



# Showcasing PHIRI use case results measuring the impact of COVID-19 on population health

Workshop – 10<sup>th</sup> Nov. 2022



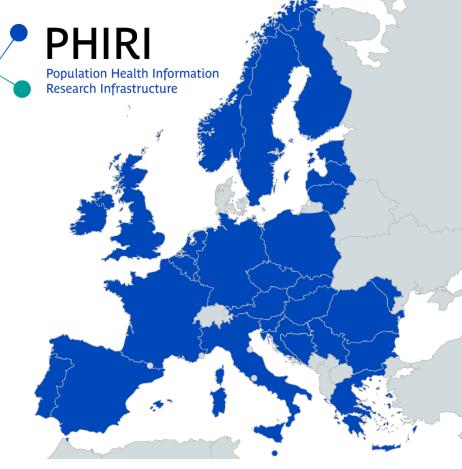




The Population Health Information Research Infrastructure for COVID-19:

- a European mechanism, that aims to
- facilitate and support data-driven population health research
- and exchange of best practices
- to support decision making

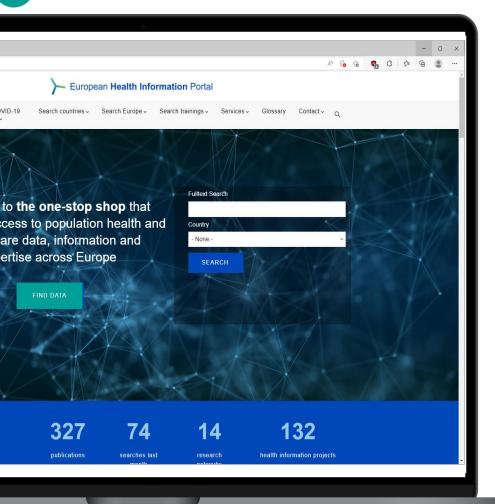




Map of PHIRI Partners

## **The European Health Information Portal**





#### www.healthinformationportal.eu

A one-stop shop that facilitates access to population health and health care data, information and expertise across Europe.



Health information (data) sources



Countries and national nodes



Research infrastructures, Research networks



Health information projects



**Publications** 



Trainings in all areas of population health



COVID-19 Policy measures



COVID-19 Rapid Exchange Forum

# PHIRI: Real-world data measuring the COVID19 indirect "impact"





Direct and indirect determinants of COVID-19 infection and outcomes in vulnerable population groups with reference to inequalities



COVID-19 related delayed care in breast cancer patients



The impact of COVID-19 on perinatal health and perinatal health inequalities



COVID-19 related changes in population mental health









# Use Case A: Indirect health effects of the COVID-19 pandemic: Insights from European countries

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With: Ronan A. Lyons, Claudia Habl, Lorenz Dolanski-Aghamanoukjan, Markus Keski-Säntti, Stefan Mathis-Edenhofer, Hanna Tolonen, Mika Gissler, Jakov Vukovic, Klea Kriz, Tamara Bubble, Enrique Bernal Delgado, Francisco Estupiñan-Romero, Javier, Gonzalez Galindo, Martin Thißen, and others...















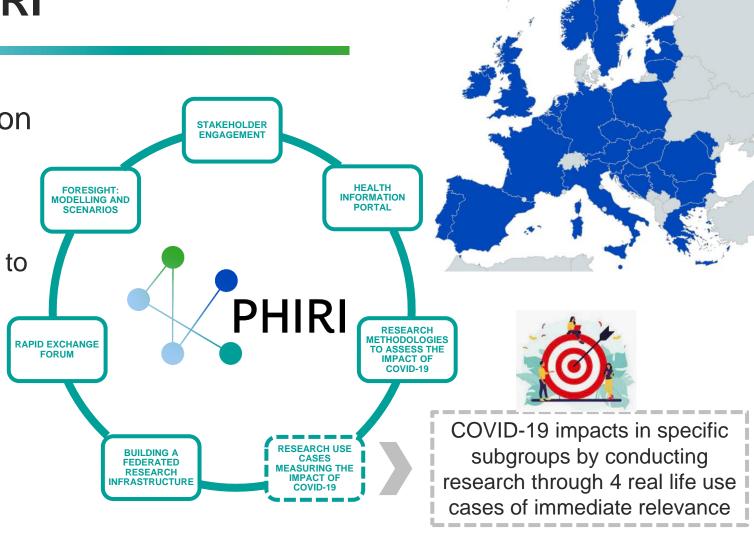


## **Background – PHIRI**

The Population Health Information Research Infrastructure for COVID-19 is:

a European mechanism, that aims to

- facilitate and support data-driven population health research
- and exchange of best practices
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#### **Use Case A**



- Research question:
  - Has the COVID19 pandemic changed existing patterns of non-COVID-19 health care utilisation for (vulnerable) populations within and between countries?
- Heart attack and strokes (Cohort 1)
- Hip and knee replacements (Cohort 2)
- Serious trauma admissions (Cohort 3)
  - **Method:** Compare age-standardised utilization rates for each month of 2020 (and possibly 2021) compared with pre-existing trends during 2017-2019, supplemented by ecological analyses and comparisons using data on infections and hospitalizations from ECDC.

#### **Objectives of the study:**

- ✓ Demonstrate how a broad variety of secondary data (e.g. administrative and survey data) can be pooled and/or reused in a distributed way across Europe
- ✓ Gain insights into the situation of socially (and potentially clinically) vulnerable groups during the COVID-19 pandemic
- ✓ Understand gaps in health system performance during crisis
- ✓ Develop learnings on system resilience and <u>inclusive</u> pandemic preparedness

#### Common data model



|          | Associated entity in ERD | Label (var_label)     | Name (var_concept)                       | Classification/Encoding        | Units       | Format     | Description                               |  |
|----------|--------------------------|-----------------------|--|--------------------------------|-------------|------------|---|--|
| basics   | patient                  | patient_id            | patient identificator                    | private key ciphering function | none        | string     | patient pseudonymized identificator       |  |
|          | patient                  | sex                   | sex                                      |                                |             |            |   |  |
|          | patient                  | age_nm                | age                                      | none                           | years       | integer    | patient's age as of 2019-01-01            |  |
|          | observation period       | period                | [time period]                            | none                           | month       | integer    | natural month                             |  |
| cohort 1 | heart event              | acute_event_heart     | major vascular event - heart attack      | ICD10:I21                      |             |            |   |  |
|          | date heart event         | date_event_heart      | date - heart attack                      | date                           | date_DMY_nr | YYYY-mm-dd |   |  |
|          | stroke event             | acute event stroke    | major vascular event - stroke            | 160-164                        |             |            |   |  |
|          | date stroke event        | date event stroke     | date - stroke                            |                                | date_DMY_nr | YYYY-mm-dd |   |  |
| cohort 2 | procedure                | ttm type cd           | type of treatment                        | types of treatment referred    | none        | integer    | type of treatment received by the patient |  |
|          | procedure                | surgery_elective_hip  | elective surgery, hip joint replacement  | OPCS codes in UK W37-W39       |             |            |   |  |
|          | procedure                | surgery elective knee | elective surgery, knee joint replacement | OPCS codes in UK W40-W42       |             |            |   |  |
| cohort 3 | condition                | acute_event_trauma    | hospital admission for trauma based on   | ICD10: S720, S721, S722, S723, | none        | string     | Based on scientific analysis by New       |  |
|          | Date of event            | date_event            | date of admission                        | date                           | date_DMY_nr | YYYY-mm-dd | date of admission                         |  |
|          | Optional:                |                       |  |                                |             |            |   |  |
| optional | patient                  | educ_cd               | highest completed education level        | quintile or top/bottom         | quintiles   | integer    | patient's highest completed education     |  |
|          | patient                  | socecon_lvl_cd        | socioeconomic level                      | quintile or top/bottom         | quintiles   | integer    | patient's socioeconomic level (quintile)  |  |
|          | patient                  | country_cd            | country (residence)                      | ISO3166                        | none        | string     | patient's country of residence            |  |
|          | patient                  | district_cd           | district (residence)                     | e.g. Eurostat NUTS             |             |            |   |  |
|          | patient                  | country_origin_cd     | country (origin)                         | ISO3166                        | none        | string     | patients' country of origin (country of   |  |

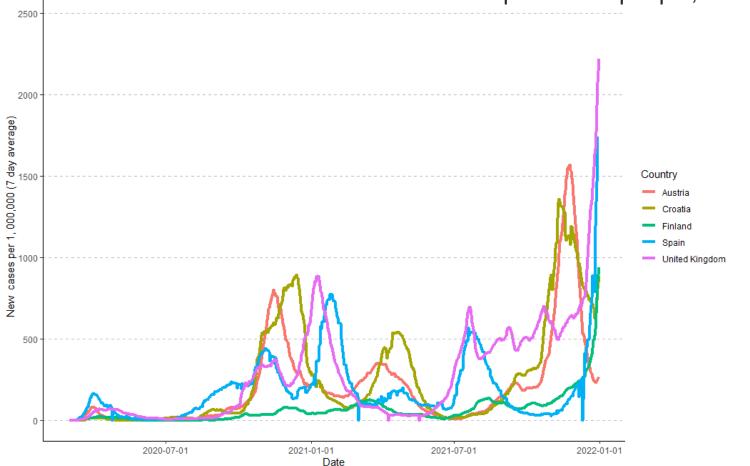




## Setting the Scene:

differing timing of patterns in infection rates between countries BERLIN | 9-12 NOVEMBER 2022





Source: Our World in Data (retrieved on 5th November 2021).





**EUROPEAN PUBLIC** 

CONFERENCE

**HEALTH** 

### **Docker output**



Use Case A: Indirect effects of COVID-19 pandemic on vulnerable populations

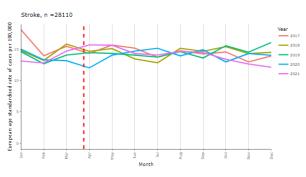
2022-07-05

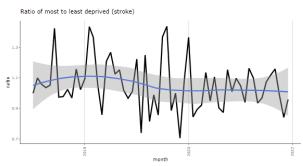
#### Use Case A on Vulnerable Populations

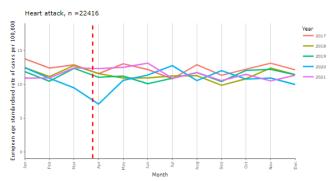
Wide variations in COVID-10 infection and outcomes exist across Europe and within countries. Use Case A explores the indirect impact of the pandemic on health care utilisation in three tracer groups of conditions - heart attack and stroke, hip and knee replacement and severe trauma. By incorporating deprivation measures, further comparisons examining differences across socioecomonic status can be investigated.

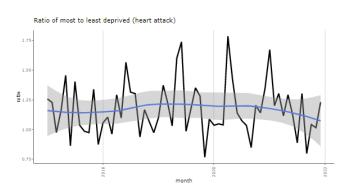
#### LOCAL ANALYSES

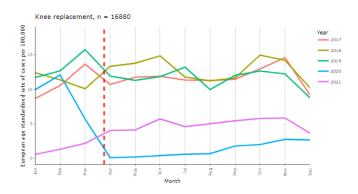
This analyses corresponds with the local part of the analysis (country-specific). Please, provide feedback on your outputs, both data quality analysis (EDA), HTML report and aggregated output to the Use Case A leaders to complete the overall analyses.

