

Air Pollution, housing and respiratory tract Infections in Children: Natlonal birth

Cohort study (PICNIC)

Pia Hardelid
University College London
GOS Institute of Child Health





## **L**

The Telegraph

More than 2,000 children die from pneumonia every day, charities warn



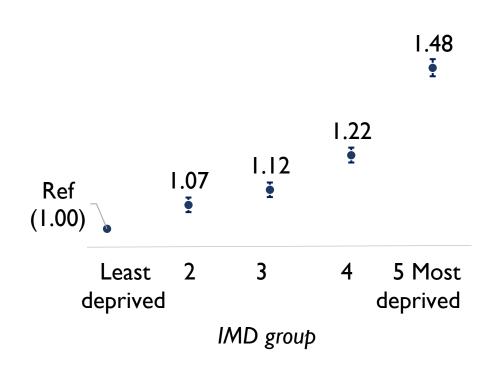
Childhood pneumonia cases up 50% in 10 years, NHS data shows



Poorest areas hardest hit by 'forgotten epidemic' of pneumonia putting a child in hospital every 10 minutes

Fig 3. Rate ratio of bronchiolitis admissions <1 year: England, 2012-2016





Lewis, De Stavola & Hardelid (J Epidemiol Comm Health, 2020)

Slide courtesy of Kate Lewis





### EUROPEAN RESPIRATORY journal

FLAGSHIP SCIENTIFIC JOURNAL OF ERS

Risk factors for admission to hospital with laboratory-confirmed influenza in young children: birth cohort study

Pia Hardelid<sup>1</sup>, Maximiliane Verfuerden<sup>1</sup>, Jim McMenamin<sup>2</sup> and Ruth Gilbert<sup>1</sup>

### Risk factor and scenario#

### Population-attributable fraction

### Age group 1: <6 months

Parity: first child	45.5 (33.5-55.3)
Maternal age: 30–39 years	18.5 (5.5–29.7)
High risk: no	4.5 [1.3–7.7]
Season of birth: April-June	65.8 (49.9–76.6)



#### RESEARCH ARTICLE

The contribution of child, family and health service factors to respiratory syncytial virus (RSV) hospital admissions in the first 3 years of life: birth cohort study in Scotland, 2009 to 2015



#### Pia Hardelid<sup>1</sup>, Maximiliane Verfuerden<sup>1</sup>, Jim McMenamin<sup>2</sup>, Rosalind L Smyth<sup>1</sup>, Ruth Gilbert<sup>1</sup>

- 1. UCL Great Ormond Street Institute of Child Health, London, United Kingdom
- 2. Health Protection Scotland, Glasgow, United Kingdom

Population attributable fraction (as a percentage of admissions prevented) by risk factor and scenario, birth cohort study, Scotland, 2009-2015

Risk factor and scenario	Population attributable fraction (95% CI)
Gestational age = post term	18.9 (13.8-23.7)
Season of birth = April-June	25.6 (21.0–30.0)
Chronic condition = No	6.5 (5.6–7.5)
Parity = 0	34.0 (31.0-36.9)
Maternal smoking during pregnancy = no	5.9 (4.2–7.7)
Maternal age = ≥ 40 years	31.4 (18.7-42.1)
Birth weight at gestational age = right for gestational age	1.6 (0.1–3.0)
Delayed infant vaccination = no <sup>a</sup>	2.5 (0.5–4.5)

## 

## Air Pollution, housing and respiratory tract Infections in Children: National birth Cohort study (PICNIC)

















#### BMJ Open Air Pollution, housing and respiratory tract Infections in Children: NatIonal birth Cohort study (PICNIC): study protocol

Graziella Favarato, 1 Tom Clemens, 2 Steven Cunningham, 3 Chris Dibben, 2 Alison Macfarlane,4 Ai Milojevic,5 Jonathon Taylor,6

Linda Petronella Martina Maria Wijlaars 0, 1 Rachael Wood 0, 7,8 Pia Hardelid 01

To cite: Favarato G, Clemens T, Cunningham S, et al. Air Pollution, housing and respiratory tract Infections in Children: National birth Cohort study (PICNIC):

study protocol RMI Open

bmiopen-2020-048038

 Prepublication history and additional supplemental material for this paper are available online. To view these files, please visit the journal online (http://dx.doi.org/10.1136/ bmiopen-2020-048038).

