

Community prevalence of SARS-CoV-2 in England during April to November 2020: Results from the ONS Coronavirus Infection Survey

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1 Sampling design

The following information on the sampling design can also be found here: <https://www.ons.gov.uk/peoplepopulationandcommunity/healthandsocialcare/conditionsanddiseases/methodologies/covid19infectionsurveyspilotmethodsandfurtherinformation#study-design-sampling>

At the start of the study at the end of April, the sample for the survey was drawn mainly from the Annual Population Survey (APS), which consists collectively of those who successfully completed the last wave of the Labour Force Survey (LFS) or local LFS boost, and who had consented to future contact regarding research.

Around 38,000 households respond to the LFS each quarter and it is the largest regular household survey in the UK. The sampling frame for the LFS is the Postal Address File of small users, which contains approximately 26 million addresses. Only private households are included in the sample. People living in care homes, other communal establishments and hospitals are not included. Only private households in England are included in the current paper.

We initially invited about 20,000 households to take part, anticipating that this would result in approximately 21,000 individuals from approximately 10,000 households participating. Since the end of May, additional households have been invited to take part in the survey each week (roughly 5,000 a week).

At the start of the study, all respondents to the COVID-19 Infection Survey were individuals who have previously participated in an Office for National Statistics (ONS) social survey, which means the number of ineligible addresses in the sample is substantially reduced. To take part, invited households opted into the survey by contacting IQVIA, a company working on behalf of the ONS, to arrange a visit.

Since the end of July, we have further expanded the survey to invite a random sample of households from AddressBase, which is a commercially available list of addresses maintained by the Ordnance Survey. In line with our plans to increase our overall sample size, we prioritised areas under government local restriction because of an outbreak of the coronavirus (COVID-19). We have invited 40,000 extra households from 14 local authorities within selected local authorities in Greater Manchester, Lancashire and West Yorkshire to participate in this study. We also boosted our sample in London, inviting 50,000 extra households to increase the household involvement rates in this area.

In August, we announced our plans to further expand the study with the aim of increasing from 28,000 people tested per fortnight in England to 150,000 people tested per fortnight by October until March 2021. A random sample of households from AddressBase was invited for this expansion.

The number of participants enrolled in each month alongside the number of subsequent visits is shown in Table S1.

2 Models estimated in the paper

Dynamic MRP

The regression model that was used for the dynamic multilevel model and post-stratification (MRP) analysis was a Bayesian multilevel generalised additive model (GAMM) with a complementary loglog link implemented using the `rstanarm` package.[1-2] Sex was modelled as a fixed effect as it has only 2 levels, while age (5 levels) and region (12 levels) were modelled as random effects. This model was implemented using the following syntax:

```
stan_gamm4(result ~ s(time, by=region, k=10) + sex,
random = ~(1|age) + (1|region),
family = binomial(link="cloglog"),
data = data, iter = 3000, cores = 4,
prior = normal(0,0.5), prior_covariance
= decov(shape = 1, scale = 1),
prior_smooth=normal(location=4),
control=list(adapt_delta=0.95))
```

Associations between variables and testing positive

To assess whether particular subgroups are more likely to test positive for SARS-CoV-2 viral RNA we performed an additional analysis including variables on which we did not post-stratify. We used the same model as for the dynamic MRP but in working aged individuals only (16-74 years inclusive as defined in the Labour Force Survey) with these additional variables included as fixed covariates and age modelled as a continuous variable, using a thin-plate spline, instead of a categorical variable. Associated results can be found in Table S1.

2 Epidemiological interpretation of the complementary log-log link function when focusing on associations between variables and testing positive for the presence of SARS-CoV-2 RNA.