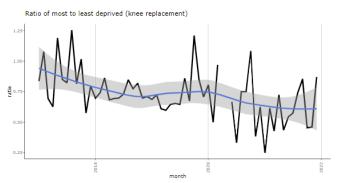










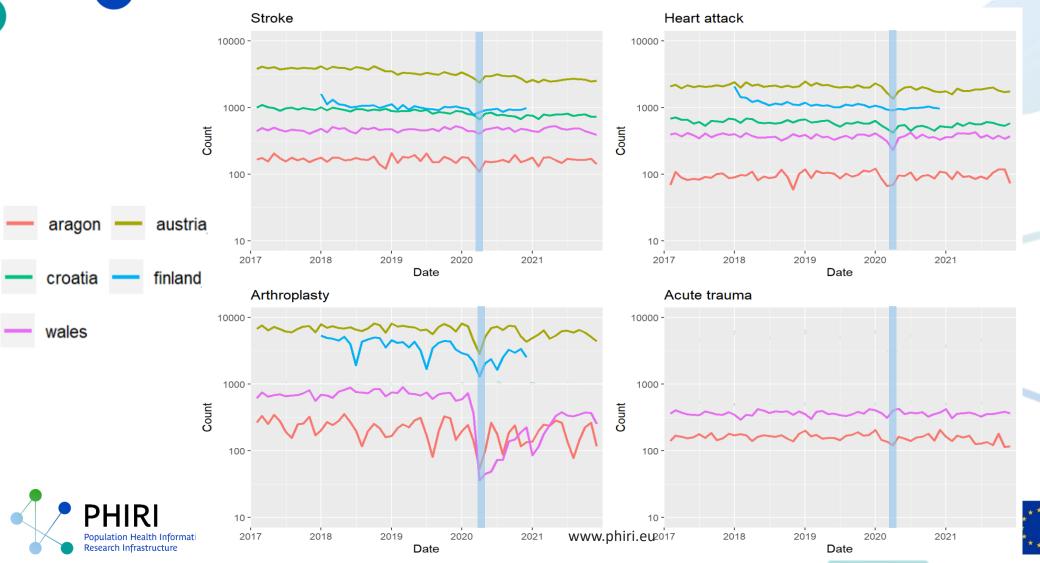






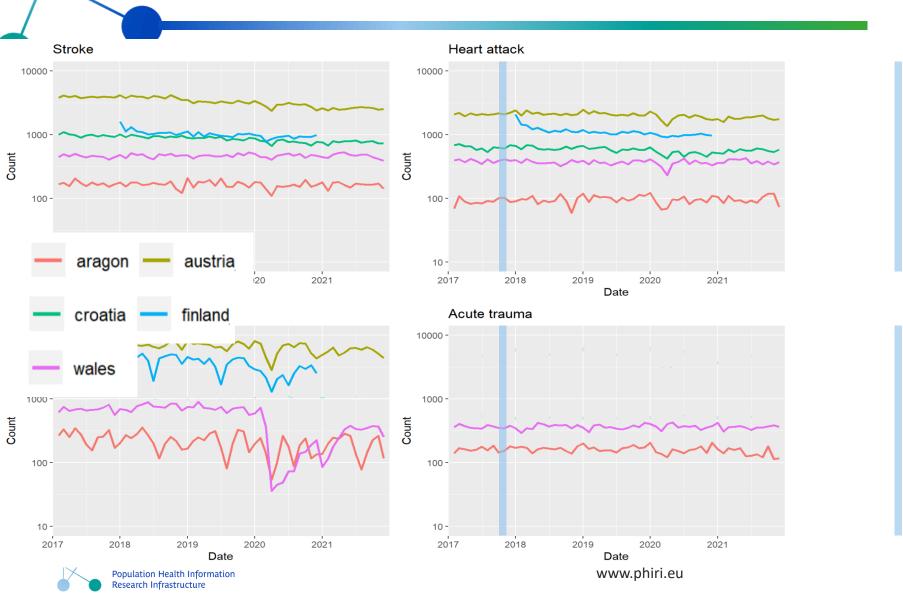
#### Raw event counts





#### Raw event counts

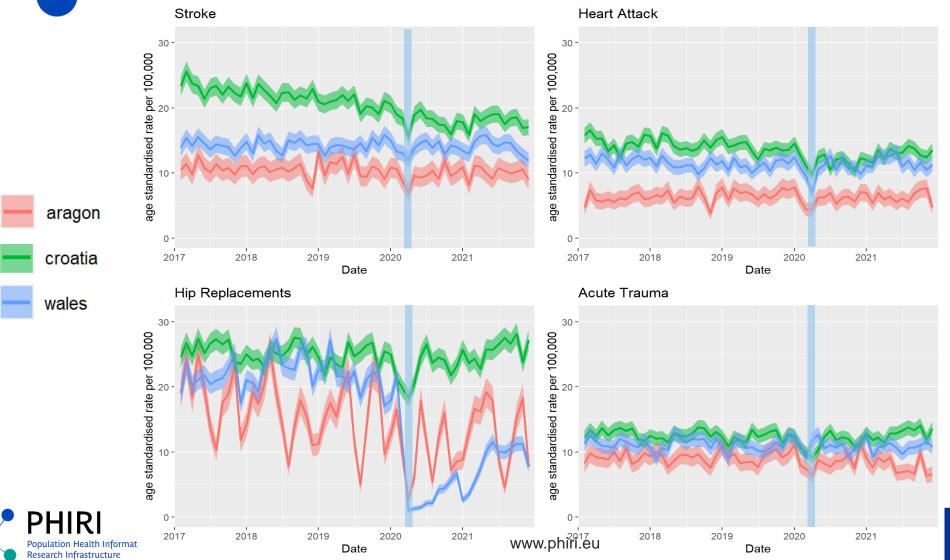




## European age standardises rates:

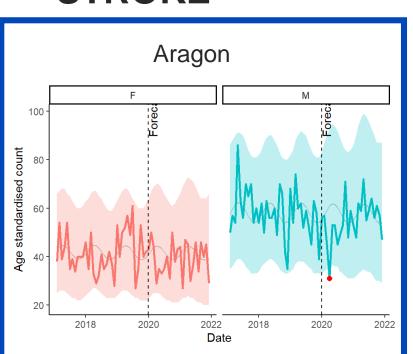
Stroke, Heart attack, Arthroplasty and Trauma



