## Frontiers in Air Quality Advances in Air Quality Toxicology

# Fumifugium 2014

### Ian Mudway



Cleaner Fuels (wood vs. coal) Separate population from pollution sources Economic benefits Green infrastructure

FUMIFUGIUM: The Inconveniencie of the AER AND SMOAK of LONDON DISSIPATED. TOGETHER With fome REMEDIES humbly PROPOSED By J. E. Efq; To His Sacred MAJESTIE, To the PARLIAMENT now Affembled.

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John Evelyn 1620 – 1706



# One very large seething Petri dish

## Know your model organism









## Think about susceptibility

Deaths Registered in London Administrative County Classified by Age (Bates, 1995)									
	< 1 Month of Age	1-12 Mo. Old	1-14 Years of Age	15-44 Years of Age	45-64 Years of Age	65-74 Years of Age	75+ Years of Age		
Week Before the Episode	16	12	10	61	237	254	335		
Week After the Episode	28	26	13	99	652	717	949		
Before/ After Episode Ratio	1.75	2.17	1.3	1.62	2.75	2.82	2.83		

# Vulnerability

"This cloud is so inextricably mixed with the naturally wholesome and excellent air that the inhabitants can breathe nothing but thick, dirty, smoggy air. This makes them <u>vulnerable</u> to thousands of diseases, <u>corrupting their lungs</u> and disordering their bodies, so that catarrh, coughs and tuberculosis."



## **Critical windows of enhanced vulnerability**



Degradation of defence mechanism with age

Development

MRC-PHE Centre for Environment & Health

Public Health

Imperial College

In utero

# Appreciate varying genetic background: An ethnically diverse city





"This plague of a smoke is truly intolerable because, although it does not kill at once, it is deadly and to slowly perish is worse than death itself. Is there any other place on Earth where such coughing and snuffling is to be heard as in the churches and assemblies of London, where the barking and spitting is incessant? Need I say more?"



## Role of air pollution in chronic disease



"There are, I must accept, certain differences in people which allow them to thrive more in some airs compared to others."



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## Impaired lung growth

#### Gauderman et al, 2004 (NEJM); southern California

- 1,759 10-yr old children, 12 communities, 8-yr follow up
- Lung function growth significantly reduced in areas with higher levels of traffic-related pollutants
  - NO<sub>2</sub>, PM<sub>2.5</sub>, elemental carbon (EC) associated with reduction in growth of FEV<sub>1</sub> of between 79.7 and 101.4 mL
  - Clinically low FEV<sub>1</sub> at age 18 positively correlated with level of exposure to NO<sub>2</sub>, PM<sub>10</sub>, PM<sub>2.5</sub>, EC

#### Gauderman et al, 2007 (Lancet); southern California

-Same study, local vs. regional pollutant levels -Living <500m from freeway associated with reduction in FEV<sub>1</sub> growth of 81mL (vs. living >1500m away) -Significantly lower attained FEV<sub>1</sub> by 18 yrs

### **Pollution in Tower Hamlets & Hackney**



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# **Modelled annual NO<sub>2</sub> concentrations**



Gehring U, et al. Environ Health Perspect. 2013;121(11-12):1357-64.

#### Context



"There are, I must accept, certain differences in people which allow them to thrive more in some airs compared to others."



# **Genetic Modification**

#### Xenobiotic metabolism:

• CYP1A1, GSTM1, GSTT1, GSTP1, EPHX1, NQO1, AhR

#### Antioxidant defence:

• GCL, ECSOD, Nrf2

#### Nitrosative stress:

• iNOS, ARG1, ARG2

#### **Onset of childhood asthma**

• GSDMA, GSDMB

#### **Random ancestry markers:**

• Adjust for population stratification and predict lung function as proxy for ethnic group



