

# **Contrasting patterns of CVD, cancers and related mortality between high-, middle-, and low-income countries**

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## **Impact of modifiable risk factors on cardiovascular disease and mortality**

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On behalf of the Prospective Urban Rural Epidemiology (PURE) Study Investigators

# Declaration of interest

- I have nothing to declare



# Declaration of Interests

PURE was supported by the Canadian Institutes of Health Research, Heart and Stroke Foundation of Canada, Ontario Ministry of Health and Long-Term Care, Astra Zeneca, Boehringer Ingelheim, Sanofi-Aventis, Servier Laboratories and Glaxo Smith Kline; plus additional sources in participating countries.

# Rationale

- Shift from communicable to non-communicable diseases and deaths globally
- Advances in prevention and treatment have reduced CV deaths in many countries
- These advances could lead to a further epidemiologic transitions in diseases and deaths from different NCDs
- Burden of some diseases & impact of risk factors may differ between countries at different economic levels

# Aims

- To describe incidence of various diseases and related deaths in countries at different economic levels (HIC, MIC, LIC)
- To document relative importance of risk factors for CVD and mortality overall and in HIC, MIC and LIC

# PURE Study

- **Total of 202,000 from 27 countries: Ongoing follow-up**
- 21 countries (N=167,000) with follow-up data
- 6 additional countries with 35,000 participants – awaiting follow-up



# Methods

- 166,762 community-dwelling adults: 35-70 years
- 21 countries (4 HIC, 12 MIC, 5 LIC)
- Follow-up: median 9.5 years
- Events documented through direct contact + administrative records
  - Vital status known in 98%
  - CVD and other events documented in 94%
- Standardized outcome definitions & event adjudication

# Baseline Characteristics

Characteristic	HIC	MIC	LIC
N	18,073	108,291	36,170
Mean age $\pm$ SD	52 $\pm$ 9.5	51 $\pm$ 9.8	49 $\pm$ 10.4
Male, %	47	41	43
INTERHEART CVD risk score*	13.1 $\pm$ 6.2	10.5 $\pm$ 5.8	7.9 $\pm$ 5.0

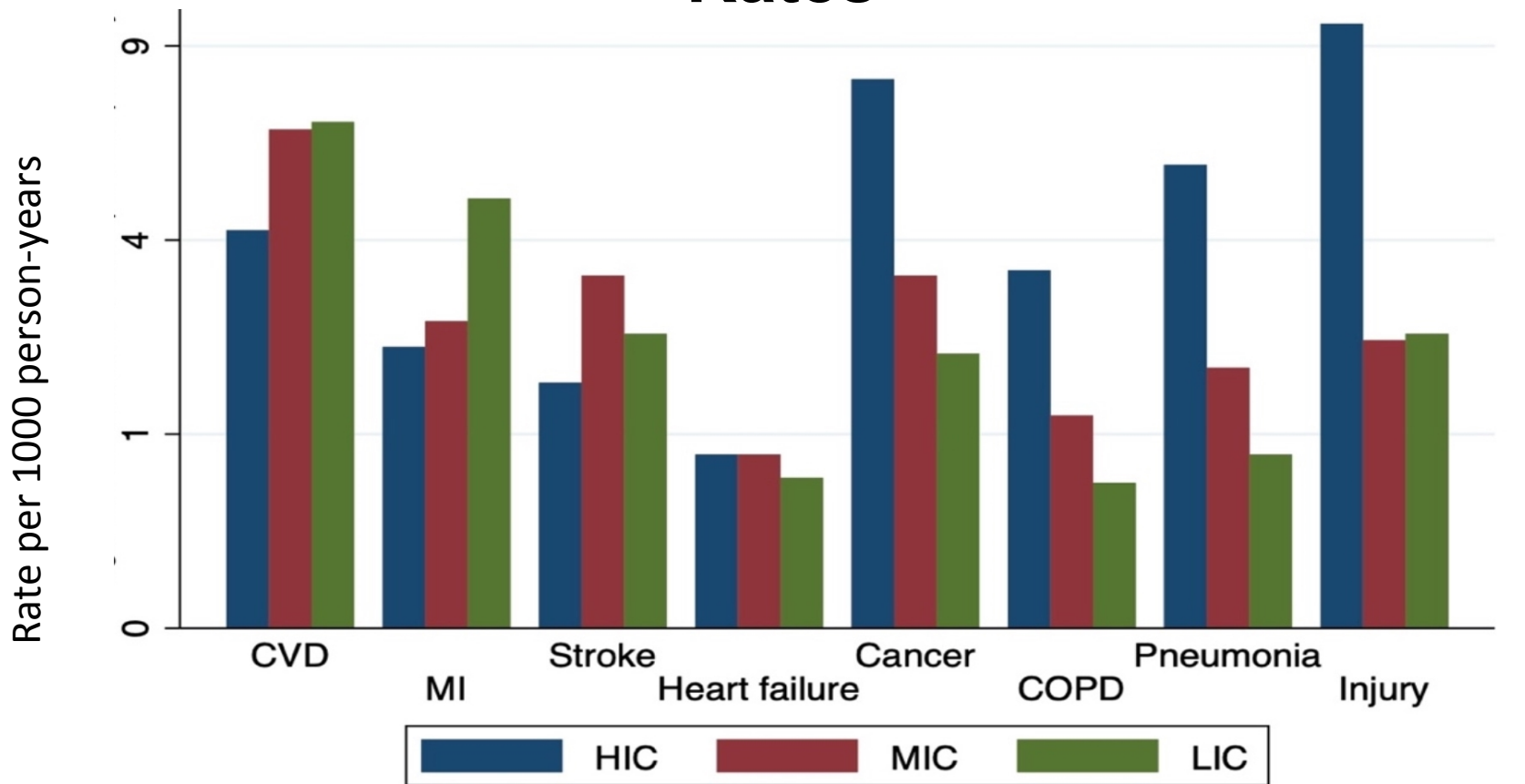
\* Age, smoking, 2<sup>nd</sup> hand smoke, diabetes, hypertension, family history, waist-hip ratio, psychosocial factors, diet, physical activity