2021;11:e048038.doi:10.1136/

Received 17 December 2020 Revised 22 March 2021 Accepted 14 April 2021

Check for updates

C Author(s) (or their

Published by BMJ.

Correspondence to

p.hardelid@ucl.ac.uk

Dr Pia Hardelid:

end of article

employer(s)) 2021, Re-use

For numbered affiliations see

permitted under CC RV

Introduction Respiratory tract infections (RTIs) are the most common reason for hospital admission among children <5 years in the UK. The relative contribution of ambient air pollution exposure and adverse housing conditions to RTI admissions in young children is unclear and has not been assessed in a UK context. Methods and analysis The aim of the PICNIC study

(Air Pollution, housing and respiratory tract Infections in Children: National birth Cohort Study) is to quantify the extent to which in-utero, infant and childhood exposures to ambient air pollution and adverse housing conditions are associated with risk of RTI admissions in children <5 years old. We will use national administrative data birth cohorts, including data from all children born in England in 2005-2014 and in Scotland in 1997-2020, created via linkage between civil registration, maternity and hospital admission data sets. We will further enhance these cohorts via linkage to census data on housing conditions and socioeconomic position and small area-level data on ambient air pollution and building characteristics. We will use time-to-event analyses to examine the association between air pollution, housing characteristics and the risk of RTI admissions in children, calculate population attributable fractions for ambient air pollution and housing characteristics, and use causal mediation analyses to explore the mechanisms through which housing and air pollution influence the risk of infant RTI admission.

Ethics, expected impact and dissemination. To date. we have obtained approval from six ethics and information governance committees in England and two in Scotland. Our results will inform parents, national and local governments, the National Health Service and voluntary sector organisations of the relative contribution of adverse housing conditions and air pollution to RTI admissions in young children. We will publish our results in open-access journals and present our results to the public via parent groups and social media and on the PICNIC website. Code and metadata will be published on GitHub.

INTRODUCTION

Upper and lower respiratory tract infections (RTIs), including croup, bronchiolitis and pneumonia, are the most common reason for

#### Strengths and limitations of this study

- ► The PICNIC study will use national, administrative data birth cohorts from England and Scotland, linked to small area-level data on environmental exposures and census data on socioeconomic position, to exand adverse housing exposures and respiratory tract infection admissions in children less than 5 years
- ▶ The national birth cohorts will include all children. born in the two countries during specified time periods, thus minimising selection bias and allowing analyses of even relatively uncommon environmental exposures and infection outcomes.
- Data on air pollution and building characteristics will be linked to maternal and child postcode histories during pregnancy and early life, thus creating longitudinal environmental exposure data at a national
- ► PICNIC will include an examination of the populationbased risk factors for SARS-CoV-2 infection in Scottish children.
- A key weakness is that only infections requiring hospital admission will be the primary outcome, thus respiratory infections not requiring healthcare contact will not be considered.

hospital admission in children aged less than 5 years old in the UK, with 170 000 admissions in England alone in 2017. RTI admission rates peak in winter months, contributing to the 'winter crisis' in the National Health Service (NHS).2 Severe RTI symptoms in infancy and early childhood have been linked to adverse respiratory health outcomes in later childhood, including asthma.3 We have previously shown that 79% of annual admissions for RTIs in infants <1 year old can be attributed to respiratory syncytial virus (RSV) and other viral infections for which no vaccines are currently available.4 Thus, alternative

http://dx.doi.org/10.1 136/bmjopen-2020-048038

BMJ





# Fury as long-awaited UK environment bill is delayed for third time

Cramped housing has helped fuel spread of Covid in England - study

Green campaigners attack further delay 'to most important piece of legislation for decades'



'The worst I've ever seen': The appalling and 'unliveable' council housing conditions some have endured during lockdown



UK broke law by 'systematically and persistently' breaching air pollution limits, top court rules