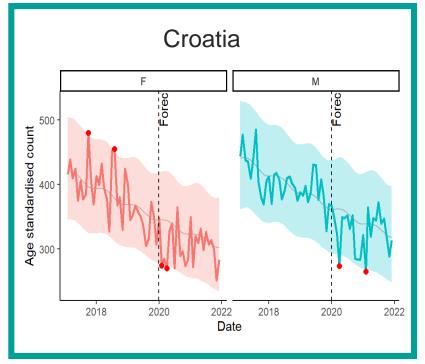


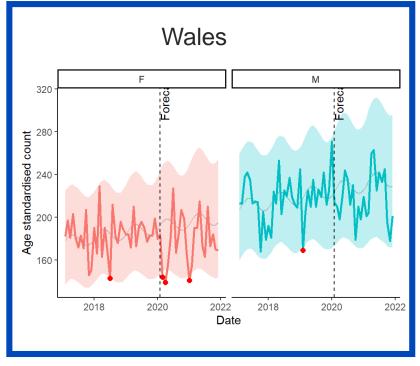


#### **STROKE**















#### **HEART ATTACK**

Aragon

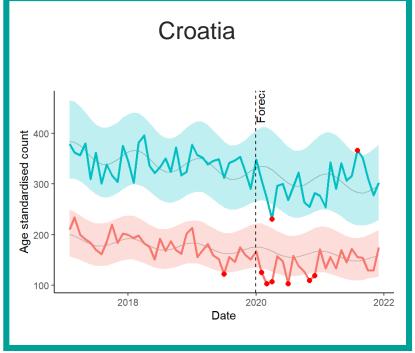
2020

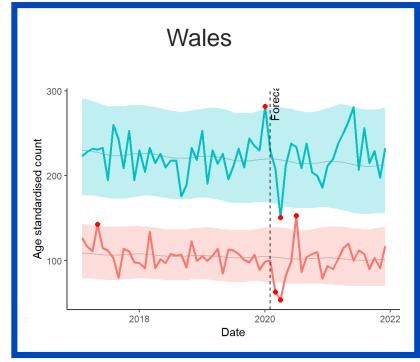
Date



2022









2018

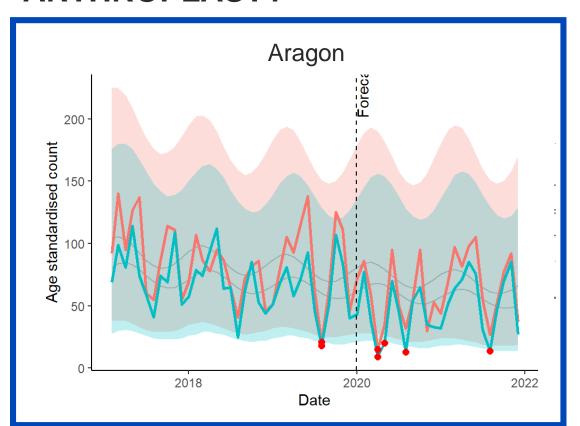
Age standardised count

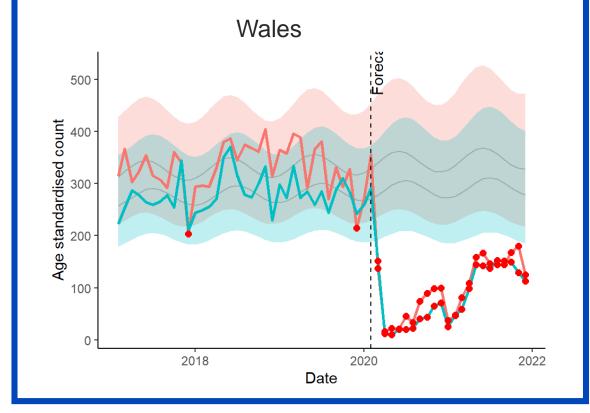




#### **ARTHROPLASTY**





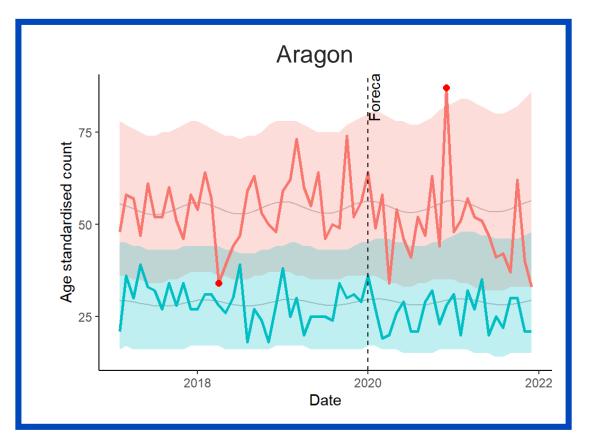


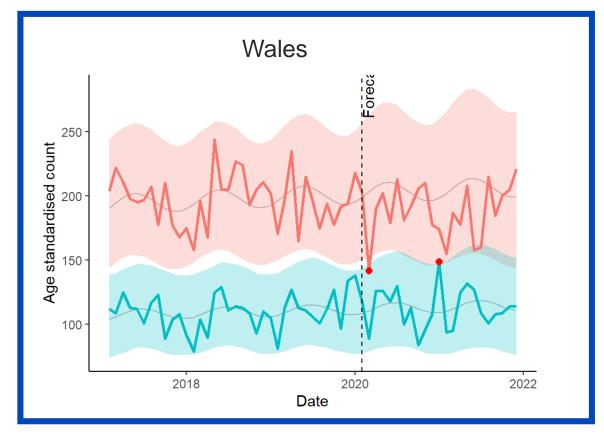




#### **ACUTE TRAUMA**





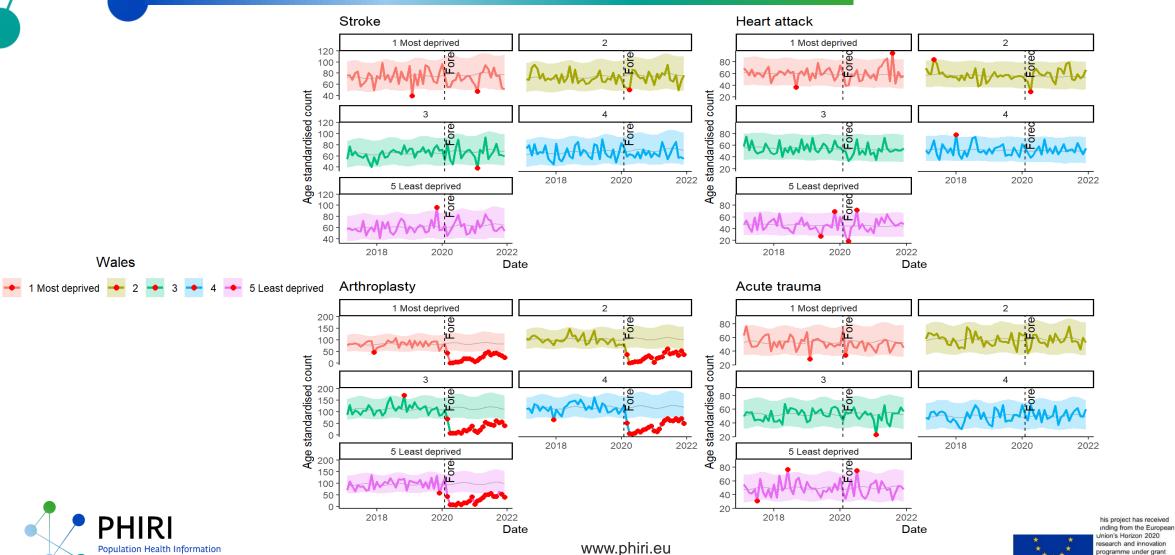








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Research Infrastructure

agreement No 101018317

#### **Further work**



- Unify analysis into one GLM, investigating demographic, geographic and epidemiological factors as contributors to changes in health care utilisation
- Incorporate additional countries into the model as they provide data
- Bring this data back to country specific policies and use this to identify which containment measures are most likely to associate with non-COVID health care utilization







## Thank you for your attention!

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SAIL DATABANK











# Was there any *delay* in breast cancer treatment because of the COVID-19 stringency measures?

Francisco Estupiñán-Romero on behalf of PHIRI Use Case B participants

















## Study design



**Design:** Observational retrospective quasi-experimental pre-post international comparison using a *federated analysis approach* 

**Cohort:** All women over 18 years old with a diagnosis of breast cancer that received hospital treatment (*i.e. surgery, radiotherapy, chemotherapy, hormonal therapy or immunotherapy*) from *Jan-2017 to Dec-2021* in each country/region

Outcome: Time from breast cancer diagnosis to first treatment

Country/region participation (N=5): Aragon (AR, Spain), Wales (WA, United Kingdom), Belgium (BE), Marche (MA, Italy), and Latvia\* (LV)





#### **Methods**



#### - Local analysis:

 Distribution of interval times from breast cancer diagnosis to first hospital treatment by treatment type

#### - Comparative analysis on local aggregated outputs:

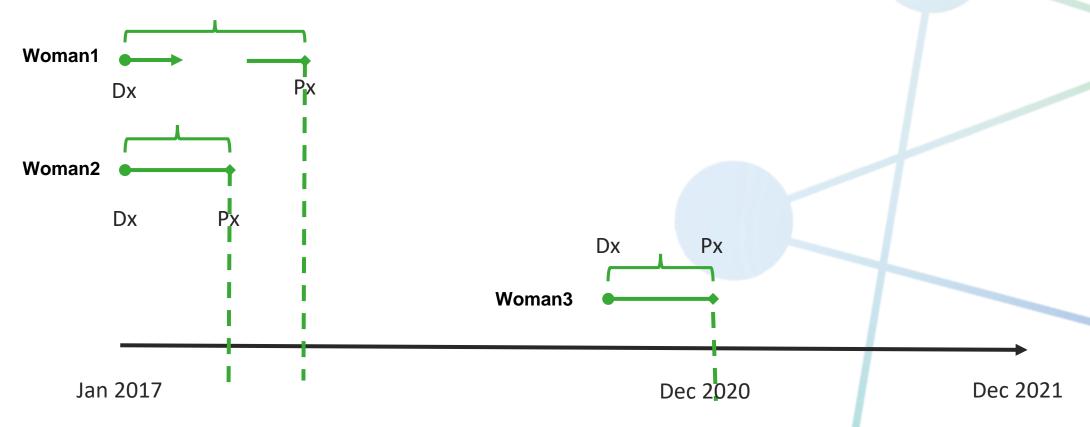
- Comparative analysis of direct standardize rates of hospital treatment
- Structural empirical breakpoint analysis of time-to-first-treatment trend by region/country
- Time series analysis of time-to-first-treatment before March 2020 with forecasting after March 2020
- Segmented regression modelling (*March 2020*) considering contextual factors (*i.e. epidemiological data, health system capacity and public health stringency measures*)
- Sensitivity analyses





# Breast cancer treatment delay: From Diagnosis to FIRST treatment





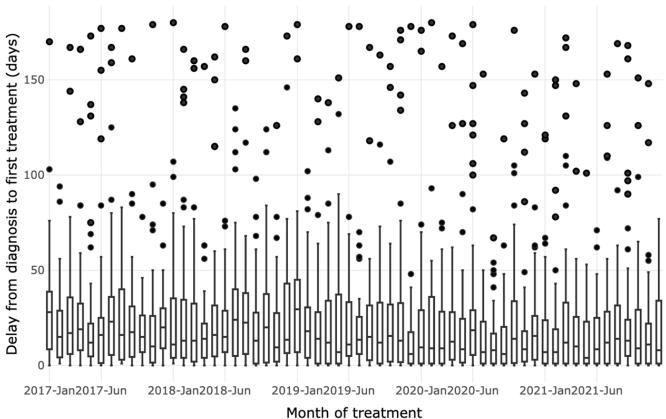


#### Results



#### Distribution of times from diagnosis to treatment for breast cancer patients

Monthly interval distribution from diagnosis to surgery (boxplots)