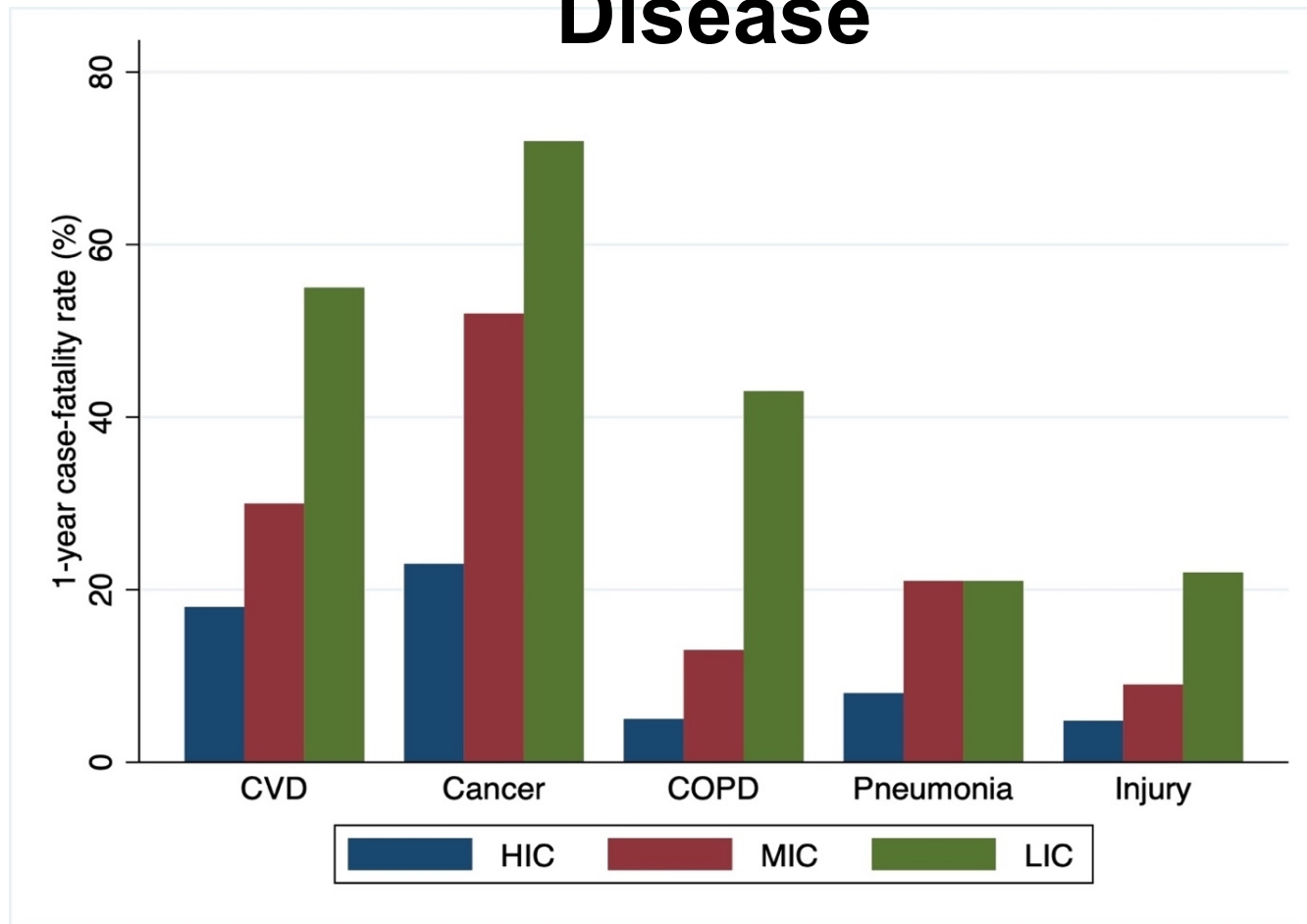
# Outcomes

Incident Event	Number
Death	11,307
CVD (MI, stroke, HF, CVD death)	9,329
Cancer	5,151
COPD	1,830
Pneumonia	2,911
Injuries with hospital admission	4,386

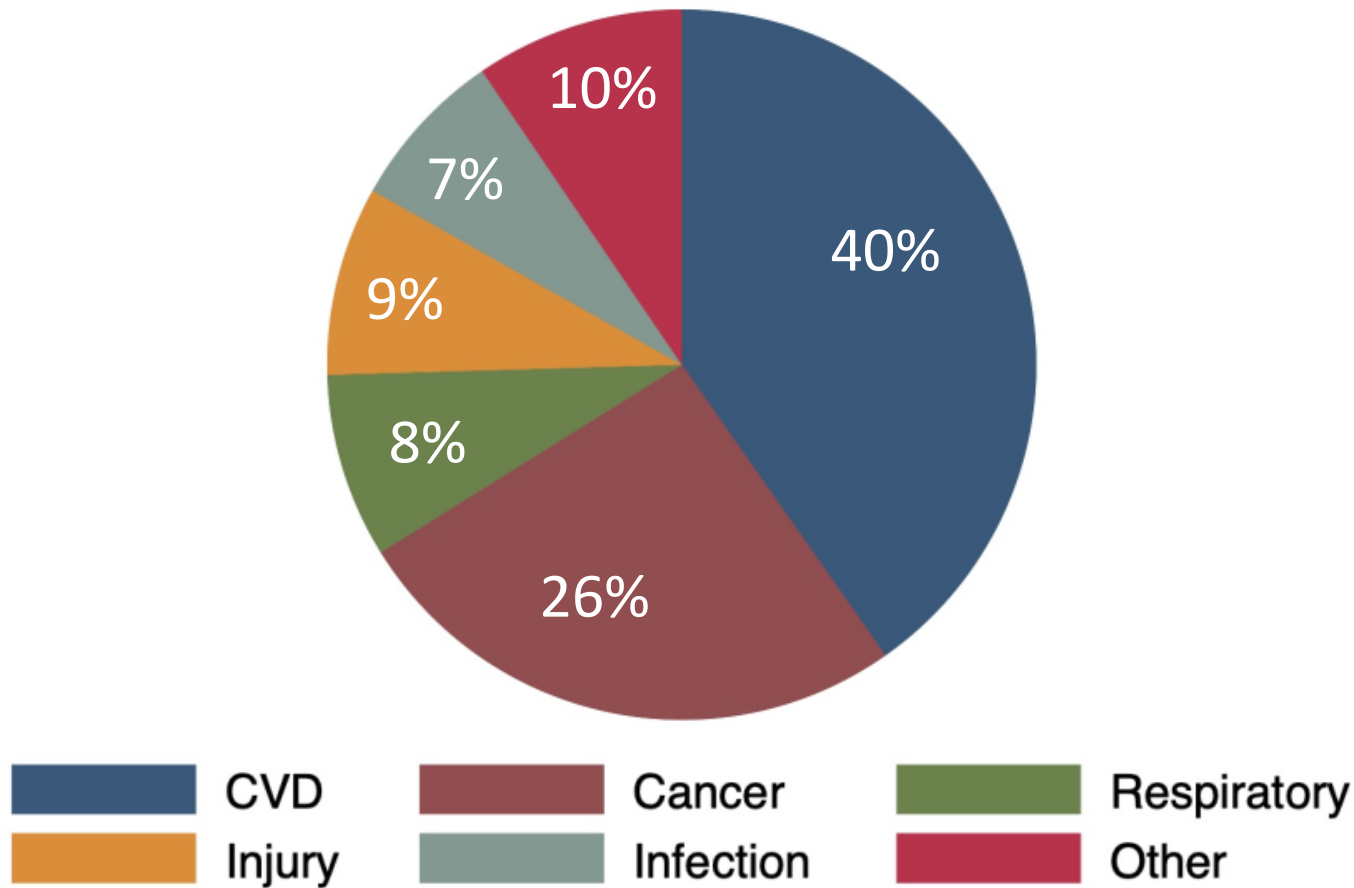
# Age- & Sex-Standardized Disease Incidence Rates



