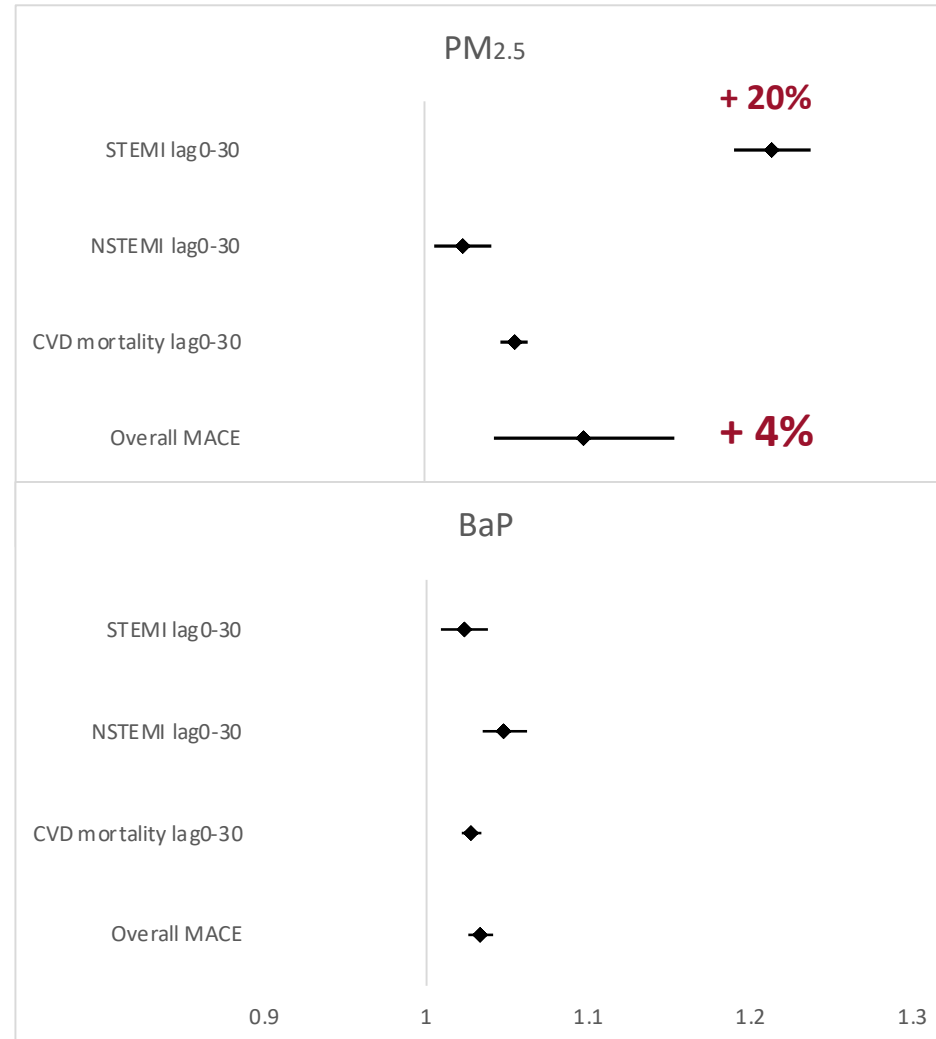
STEMI (RR=**1.024**, 95%CI 1.009 - 1.038), P=0.005