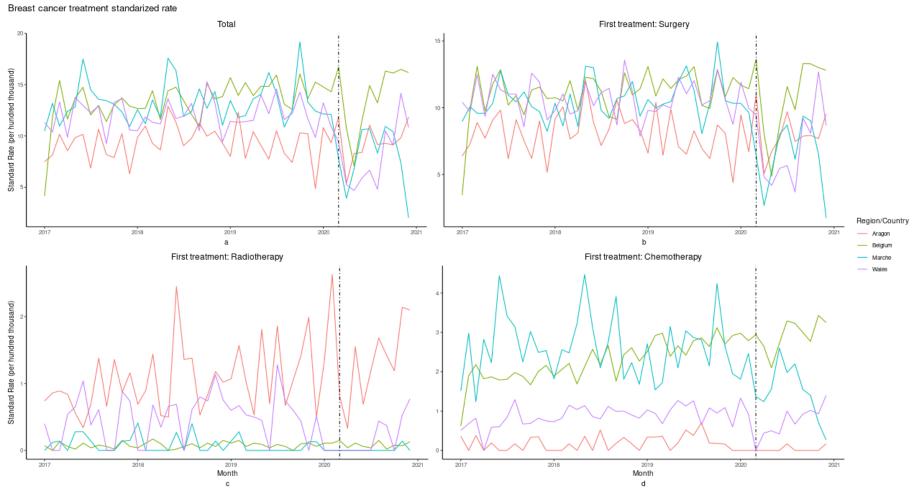
DOI 10.5281/zenodo.6724454





# Direct standardized rates of treatment for breast cancer by treatment type and region/country







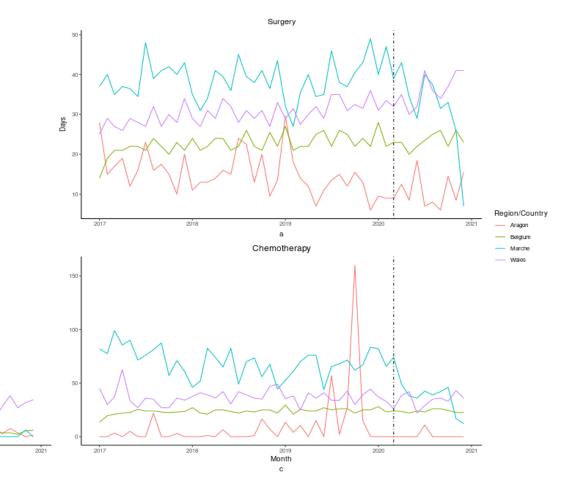


#### Median time-to-first-treatment by region/country (trend)



Median time-to-first treatment

Radiotherapy

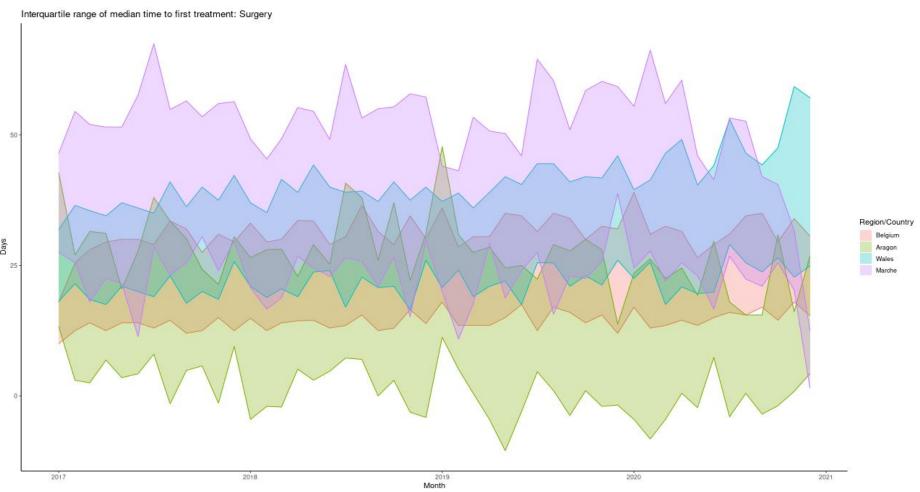






#### IQR time-to-first-treatment (surgery) by region/country



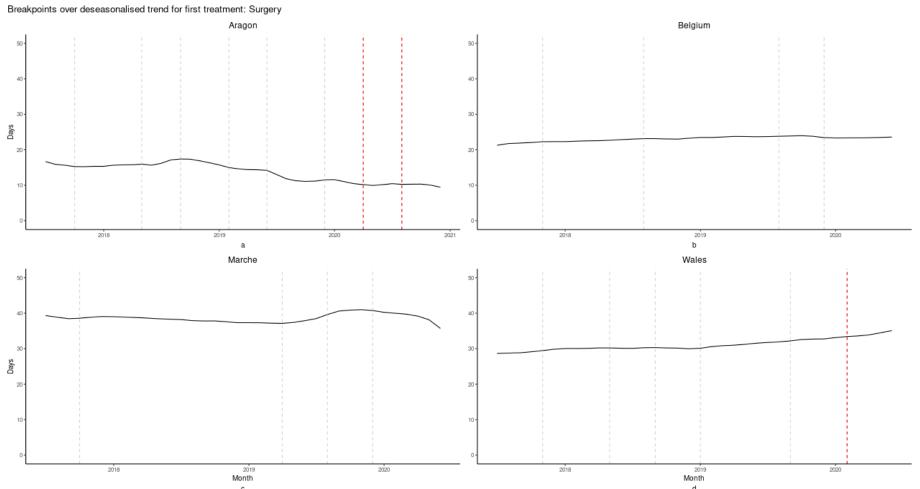






# Structural break point analysis of the destationalised trend of median time-to-first-treatment (surgery)



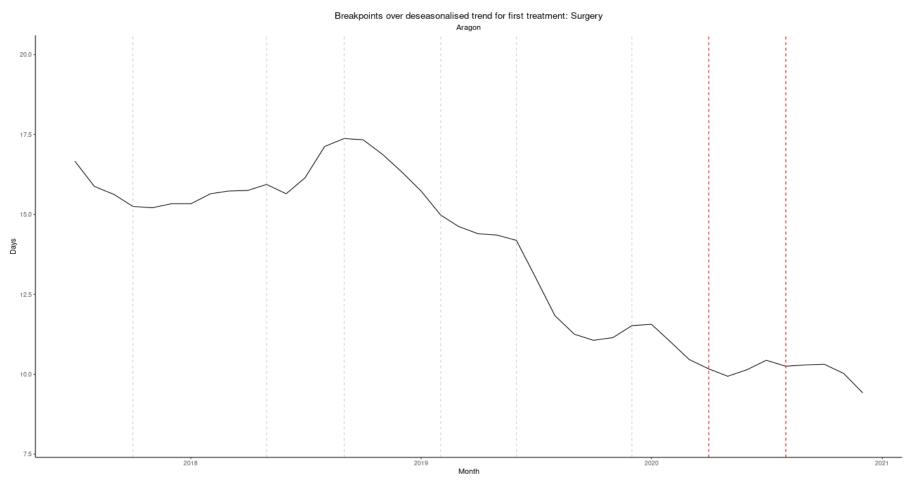






# Structural break point analysis of the destationalised trend of median time-to-first-treatment (surgery)





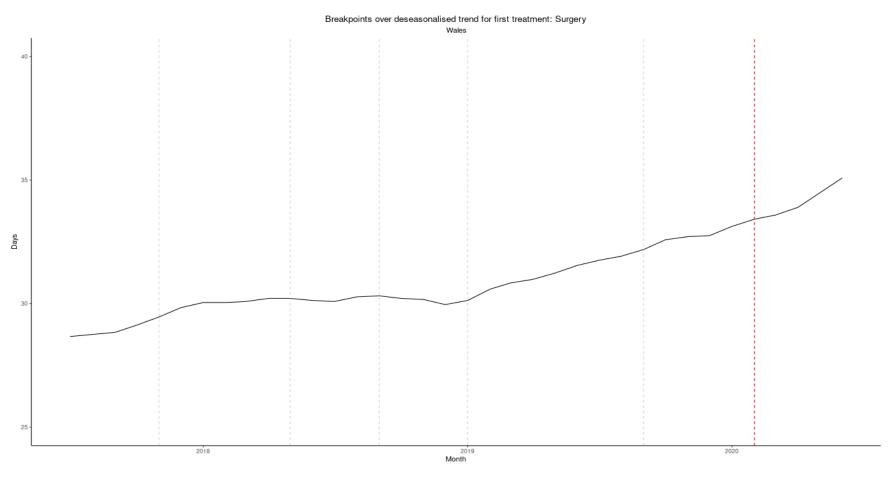






# Structural break point analysis of the destationalised trend of median time-to-first-treatment (surgery)





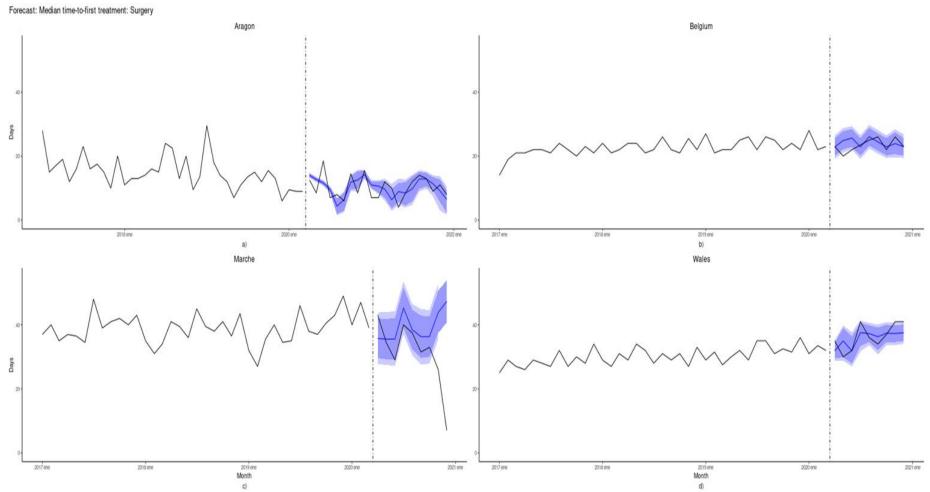






## Time series analysis and forecast from March 2020 onwards

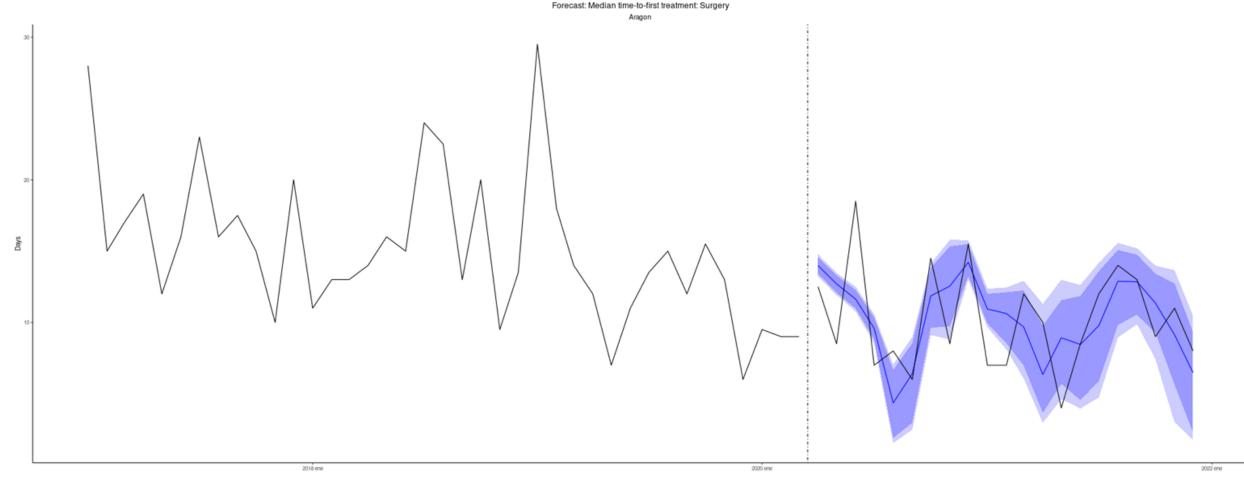






## Time series analysis and forecast from March 2020 onwards





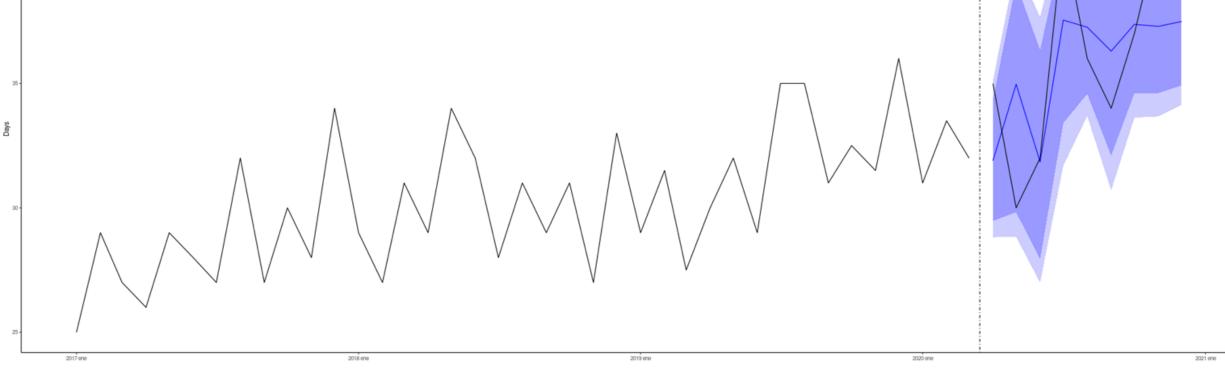




#### Time series analysis and forecast from March 2020 onwards







Forecast: Median time-to-first treatment: Surgery





# Interrupted time series analysis considering contextual factors - March 2020 onwards- (Aragon | All regions)



```
Generalized linear mixed model fit by maximum likelihood (Laplace Approximation) ['glmerMod']
Generalized linear mixed model fit by maximum likelihood (Laplace Approximation) ['glmerMod']
                                                                                                                    Family: Gamma ( log )
 Family: Gamma ( log )
                                                                                                          (1 | i) Formula: median_time_to_surgery ~ Region + (Region | i)
Formula: median_time_to_surgery ~ period + moda_stringency_index + median_hosp_admissions_per_1E5 +
                                                                                                                      Data: data regressionALL
   Data: data_regressionAR
                                                                                                                                BIC logLik deviance df.resid
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                   logLik deviance df.resid
                                                                                                                            1335.3 -627.8 1255.6
                                                                                                                     1285.6
   349.5
            362.1
                    -168.8
                              337.5
                                                                                                                   Scaled residuals:
Scaled residuals:
                                                                                                                                10 Median
             1Q Median
                                    Max
                                                                                                                   -3.5434 -0.4138 -0.0669 0.3832 4.5390
-1.8299 -0.6617 -0.2340 0.5821 3.2664
                                                                                                                   Random effects:
Random effects:
                                                                                                                                          Variance Std.Dev. Corr
                                                                                                                    Groups
                      Variance Std.Dev.
 Groups
                                                                                                                             (Intercept) 0.008399 0.09164
          (Intercept) 0.0000
                              0.0000
                                                                                                                             RegionBelgium 0.009899 0.09949 -1.00
 Residual
                      0.1028 0.3206
                                                                                                                             RegionMarche 0.001599 0.03999 -1.00 1.00
Number of obs: 60, groups: i, 2
                                                                                                                             RegionWales 0.018213 0.13496 -1.00 1.00 1.00
                                                                                                                    Residual
                                                                                                                                          0.048470 0.22016
Fixed effects:
                                                                                                                   Number of obs: 204, groups: i, 2
                                Estimate Std. Error t value Pr(>|z|)
                                                                                                                   Fixed effects:
(Intercept)
                                2.913077 0.090727 32.108
                                                              <2e-16 ***
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period
                               -0.009807
                                           0.003704
                                                     -2.648
                                                              0.0081 **
                                                                                                                   (Intercept)
                                                                                                                                 2.49128
                                                                                                                                            0.16991 14.662 < 2e-16 ***
moda_stringency_index
                               -0.001349
                                           0.002388 -0.565
                                                              0.5722
                                                                                                                   RegionBelgium 0.64561
                                                                                                                                            0.18950 3.407 0.000657 ***
median_hosp_admissions_per_1E5 -0.004280
                                          0.007790 -0.549
                                                                                                                   RegionMarche
                                                                                                                                 1.04655
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                                                                                                                   RegionWales
                                                                                                                                 1.02097
                                                                                                                                            0.25222 4.048 5.17e-05 ***
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' '1
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Correlation of Fixed Effects:
            (Intr) period md st
                                                                                                                   Correlation of Fixed Effects:
            -0.829
period
                                                                                                                              (Intr) RanBla RanMrc
md_strngnc___0.335 -0.631
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mdn h 1E5 0.099 -0.121 -0.456
                                                                                                                   RegionMarch -0.845 0.818
optimizer (Nelder_Mead) convergence code: 0 (OK)
                                                                                                                   RegionWales -0.982 0.954 0.818
boundary (singular) fit: see help('isSingular')
                                                                                                                   optimizer (Nelder_Mead) convergence code: 0 (OK)
                                                                                                                   Model failed to converge with max[grad] = 0.112076 (tol = 0.002, component 1)
```