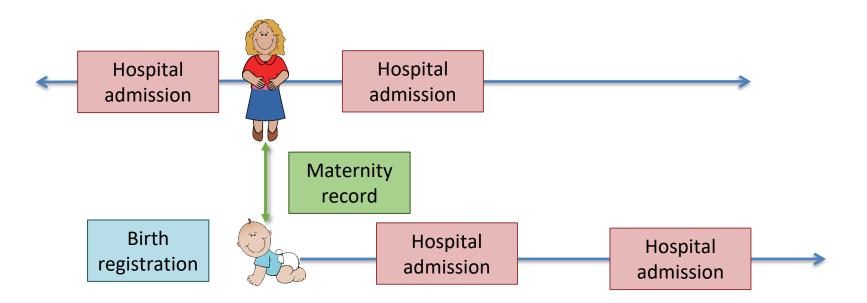


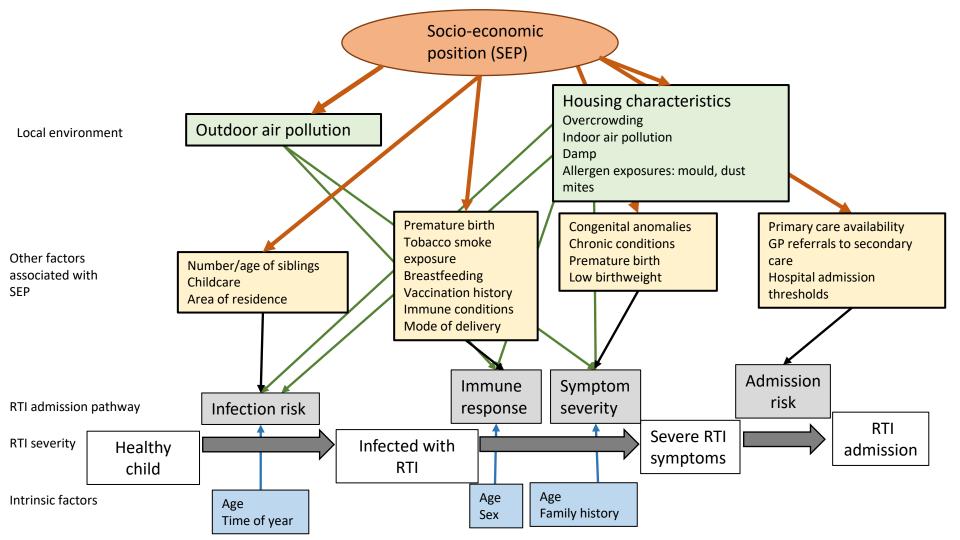
Pregnant mum 'living out of bags' in cramped house plagued by mould and mice

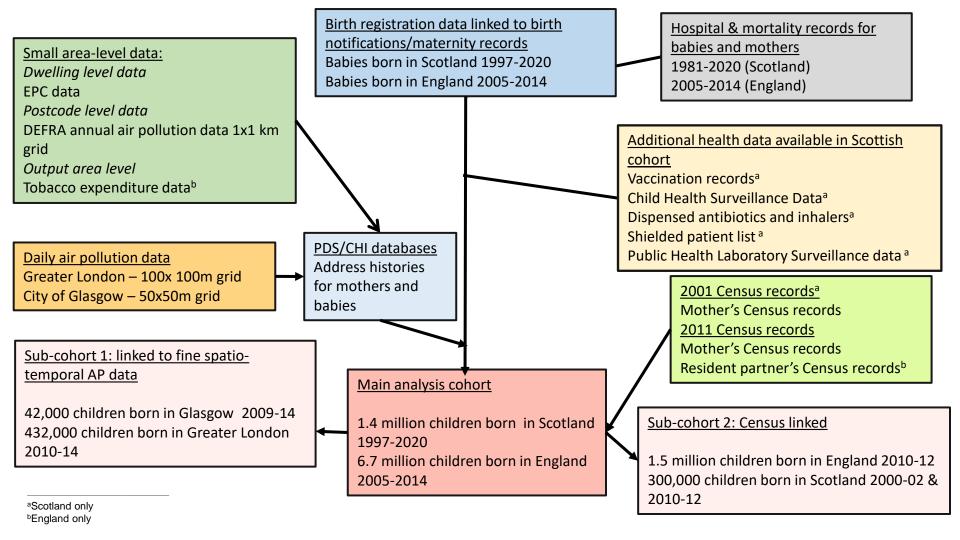
## **PICNIC Aim:**



'To determine the contribution of exposure to air pollution and adverse housing conditions during pregnancy, infancy and early childhood to hospital admissions for respiratory tract infections (RTIs) in children less than 5 years old in England and Scotland'







