

Supporting Information

Predictions of COVID-19 dynamics in the UK: short-term forecasting and analysis of potential exit strategies

Matt J. Keeling^{1*}, Edward M. Hill^{1‡}, Erin E. Gorsich¹, Bridget Penman¹, Glen Guyver-Fletcher^{1,2}, Alex Holmes^{1,3}, Trystan Leng^{1,3}, Hector McKimm^{1,4}, Massimiliano Tamborrino^{1,4}, Louise Dyson^{1‡}, Michael J. Tildesley^{1‡}.

1 The Zeeman Institute for Systems Biology & Infectious Disease Epidemiology Research, School of Life Sciences and Mathematics Institute, University of Warwick, Coventry, United Kingdom.

2 Midlands Integrative Biosciences Training Partnership, School of Life Sciences, University of Warwick, Coventry, United Kingdom.

3 Mathematics for Real World Systems Centre for Doctoral Training, Mathematics Institute, University of Warwick, Coventry, United Kingdom.

4 Department of Statistics, University of Warwick, Coventry, United Kingdom.

‡These authors contributed equally to this work.

* M.J.Keeling@warwick.ac.uk

Table of Contents

1	Model description	2
2	Public health measurable quantities	5
3	Key inferred parameters	7
4	Modelling social distancing	8
5	QALY losses	9
6	Additional tables	10
7	Additional figures	11

1 Model description

We present here the system of equations that account for both household saturation of transmission and household quarantining. As described in the main text, individuals may be susceptible (S), exposed (E), with detectable infection (D), or undetectable infection (asymptomatic, U). Undetectable infections are assumed to transmit infection at a reduced rate given by τ . We let superscripts denote the first infection in a household (F), a subsequent infection from a detectable/symptomatic household member (SD) and a subsequent infection from an asymptomatic household member (SU). A fraction (H) of the first detected case in a household is quarantined (QF), as are all their subsequent household infections (QS).

Model equations

The full equations are given by

$$\begin{aligned}
\frac{dS_a}{dt} &= -(\lambda_a^F + \lambda_a^{SD} + \lambda_a^{SU} + \lambda_a^Q) \frac{S_a}{N_a}, \\
\frac{dE_a^F}{dt} &= \lambda_a^F \frac{S_a}{N_a} - \epsilon E_a^F, \\
\frac{dE_a^{SD}}{dt} &= \lambda_a^{SD} \frac{S_a}{N_a} - \epsilon E_a^{SD}, \\
\frac{dE_a^{SU}}{dt} &= \lambda_a^{SU} \frac{S_a}{N_a} - \epsilon E_a^{SU}, \\
\frac{dE_a^Q}{dt} &= \lambda_a^Q S - \epsilon E_a^Q, \\
\frac{dD_a^F}{dt} &= d_a(1 - H)\epsilon E_a^F - \gamma D_a^F, \\
\frac{dD_a^{SD}}{dt} &= d_a\epsilon E_a^{SD} - \gamma D_a^{SD}, \\
\frac{dD_a^{SU}}{dt} &= d_a(1 - H)\epsilon E_a^{SU} - \gamma D_a^{SU}, \\
\frac{dD_a^{QF}}{dt} &= d_a H \epsilon E_a^F - \gamma D_a^{QF}, \\
\frac{dD_a^{QS}}{dt} &= d_a H \epsilon E_a^{SU} + d_a \epsilon E_a^Q - \gamma D_a^{QS}, \\
\frac{dU_a^F}{dt} &= (1 - d_a)\epsilon E_a^F - \gamma U_a^F, \\
\frac{dU_a^S}{dt} &= (1 - d_a)\epsilon(E_a^{SD} + E_a^{SU}) - \gamma U_a^S, \\
\frac{dU_a^Q}{dt} &= (1 - d_a)\epsilon E_a^Q - \gamma U_a^Q,
\end{aligned}$$

with the forces of infection obeying

$$\begin{aligned}\lambda_a^F &= \sigma_a \sum_b (D_b^F + D_b^{SD} + D_b^{SU} + \tau(U_b^F + U_b^S)) \beta_{ba}^N, \\ \lambda_a^{SD} &= \sigma_a \sum_b D_b^F \beta_{ba}^H, \\ \lambda_a^{SU} &= \sigma_a \tau \sum_b U_b^F \beta_{ba}^H, \\ \lambda_a^Q &= \sigma_a \sum_b D_b^{QF} \beta_{ba}^H,\end{aligned}$$