# Interrupted time series analysis considering contextual factors - March 2020 onwards- (Aragon | All regions)



| Name        | Model    | AIC      | AIC_wt       | BIC      | BIC_wt       | RMSE     | Sigma     | AICc     | AICc_wt | R2_conditional | R2_marginal | ICC       | R2_Nagelkerke |
|-------------|----------|----------|--------------|----------|--------------|----------|-----------|----------|---------|----------------|-------------|-----------|---------------|
| 1 model1ALL | glm      | 1303.884 | 7.338673e-05 | 1320.474 | 9.993639e-01 | 4.809907 | 0.2534666 | NA       | NA      | NA             | NA          | NA        | 0.7314282     |
| 2 model2ALL | glmerMod | 1285.585 | 6.903376e-01 | 1335.357 | 5.860863e-04 | 4.270617 | 0.2200435 | 1288.139 | NA      | 0.8023367      | 0.7877803   | 0.0685913 | NA            |
| 3 model3ALL | glmerMod | 1287.189 | 3.095891e-01 | 1340.279 | 5.002246e-05 | 4.313918 | 0.2198807 | 1290.098 | NA      | 0.8068935      | 0.7931191   | 0.0665813 | NA            |

Model1 ALL: glm(median\_time\_to\_surgery ~ region)

- Model2ALL: glme(median\_time\_to\_surgery ~ region + (region | i))

- Model3ALL: glmer(median\_time\_to\_surgery ~ region + period + (region | i))

i = binary variable considering before/after March 2020

period = months [1, 48]





## Challenges



#### Local analysis:

- Data availability and quality issues in the case of Latvia
- Lack of availability of socioeconomic status for most participants

### - Comparative analysis on local aggregated outputs:

- Some restrictive aggregation decisions on local analysis output may limit the scope of comparative analysis
- Lack of availability or difficulty to access reliable international data on healthcare reorganization apart from qualitative information





### Some lessons learned



- We can observe an acute decrease in treatments just after March 2020 in all country/regions
- There are changes in the destationalised trend of time-to-first-treatment (surgery) in relation with the surge of COVID-19 cases in March 2020 in some of the participating regions/countries
- Those changes are statistically significant in the case of Aragon (Spain) and Wales (UK), although not clinically relevant and seemingly part of a broader trend
- Changes are significantly different depending on the region/country (both in magnitude and direction)
- Changes seemed not to be associated with the epidemiological factors, nor other factors measured monthly at region/country level









# Thank you!

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♥ @PHIRI4EU

in /company/phiri























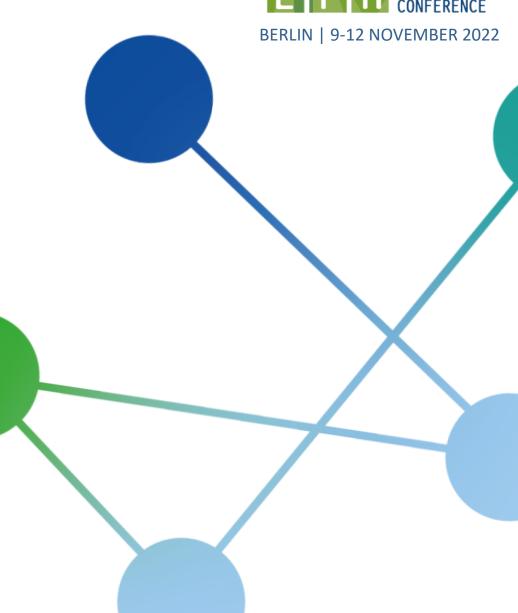
## USE CASE C Indirect effets of the COVID-19 pandemic on perinatal health

Jennifer Zeitlin for the Euro-Peristat Network









## **COVID-19** and perinatal health



- Pregnant women and newborns are vulnerable populations
  - ➤ <u>Direct effects</u> linked to infection by COVID-19
    - Specificities of their immune systems
    - maternal-fetal transmission (fetal development/newborn health)
  - ➤ <u>Indirect effects</u> disruptions associated with the pandemic and its mitigation measures





## Indirect effects: two main pathways



#### **Disrupted maternity care**

- Reluctance to go to the hospital or to ask women to come into hospital → delayed responses to danger signs.
- <u>Use of telemedicine</u> or other changes in care provision
- Transmission reduction measures (maternalbaby separation, caesarean, restricted breastfeeding) → adverse psychosocial and health consequences

# Changes to wellbeing, lifestyle and environment

- Increased stress, anxiety, depression
- Economic hardship,
- Lifestyle changes (activity, diet
- Environment (air pollution)

> Describing these effects is essential for shaping and evaluating pandemic strategies now and for the future





## Indirect effects: a population approach is essential



- Pregnant women and newborns are generally in good health → require large population-based samples.
- Must consider seasonal effects and secular trends
- Comprehensive coverage including disadvantaged populations

> Perinatal outcomes are sensitive to changes in socioeconomic circumstances and social disadvantage





## Unexpected decrease in preterm birth rates?



- Due to fewer indicated preterm births?
- Due to positive effects of the lockdowns (less physical activity, pollution?)

#### Did Lockdowns Lower Premature Births? A New Study Adds Evidence

Dutch researchers say the "impact was real," adding to hopes that doctors will learn more about factors contributing to preterm birth.





A preterm newborn was wheeled to meet his mother in a hospital in Istanbul. A large study of babies born in the Netherlands links the lockdowns with fewer preterm births. Chris Mcgrath/Getty Images



## Confinement : les naissances prématurées en baisse

Le Covid-19 aura au moins eu une conséquence heureuse. Le nombre de naissances prématurées est en sensible baisse dans plusieurs départements. Le calme des deux confinements successifs s'avère très positif pour les femmes enceintes et leurs enfants.









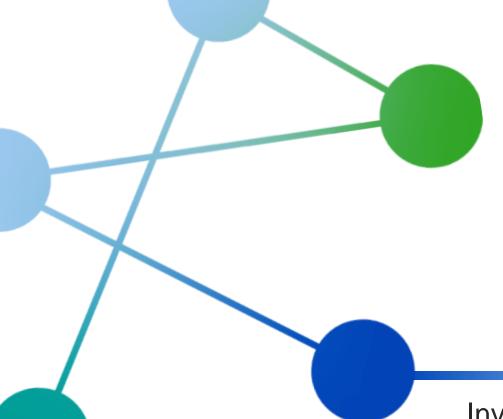


Société

Corona Baby: jusqu'à 80% de bébés prématurés en moins, le mystère du confinement 5









## **Objective**

Investigate the indirect impact of the COVID-19 pandemic in 2020 on perinatal health in Europe using population-based national data





## Key perinatal health indicators



#### Stillbirth (baby born without signs of life)

- 3-4 per 1000 births (15-18,000 babies per year in Europe)
- High health and psychological burden for parents, costs for families and society

#### Preterm birth (birth before 37 weeks of gestation)

- Affects about 350,000 births per year in Europe, few effective prevention strategies
- Principal cause of infant death
- Long-term neurodevelopment impairment and other health problems among survivors
- Consequences are lifelong and perpetuate health inequalities





## The EURO-PERISTAT network



- Aim: to monitor and evaluate perinatal health in Europe based on valid and reliable indicators
- Health Monitoring Programme in 2000
- Routine national statistics
- Data collected using a common protocol
- 3 European reports, scientific publications
- 31 participating countries





www.europeristat.com



Core indicators to the health and care of pregnant women and babies in Europe in 2015







## Methods: Common data model



- Based on the Euro-Peristat indicators and a consensus process
- 22 variables to produce the core indicators and for the PHIRI use case
- Health indicators collected by year and by month
- Population: all live births and stillbirths with a gestational age ≥
   22 weeks of gestation
- Study period: 2015 to 2020





## **Methods: Participating countries**



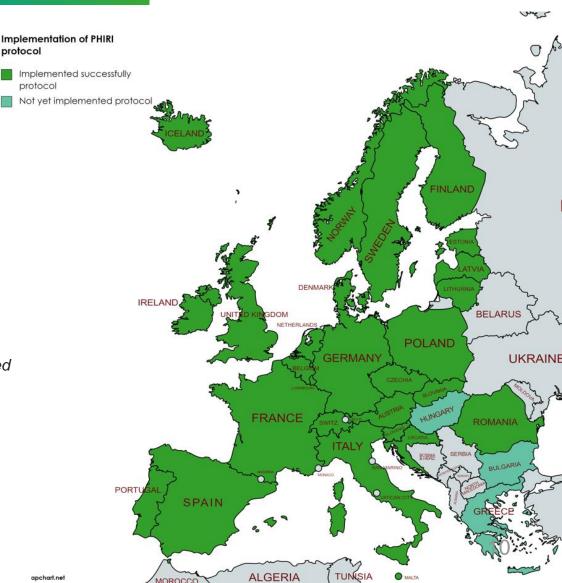
#### 28 countries

Austria, Belgium, Croatia, Cyprus, Czech Republic, Denmark, Estonia, Finland, France, Germany, Iceland, Ireland, Italy, Latvia, Lithuania, Luxembourg, Malta, Netherlands, Norway, Poland, Portugal, Slovakia, Slovenia, Spain, Sweden, Switzerland, Romania\*, UK (MBRRACE, and UK nations constituents: England and Wales, Northern Ireland, Scotland, Wales)

\* Recently added

protocol

 >27 million total births, 1.9M preterm births, 15K stillbirths, 10K neonatal deaths



