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England

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# Attribution of all-cause mortality associated with long-term average concentrations of NO<sub>2</sub>

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*Associations of long-term average concentrations of nitrogen dioxide with mortality*

A report by the Committee on the Medical Effects of Air Pollutants

Chairman: Professor Frank Kelly

Chairman of the QUARK Working Group on Nitrogen Dioxide:  
Professor Roy Harrison

## Request from Defra:

- How to quantify the benefits of reducing long-term average concentrations of NO<sub>2</sub>.
- To support the development of Air Quality Plans to reduce NO<sub>2</sub> concentrations

COMEAP 2018

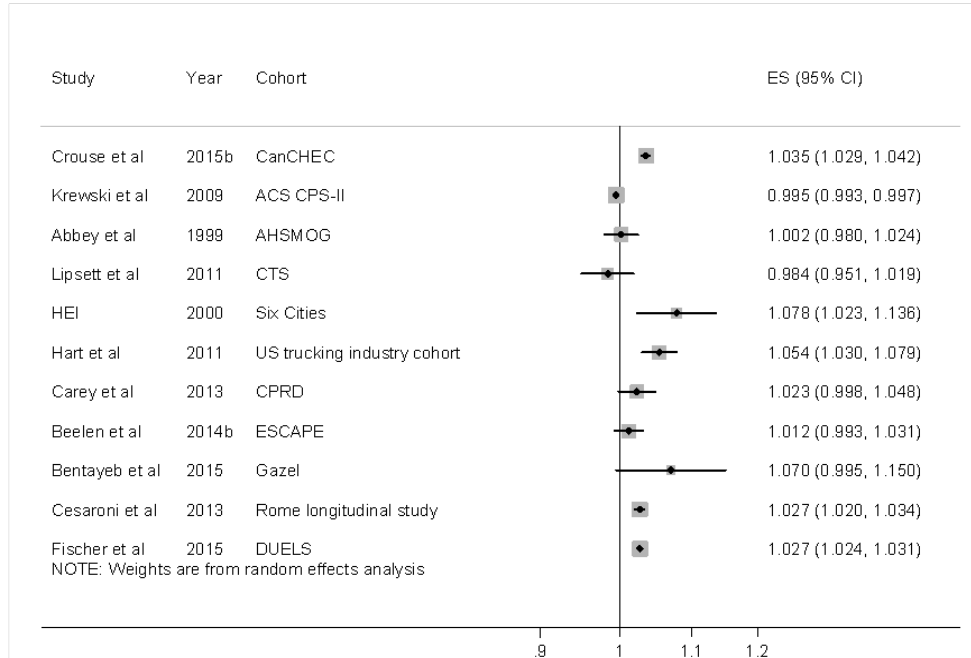
[www.comeap.org.uk](http://www.comeap.org.uk)

# Mortality estimates for NO<sub>2</sub> - HRAPIE

Pollutant metric	Health outcome	Group	RR (95% CI) per 10 µg/m <sup>3</sup>	Range of concentration
NO <sub>2</sub> , annual mean	Mortality, all (natural) causes, age 30+ years	B*	1.055 (1.031–1.080)	>20 µg/m <sup>3</sup>

Source of background health data	Source of CRF	Comments
MDB (WHO, 2013c), rates for deaths from all natural causes (ICD-10 chapters I–XVIII, codes A–R) in each of the 53 WHO Regional Office for Europe countries, latest available data	Meta-analysis of all (11) cohort studies published before January 2013 by Hoek et al. (2013); RR based on single-pollutant models	Some of the long-term NO <sub>2</sub> effects may overlap with effects from long-term PM <sub>2.5</sub> (up to 33%); this is therefore recommended for quantification under Group B to avoid double counting in Group A analysis

# New meta-analysis



Random effects  
summary HR:  
1.023 (95% CI  
1.008, 1.037)  
per 10 µg/m<sup>3</sup> NO<sub>2</sub>

HRs (95% CI) per 10 µg/m<sup>3</sup> for cohort studies reporting associations between NO<sub>2</sub> and all-cause mortality

To note:

**All-cause** mortality only considered (not cause-specific mortality)

Cut-off date for literature review – **October 2015** (new studies now available)