NSTEMI (OR=**1.048**, 95%CI 1.035 - 1.062), P<0.001

CVD mortality (OR=**1.028** 95%CI 1.022 - 1.034), P<0.001

Overall (OR=**1.033**, 95%CI 1.026 - 1.041), P<0.001

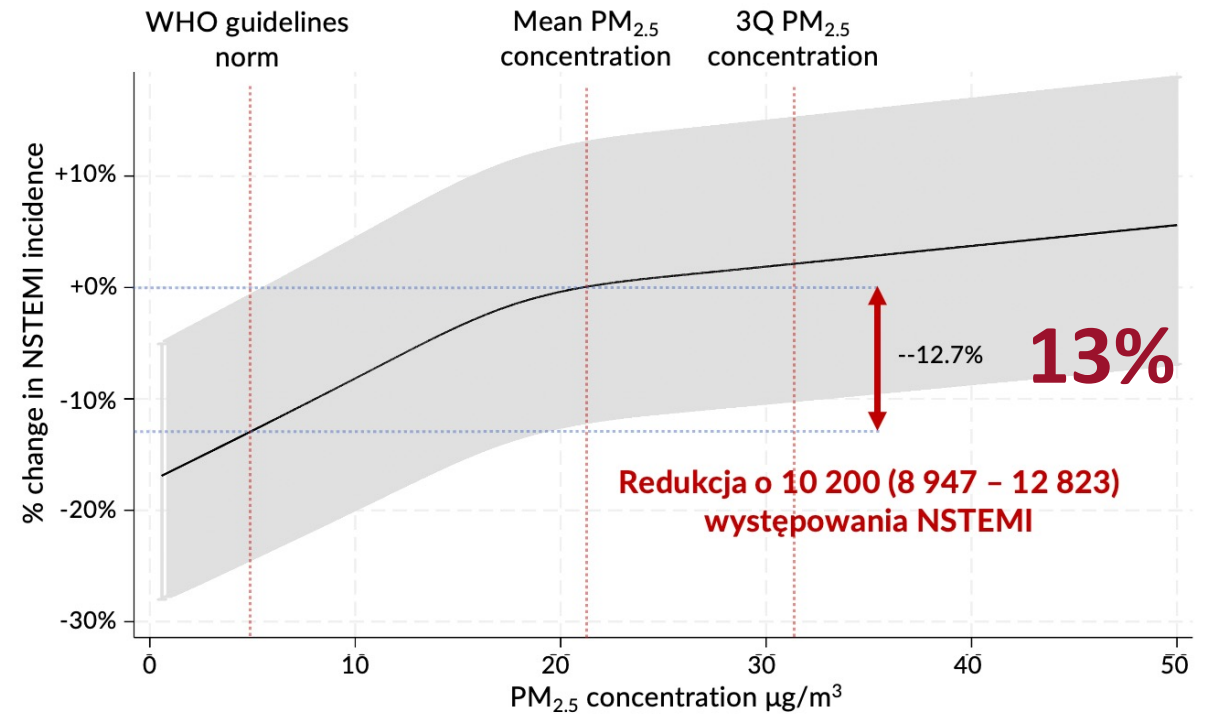
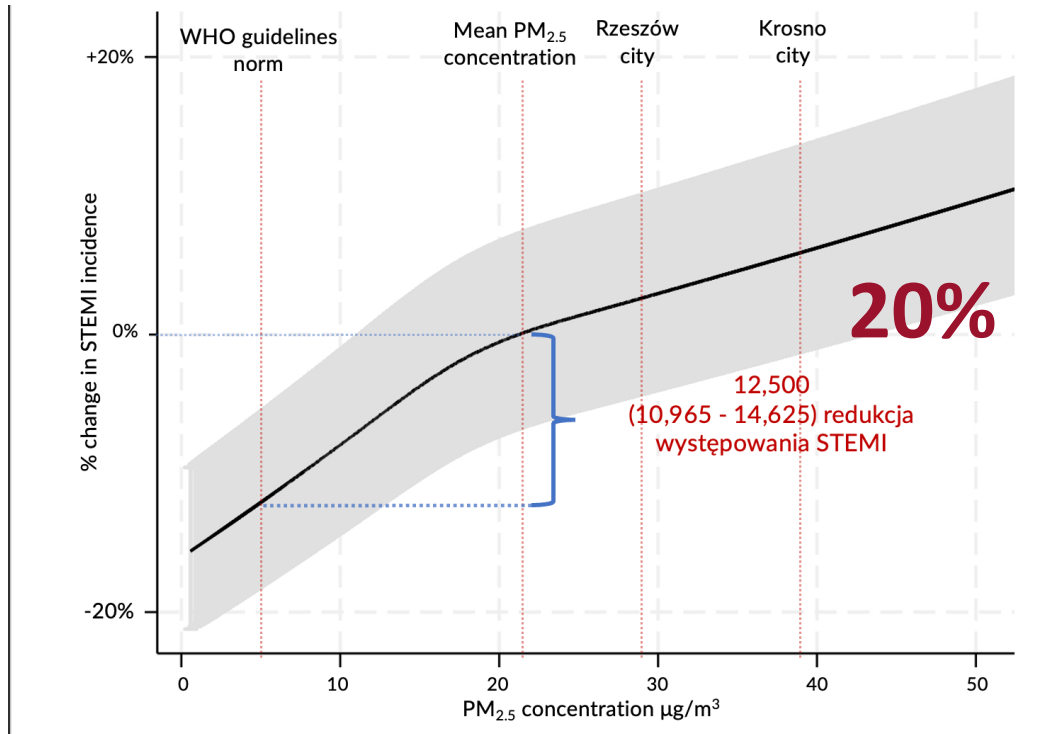
na wzrost o IQR BaP ng/m³



Kobiety/mężczyźni 65-/65+

Outcome	Kobiety	mężczyźni
STEMI lag0-30	+1,2%	+4.9%
NSTEMI lag0-30	+5.7%	+12.6%
CVD mortality lag0-30	+3.4%	+5.7%
Overall MACE	+4.8%	+8.7%
STEMI lag0-30	-0,01%	+0.2%
NSTEMI lag0-30	+1.2%	+0.5%
CVD mortality lag0-30	+1.5%	+1.7%
Overall MACE	+0,9%	+0,7%