## Quantifying exposure BIOMARKERS

1

2

3

6

7

Air in

1 IA Traffic related met   1 2 IIA IIIA   1 2 IIA IIIA   1 8 9 10   1 1008 2 IIIA IIIA   1 8 9 10 IIIA   1 12 3 4 5 6 7 VIIIB 11 12 13   22.99 24.31 IIIB 1VB VB VIB VIB 11 12 13   2000 Augustum IIIB 1VB VIB VIIIB 11 12 13   2000 IIIB VIB VIB IIIB 12 13   2000 IIIB IVB VIB IIIB 12 13   2000 IIIB IIIB IIIB IIIA IIIB IIIB IIIB IIIB IIIB IIIB IIIA	14 15 16 1 VA VA VIA V	He
3   4   5   5     11   12   3   4   5   6   7   VIIIB   11   12   13     22.99   24.31   3   4   5   6   7   VIIIB   11   12   26.98     Sodium   Magnetium   IIIB   IVB   VB   VIIB   IB   IIB   IIB   All	SCONT CONTRACTOR OF STREET	17 7IIA 4.00 Helium
11   12   3   4   5   6   7   VIIIB   11   12   26.98     Sodum   Magnesium   IIIB   IVB   VB   VIB   VIIB   IB   IIB   IIB   Auminum	6 7 8 12.01 14.01 16.00 11	F 10 9.00 20.18 Neon
K C C T' N C N T C N' C Z C	14 15 16 28.09 30.97 32.07 33	Cl Ar 17 18 15.45 39.95 hiorine Argon
K   Ca   Sc   Ti   V   Cr   Mn   Fe   Co   Ni   Cu   Zn   Ga     19   20   21   22   23   24   25   26   27   28   29   30   31     39.10   40.08   44.96   47.88   50.94   52.00   54.94   55.85   58.93   58.69   63.55   65.39   69.72     Pressum   Cutation   Scandium   Titanium   Viewfelium   52.000   Titanium   Titani	32 33 34 72.61 74.92 78.96 79	Br 35 36 99.90 83.80 Krypton
Rb   Sr   Y   Zr   Nb   Mo   Tc   Ru   Rh   Pd   Ag   Cd   In     37   38   39   40   41   42   43   44   45   46   47   48   49   92.91   101.07   102.91   106.42   107.87   112.41   114.82     Mobdum   Yttruin   Zircosium   Wohm   Wohm   Russement   Rindium   Rindium   Rindium   Silver   Cadmient   Indum	50 51 52 1 18.71 121.76 127.60 12	I 53 26.90 26.90 Xenon
Cs   Ba   La   Hf   Ta   W   Re   Os   Ir   Pt   Au   Hg   Tl     55   56   57   72   73   74   75   76   77   78   79   80   81     132.91   137.33   138.91   178.49   180.95   138.85   190.2   192.22   195.08   Gold   200.59   204.38     Cesum   Bartimum   Lambarum   Hathum   Trathum   Trathum   Trathum   Gold   Mercary   Tallum	82 83 84 2 207.2 208.98 (209) (2	At Rn 85 210) (222) statine Radon
Fire   Ra   Acc   Rf   Db   Sg   Bh   Hs   Mt   Inamed   Unnamed   Discovery   Unnamed   Discovery   Inamed   Disco	nnamed Unnamed	Unnamed Discovery



#### **Biomarkers out**



# PM<sub>10</sub> Metals (Fe and Cu)





# PM<sub>10</sub> Metals (Ba and Mn)



# PM<sub>10</sub> Metals (Cr and V)



Cleaner Fuels (diesel) Separate the vulnerable population from pollution sources **Economic benefits** Green infrastructure Moral imperative

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"I have made this proposal for refining the quality of air in London because I believe that we should do whatever we can to enhance the honour of our nation that is capable of such greatness. <u>It troubles me</u> <u>that the health and happiness of so many people should suffer from</u> <u>the greed of a few in a city that contains enough to make its people</u> <u>the happiest on Earth</u>. Money has blinded people to the thing which keeps them alive and which can, for their own sake, be improved so easily."

# **Omnia explorate; meliora retinete**