where β_{ba}^H (with the subscript ba corresponding to transmission from age group b towards age group a) is household transmission and $\beta_{ba}^N = \beta_{ba}^S + \beta_{ba}^W + \beta_{ba}^O$ is all other transmission locations, comprising school-based transmission (β_{ba}^S), work-place transmission (β_{ba}^W) and transmission in all other locations (β_{ba}^O). σ_a corresponds to the age-dependent susceptibility of individuals to infection, d_a the age-dependent probability of displaying symptoms (and hence being detected), and τ represents reduced transmission of infection by undetectable individuals compared to detectable infections.

Amendments to within-household transmission

Given the novelty of the additional household structure that is included in this model, we clarify in more detail here the action of this formulation. We give a simpler set of equations (based on a standard *SIR* model) that contains a similar household structure; in particular, we take the standard *SIR* model and split the infected class into those first infected within a household (I_F) and subsequent infections (I_S):

$$\begin{aligned}\frac{dS}{dt} &= -\beta^H S I_F - \beta^O S (I_F + I_S) \\ \frac{dI_F}{dt} &= \beta^O S (I_F + I_S) - \gamma I_F S \\ \frac{dI_S}{dt} &= \beta^H S I_F - \gamma I_S \\ \frac{dR}{dt} &= \gamma (I_F + I_S)\end{aligned}$$

where the transmission rate is also split into within household transmission β^H and all other transmission β^O (i.e out-of-household transmission). Again, we make the assumption that only the first infection in any household generates infections within the household. We compare this to the *SIR* model without this additional structure:

$$\begin{aligned}\frac{dS}{dt} &= -\hat{\beta}^H S I - \hat{\beta}^O S I \\ \frac{dI}{dt} &= \hat{\beta}^H S I + \hat{\beta}^O S I - \gamma I \\ \frac{dR}{dt} &= \gamma I\end{aligned}$$

where we retain the split in transmission type.

The early growth rate of the two models are $\hat{r} = \hat{\beta}^H + \hat{\beta}^O - \gamma$ for the simple *SIR* model, and $r = \frac{1}{2} \left[\beta^O - 2\gamma + \sqrt{\beta^O^2 + 4\beta^O \beta^H} \right]$ for the household structured version. From this simple comparison, it is clear that for the simple model the growth rate can remain positive even when control measures

substantially reduce transmission outside the home ($\widehat{\beta}^O$ gets reduced), whereas in contrast for the structured version there is always a threshold level of transmission outside the household ($\beta_c^O = \gamma^2/(\beta^H + \gamma)$) that is needed to maintain positive growth.

For both the simple model given here and the full COVID-19 model, the inclusion of this additional household structure reduces the amount of within-household transmission compared to a model without this structure — as only the initial infection in each household (I_F) generates secondary within-household cases. It is therefore necessary to rescale the household transmission rate β^H to obtain the appropriate average within-household attack rate. For the full COVID-19 model, we find that a simple multiplicative scaling to the household transmission ($\beta^H \rightarrow z\beta^H$, $z \approx 1.3$) generates a comparable match between the new model and a model without this household structure — even when age structure is included.

Relationship between age-dependent susceptibility and detectability

We interlink age-dependent susceptibility, σ_a , and detectability, d_a , by a quantity Q_a . Q_a can be viewed as the scaling between force of infection and symptomatic infection. Taking a next-generation approach, the early dynamics would be specified by:

$$R_0 D_a = d_a \sigma_a \beta_{ba}^N (D_a + \tau U_a) / \gamma \quad R_0 U_a = (1 - d_a) \sigma_a \beta_{ba}^N (D_a + \tau U_a) / \gamma$$

where D_a measures those with detectable infections, which mirrors the early recorded age distribution of symptomatic cases. Explicitly, we let $d_a = \frac{1}{\kappa} Q_a^{(1-\alpha)}$ and $\sigma_a = \frac{1}{k} Q_a^\alpha$. As a consequence, $Q_a = \kappa k d_a \sigma_a$; where the parameters κ and k are determined such that the oldest age groups have a 90% probability of being symptomatic ($d_{>90} = 0.90$) and such that the basic reproductive ratio from these calculations gives $R_0 = 2.7$.