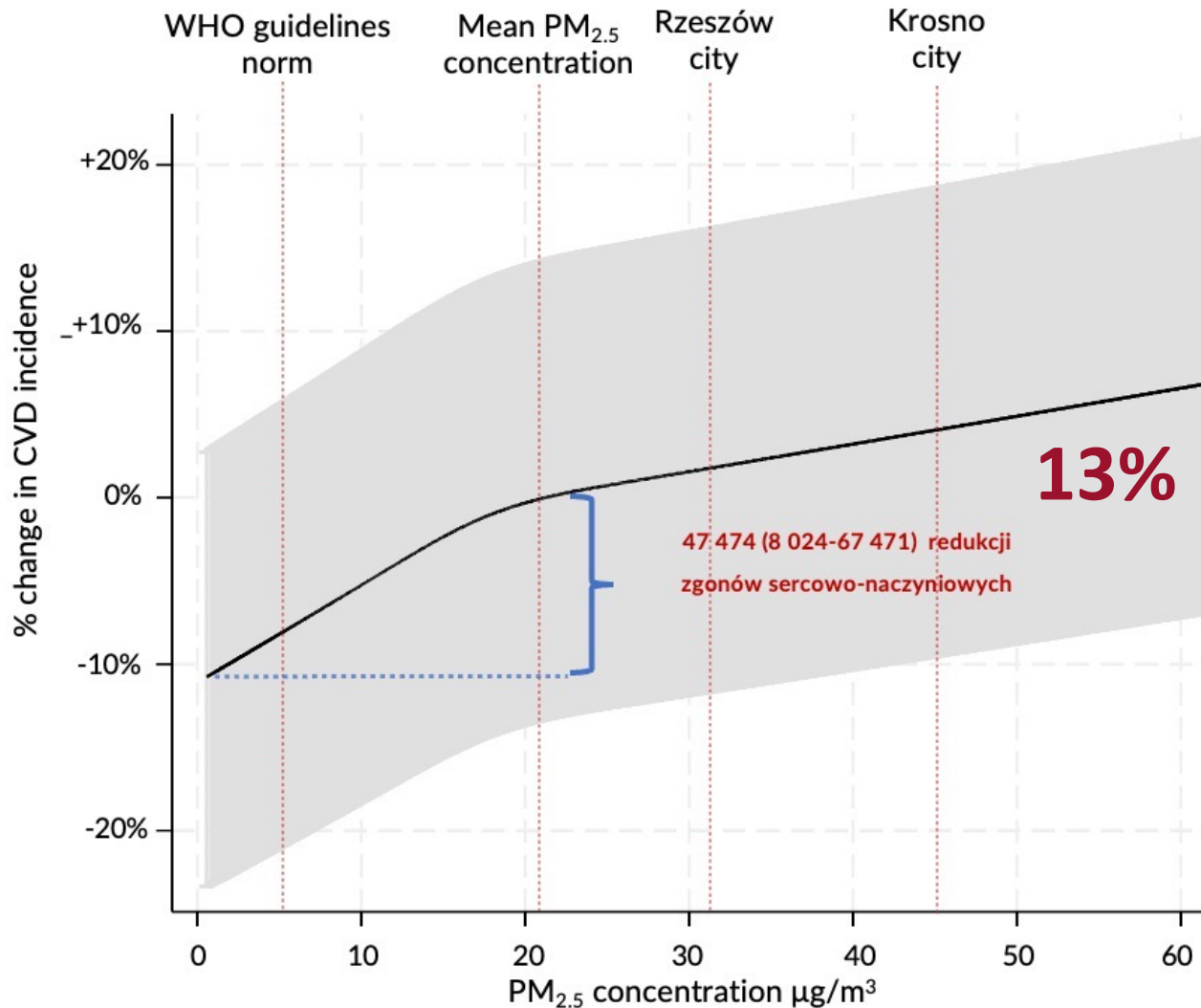
Gdyby, w analizowanym okresie, dzienne stężenia $PM_{2.5}$ nie przekraczały dopuszczalnej normy WHO:



Funkcje narażenie-odpowiedź: % wzrost przyjęć do STEMI i NSTEMNI na rosnące poziomy $PM_{2.5}$, modele specyficzne dla prowincji skorygowane o trend, temperatura powietrza: naturalne splajny dla opóźnienia 0-1 (df=6), opóźnienie 0-6 (df=3), wilgotność względna (df=3) ciśnienie atmosferyczne (df=3) i zmienne wskaźnikowe dla gminy, dnia tygodnia, świąt państwowych oraz okresów epidemii grypy.i SARS-CoV-2

Zero na osi Y reprezentuje średni efekt, a część krzywej poniżej/powyżej zera oznacza mniejsze/większe oszacowanie niż średni efekt.

Gdyby, w analizowanym okresie, codzienne stężenia $PM_{2.5}$ nie przekraczały dopuszczalnej normy WHO:



Funkcje narażenie-odpowiedź: % wzrost zgonów CVD na rosnące poziomy $PM_{2.5}$ (Lag 0), modele specyficzne dla prowincji skorygowane o trend, temperatura powietrza: naturalne splajny dla opóźnienia 0-1 (df=6), opóźnienie 0-6 (df=3), wilgotność względna (df=3) ciśnienie atmosferyczne (df=3) i zmienne wskaźnikowe dla gminy, dnia tygodnia, świąt państwowych oraz okresów epidemii grypy i SARS-Ciov-2

Zero na osi y reprezentuje średni efekt, a część krzywej poniżej/powyżej zera oznacza mniejsze/większe oszacowanie niż średni efekt.