# One-Year Case-Fatality after Incident Disease

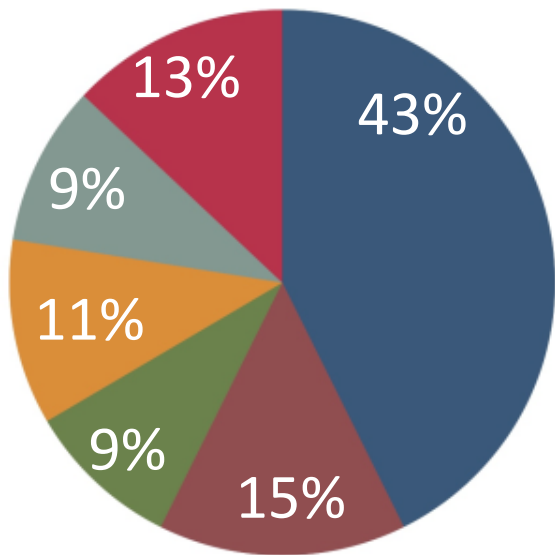


# Causes of Death: Overall

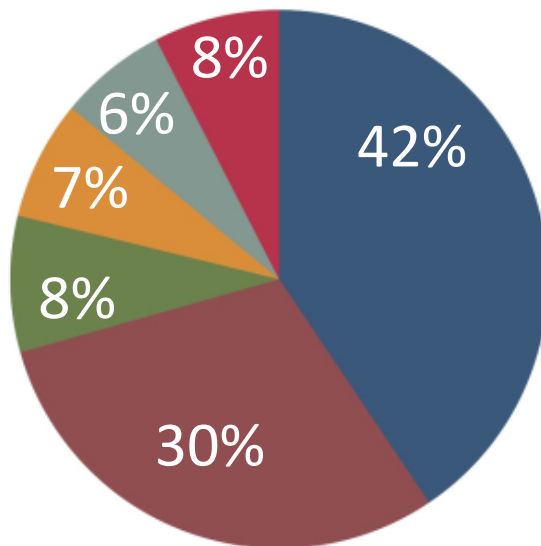


# Causes of Death by Country Income Level

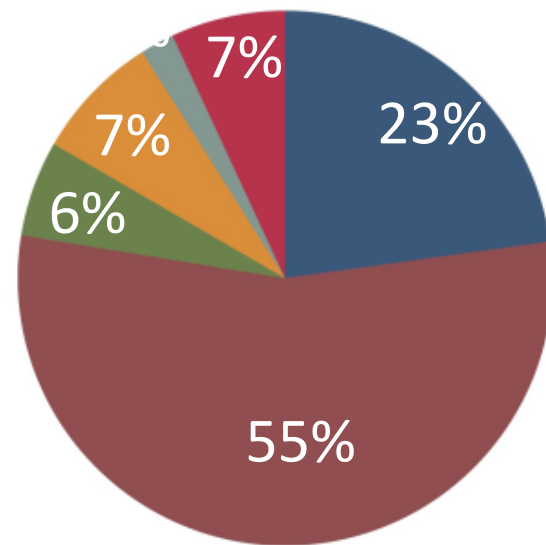
LIC



MIC



HIC



CVD



Cancer



Respiratory



Injury



Infection



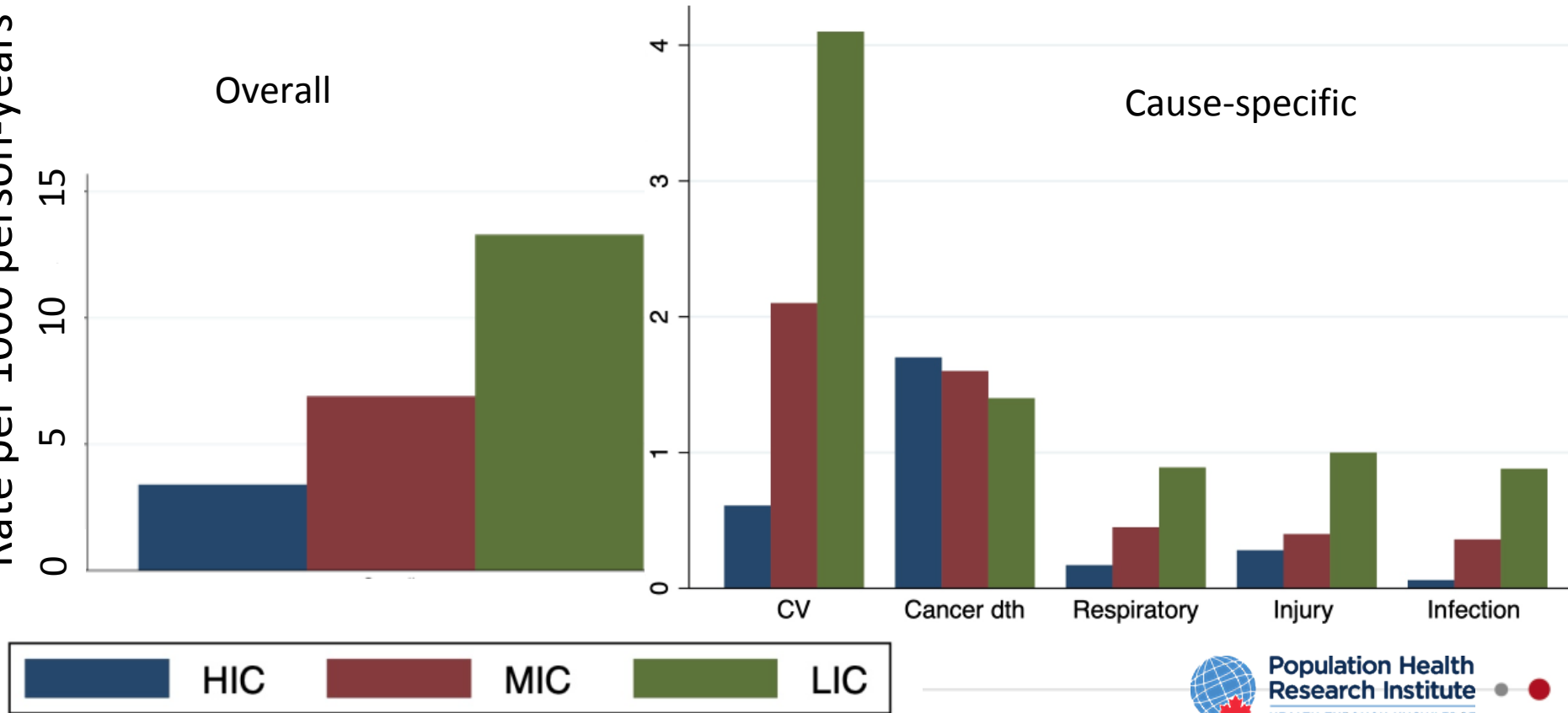
Other

# Age- & Sex-Standardized Mortality Rates

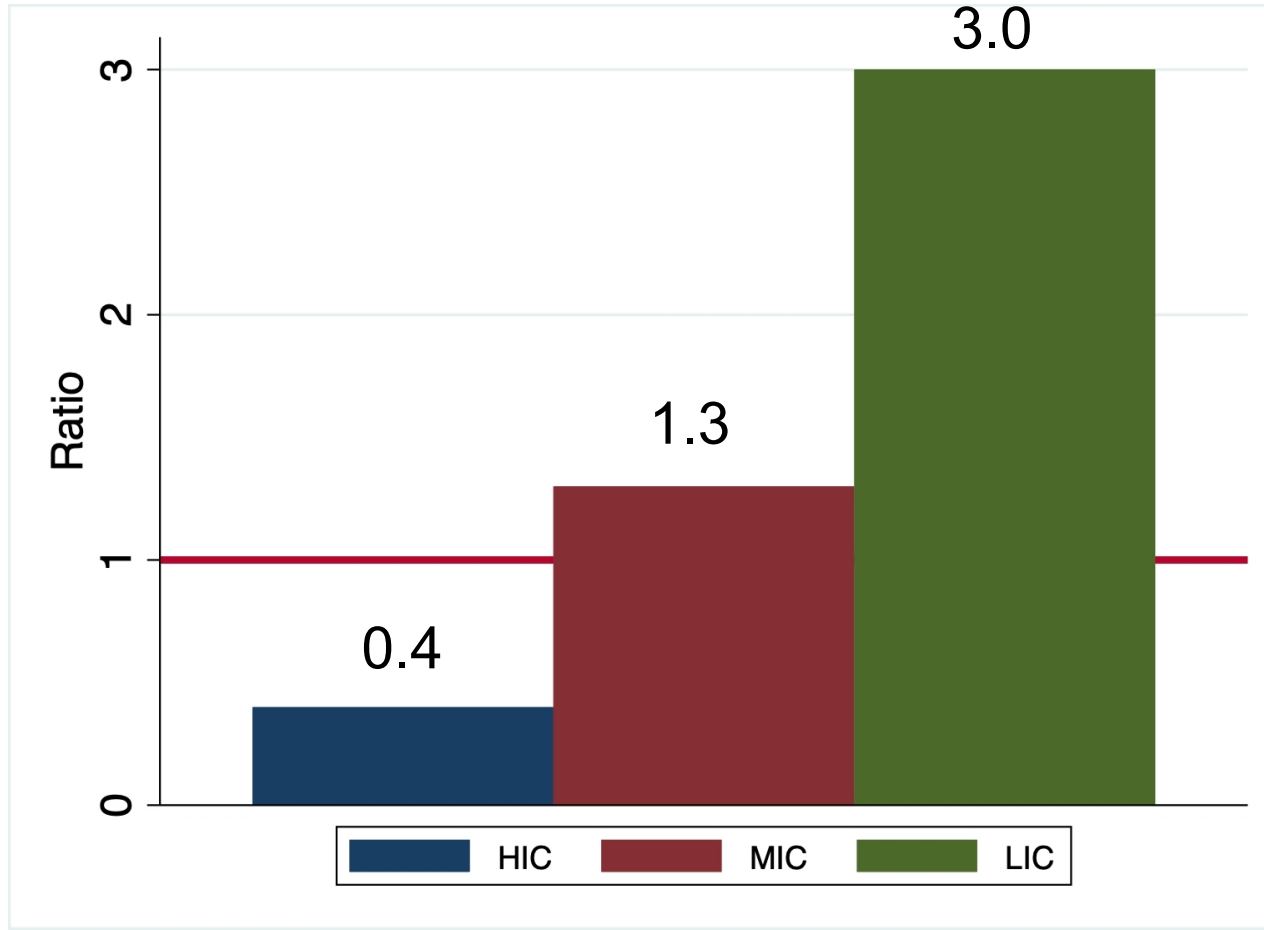
Rate per 1000 person-years

Overall

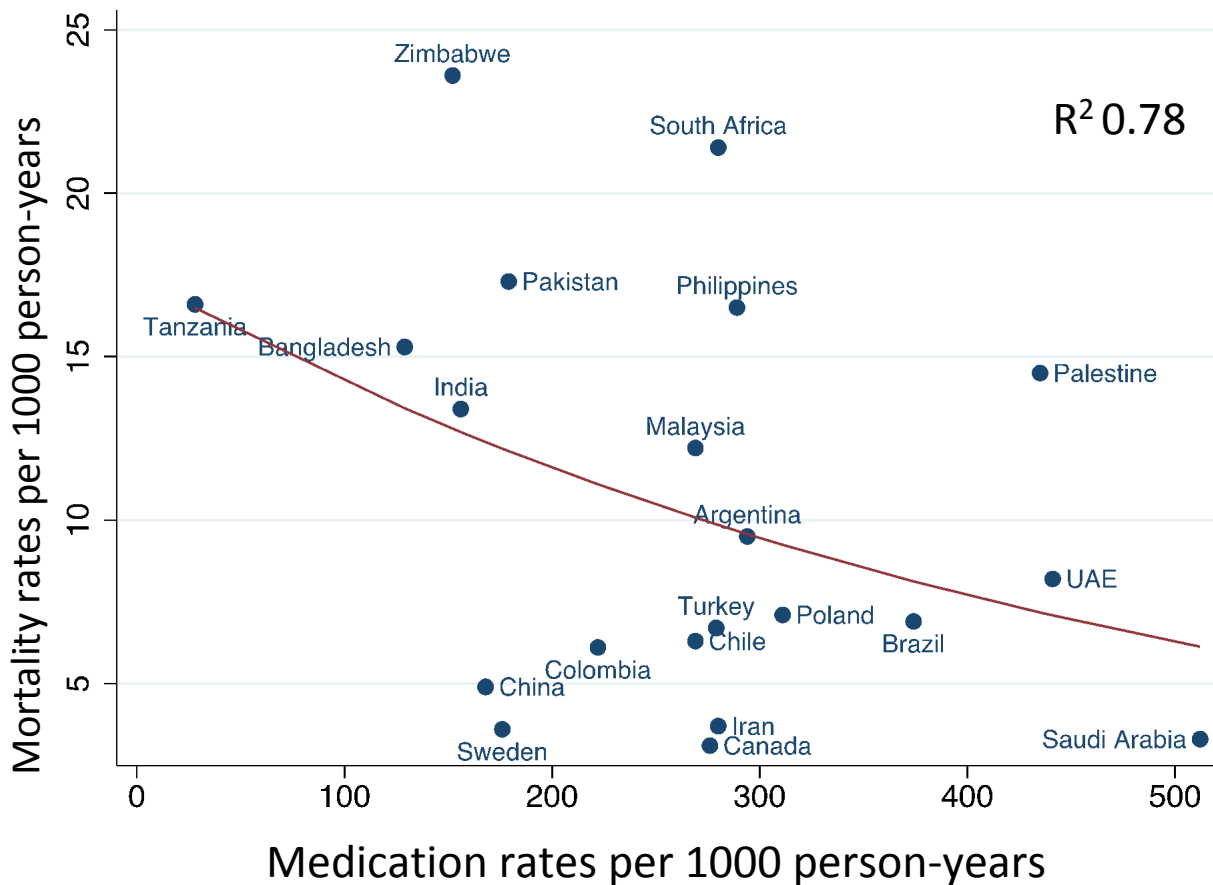
Cause-specific



# Ratio of CV deaths to Cancer deaths



# Age & Sex-Std CV Medication vs Mortality Rates





# Impact of modifiable risk factors on CVD and mortality

## Objectives:

- Quantify risk for CVD and death associated with 14 common, modifiable risk factors
- Document variations in the importance of risk factors by HIC, MIC, LIC

# Methods

**Population:** 152,722 individuals without CVD

## **Risk Factors:**

- ***Behavioral:*** tobacco, diet, alcohol, physical activity, and sodium
- ***Metabolic:*** HTN, DM, obesity, and lipids
- ***Psychosocial:*** Education, depression
- **Grip Strength**
- ***Air pollution:*** household (solid fuel use), and PM 2.5 ambient air pollution

**Analysis:** Mutually adjusted risk (hazard ratios) and population level burden (population attributable fractions [PAF]) associated with each risk factor