2 Public health measurable quantities

The main model equations focus on the epidemiological dynamics, allowing us to compute the number of symptomatic and asymptomatic infectious individuals over time. However, these quantities are not measured - and even the number of confirmed cases (the closest measure to symptomatic infections) is highly biased by the testing protocols at any given point in time. It is therefore necessary to convert infection estimates into quantities of interest that can be compared to data. We considered five such quantities which we calculated from the number of newly detectable symptomatic infections on a given day nD_d .

1. **Hospital Admissions:** We assume that a fraction $P_a^{D \rightarrow H}$ of detectable cases will be admitted into hospital after a delay q from the onset of symptoms. The delay, q , is drawn from a distribution $D_q^{D \rightarrow H}$ (note that $\sum_q D_q^{D \rightarrow H} = 1$.) Hospital admissions on day d of age a are therefore given by

$$H_a(d) = P_a^{D \rightarrow H} \sum_q D_q^{D \rightarrow H} nD_{d-q}$$

2. **ICU Admissions:** Similarly, a fraction $P_a^{D \rightarrow I}$ of detectable cases will be admitted into ICU after a delay, drawn from a distribution $D_q^{D \rightarrow I}$ which determines the time between the onset of symptoms and admission to ICU. ICU admissions on day d of age a are therefore given by

$$ICU_a(d) = P_a^{D \rightarrow I} \sum_q D_q^{D \rightarrow I} nD_{d-q}$$

3. **Hospital Beds Occupied:** Individuals admitted to hospital spend a variable number of days in hospital. We therefore define two weightings, which determine if someone admitted to hospital still occupies a hospital bed q days later (T_q^H) and if someone admitted to ICU occupies a hospital bed on a normal ward q days later ($T_q^{I \rightarrow H}$). Hospital beds occupied on day d of age a are therefore given by

$$H_a^o(d) = \sum_q H_a(d-q)T_q^H + \sum_q ICU_a(d-q)T_q^{I \rightarrow H}$$

4. **ICU Beds Occupied:** We similarly define T_q^I as the probability that someone admitted to ICU is still occupying a bed in ICU q days later. ICU beds occupied on day d of age a are therefore given by

$$ICU_a^o(d) = \sum_q ICU_a(d-q)T_q^I$$

5. **Number of Deaths:** The mortality ratio $P_a^{H \rightarrow Death}$ determines the probability that a hospitalised case of a given age, a , dies after a delay, q drawn from a distribution, $D_d^{H \rightarrow Death}$ between hospitalisation and death. The number of deaths on day d of age a are therefore given by

$$Deaths_a(d) = P_a^{H \rightarrow Death} \sum_q H_a(d-q)D_d^{H \rightarrow Death}$$

These nine distributions are all parameterised from individual patient data as recorded by the COVID-19 Hospitalisation in England Surveillance System (CHESS) [1].

However, these distributions all represent a national average and do not therefore reflect regional differences. We therefore define regional scalings of the three key probabilities ($P_a^{D \rightarrow H}$, $P_a^{D \rightarrow I}$ and

$P_a^{H \rightarrow Death}$) and two additional parameters that can stretch (or contract) the distribution of times spent in hospital and ICU. These five regional parameters are necessary to get good agreement between key observations in all regions and may reflect both differences in risk groups (in addition to age) between regions or differences in how the data are recorded between devolved nations. We stress that these parameters do not (of themselves) influence the epidemiological dynamics, but do strongly influence how we fit to the evolving dynamics.

3 Key inferred parameters

We show the distribution of key parameters from the MCMC process in Fig A. We predict a moderate α and low τ ; the moderate α translates into a sharp change in susceptibility with age (σ_a) but a more steady change in the probability of symptoms. We also show the impact of restrictions (ϕ) for the ten regions within the model.