### **Ambient air pollution data sources - DEFRA**



DEFRA	CERC
PM10, PM2.5, NO2, NOx, CO,	DM10 DM2 F NO2 O2
SO2, O3, benzene	PM10, PM2.5, NO2, O3
1x1km grid*	100x100m grid*
Annual	Daily
2001-2018	2010-2014
England & Scotland	Greater London
x & y co-ordinates of each grid	x & y co-ordinates of each grid
cell	cell
Postcode*	Postcode*
	PM10, PM2.5, NO2, NOx, CO, SO2, O3, benzene  1x1km grid*  Annual  2001-2018  England & Scotland  x & y co-ordinates of each grid cell

<sup>\*</sup>All based on the Ordnance Survey (OS) National Grid reference system (also known as British National Grid)

# Ambient air pollution data sources: Cambridge Environmental Research Consultants (CERC): ADMS-Urban

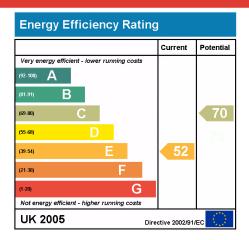


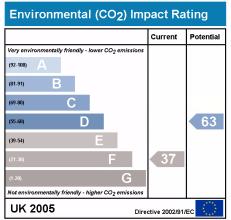
	DEFRA	CERC
Air pollutants	PM10, PM2.5, NO2, NOx, CO, SO2, O3, benzene	PM10, PM2.5, NO2, O3
Spatial resolution	1x1km grid*	100x100m grid* / 50x50m grid
Temporal resolution	Annual	Daily
Coverage (years)	2001-2018	2010-2014 / 2009-2014
Coverage (geography)	England & Scotland	Greater London / Glasgow
Geographical identifier in dataset	x & y co-ordinates of each grid cell	x & y co-ordinates of each grid cell
Linking variable (to mother-birth cohorts)	Postcode*	Postcode*
*All based on the Ordnance Survey (OS) National Grid reference system (also known as British		

<sup>\*</sup>All based on the Ordnance Survey (OS) National Grid reference system (also known as British National Grid)

### **Energy Performance Certificates**







- Data collected by surveyor when a property is let, sold or constructed in the UK (valid for 10 years)
- EPC data for PICNIC includes:
  - directly observed variables e.g. property type, number of rooms, number of heated rooms, total floor area, mains gas connection..
  - Inferred variables: infiltration rate
  - Modelled variable: indoor/outdoor ratio for PM<sub>2.5</sub> and NO<sub>2</sub>
- EPC data linked at postcode level in England, UPRN (dwelling level) in Scotland.





#### Household Questionnaire England





#### Complete online www.census.gov.uk Your personal internet access code is: **OR** fill in this paper questionnaire and post it back using the pre-paid envelope supplied. If your address is incorrect or missing, enter your correct address here: A message to everyone - act now Everyone should be included in the census - all It is used to help plan and fund services for your Declaration afterwards. You can fill it in online or on paper. This questionnaire has been completed to the best of my knowledge and belief. Signature don't participate or if you supply false information.

Telephone number

We may contact you if we need to collect

If you have lost your envelope, please return to:

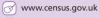
FREEPOST 2011 Census, Processing Centre, UK

missing information.

lices of Deprivation

ble Deprivation 2020

#### Where can you get help?



Census helpline 0300 0201 101

Text Relay 18001 0300 0201 160

Halp is available in large print and Praille

# **Examples of public, parent and charity engagement**



- Parents in Bradford consulted on the prevention of respiratory infections in children
- Parents' & Carers' Advisory Group (PCAG) at Great Ormond Street Hospital/UCL Institute of Child Health
- Shelter Birmingham (Visit to coffee morning)
- National Children's Bureau Parents Research Advisory group
- Conversations with British Lung Foundation & Clean Air Parents' Network / Mums for Lungs



Credit Alexander Drummer (unsplash)