YLL= 893 lat

(395 lat – 1,086 lat)/100,000 mieszkańców

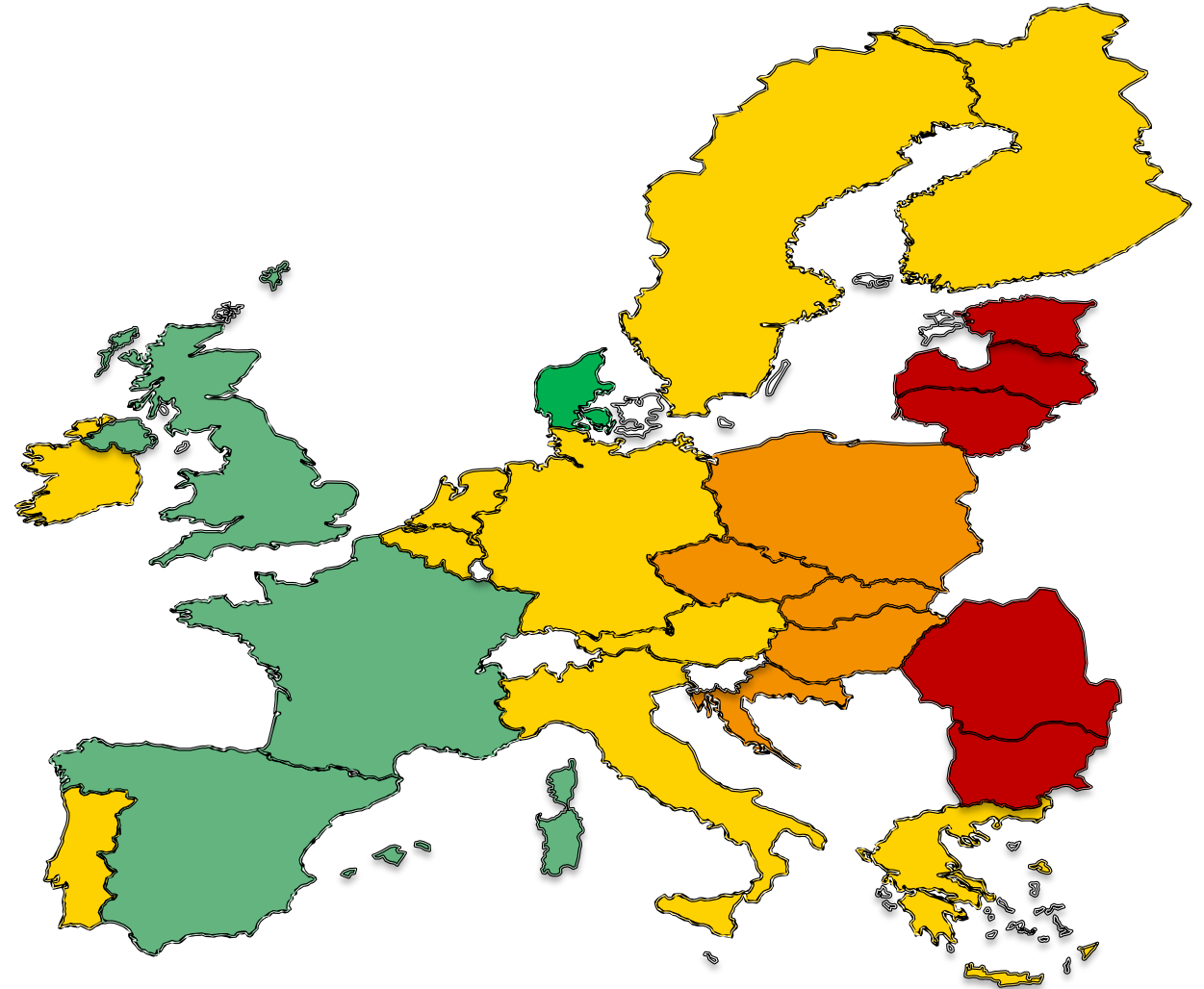


71,440 lat

w całej analizowanej populacji

Why do we need a new indicator?

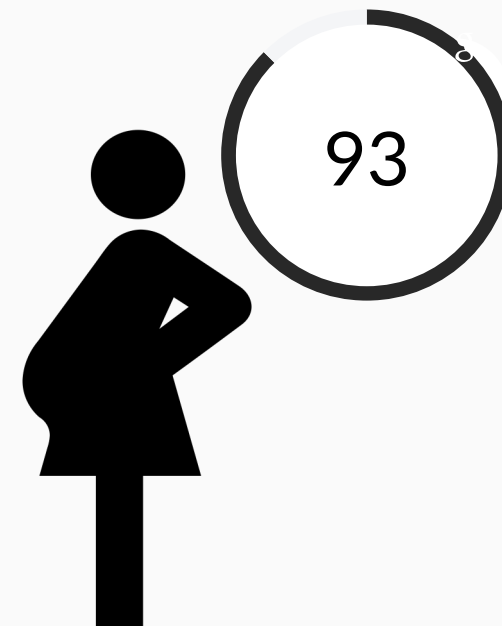
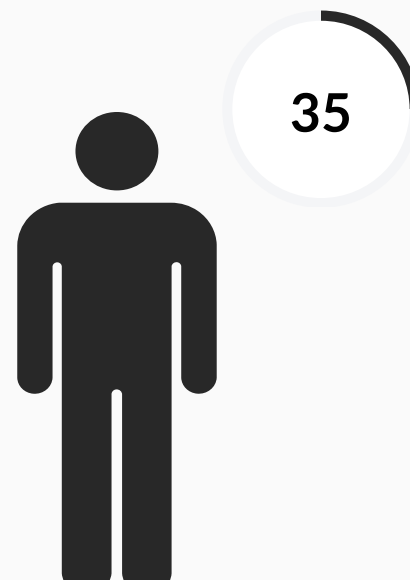
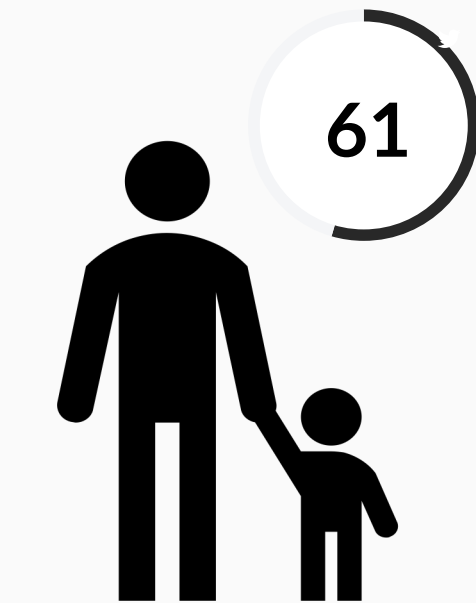
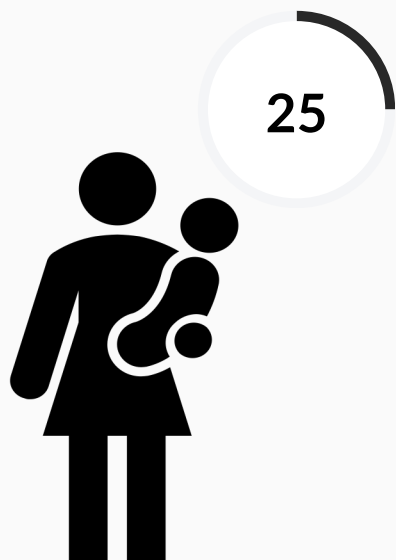
- Clinical risk stratification focuses on patient-centric risk factors, without consideration of environmental pollution
- 14% people with fatal ACS and 50% patients with ACS have none or only one classical risk factor
- Risk scores are not validated for air pollution so they may under- or overestimate the true risk
- Environmental cardiology offers opportunities for primary prevention and can help address health disparities



ePM-years Index

$$\Sigma = \max \left[\frac{c^1 + c^2 + c^3 \dots + c^N}{C_w} - t; 0 \right]$$

C^1 - mean yearly concentration in the first year of observation
 C^N - mean yearly concentration in the last year of observation
 C_w - World Health Organization annual norm form $PM_{2.5}$
 t - time of observation (years)