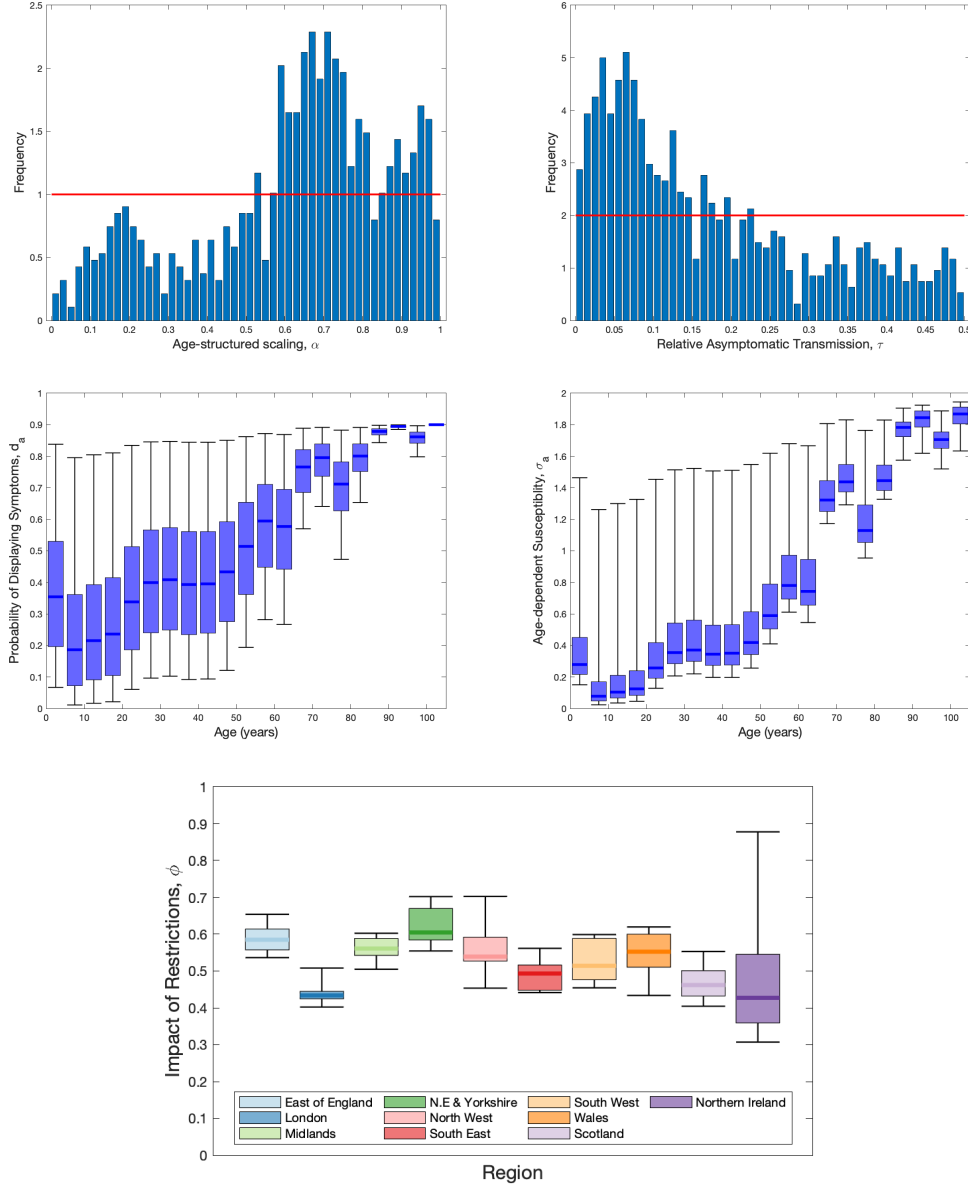


Fig A: Key parameters inferred by the MCMC process. The top two figures show the frequency distribution of α and τ which control the age-structured dynamics; the red line shows the uninformative prior ($[0, 1]$ and $[0.5, 0.5]$) respectively. The middle row shows the results of the inferred α value, giving the distributions of d_a and σ_a . The lower figure shows the impact of control measures ϕ in each of the ten regions. Throughout, error bars give the 95% credible interval, the box is the 50% credible interval and the line is the median value. (Predictions were produced on 23rd April, using data until 21st April).

4 Modelling social distancing

Age-structured contact matrices for the United Kingdom were obtained from Prem et al. [2] and used to provide information on household transmission (β_{ba}^H , with the subscript ba corresponding to transmission from age group b towards age group a), school-based transmission (β_{ba}^S), work-place transmission (β_{ba}^W) and transmission in all other locations (β_{ba}^O). We assumed that the suite of social-distancing and lockdown measures acted in concert to reduce the work, school and other matrices while increasing the strength of household contacts.

We capture the impact of social-distancing by defining new transmission matrices (B_{ba}) that represent the potential transmission in the presence of extreme lockdown. In particular, we assume that:

$$B_{ba}^S = q^S \beta_{ba}^S, \quad B_{ba}^W = q^W \beta_{ba}^W, \quad B_{ba}^O = q^O \beta_{ba}^O,$$

while household mixing B^H is increased by up to a quarter to account for the greater time spent at home. We take $q^S = 0.05$, $q^W = 0.2$ and $q^O = 0.05$ to approximate the reduction in attendance at school, attendance at workplaces and engagement with shopping and leisure activities during the lock-down, respectively.