Our regression model operates at the individual level; in particular, we assume that there are n swabs taken, and the i -th of these is associated with time t_i , English region e_i , and a vector of other covariates x_i . These covariates are as detailed in the main text: age; work etc. The probability of the i -th swab being positive is then given by a generalised linear model (generalised additive model).

$$\pi_i = p(x_i, e_i, t_i) = g^{-1}(s_{e_i}(t_i) + \beta \cdot x_i + \zeta_{e_i}). \quad (1)$$

Here, g is the link function of the GAMM, s_{e_i} is the time smoother for the region e , β is the vector of regression coefficients, and ζ_e is the random effect for region e , with these effects assumed i.i.d. with $\zeta_e \sim N(0, \sigma^2)$. The likelihood function for this model given observations $y_i = 1$ for a positive swab and $y_i = 0$ for negative, is then

$$L = \prod_{i=1}^n \pi_i^{y_i} (1 - \pi_i)^{1-y_i}. \quad (2)$$

Now suppose that the individual i acquires infection at a rate $\lambda_i(t)$, known as the *force of infection* in infectious disease modelling. The Chapman-Kolmogorov equation, using dots for time derivatives, is:

$$\dot{\pi}_i(t) = (1 - \pi_i) \lambda_i(t). \quad (3)$$

This has solution

$$\pi_i = 1 - \exp\left(-\int_{u=0}^t \lambda_i(u) du\right). \quad (4)$$

If we choose a complementary log-log link function

$$g(x) = \log(-\log(1 - x)) \quad (5)$$

in (1), and assume that

$$\lambda_i(t) = \phi_i \lambda_{e_i}(t). \quad (6)$$

in (5), then we get

$$\pi_i = 1 - \exp\left(-\phi \int_{u=0}^t \lambda_{e_i}(u) du\right) = 1 - \exp\left(-\exp(\beta \cdot x_i) \exp(s_{e_i}(t_i)) \exp(\zeta_{e_i})\right). \quad (7)$$

This implies that

$$\phi_i \propto \prod_a e^{\beta_a x_{ia}}, \quad (8)$$

and so we quote the value of $\exp(\beta_a)$ for the a -th covariate, with 1 as reference, since this is interpretable as the relative exposure to infectious risk.

3 Figures and tables

Table S1. Enrollment by month and % of participants with follow-up visits

Recruitment month	Number of participants with a first visit	Number of participants with at least 2 visits (%)	Number of participants with at least 3 visits (%)	Number of participants with at least 4 visits (%)	Number of participants with at least 5 visits (%)
April	2,440	2,421 (99%)	2,411 (99%)	2,398 (98%)	2,376 (97%)
May	19,575	19,377 (99%)	19,267 (98%)	19,156 (98%)	18,804 (97%)
June	15,987	15,811 (99%)	15,737 (98%)	15,608 (98%)	15,288 (96%)
July	16,194	16,041 (99%)	15,896 (98%)	15,567 (96%)	14,784 (91%)
August	36,395	35,755 (98%)	34,977 (96%)	32,917 (90%)	27,499 (76%)
September	94,120	90,112 (96%)	84,601 (90%)	70,090 (75%)	40,606 (43%)
October	93,064	70,918 (76%)	42,437 (46%)	15,977 (17%)	2,648 (3%)