ESC Congress 2023
Amsterdam, October 11-15

32 y.o Białystok city

52 y.o Katowice city

51 y.o Białystok city

31 y.o Kraków city

ePM-years Index

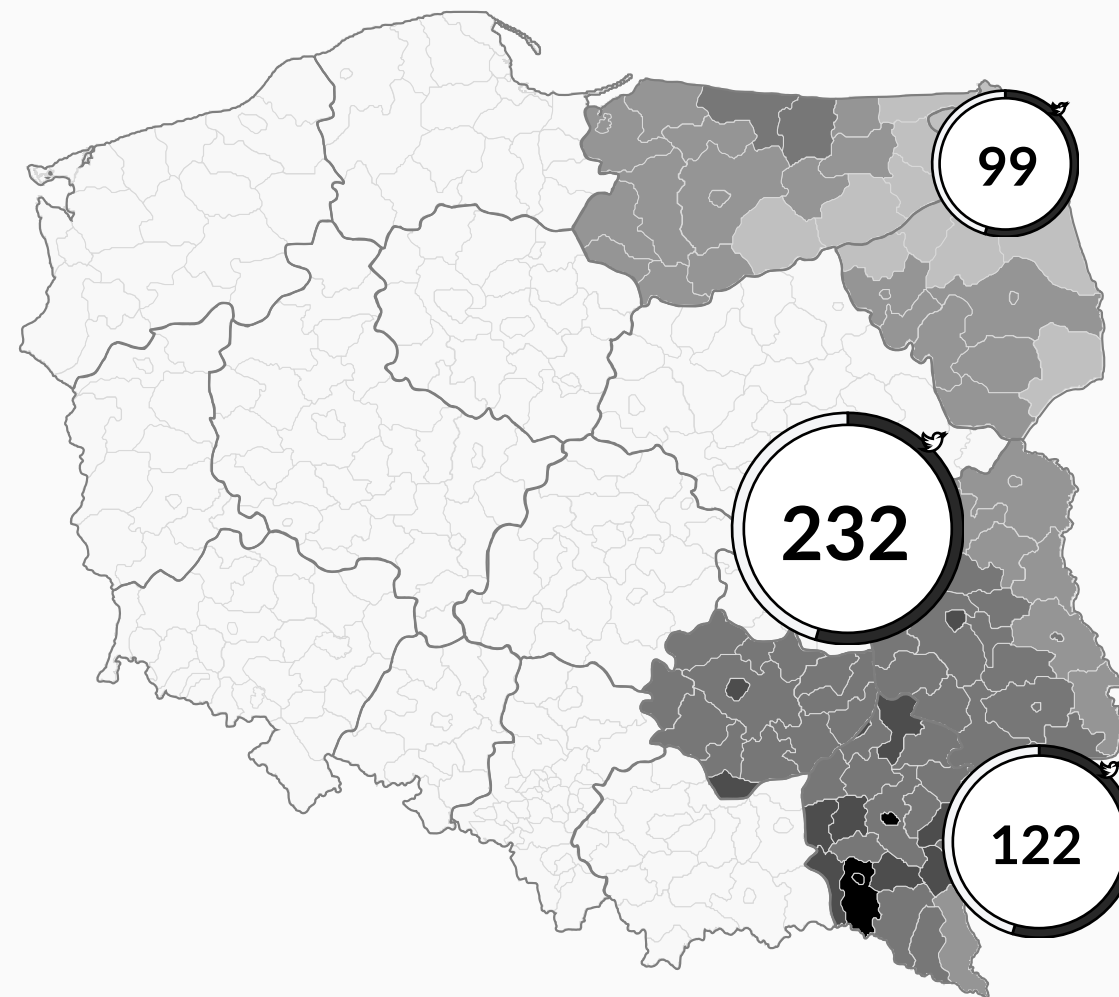
$$\Sigma = \max \left[\frac{c^1 + c^2 + c^3 \dots + c^N}{C_w} - t; 0 \right]$$

C^1 - mean yearly concentration in the first year of observation

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t - time of observation (years)