For a given compliance level, ϕ , we generate new transmission matrices as follows:

$$\begin{aligned} \widehat{\beta}_{ba}^H &= (1 - \phi)\beta_{ba}^H + \phi B_{ba}^H \\ \widehat{\beta}_{ba}^S &= (1 - \phi)\beta_{ba}^S + \phi B_{ba}^S \\ \widehat{\beta}_{ba}^W &= (1 - \theta) [(1 - \phi)\beta_{ba}^W + \phi B_{ba}^W] + \theta ((1 - \phi) + \phi q^W) ((1 - \phi) + \phi q^O) \beta_{ba}^W \\ \widehat{\beta}_{ba}^O &= \beta_{ba}^O ((1 - \phi) + \phi q^O)^2 \end{aligned}$$

As such, home and school interactions are scaled between their pre-lockdown values (β) and post-lockdown limits (B) by the scaling parameter ϕ . Work interactions that are not in public-facing ‘industries’ (a proportion $1 - \theta$) were also assumed to scale in this manner; while those that interact with the general populations (such as shop-workers) were assumed to scale as both a function of their reduction and the reduction of others. We have assumed $\theta = 0.3$ throughout. Similarly, the reduction in transmission in other settings (generally shopping and leisure) has been assumed to scale with the reduction in activity of both members of any interaction, giving rise to a squared term.

5 QALY losses

The loss of quality adjusted life years (QALYs) due to deaths was based on the quality adjusted life expectancy by age, and modified for the relative life-expectancy of individuals that die:

$$\text{Fatal case QALY loss} = \sum_{a=1}^{21} (D(a) \times E(a)),$$

whereby $D(a)$ is the number of deaths in age bracket a , and $E(a)$ is the discounted quality adjusted value of the remaining life expectancy, $L(a)$, of individuals in age group a . This quality adjusted life expectancy is given by:

$$E(a) = \sum_{i=1}^{L(a)} \frac{Q_w(\hat{a} + i)}{(1 + d)^i}$$

where $Q_w(a)$ is the age-specific quality of life weight at age a , \hat{a} is the average age (in years) of an individual in age-group a , and d the discount rate (set at 0.035, corresponding to 3.5% per annum); the values of $L(a)$ are rounded to full years.

For parameterising the age-specific quality of life weights, Q_w , we obtained age-specific EQ-5D index population norms estimates for England from two literature sources. We took childhood estimates (which we used to cover 0–19 years of age) from Table 3 of [3], and values for those aged 20 and above were sourced from Table 3.6 of [4]. A complete listing of age-specific quality of life weights values by age is presented in Table A.

Table A: EQ-5D index population norms for England.

Age group (yrs)	EQ-5D index population norms scale
<20	0.948
20–24	0.929
25–34	0.919
35–44	0.893
45–54	0.855
55–64	0.810
65–74	0.773
75+	0.703

6 Additional tables

Table B: UK population aggregated to ten regions (rounded to nearest 10,000). For our intervention scenario in which regional ICU occupancy triggered the reintroduction and relaxation of social distancing measures within that region, the final column lists each of the regional ICU bed occupancy thresholds (equating to 45 occupied ICU beds per one million population).

Region	Population (millions)	Intervention trigger threshold (ICU bed occupancy)
Wales	3.14	142
Scotland	5.44	245
Northern Ireland	1.88	85
East of England	6.20	280
London	8.91	401
Midlands	10.70	482
North East and Yorkshire	8.14	367
North West England	7.29	328
South East England	9.13	412
South West England	5.60	252

7 Additional figures

Here we show the model fit to the available data on hospital bed occupancy, hospital admissions and ICU bed occupancy, which are combined with the data on deaths (see main text Fig 2). It is clear that due to their greater number fitting to hospital occupancy dominates the fitting process.