**NH<sub>3</sub>**

**NM VOC**

# Health Matters

**Air pollution:** sources, impacts and actions

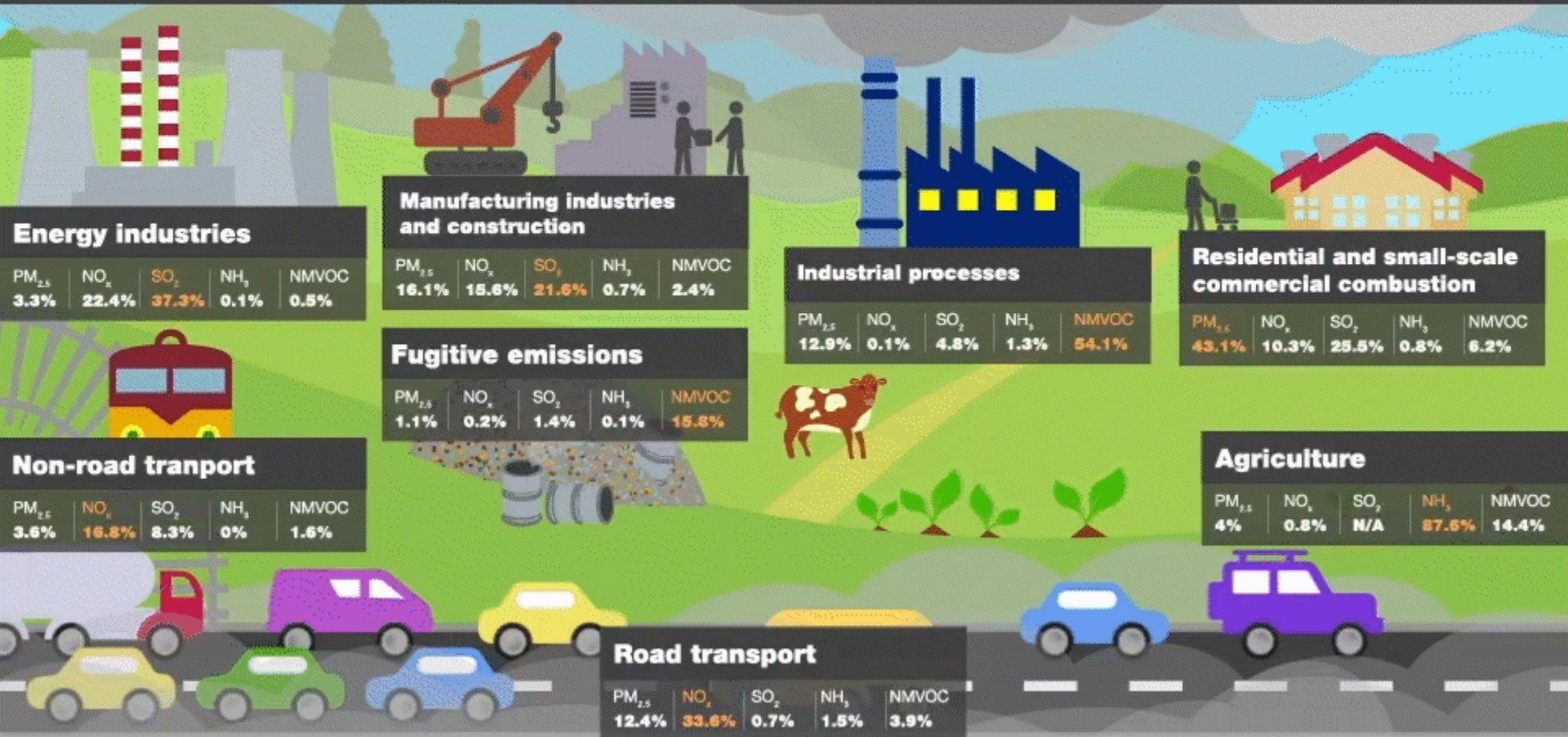
**PM<sub>2.5</sub>**

**SO<sub>2</sub>**

**NO<sub>x</sub>**



# Sources of air pollution



**Pollution substances:**

SO<sub>2</sub> - Sulphur dioxide  
 NO<sub>x</sub> - Nitrogen oxides

NH<sub>3</sub> - Ammonia  
 PM<sub>2.5</sub> - Primary particulate matter

NMVOCS - Non-methane volatile organic compounds

# **Report of a Workshop to Identify Needs for Research on the Health Effects of Nitrogen Dioxide - London, 2-3 March 2011**

**Workshop convened by the Air Pollution Group, Health Protection Agency. Commissioned by the Department of Health's Policy Research Programme**

## **ABSTRACT**

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It is clear from epidemiological and laboratory studies that air pollution has adverse effects on health. However, understanding whether ambient concentrations of nitrogen dioxide (NO<sub>2</sub>) have direct adverse effects on health has proved to be difficult, because levels in ambient air correlate closely with those of other pollutants, notably particles. This difficulty arises because NO<sub>2</sub> and particles have similar sources, such as traffic.