\*analyses performed separately

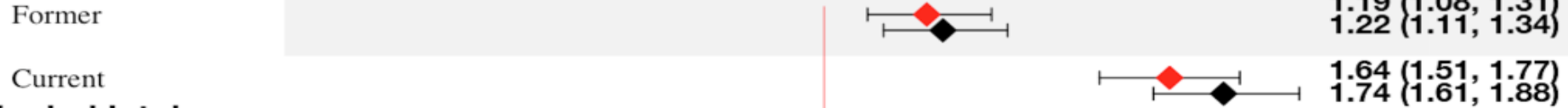


# Behavioral Risk Factors: CVD♦ & Death♦

HR

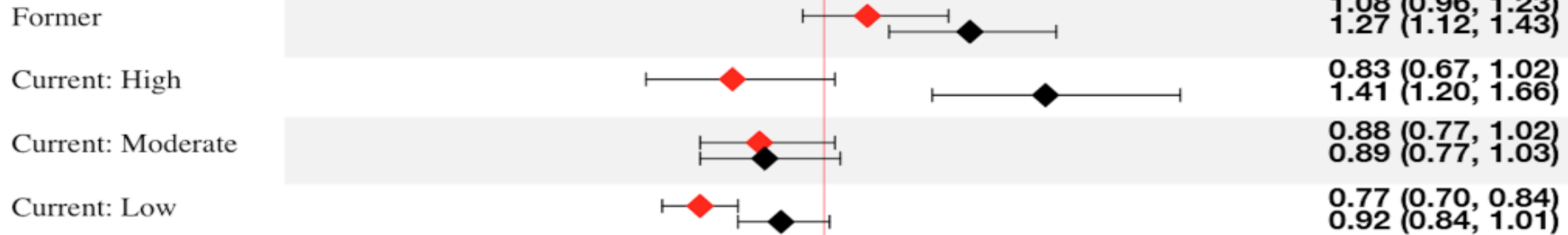
## Tobacco Use

Ref: Never



## Alcohol Intake

Ref: Never



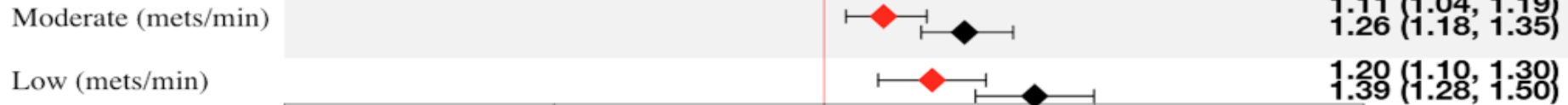
## Diet Quality

Ref:  $\geq 5$



## Physical Activity

Ref: Highly Active



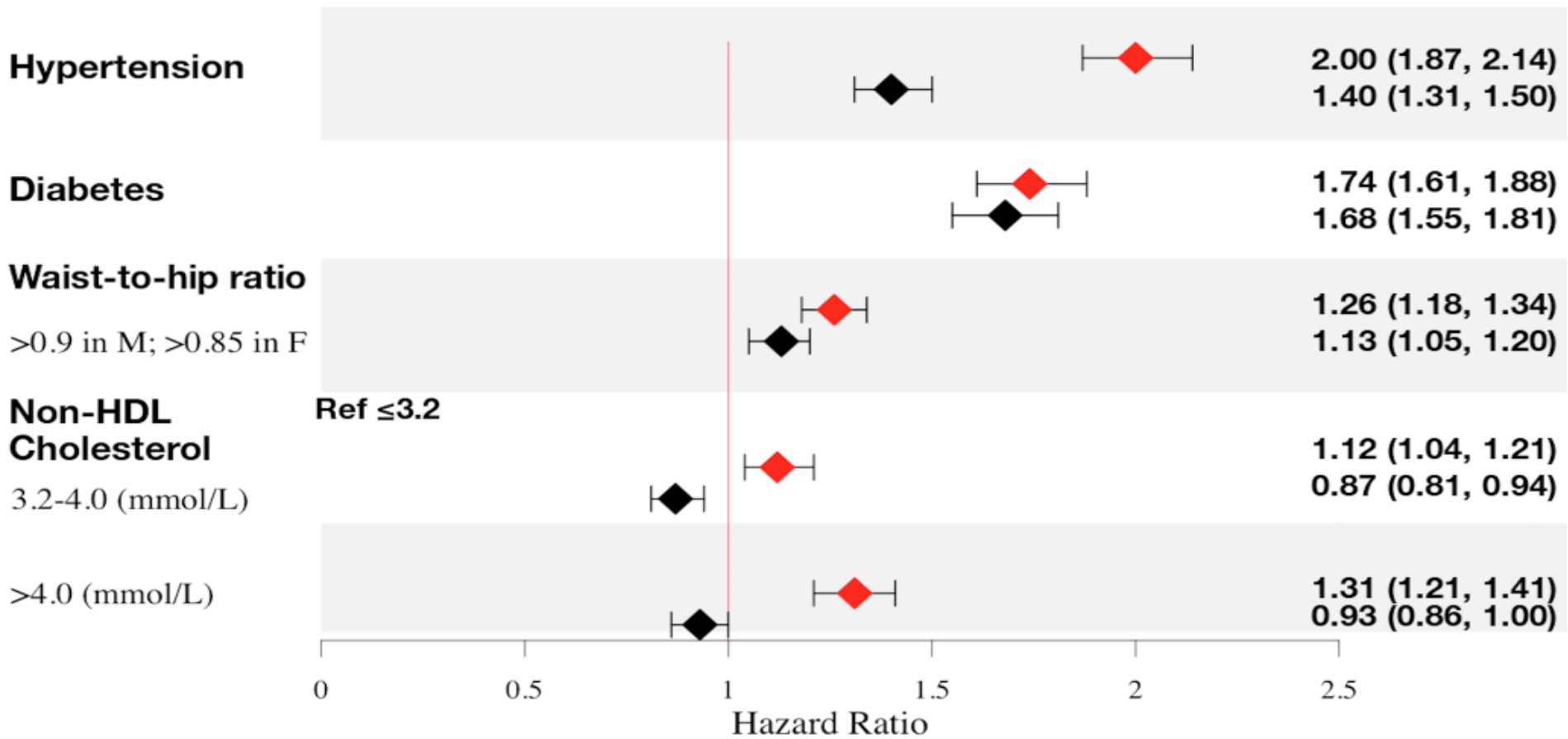
0 0.5 1 1.5 2  
Hazard Ratio



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# Metabolic Risk Factors: CVD ♦ & Death ♦

HR





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# Other Risk Factors: CVD ◆ & Death ◆

HR

## Education

Ref: Trade/college/uni

Secondary

Primary

## Depression

## Household Air Pollution

## Grip Strength

Ref: Q5

Q4

Q3

Q2

Q1

0

0.5

1

1.5

2

Hazard Ratio

1.11 (1.01, 1.22)

1.15 (1.03, 1.29)

1.37 (1.23, 1.52)

1.55 (1.39, 1.74)

1.17 (1.05, 1.29)

1.31 (1.19, 1.43)

1.09 (1.00, 1.19)

1.24 (1.14, 1.36)

1.12 (1.01, 1.24)

1.09 (0.97, 1.23)

1.18 (1.07, 1.31)

1.16 (1.04, 1.30)

1.21 (1.09, 1.35)

1.25 (1.11, 1.40)