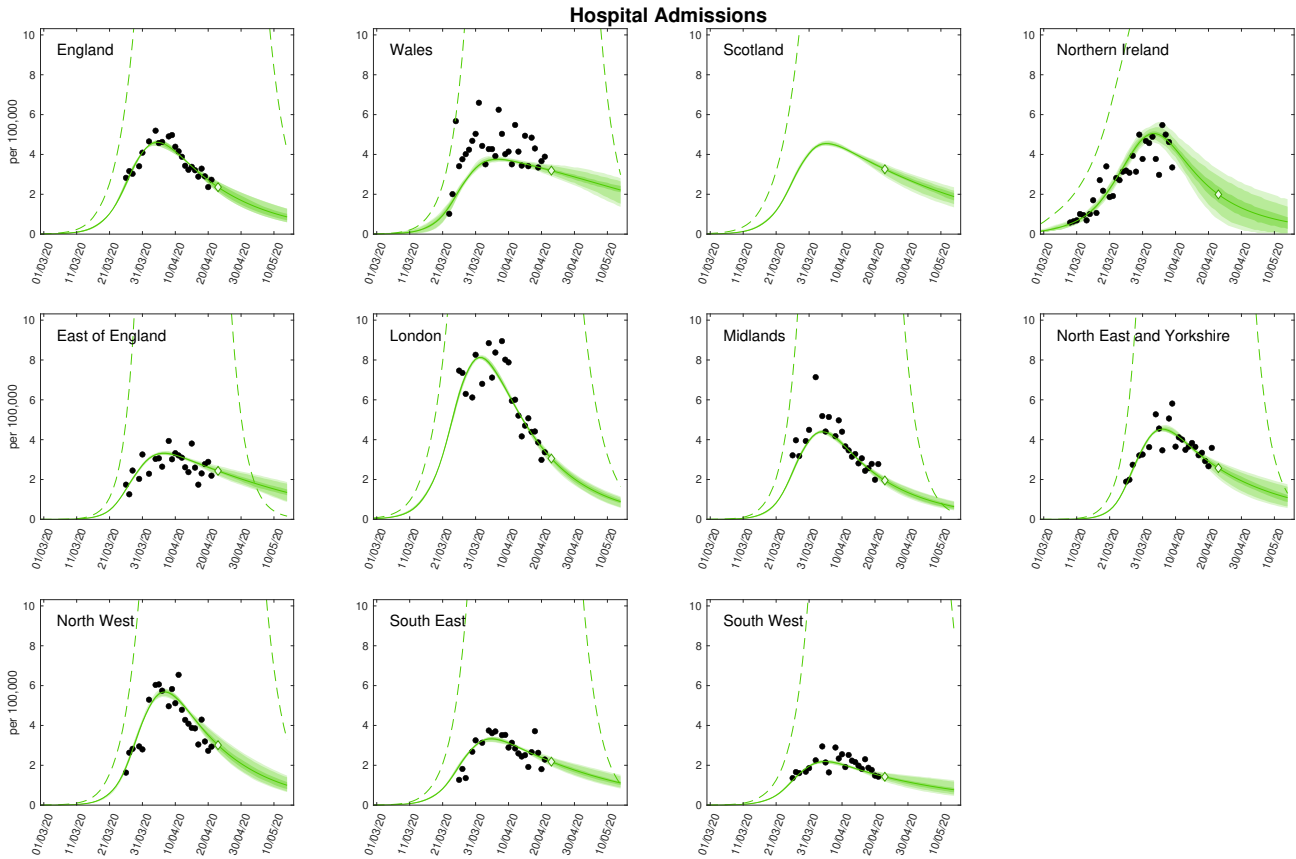


Fig B: Regional projections for hospital admissions per 100,000 with and without imposition of lockdown. In each panel: filled markers correspond to observed data, solid lines correspond to the mean outbreak over a sample of posterior parameters; shaded regions depict prediction intervals, with darker shading representing stricter confidence (dark shading - 50%, moderate shading - 90%, light shading - 99%); dashed lines illustrate the mean projected trajectory had no lockdown measures being introduced (predictions were produced on 23rd April, using data until 21st April).