Table S2. Characteristics of participants

	Individuals (n=280,327) N (%) ^a	Samples (n=1,119,170) N (%) ^a	Private households in England (n = 54,524,766) N (%)
<i>Age (years)</i>			
2-11	22,317 (8.0%)	93,125 (7.8%)	6,934,338 (12.7%)
12-16	14,692 (5.2%)	61,789 (5.2%)	3,293,437 (6.0%)
17-24	16,563 (5.9%)	64,418 (5.4%)	4,933,547 (9.1%)
25-34	28,619 (10.2%)	112,812 (9.5%)	7,526,664 (13.8%)
35-49	55,226 (19.7%)	230,480 (19.3%)	10,813,332 (19.8%)
50-69	92,629 (33.0%)	403,707 (33.9%)	13,621,021 (25.0%)
70+	50,281 (17.9%)	224,839 (18.9%)	7,402,427 (13.6%)
<i>Sex</i>			
Male	133,371 (47.6%)	566,252 (47.5%)	26,959,721 (49.4%)
Female	146,956 (52.4%)	624,918 (52.5%)	27,565,045 (50.6%)
<i>Region</i>			
North East	13,157 (4.7%)	59,616 (5.0%)	2,575,735 (4.7%)
North West	39,938 (14.2%)	174,819 (14.7%)	7,083,473 (13.0%)
Yorkshire and The Humber	27,894 (10.0%)	123,105 (10.3%)	5,315,017 (9.8%)
East Midlands	21,830 (7.8%)	98,399 (8.3%)	4,692,054 (8.6%)
West Midlands	25,327 (9.0%)	114,562 (9.6%)	5,764,018 (10.6%)
East of England	33,274 (11.9%)	134,410 (11.3%)	6,046,125 (11.1%)
London	51,113 (18.2%)	181,079 (15.2%)	8,719,114 (16.0%)
South East	41,781 (14.9%)	186,857 (15.7%)	8,855,708 (16.2%)
South West	26,013 (9.3%)	118,323 (9.9%)	5,473,522 (10.0%)
<i>Ethnicity^b</i>			
White	257,413 (91.8%)	1,102,405 (92.5%)	82.6% ^b
Asian	11,748 (4.2%)	46,381 (3.9%)	9.9% ^b
Black	2,715 (1.0%)	10,790 (0.9%)	3.8% ^b
Mixed	5,115 (1.8%)	20,433 (1.7%)	2.7% ^b
Other	2,763 (1.0%)	10,178 (0.9%)	1.2% ^b
Not reported	573 (0.2%)	983 (0.08%)	NA
<i>Household size^b</i>			
1	44,914 (16.0%)	118,122 (15.8%)	16% ^c
2	119,050 (42.5%)	509,332 (42.8%)	33% ^c
3	45,273 (16.2%)	189,739 (15.9%)	19% ^c
4	49,259 (17.6%)	211,404 (17.7%)	21% ^c
5+	21,831 (7.7%)	92,573 (7.8%)	12% ^c

^a The methods that we used – Bayesian dynamic multilevel regression and poststratification – adjust for residual non-representativeness in terms of age, sex, and region.

^b Ethnicity distribution in the target population was estimated by combining ONS estimates of age-sex-region specific number of individuals living in private households with estimates of the ethnicity distribution within those same age-sex-region categories in the overall population (including those not living in private households) obtained from the Ethpop database: Wohland P, Burkitt M, Norman P, Rees P, Boden P and Durham H, ETHPOP Database, ESRC Follow on Fund "Ethnic group population trends". www.ethpop.org. Date of extraction 21 09 2020.

^c Household size distribution is provided for the overall population (including those not living in private households) in absence of data on household size of the target population. Therefore the 'reference data' may have too many large household sizes (e.g. student halls).

Table S3. Risk factors for testing positive for SARS-CoV-2 between 26 April and 28 June 2020.

Factor	Number of visits in sample (number positive)	Relative exposure to SARS-CoV-2 (95% CrI)^a
Male	44,308 (53)	Ref.
Female	49,308 (57)	0.84 (0.57 - 1.25)
<i>Work location</i>		
Working from home	22,392 (11)	Ref.
Working outside of your home	20,621 (45)	2.47 (1.40 - 4.55)
Both	4,370 (5)	1.43 (0.53 - 3.54)
Not applicable	46,233 (49)	2.09 (1.20 - 3.75)
<i>Job with direct contact with patients or care home residents</i>		
Non-patient facing	89,643 (87)	Ref.
Patient facing	3,973 (23)	4.06 (2.37 - 6.72)
Non-resident facing	92,690 (105)	Ref.
Care home resident facing	926 (5)	2.35 (0.85 - 5.27)
<i>Ethnicity</i>		
White	88,793 (93)	Ref.
Asian	2,514 (8)	1.89 (0.87 - 3.64)
Black	788 (2)	1.04 (0.28 - 3.07)
Mixed	1,068 (0)	0.46 (0.09 - 1.84)
Other	453 (7)	7.50 (2.86 - 16.50)
<i>Household size</i>		
1	13,096 (16)	Ref.
2	40,426 (28)	0.62 (0.35 - 1.09)
3	17,125 (33)	1.51 (0.83 - 2.73)
4	16,704 (21)	1.36 (0.68 - 2.63)
5 or more	6,261 (12)	1.25 (0.51 - 2.88)
<i>Number of children in household</i>		
0	64,917 (70)	Ref.
1	11,206 (22)	1.01 (0.59 - 1.73)
2	10,083 (8)	0.44 (0.20 - 0.94)
3 or more	2,593 (8)	1.65 (0.64 - 4.17)