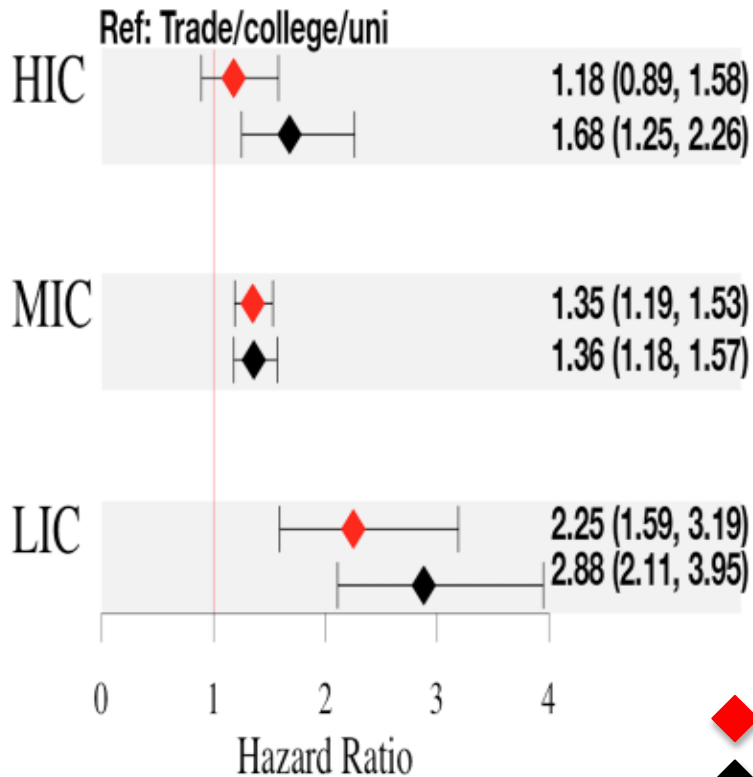
1.36 (1.21, 1.52)

1.60 (1.42, 1.79)