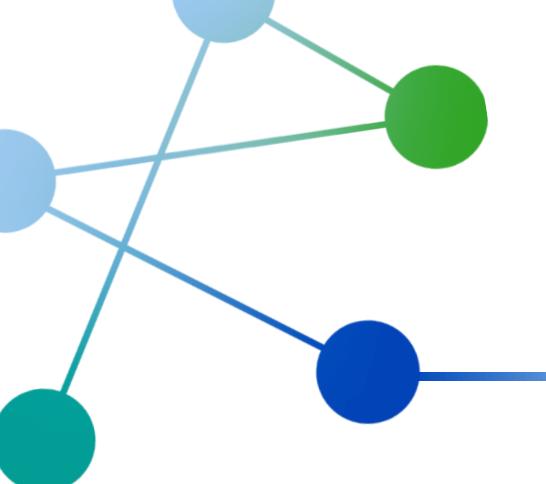
## **Methods: Analysis strategy**



- Describe perinatal health trends in 5 years preceding the pandemic
- What "should" have happened as opposed to what "did" happen.
- Estimate relative risk of observed to expected outcome
- Linear models comparing 2020 and the period March-September
- AutoRegressive Integrated Moving Average (ARIMA) models in each country to confirm linear trends and do more detailed country analyses
- Analyse country-level estimates using random-effects metaanalysis (DerSimonian and Laird method)







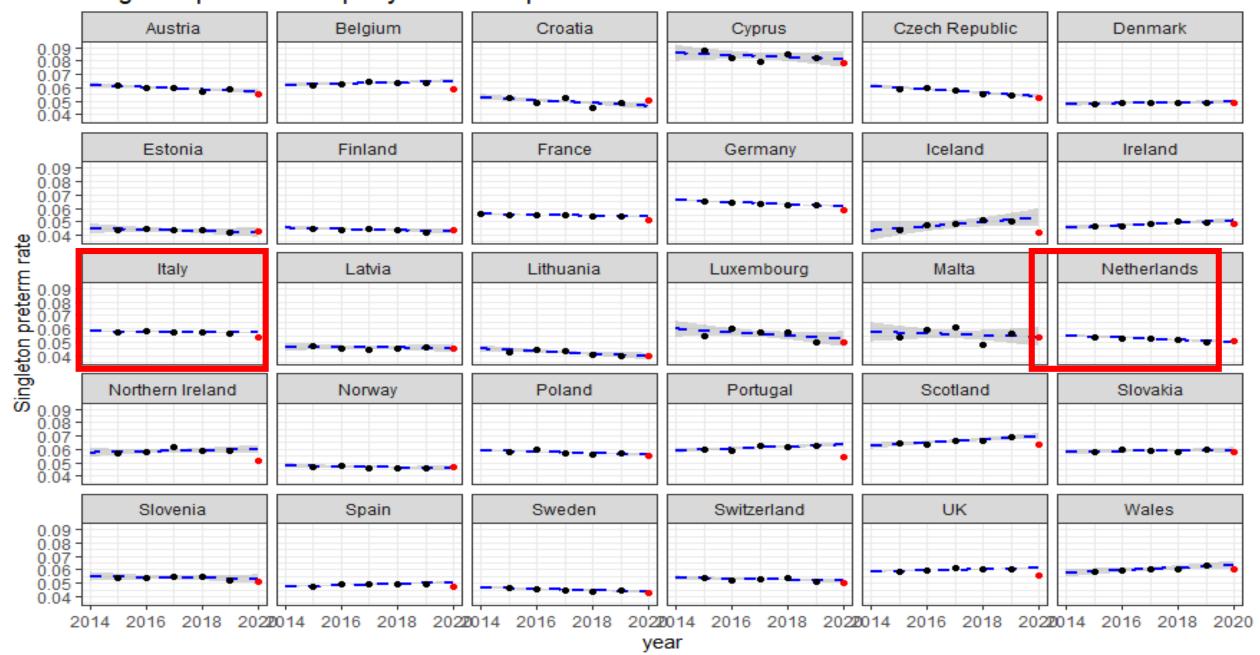


## Results





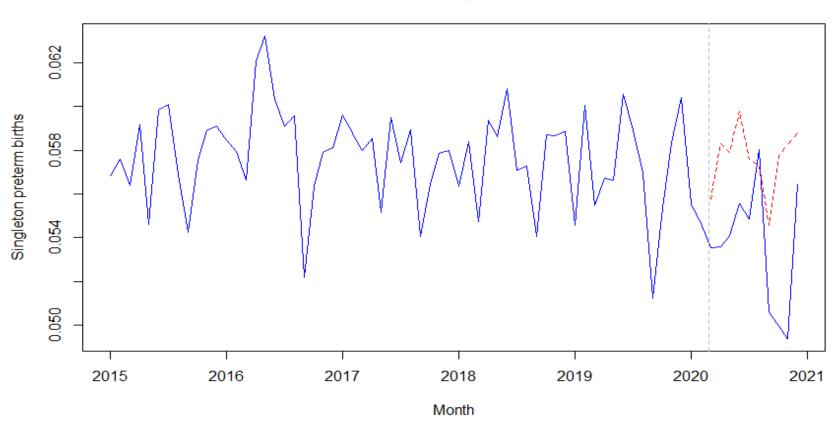
#### Singleton preterm rate per year in Europe



## Time series (ARIMA\*) models using monthly data







\* Autoregressive integrated moving average

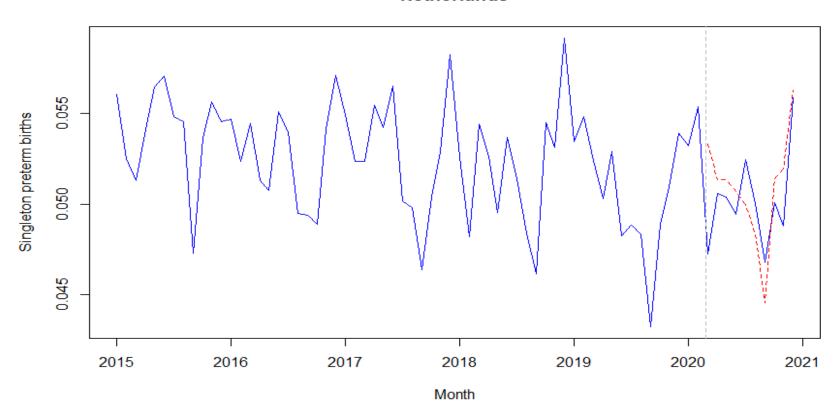




## Time series (ARIMA\*) models using monthly data



#### Netherlands



\* Autoregressive integrated moving average





## Singleton preterm birth rate

#### **Pooled estimate**

RR=0.96 (0.96 to 0.98) = 4% decrease in preterm birth

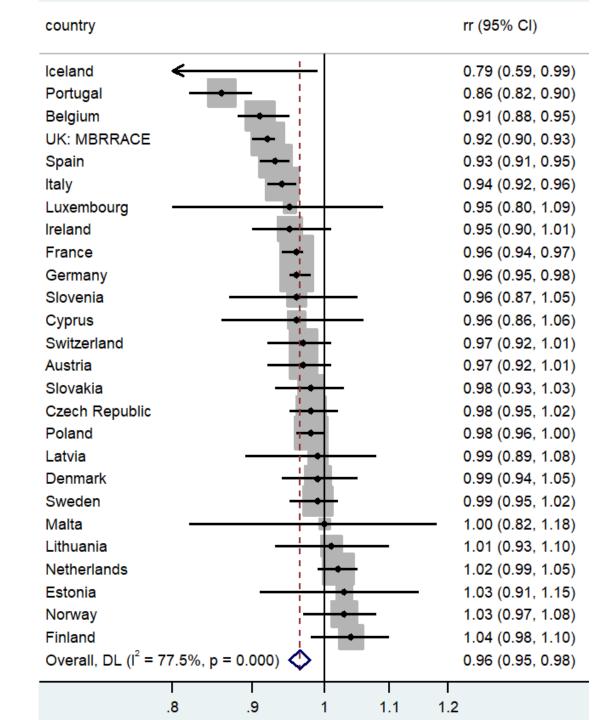
#### **High heterogeneity**

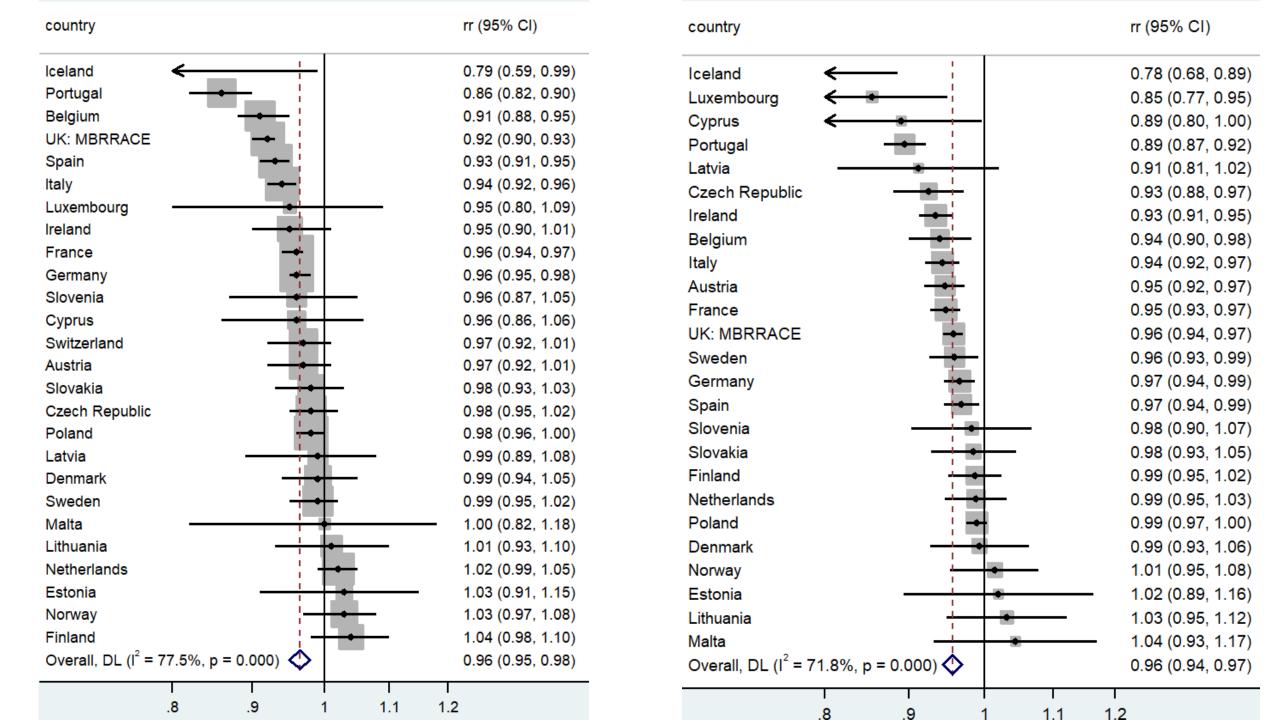
 $I^2 = 77.5\%$  (proportion of total variation in effect estimate due to between-study heterogeneity)

Range of effects = 10% decrease in preterm birth to moderate increase of 3 to 4%.