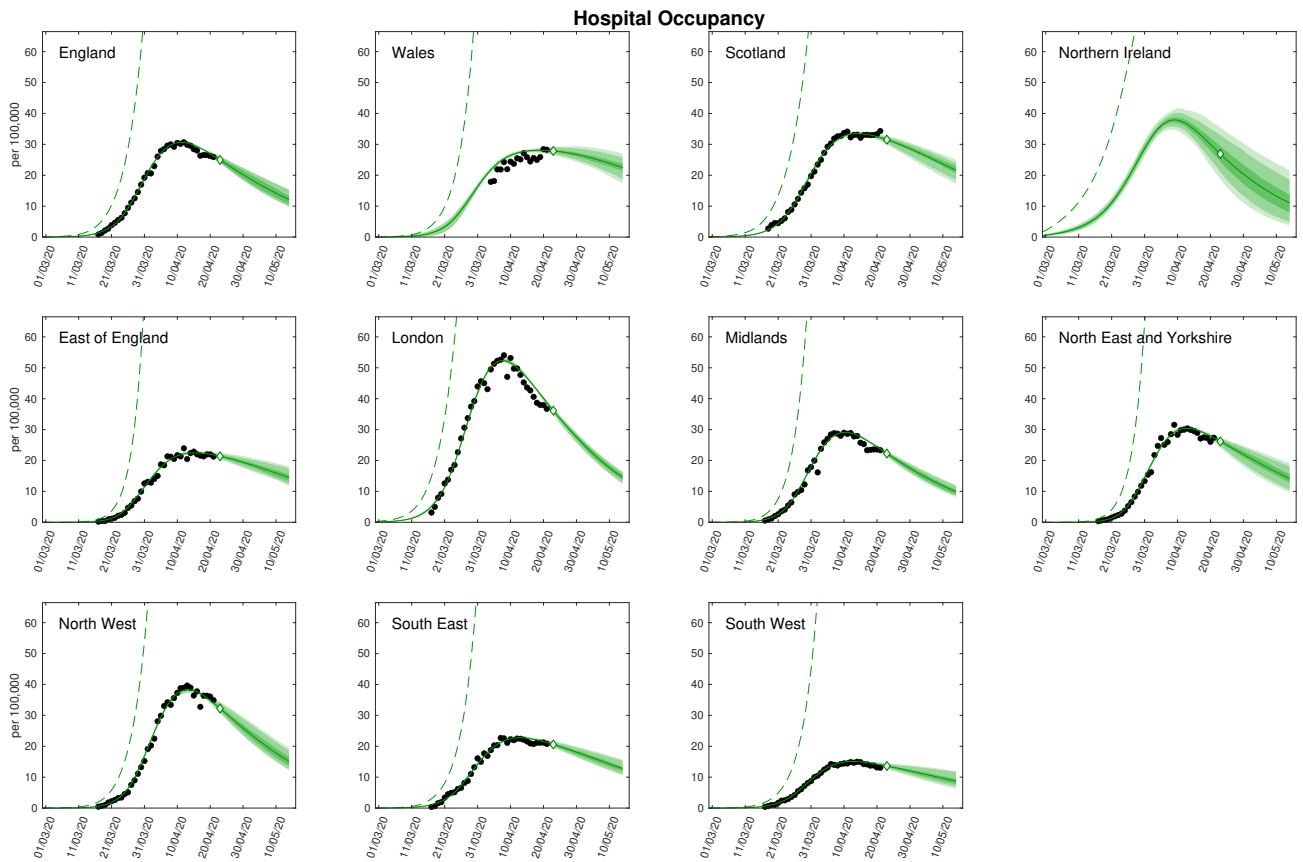


Fig C: Regional projections for hospital occupancy per 100,000 with and without imposition of lockdown. In each panel: filled markers correspond to observed data, solid lines correspond to the mean outbreak over a sample of posterior parameters; shaded regions depict prediction intervals, with darker shading representing stricter confidence (dark shading - 50%, moderate shading - 90%, light shading - 99%); dashed lines illustrate the mean projected trajectory had no lockdown measures being introduced (predictions were produced on 23rd April, using data until 21st April).