# Significant Variations by HIC, MIC & LIC

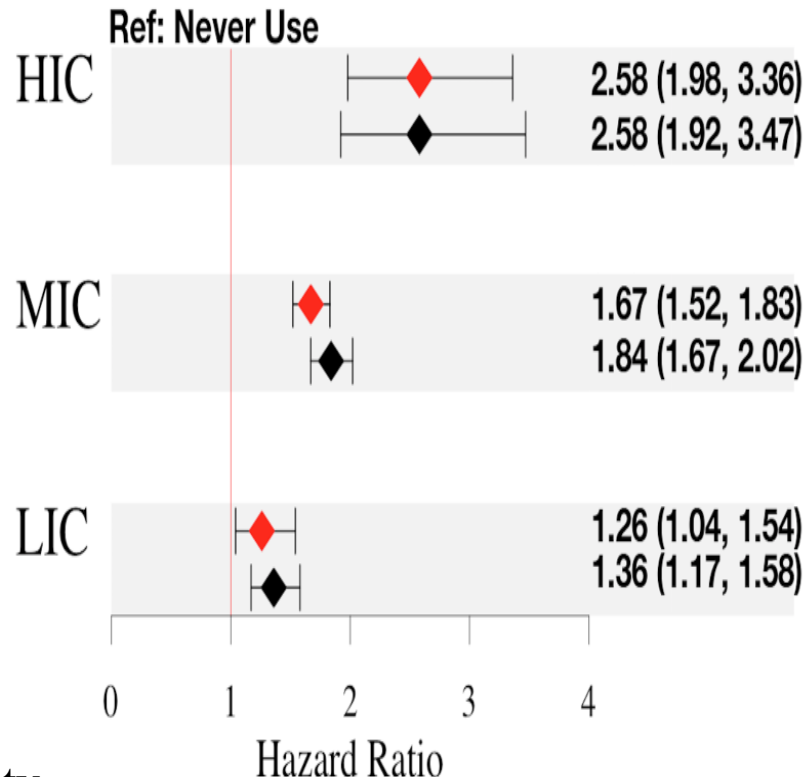
## Primary Education

$P_{\text{interaction}} = 0.005$



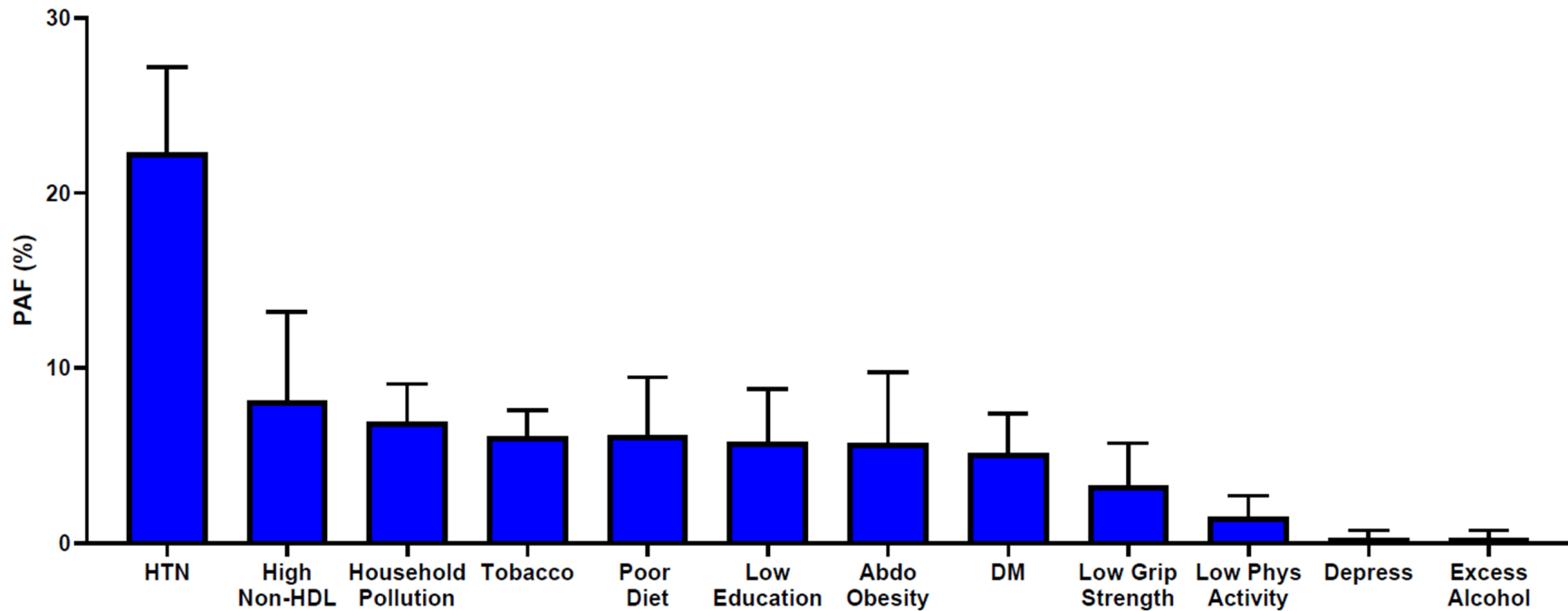
## Tobacco Use

$P_{\text{interaction}} = 0.0169$

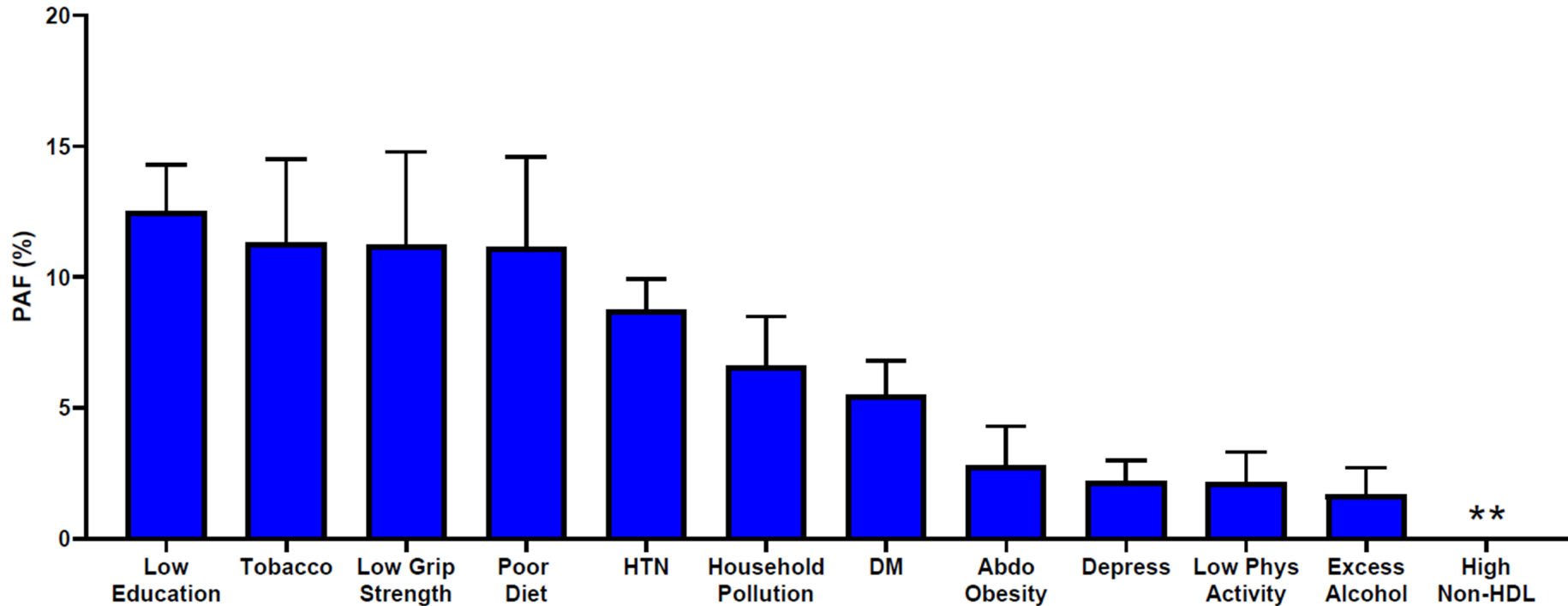


 CVD  
 Mortality

# Population Attributable Fraction: CVD

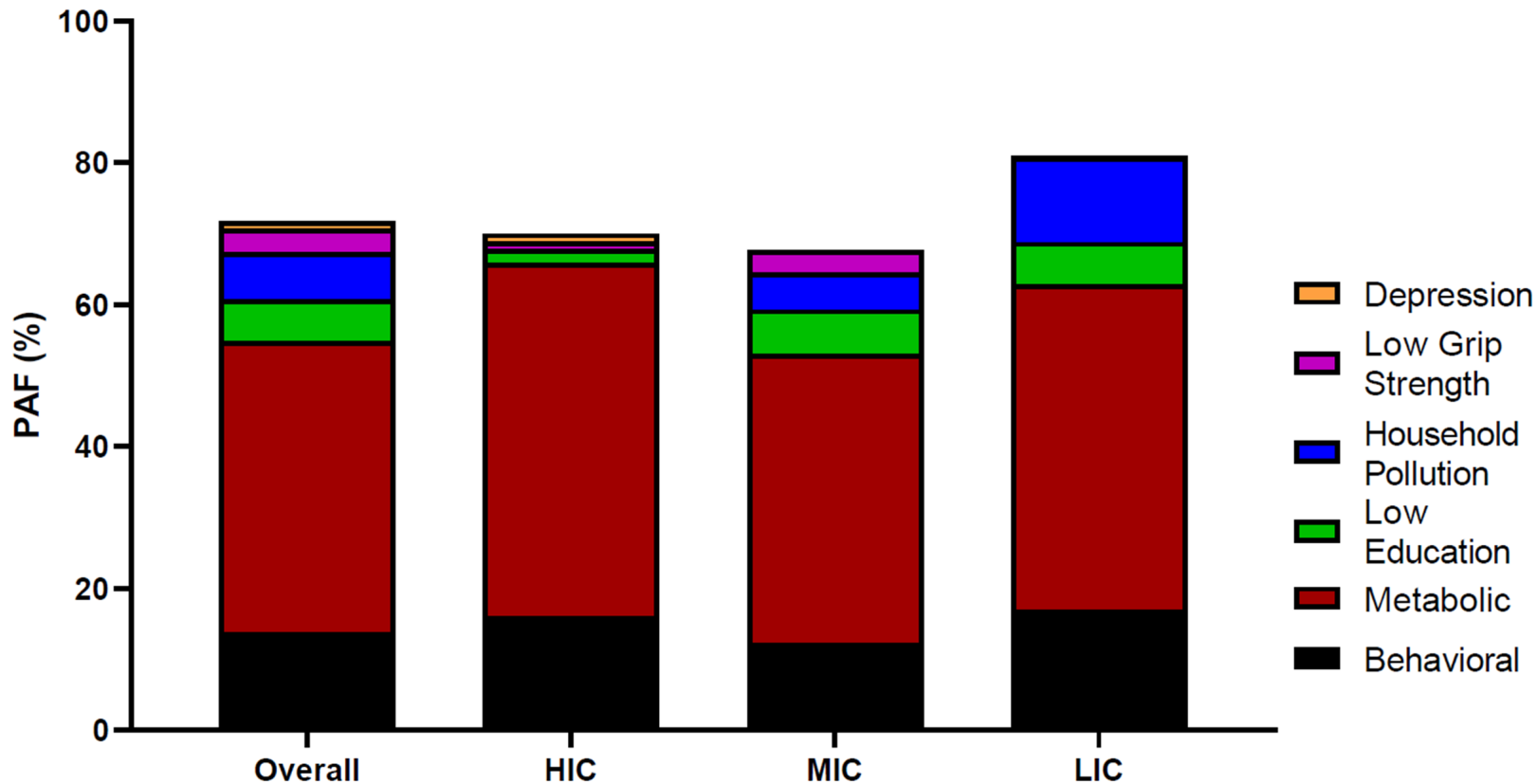


# Population Attributable Fraction: Mortality

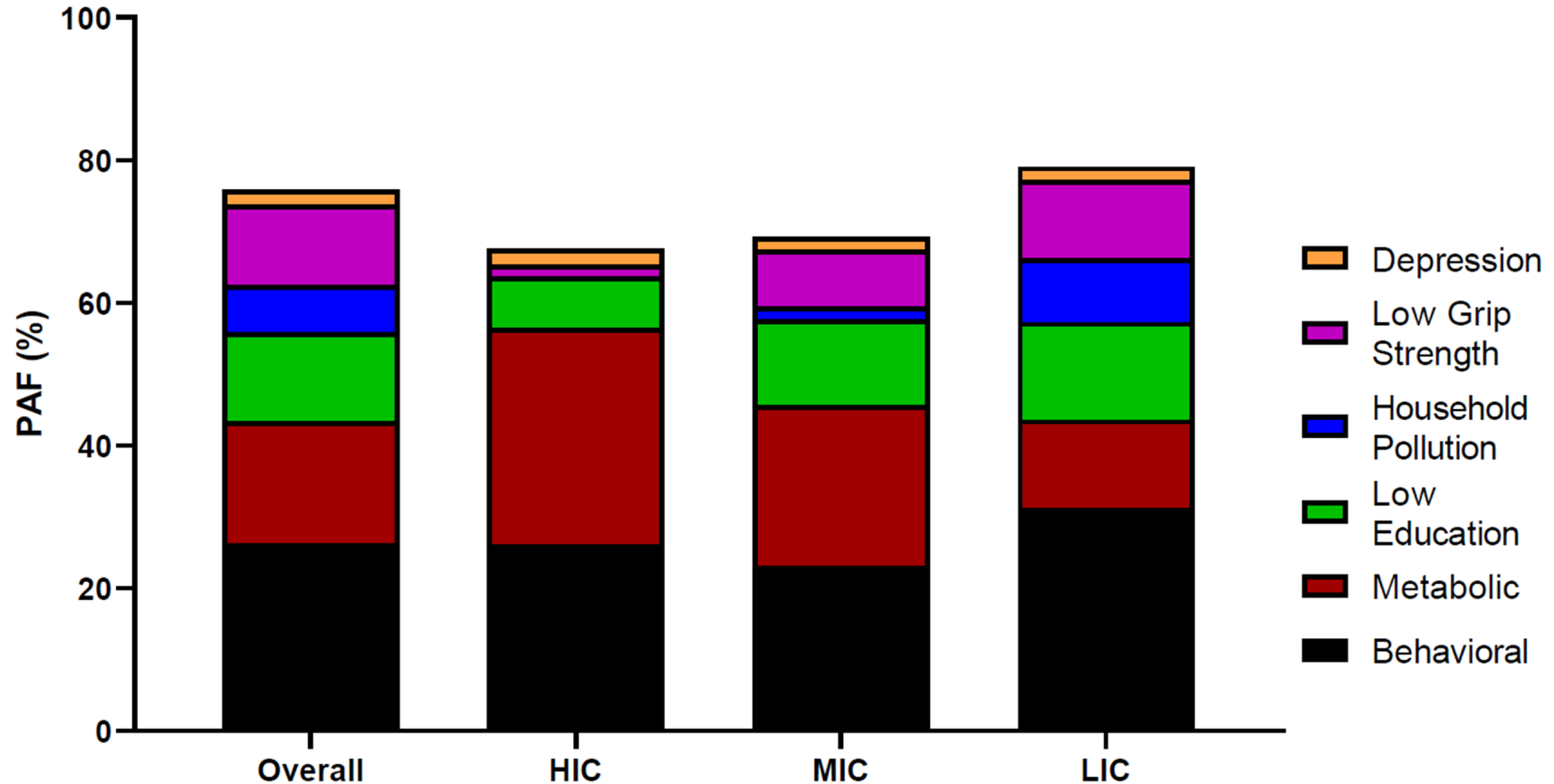




# Population Attributable Fraction: CVD



# Population Attributable Fraction: Mortality



# Ambient Air Pollution: CVD & Death per 10 $\mu\text{g}/\text{m}^3$

	HR (per 10 $\mu\text{g}/\text{m}^3$ in PM <sub>2.5</sub> )	PAF (>10 $\mu\text{g}/\text{m}^3$ in PM <sub>2.5</sub> )
CV Mortality	1.03 (1.00-1.05)	8.7%
CVD	1.05 (1.03-1.07)	13.9%
MI	1.03 (1.00-1.06)	8.7%
Stroke	1.08 (1.05-1.11)	21.1%

PM 2.5 in Canada & Europe is 10-20  $\mu\text{g}/\text{m}^3$ , 50  $\mu\text{g}/\text{m}^3$  in S Asia & 70  $\mu\text{g}/\text{m}^3$  in

China

Together with

ESC Congress  
Paris 2019

World Congress  
of Cardiology



Population Health  
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HEALTH THROUGH KNOWLEDGE

# Conclusions (1)

- CVD is more common in poorer countries
- CVD accounts for 40% of deaths globally (23% HIC, 42% MIC, 43% LIC)
- Cancer deaths more frequent than CVD deaths in several HIC and some UMIC
- Differences in CVD rates not primarily due to differences in metabolic risk factors, but may be due to differences in smoking, diet, air pollution and access to care

# Conclusions (2)

- Modifiable risk factors explain about 70% of the risk of CVD and deaths
- HR of education & tobacco vary between HIC, MIC & LIC
- Importance of education, grip strength and air pollution previously under-appreciated
- Given the higher prevalence of low education, air pollution, and poor diet in LIC & MIC, their impacts are larger in these countries