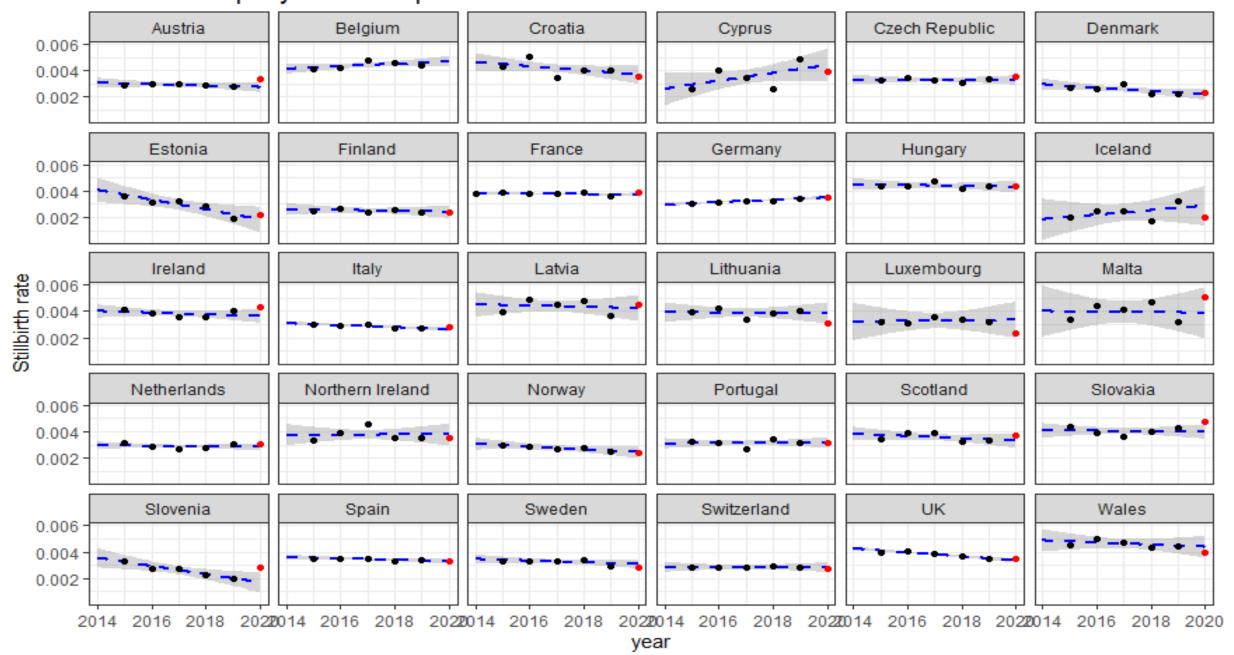
Countries with stronger effects: Portugal—Belgium — - UK - Spain — Italy — France

Countries with no effects: Nordic and Baltic countries, Netherlands





#### Stillbirth rate per year in Europe



## Stillbirth rate

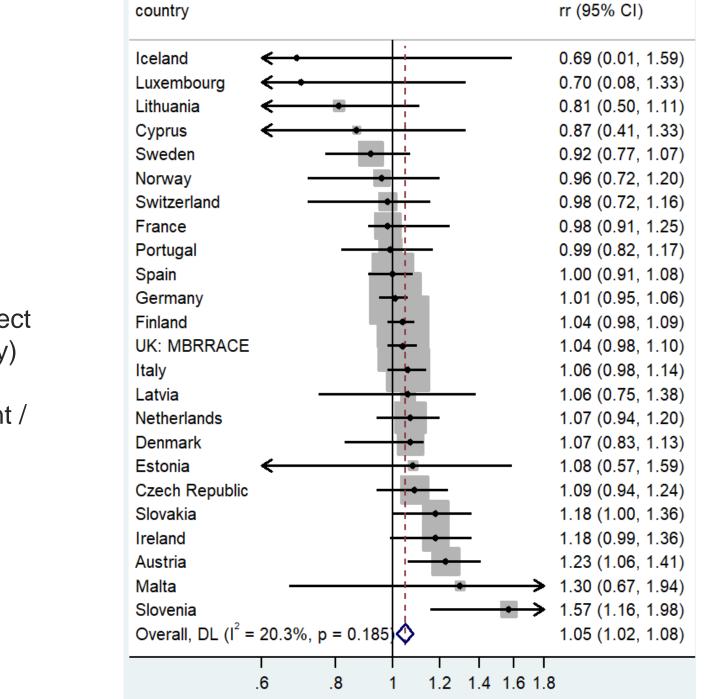
#### **Estimate of pooled effect**

RR=1.05 (1.02 to 1.08) = 5% increase in stillbirth

#### Lower heterogeneity

 $I^2 = 20.3\%$  (proportion of total variation in effect estimate due to between-study heterogeneity)

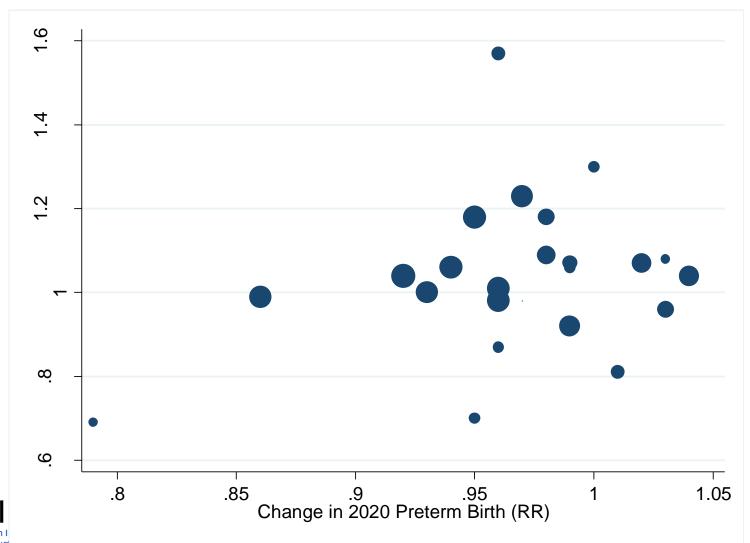
Range of effects = No decreases significant / Austria higher stillbirth rates





# Association between changes in stillbirth and preterm birth









## **Summary and discussion**



- COVID-19 in 2020 did not impact all countries in the same way.
- A puzzling decline in preterm birth in some countries
  - PTB meta-analyses— OR: 0.91 (0.84-0.99), 12 studies 01/2021
  - OR: 0.94 (0.91–0.98), 28 studies 05/2021;
  - OR 0.99 (0.95–1.03) 8 population studies.
- Increase in stillbirth in many countries, not more pronounced in countries with decline in preterm birth rate
- Our study confirms this high variability in indirect effects of the epidemic on pregnant women and babies in European countries
  - some policies more protective of pregnant women and newborns than other?
  - Women's experiences of the pandemic different?
  - Interaction with other health system factors?

Chmielewska et al. Lancet Global Health (2021), Yang et al. Acta Obstetr. (2022)





## Conclusions: population birth data



Shows the feasibility and value of bringing population birth data together in Europe



DOI: 10.1111/1471-0528.16946

www.bjog.org

**Commentary** 

## Population birth data and pandemic readiness in Europe

#### Euro-Peristat Research Network\*

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#### **EURO-PERISTAT COUNTRY TEAMS**



Austria

Belgium



Croatia



Cyprus



Czech Rep.



BERLIN | 9-12 NOVEMBER 2022

Denmark



Hungary



Luxembourg



Romania



Catania



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^-----

Thank you for the input and effort

of the Euro-Peristat data hubs

Support from the team at WP7
Other use cases in WP6









Greece



Lithuania



Portugal



Switzerland



UK



https://www.europeristat.com/index.php/our-network/country-teams.html









# Monitoring COVID-19 related changes in population mental health

<u>C Rodríguez-Blázquez</u><sup>1</sup>, S Aldridge<sup>2</sup>, E Bernal-Delgado<sup>3</sup>, L Dolanski-Aghamanoukjan<sup>4</sup>, F Estupiñán-Romero<sup>3</sup>, C Garriga<sup>1</sup>, M Gissler<sup>5,6</sup>, RA Lyons<sup>2</sup>, S Sagerschnig<sup>4</sup>, H Tolonen<sup>5</sup>

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 Austrian National Public Health Institute, Vienna, Austria.
 THL Finnish Institute for Health and Welfare. Helsinki. Finland.
 Karolinska Institute. Stockholm. Sweden.



## **Background**



The COVID-19 pandemic, and its consequences in terms of control measures and restrictions to normal life, has impacted the population mental health.

Table 3: Proportion of respondents reporting having negative feelings by age and gender, EU27 (%)

|       |             |       | Summer 2020 |           | Spring 2021 |        |           |  |
|-------|-------------|-------|-------------|-----------|-------------|--------|-----------|--|
|       |             | Tense | Lonely      | Depressed | Tense       | Lonely | Depressed |  |
| Men   | 18–34 years | 34    | 25          | 21        | 46          | 35     | 34        |  |
|       | 35–49 years | 30    | 21          | 19        | 41          | 31     | 32        |  |
|       | 50+ years   | 22    | 18          | 15        | 28          | 26     | 23        |  |
| Women | 18–34 years | 45    | 30          | 28        | 52          | 38     | 40        |  |
|       | 35–49 years | 38    | 22          | 27        | 49          | 34     | 39        |  |
|       | 50+ years   | 24    | 18          | 17        | 35          | 30     | 29        |  |

Notes: Green = lowest value, red = highest value. All differences between the two time periods are statistically significant. Any discrepancies between the figures in the text and table are due to rounding.

Source: Living, working and COVID-19 e-survey data. Mental health and trust decline across EU as pandemic enters another year. EuroFound, 2021.





# Multidisciplinary research priorities for the COVID-19 pandemic: a call for action for mental health science



Emily A Holmes\*, Rory C O'Connor\*, V Hugh Perry, Irene Tracey, Simon Wessely, Louise Arseneault, Clive Ballard, Helen Christensen, Roxane Cohen Silver, Ian Everall, Tamsin Ford, Ann John, Thomas Kabir, Kate King, Ira Madan, Susan Michie, Andrew K Przybylski, Roz Shafran, Angela Sweeney, Carol M Worthman, Lucy Yardley, Katherine Cowan, Claire Cope, Matthew Hotopft, Ed Bullmore†

Lancet Psychiatry 2020; 7: 547–60 https://doi.org/10.1016/S2215-0366(20)30168-1

Bidirectional associations between COVID-19 and psychiatric disorder: retrospective cohort studies of 62354 COVID-19 cases in the USA

Maxime Taquet, Sierra Luciano, John R Geddes, Paul J Harrison

Lancet Psychiatry 2021; 8: 130-40 https://doi.org/10.1016/ S2215-0366(20)30462-4

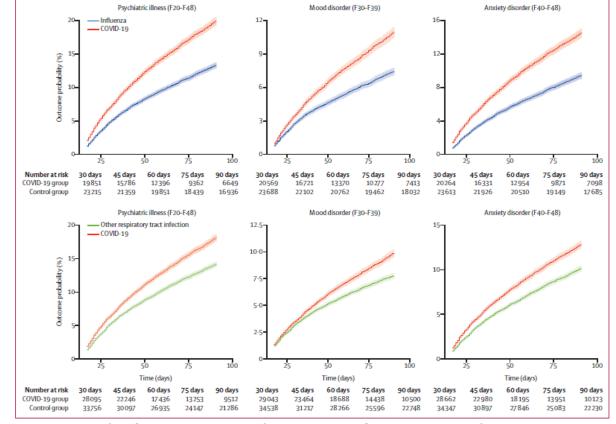




Figure 2: Kaplan-Meier curves for any (first or recurrent) psychiatric diagnoses after COVID-19 compared with influenza and other respiratory tract infections

Curves for the other control health events are presented in the appendix (p 29). Shaded areas represent 95% Cls. The number of subjects within each cohort corresponds to all those who did not have
the outcome before the follow-up period.