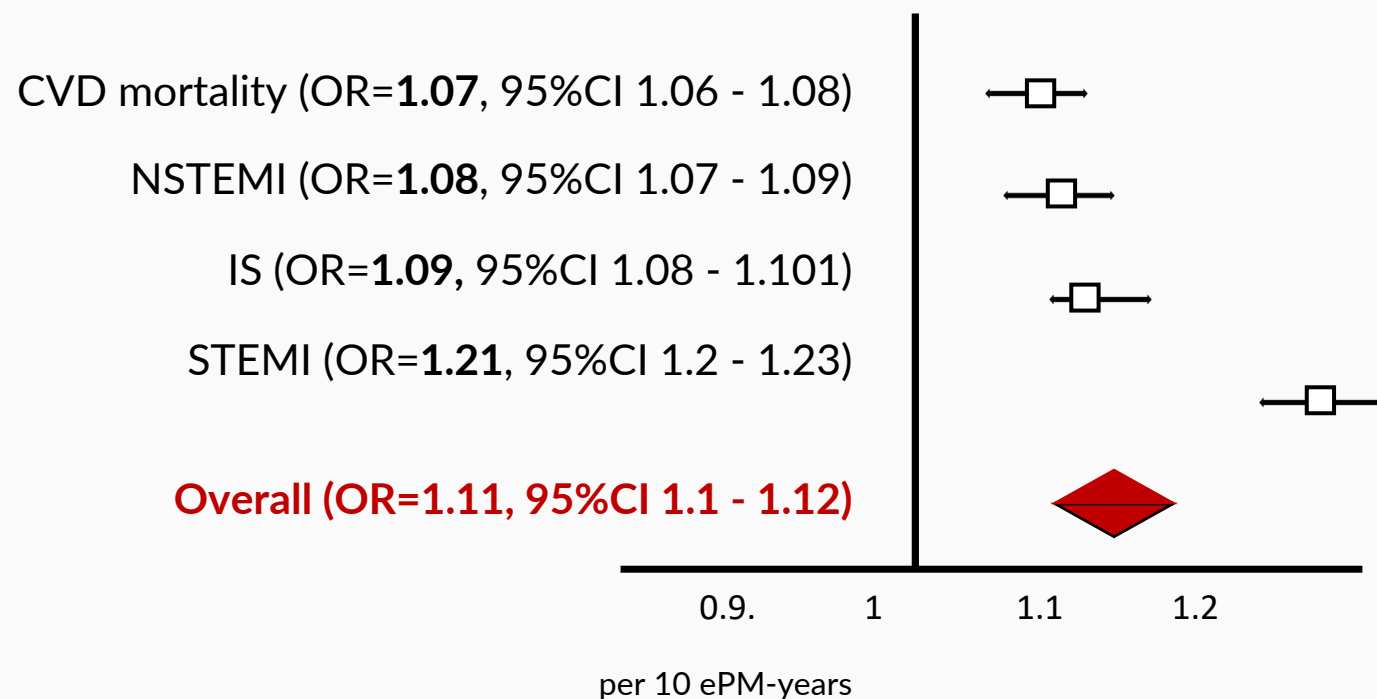
What it delivers:

- CVD risk reclassification
- identifies the cohorts at high risk
- more personal information and role in decision-making for those at „intermediate” risk
- allows for modification of intensity of monitoring or early preventive strategies
- support decisions making regarding the location of the public health as well as personal investments
- Informs insurance companies about patients risks

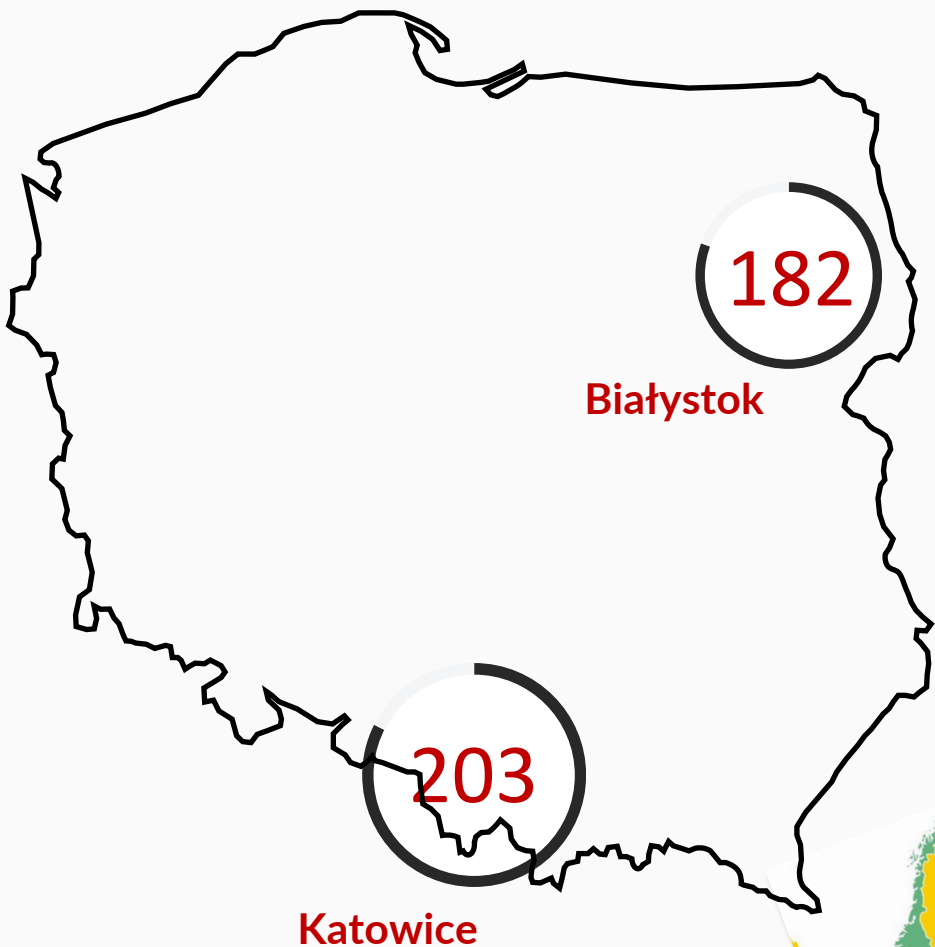


ePM-years Index

- ✓ is an independent predictor of major cardiovascular events
- ✓ may be used as a simple index for risk stratification
- ✓ should be incorporated into prediction models