## Information governance approvals



#### England

- 1. NHS London Queen's Square Ethics Committee (reference: 18/LO/1514)
- 2. Confidentiality Advisory group (reference: 18/CAG/0159)
- 3. Administrative Data Research Network (reference PROJ-194 note committee is now defunct)
- 4. ONS Research Accreditation Panel (reference 2019/020)
- 5. National Statistician's Data Ethics Advisory Committee (reference: 18(07))
- 6. Independent Group Advising on Release of Data (NHS Digital, reference DARS-NIC-234656)

#### Scotland

- 1. Public Benefit and Privacy Committee- Health and Social Care (reference 1819-0049)
- 2. Public Benefit and Privacy Committee- Statistics (reference 1819-0049)
- 3. University of Edinburgh School of Geosciences Ethics Committee (reference 2020-401)

### Risks of SARS-CoV-2 infection in children in Scotland

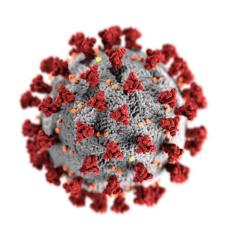


### **Objectives**

- 1) determine the population-based risks of:
  - a) SARS-CoV-2 test
  - b) SARS-CoV-2 infection
  - c) COVID19-related hospital admission
  - d) Multisystem Inflammatory Syndrome in Children
  - e) COVID-related intensive care unit stay

In children and young people in Scotland

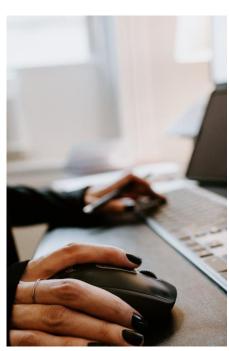
2) establish how these risks vary according to clinical, family and socio-demographic risk factors



## **Progress to date**



- All approvals are now in place
- Air pollution data downloaded and mapped to postcodes
- EPC data mapped to postcodes for England & uploaded to ONS SRS
- Comparison of air pollution exposures during pregnancy according to different spatio-temporal resolution work in progress
- Cleaning and validation of English & Scottish birth cohorts
- Epidemiology of SARS-CoV-2 in children and young people in Scotland study in write up phase





## Factors associated with amoxicillin prescribing in primary care among young children in Bradford, England: cohort study

- One third of children <5 years old are prescribed at least one antibiotic annually
- 75% of antibiotics prescribed to children in primary care in the UK are for respiratory infections (pneumonia, tonsillitis, sinusitis)
- What child, family and environmental factors are associated amoxicillin prescribing in young children?





## **Data sources**



- Born in Bradford Allergy and Infection (ALL-IN) study
- Data from ~2500 children from:
  - Maternity records
  - Baseline questionnaire during pregnancy
  - Congenital anomaly register
  - Parental questionnaires at age 12 and 24 months
  - Electronic primary care records
  - DEFRA air pollution modelled PM<sub>2.5</sub> annual averages at 1x1km grid linked to child LSOA at birth/1 year of age



https://borninbradford.nhs.uk/

## **Exposures and outcomes**



### **Exposure variables**

<u>Child</u>: Sex, annual quarter of birth, prematurity, mode of delivery, congenital anomalies, prematurity, ethnic group

<u>Maternal/family</u>: socio-economic status, maternal smoking during pregnancy, formal childcare, breastfeeding

<u>Environment</u>: overcrowding, mould/damp, gas cooking, quartile of PM2.5 exposure (in relation to Bradford annual average)

### **Outcome variables**

- -One or more amoxicillin prescription (by year of age)
- -One or more consultations for lower respiratory infections (by year of age)
- -One or more consultations for upper respiratory infections (by year of age)

## Results – amoxicillin (odds ratios & 95% CI)



	Age 1	Age 2
	Mutually adjusted: n=2,450	Mutually adjusted: n=2,476
Mother's ethnicity		
White British	1	1
Pakistani, UK born	1.44 (1.06-1.94)	1.46 (1.10-1.94)
Pakistani, not UK born	1.42 (1.07-1.90)	1.56 (1.19-2.04)
Other	0.70 (0.52-0.96)	0.98 (0.74-1.31)
Socio-economic position		
Least deprived and most educated	1	1
<b>Employed not materially deprived</b>	0.79 (0.59-1.06)	1.13 (0.85-1.50)
Employed no access to money	0.92 (0.69-1.22)	1.11 (0.85-1.46)
Benefits but coping	0.92 (0.70-1.21)	1.26 (0.97-1.64)
Most deprived	1.36 (1.00-1.86)	1.26 (0.93-1.70)