# Use Case D: COVID-19 Related changes in population mental health



#### **Objective:**

To measure changes in population mental health associated with the COVID-19 pandemic in several European countries







# Use Case D: COVID-19 related changes in population mental health

Objective: To measure changes in population mental health associated with the COVID-19 pandemic in several European countries





## Methods



#### Two layer questionnaire:

- 1. Aims to get a better sense on the actual access to data and data hubs' capabilities
- 2. A second round survey, specific of each use case, to start off the process of harmonising data throughout a common data model







## **Research Question**



# Has the mental health status (depression/anxiety) of the general population changed during the COVID-19 pandemic?

This RQ was addressed using electronic health records (EHR):

• Indicators such as prescription of antidepressants and anxiolytics, visits to primary care or specialist care with an episode of depression/anxiety, etc.





## Common data model according to the research question



|   |                     |                                   |  |                                |           | ,       |   |  |  |  |
|---|---------------------|-----------------------------------|--|--------------------------------|-----------|---------|---|--|--|--|
| Data model entity                       | Variable Variable   |                                   |  |                                |           |         |   |  |  |  |
| Associated entity in ER                 | D Label (var_label) | Name (var_concept)                | Level<br>(required/recommended<br>/optional) | Classification/Encoding        | Units     | Format  | Description                                       |  |  |  |
| patient                                 | patient_id          | patient identificator             | required                                     | private key ciphering function | none      | string  | patient pseudonymized identificator               |  |  |  |
| patient                                 | sex                 | patient's sex                     | required                                     |                                | none      | integer | patient's sex                                     |  |  |  |
| patient                                 | age_nm              | age                               | required                                     | none                           | years     | integer | patient's age at the moment                       |  |  |  |
| patient                                 | socecon_lvl_cd      | socioeconomic level               | optional                                     | quintile                       | quintiles | integer | patient's socioeconomic level (quintile)          |  |  |  |
| patient                                 | country_cd          | country (residence)               | required                                     | ISO3166                        | none      | string  | patient's country of residence                    |  |  |  |
| patient                                 | country_origin_cd   | country (origin)                  | recommended                                  | ISO3166                        | none      | string  | patient's country of origin                       |  |  |  |
| patient                                 | diagnosis           | diagnosis                         | required                                     | icd-10/icd-10_mc/icd-9-mc/SN0  | none      | string  | patient diagnosis                                 |  |  |  |
| date                                    | dx_date             | date of diagnosis                 | required                                     | dd-mm-yyyy                     | date      | integer | date of diagnosis                                 |  |  |  |
| prescription                            | drug                | drug                              | recommended                                  | ATC                            | none      | string  | patient's prescription                            |  |  |  |
| date                                    | drug_date           | date of prescription              | recommended                                  | dd-mm-yyyy                     | date      | integer | date of prescription                              |  |  |  |
| visit (contact w<br>healthcare service) | prim_visit          | number of visits to primary care  | recommended                                  | ICPC                           | visits    | integer | number of visits to primary care                  |  |  |  |
| visit (contact w<br>healthcare service) | hosp_visit          | number of visits to hospital unit | recommended                                  | ICPC                           | visits    | integer | number of visits to hospital unit                 |  |  |  |
| visit (contact w<br>healthcare service) | emer_visit          | number of visits to emergency     | recommended                                  | ICPC                           | visits    | integer | number of visits to emergency or unplanned visits |  |  |  |
| observation period                      | visit_date          | date of visit                     | recommended                                  | dd-mm-yyyy                     | date      | integer | date of visit                                     |  |  |  |



Open Access

# PHIRI - WP6 - Use Case D scripts for local analyses (R Markdown)

(b) Sarah Aldridge; (b) Javier González-Galindo; (b) Francisco Estupiñán-Romero; (b) Cesar Garriga

Contact person(s)

(b) Carmen Rodríguez-Blazquez

#### Researcher(s)

(b) Enrique Bernal-Delgado; (b) Maria João Forjaz; (b) Hannah Tolonen; (b) Mike Gissler

The PHIRI Federated Research Infrastructure (FRI) is supported by a containerized reproducible solution for data analysis to be deployed on-premises by each participant partner (a.k.a PHIRI-app).

This solution is based on the identification of the relevant data sources for each cases study (including the demonstration pilot), the development of the common data models and the **analytical pipelines**, and enables the FAIR reporting of the rapid cycle outputs.

The R Markdown script is provided, integrated within the PHIRI-app, for PHIRI Use Case D local analyses.

Here, the R Markdown script is provided with:

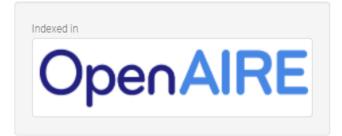
- a) synthetic dataset build following the specifications from the Use Case D Common Data Model,
- b) instructions on where to find the synthetic dataset within the Use Case D Common Data Model description (HTML), and
- c) an HTML interactive report produced by performing the analyses proposed within the R Markdown using the synthetic dataset above.

These elements are provided to facilitate collaboration on testing and improving the Use Case D analytical pipeline within the scope of PHIRI WP6.

If you wish to contribute to the development of the PHIRI - Use Case D analysis, please contact the WP6 Coordinator through the PHIRI website.

The script (software) is offered "as-is", without warranty, and disclaiming liability for damages resulting from using it. Software is released under the CC-BY-4.0 licence, which gives you permission to use the content for almost any purpose (but does not grant you any trademark permissions), so long as you note the license and give credit.

Aldridge et al. (2022). PHIRI - WP6 - Use Case D scripts for local analyses (R Markdown) (1.1.0). Zenodo. https://doi.org/10.5281/zenodo.6377112



#### Publication date:

March 22, 2022

#### DOI:

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#### Grants:

#### European Commission:

 PHIRI - Population Health Information Research Infrastructure (101018317)

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#### Versions

Version 1.1.0

10.5281/zenodo.6377112

Mar 22, 2022

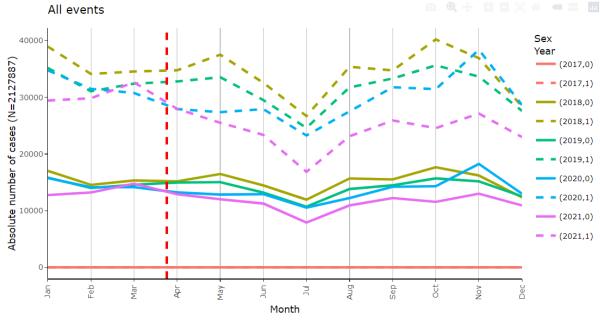
## Results by country

#### Results from 6 datahubs:

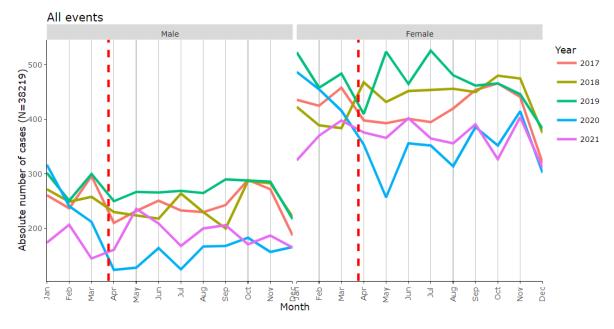
- Aragon (Spain)
- Austria
- Croatia
- Finland
- Romania
- Wales (UK)



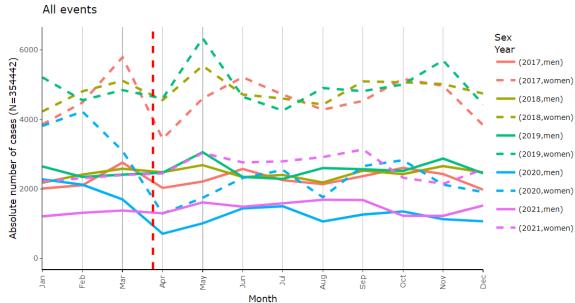
#### **Finland**



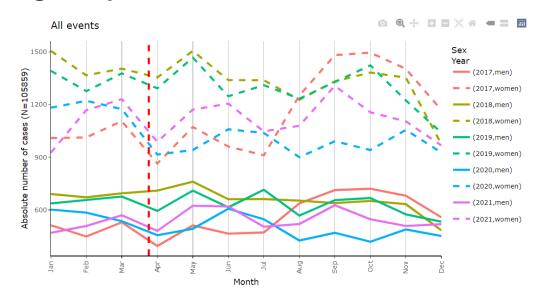
#### Wales (UK)



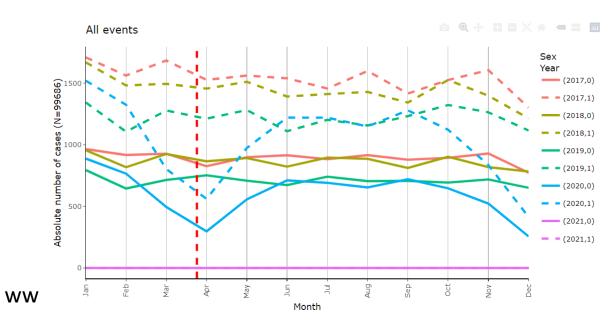
#### Romania



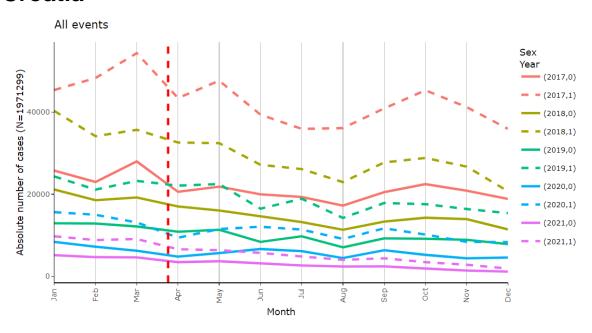
#### Aragon, Spain



#### **Austria**



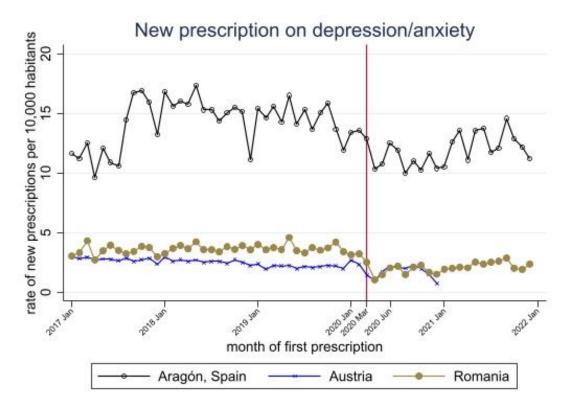
#### Croatia



## Results: cross-country comparison



Preliminary analysis of new prescriptions







## **Conclusions**



- Results showed a decrease in diagnoses of depression and anxiety and drug prescriptions in 2020 in participant countries compared with previous years.
- EHR for the secondary use can be retrieved in a common way across Europe to analyze and compare the impact of COVID-19 in population mental health in European countries.
- The use case D facilitates research by making scalable, reproducible methods available within PHIRI.
- Lessons learned during this exercise could be used to overcome difficulties and allow for cross-country comparisons.







# Thank you for your attention!

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