EP-PARTICLES COHORT



41 y.o. healthy man, non-smoker

	Białystok	Katowice
SCORE2	<1%	<1%
ePM-years Index	159	173

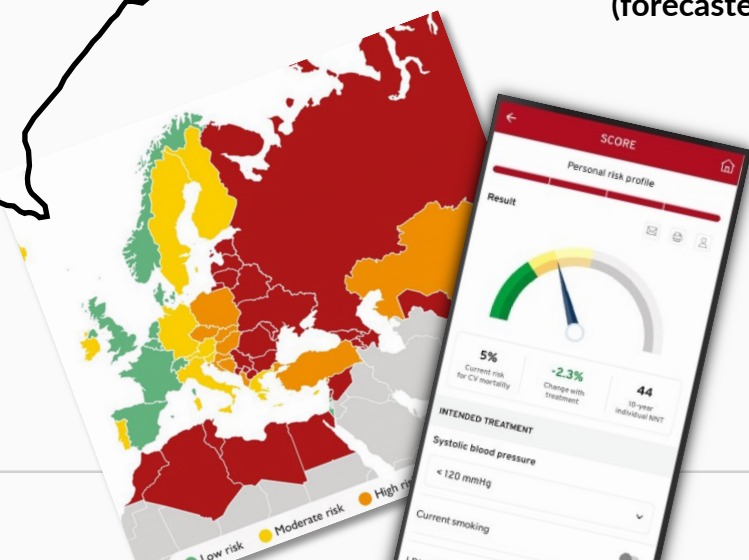
51 y.o. healthy man, non-smoker

	Białystok	Katowice
SCORE2	<1.5%	<1.5%
ePM-years Index (forecasted)	182	203

2020



2030

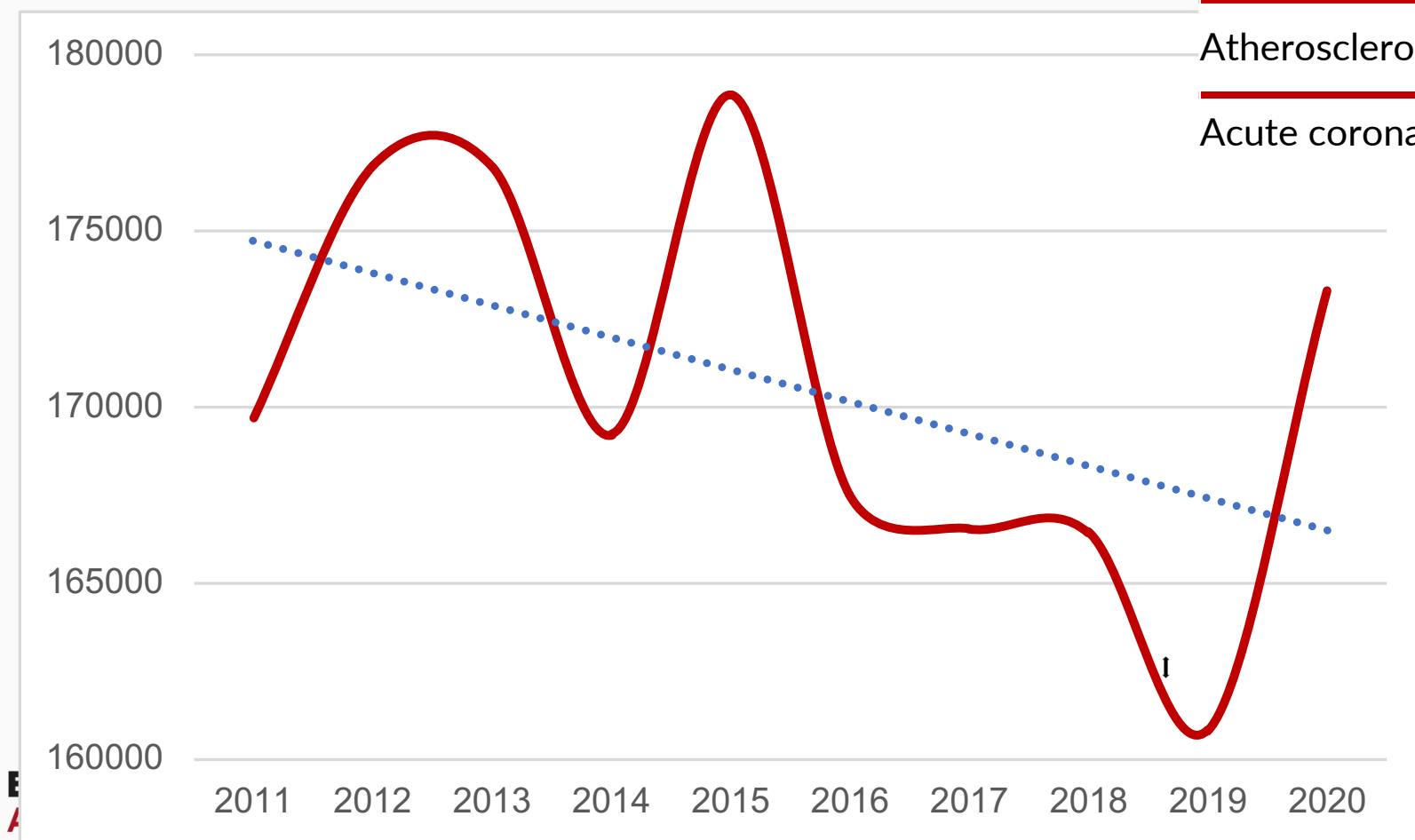


EUROsmog COHORT

CVD-mortality (N= 1,706,111)