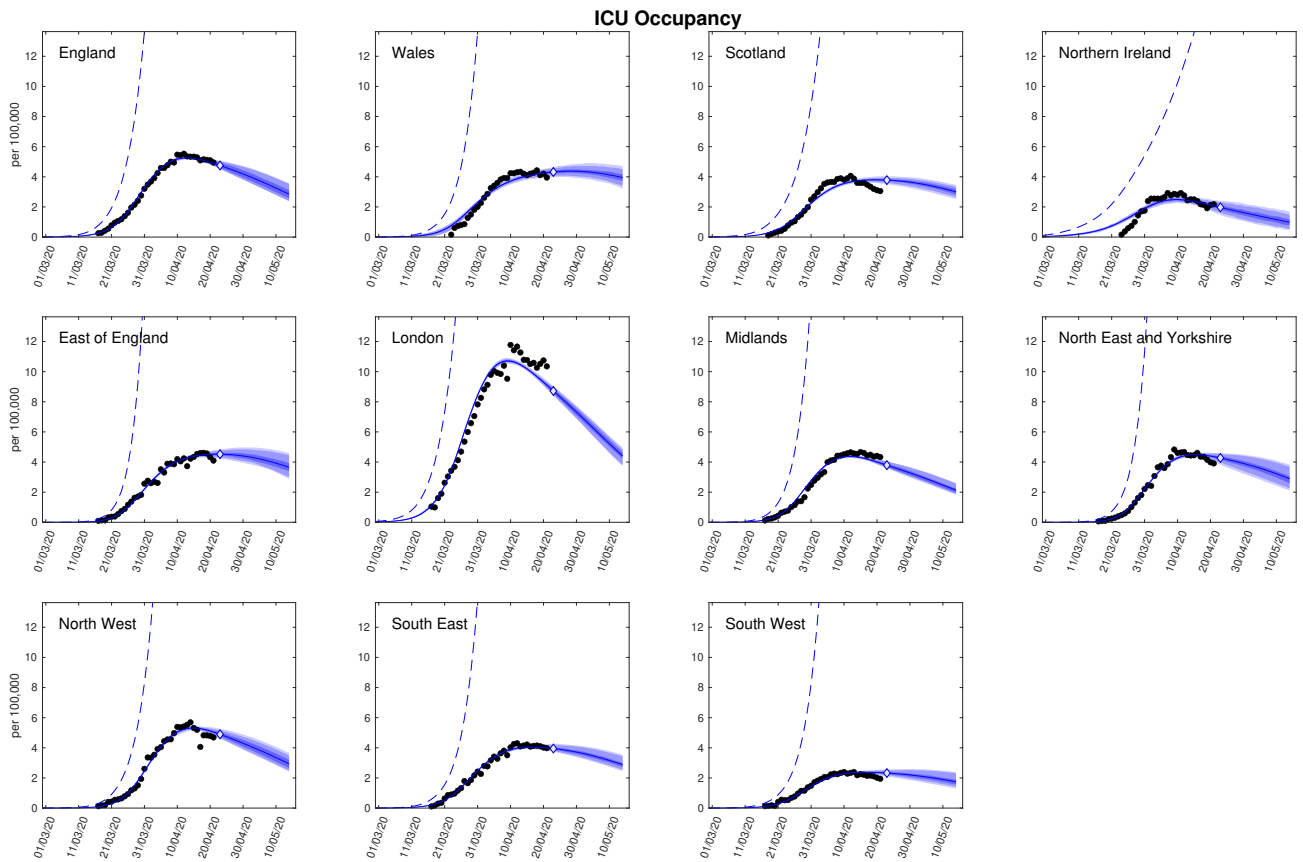


Fig D: Regional projections for ICU bed occupancy per 100,000 with and without imposition of lockdown. In each panel: filled markers correspond to observed data, solid lines correspond to the mean outbreak over a sample of posterior parameters; shaded regions depict prediction intervals, with darker shading representing stricter confidence (dark shading - 50%, moderate shading - 90%, light shading - 99%); dashed lines illustrate the mean projected trajectory had no lockdown measures being introduced (predictions were produced on 23rd April, using data until 21st April).

Reference

1. Public Health England. COVID-19 Hospitalisation in England Surveillance System (CHESS) – daily reporting.; 2020. Available from: <https://www.england.nhs.uk/coronavirus/wp-content/uploads/sites/52/2020/03/phe-letter-to-trusts-re-daily-covid-19-hospital-surveillance-11-march-2020.pdf>.
2. Prem K, Cook AR, Jit M. Projecting social contact matrices in 152 countries using contact surveys and demographic data. *PLOS Comput Biol.* 2017;13(9):e1005697. doi:10.1371/journal.pcbi.1005697.
3. Kwon J, Kim SW, Ungar WJ, Tsiplova K, Madan J, Petrou S. A Systematic Review and Meta-analysis of Childhood Health Utilities. *Med Decis Mak.* 2018;38(3):277–305. doi:10.1177/0272989X17732990.
4. Janssen B, Szende A. Population Norms for the EQ-5D. In: *Self-Reported Popul. Heal. An Int. Perspect. based EQ-5D.* Dordrecht: Springer Netherlands; 2014. p. 19–30.