# Implications

## Reduction of CVD and mortality requires:

- modifying risk factors which have larger effects in specific contexts, with continuing emphasis on low cost proven treatments, control of hypertension and tobacco control
- improving health care & reducing indoor and outdoor air pollution, particularly in poorer countries

# Published today in the Lancet

Variations in common diseases, hospital admissions, and deaths in middle-aged adults in 21 countries from five continents (PURE): a prospective cohort study

*Gilles R Dagenais\*, Darryl P Leong\*, Sumathy Rangarajan, Fernando Lanas, Patricio Lopez-Jaramillo, Rajeev Gupta, Rafael Diaz, Alvaro Avezum, Gustavo B F Oliveira, Andreas Wielgosz, Shameena R Parambath, Prem Mony, Khalid F Alhabib, Ahmet Temizhan, Noorhassim Ismail, Jephath Chifamba, Karen Yeates, Rasha Khatib, Omar Rahman, Katarzyna Zatonska, Khawar Kazmi, Li Wei, Jun Zhu, Annika Rosengren, K Vijayakumar, Manmeet Kaur, Viswanathan Mohan, Afzal Hussein Yusufali, Roya Kelishadi, Koon K Teo, Philip Joseph, Salim Yusuf*

Modifiable risk factors, cardiovascular disease, and mortality in 155722 individuals from 21 high-income, middle-income, and low-income countries (PURE): a prospective cohort study

*Salim Yusuf\*, Philip Joseph\*, Sumathy Rangarajan, Shofiqul Islam, Andrew Mente, Perry Hystad, Michael Brauer, Vellappillil Raman Kutty, Rajeev Gupta, Andreas Wielgosz, Khalid F AlHabib, Antonio Dans, Patricio Lopez-Jaramillo, Alvaro Avezum, Fernando Lanas, Aytekin Oguz, Iolanthe M Kruger, Rafael Diaz, Khalid Yusoff, Prem Mony, Jephath Chifamba, Karen Yeates, Roya Kelishadi, Afzal Hussein Yusufali, Rasha Khatib, Omar Rahman, Katarzyna Zatonska, Romaina Iqbal, Li Wei, Hu Bo, Annika Rosengren, Manmeet Kaur, Viswanathan Mohan, Scott A Lear, Koon K Teo, Darryl Leong, Martin O'Donnell, Martin McKee, Gilles Dagenais*