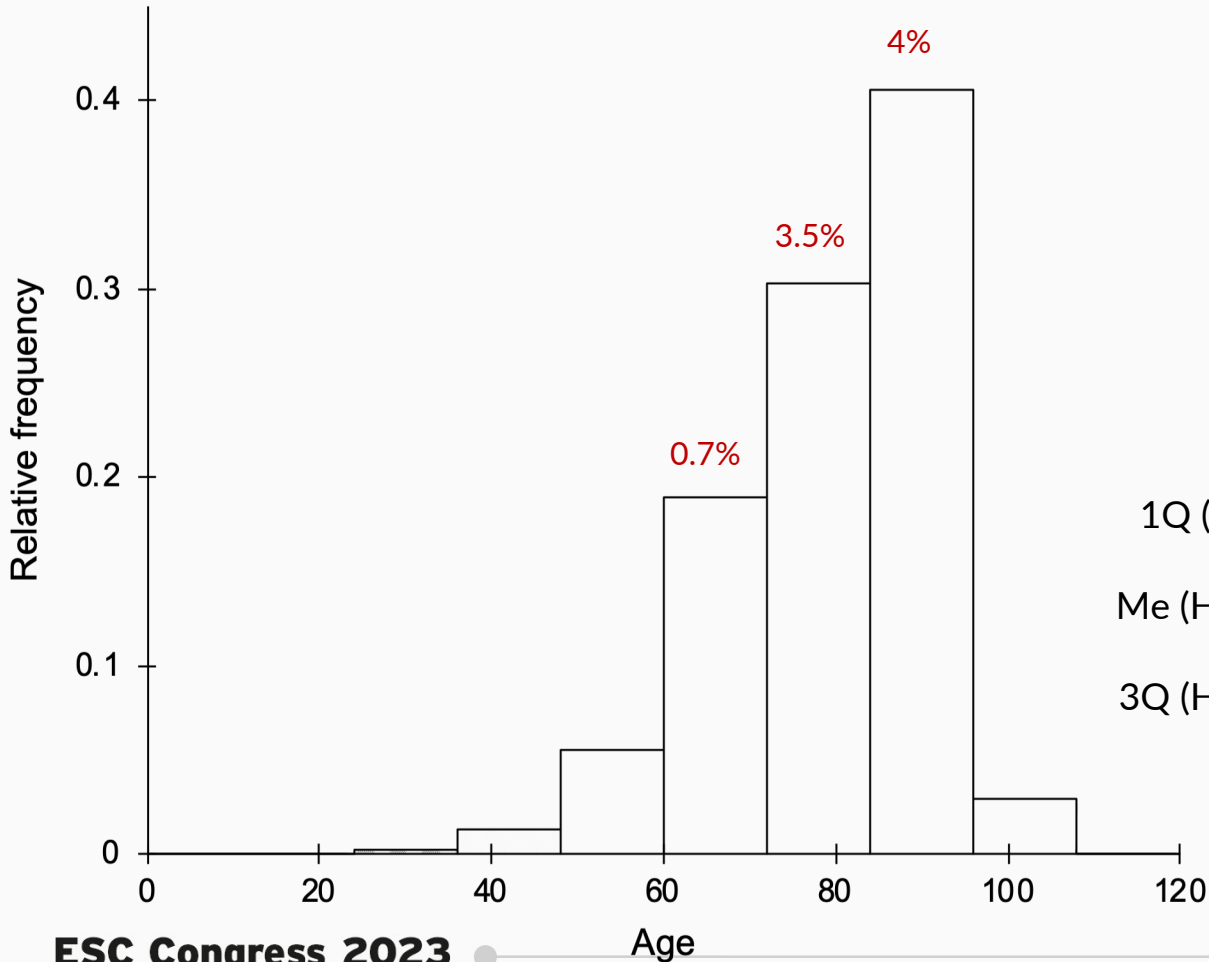
Me 81 [71-87], ♂ 47%

Heart failure	22.56%
Ischemic stroke	7.57%
Atherosclerosis	18.84%
Acute coronary syndromes	24.27%



CVD-mortality (N= 1,706,111). Me 81 [71-87], ♂ 47%

What we are doing at the moment is NOT effective

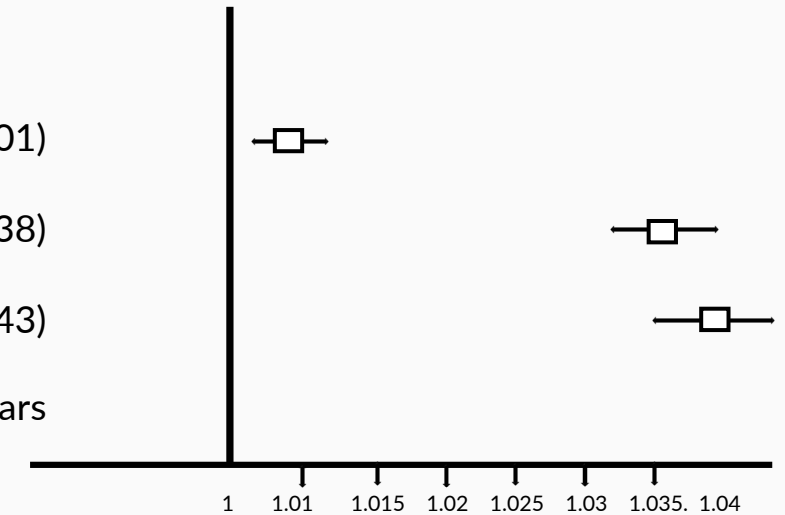


1Q (HR=1.007, 95%CI 1.003 - 1.01)

Me (HR=1.035, 95%CI 1.033 - 1.038)

3Q (HR=1.041, 95%CI 1.039 - 1.043)

per 10 ePM-years





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