# COMEAP (2015) Statement on NO<sub>2</sub>

- Strengthening evidence for associations with NO<sub>2</sub>
- Associations robust to adjustment for other pollutants incl PM
- May act as a marker for TRAP to some extent, but evidence suggests that it would be sensible to regard NO<sub>2</sub> as causing some of the health impact reported in epidemiological studies

## **Other assessments:**

WHO REVIHAAP (2013): NO<sub>2</sub> might represent TRAP but

- Reasonable to infer that NO<sub>2</sub> has some direct effects following short-term exposure (associations robust to adjustment, some mechanistic support for causality - particularly for respiratory outcomes)
- Harder to judge independent effects of NO<sub>2</sub> in studies of long-term exposure. Short-term associations and mechanistic evidence, particularly on respiratory effects, suggest a causal relationship

USEPA ISA (2016)

Health Canada (2016)



# Unadjusted coefficient for NO<sub>2</sub>

Reflects:

- any causal effect of NO<sub>2</sub>
- and also, to some extent, the effects of other pollutants correlated with NO<sub>2</sub>

Correlated pollutants:

- PM<sub>2.5</sub>
- other components of the air pollution mixture eg
  - Ultrafine particles
  - Black Carbon
  - Volatile Organic Compounds etc

# Independence from PM mass

Using HRs (per IQR) from studies reporting results from single and two/multi-pollutant models for NO<sub>2</sub> and PM:

Compared:

- HRs from single-pollutant models with
  - Combined adj HRs (NO<sub>2</sub> adj for PM; PM adj for NO<sub>2</sub>)
- The combined effect using coefficients each adjusted for the effects of the other, is either similar to or only little higher than what would be estimated for either PM<sub>2.5</sub> or NO<sub>2</sub> alone

## Hazard ratios (HRs) expressed per interquartile range (IQR) from single and two pollutant models for NO<sub>2</sub> and PM<sub>2.5</sub>

Study (additional sig figs for HRs obtained from the authors)	Corr NO <sub>2</sub> / PM <sub>2.5</sub>	NO <sub>2</sub> IQR (µg/m <sup>3</sup> )	PM <sub>2.5</sub> IQR (µg/m <sup>3</sup> )	NO <sub>2</sub> adj PM <sub>2.5</sub>	PM <sub>2.5</sub> adj NO <sub>2</sub>	Combined NO <sub>2</sub> adj / PM adj	PM <sub>2.5</sub> Single	NO <sub>2</sub> Single
Cesaroni et al 2013	0.79	10.7	5.7	1.026	1.004	<b>1.030</b>	1.023	<b>1.029</b>
Carey et al 2013 pers comm	0.85	10.7	1.9	1.001	1.023	<b>1.024</b>	<b>1.023</b>	1.022
Beelen et al 2014 14 cohorts	0.2- <0.7	10.0	5.0	1.007	1.060	<b>1.067</b>	<b>1.070</b>	1.015
Fischer et al 2015 PM <sub>10</sub>	0.58 (with PM <sub>10</sub> )	10.0	2.4	1.019	1.010	<b>1.029</b>	1.019	<b>1.027</b>
HEI 2000 41 cities	-0.08	81.4 (min,max)	24.5 (min,max)	0.90	1.22	<b>1.09</b>	1.15	0.95
Jerrett et al 2013	0.55	7.7	5.3	1.025	1.015	<b>1.040</b>	<b>1.032</b>	<b>1.031</b>

# Coefficient for $\text{NO}_2$ adjusted for $\text{PM}_{2.5}$

Excludes, as far as possible:

- effects associated with  $\text{PM}_{2.5}$  concentrations, and other components of the air pollution mixture that are more closely correlated with  $\text{PM}_{2.5}$  concentrations than with  $\text{NO}_2$  concentrations

Reflects:

- any causal effect of  $\text{NO}_2$  and also, to some extent, of other pollutants closely correlated with  $\text{NO}_2$
- The extent to which the effect is likely to be causally related to  $\text{NO}_2$  is unclear

# Uncertainty related to causality

## **A key point of COMEAP's discussions:**

Uncertainty in the extent to which the effects associated with NO<sub>2</sub> in epidemiological studies are caused by NO<sub>2</sub> itself

- How to take the uncertainty into account quantitatively when predicting the benefits that would be expected from interventions (eg in cost-benefit assessments)
- Some Members doubtful that the evidence is sufficient to allow a robust recommendation for quantification
- The majority thought it important to attempt to estimate the possible mortality benefit from reducing NO<sub>2</sub> concentrations

# COMEAP – HIA Recommendations

## **HIA of reductions in all traffic-related pollutants:**

- Use summary unadjusted  $\text{NO}_2$  coefficient:
  - 1.023 (95% CI 1.008, 1.037) per  $10 \mu\text{g}/\text{m}^3$
- (Alternatively, could assess based on  $\text{PM}_{2.5}$  reductions)