#### Overall rates of amoxicillin prescribing:

Age 1: 710/1000 child years Age 2: 780/1000 child years

# Results – amoxicillin: (odds ratios & 95% CI) environment variables



	Age 1	Age 2
Quartile of PM2.5 in relation to		
Bradford level		
1st/2nd quartile of PM2.5		1
3rd quartile of PM2.5		0.94 (0.76-1.16)
4th quartile of PM2.5		0.97 (0.75-1.26)
Mould or visible damp spots		
No mould or damp	1	
Mould or damp	0.98 (0.80-1.21)	
Number of people in household		
2-5 people	1	
6 or more people	1.41 (1.14-1.74)	

# Results – lower respiratory infections (odds ratios & 95% Cls)

	Age 1	Age 2
Mother's ethnicity		
White British	1	1
Pakistani, UK born	1.21 (0.88-1.65)	1.40 (1.01-1.95)
Pakistani, not UK born	1.43 (1.06-1.91)	1.61 (1.17-2.21)
Other	0.76 (0.54-1.06)	1.15 (0.83-1.61)
Socio-economic position		
Least deprived and most educated	1	1
Employed not materially deprived	1.27 (0.91-1.76)	1.41 (1.02-1.95)
Employed no access to money	1.55 (1.13-2.12)	1.01 (0.73-1.39)
Benefits but coping	1.42 (1.06-1.91)	1.19 (0.88-1.60)
Most deprived	1.84 (1.32-2.58)	1.47 (1.05-2.08)

#### **Overall rates of LRI consultations:**

Age 1: 458/1000 child years

Age 2: 409/1000 child years

# Results – lower respiratory infections Environment variables (odds ratios & 95% CIs)



	Age 1	Age 2
Quartile of PM2.5 in relation to		
Bradford level		
1st-2nd quartile of PM2.5		1
3rd quartile of PM2.5		1.06 (0.83-1.35)
4th quartile of PM2.5		1.09 (0.81-1.46)
Mould or visible damp spots		
No mould or damp	1	
Mould or damp	0.84 (0.67-1.06)	

## Summary – factors associated with amoxicillin prescribing in BiB ALL-IN

- 43% and 47% of children were prescribed amoxicillin in their  $1^{st}$  /  $2^{nd}$  year of life respectively
- Ethic group was consistently associated with amoxicillin prescribing and lower respiratory infection consultations; socio-economic status associated with lower respiratory infection consultations and amoxicillin prescriptions during first year
- Overcrowding was only environmental variable associated with amoxicillin prescribing
- Study limited to one specific area of England with particular ethnic group composition;
   need national studies with more variation in environmental exposures across different families in order to detect these effects

## **Lessons learned from PICNIC study: so far**



- To examine questions about the impact of ambient air pollution and child health, we need national administrative data linked to environmental data: they are inclusive, large and geographically varied
- These datasets are extremely difficult to access, with applications to multiple data providers take years
- This is despite low risk of disclosure and evidence of public support



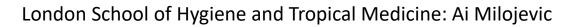
Photo credit: Kiana Bosman, unsplash



### **PICNIC** project partners

UCL GOS ICH: Pia Hardelid, Graziella Favarato, Linda Wijlaars, Bianca De Stavola







University of Edinburgh: Steve Cunningham, Chris Dibben, Tom Clemens

Tampere University: Jonathon Taylor





City, University of London: Alison Macfarlane

Charity collaborators: Shelter, British Lung Foundation/Clean Air Parents Network



Public Health Scotland: Rachael Wood, Lynda Fenton, Jim McMenamin



Funding: Medical Research Council

### **ALL-IN Amoxicillin study: team**



UCL GOS ICH: Faith Miller, Graziella Favarato

Bradford Institute for Health Research: Dan Mason, John Wright

London School of Hygiene and Tropical Medicine: Lucy Pembrey

UCL Institute of Health Informatics: Laura Shallcross

University of Edinburgh: Imad Adamestam

Funding: Medical Research Council (via Studentship to FM)







https://www.ucl.ac.uk/child-health/picnic-study