## **HIA of interventions which target $\text{NO}_x$ emissions**

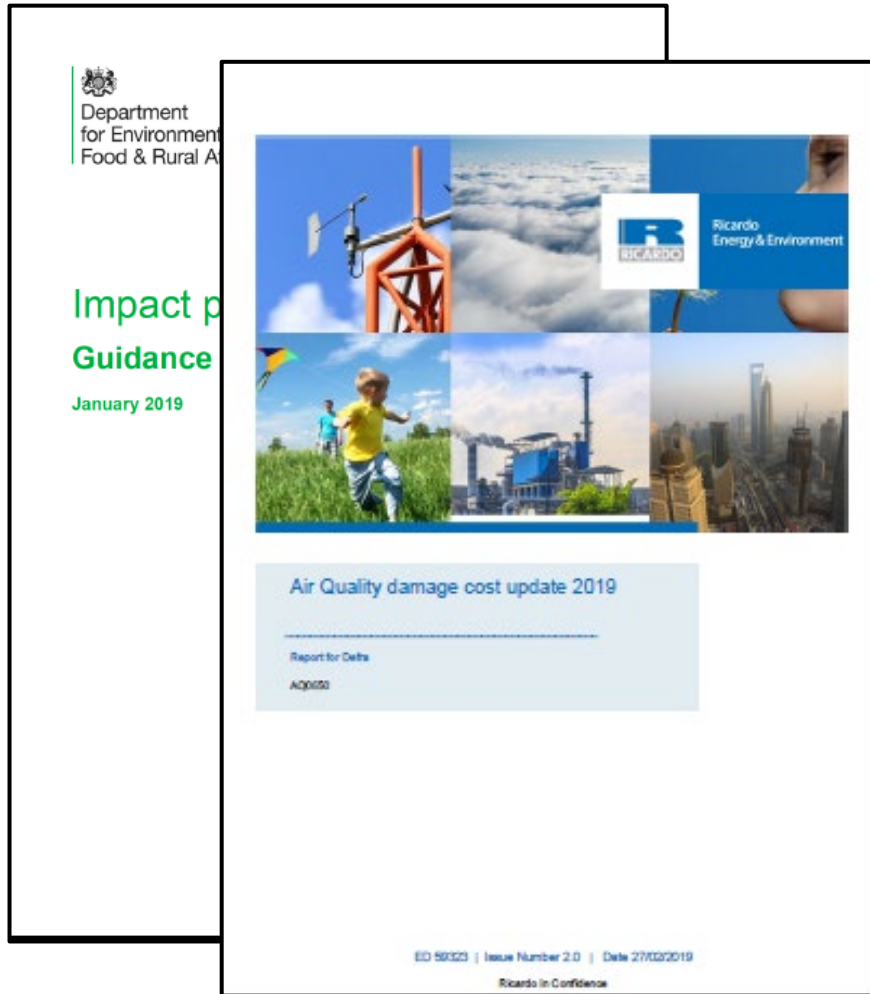
- Use 25 – 55% of summary unadjusted coefficient
  - 1.006 – 1.013 per  $10 \mu\text{g}/\text{m}^3$  – “central range”

Informal expert judgement approach equivalent to:

- Reduction of approx 20% on adjustment for PM
- 30 - 70% of adjusted coefficient may be causal

**Advised:** assessment on the basis of reductions of both  $\text{NO}_2$  and PM (unadjusted) is likely to result in an overestimate

# Implementation in CBA



In practice:

Modelling combines predicted changes in pollutant concentrations arising from a range of proposed interventions

- CBA guidance includes mortality benefits associated with reductions in both
  - NO<sub>2</sub> (reduced coefficient)
  - PM<sub>2.5</sub> (unadjusted)
- Analysts should
  - Acknowledge this limitation
  - Undertake sensitivity analyses

# COMEAP: Research recommendations

- Studies to reduce uncertainties in understanding the effects of long-term exposure to NO<sub>2</sub> on health
  - Toxicological, epidemiological, exposure errors
- Studies to improve quantification of effects associated with air pollution mixtures
  - Develop multi-pollutant and statistical methods
- Investigation of reasons for between-study variability in reported associations



# Summary and discussion points

Science-policy interface:

- NO<sub>2</sub> has been less-studied than PM<sub>2.5</sub>
- But evidence for independent effects of NO<sub>2</sub> grows
- Nonetheless, there is more certainty about the health benefits of interventions that reduce both PM<sub>2.5</sub> and NO<sub>2</sub> (and other co-pollutants)
- Communicating uncertainty is important
  - to allow informed policy decisions to be taken
- New studies and method developments will continue to inform scientific thinking and policy development

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Thank you !