^aA relative exposure of 1 is the reference value (no effect).

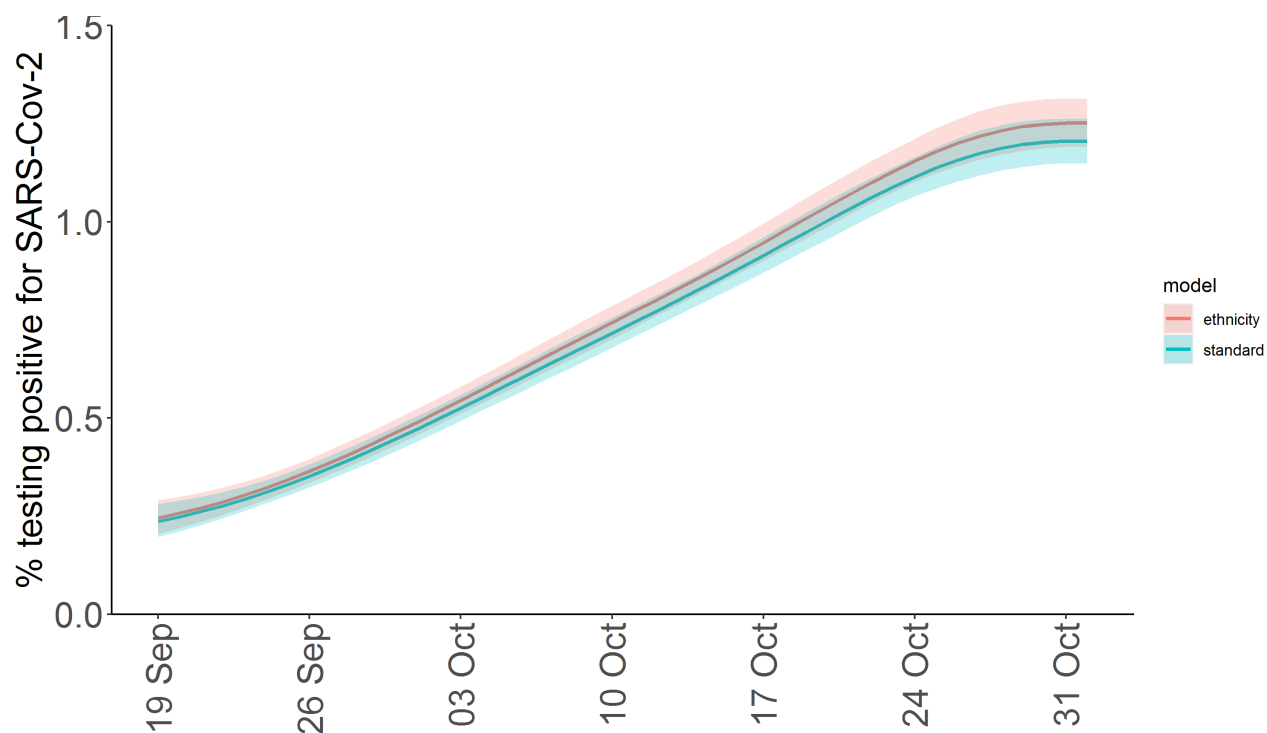


Figure S1. Comparison of multi-level regression with post-stratification model without accounting for ethnicity (standard) and with post-stratification for ethnicity (ethnicity). The shaded area falls within the 95% credible intervals.

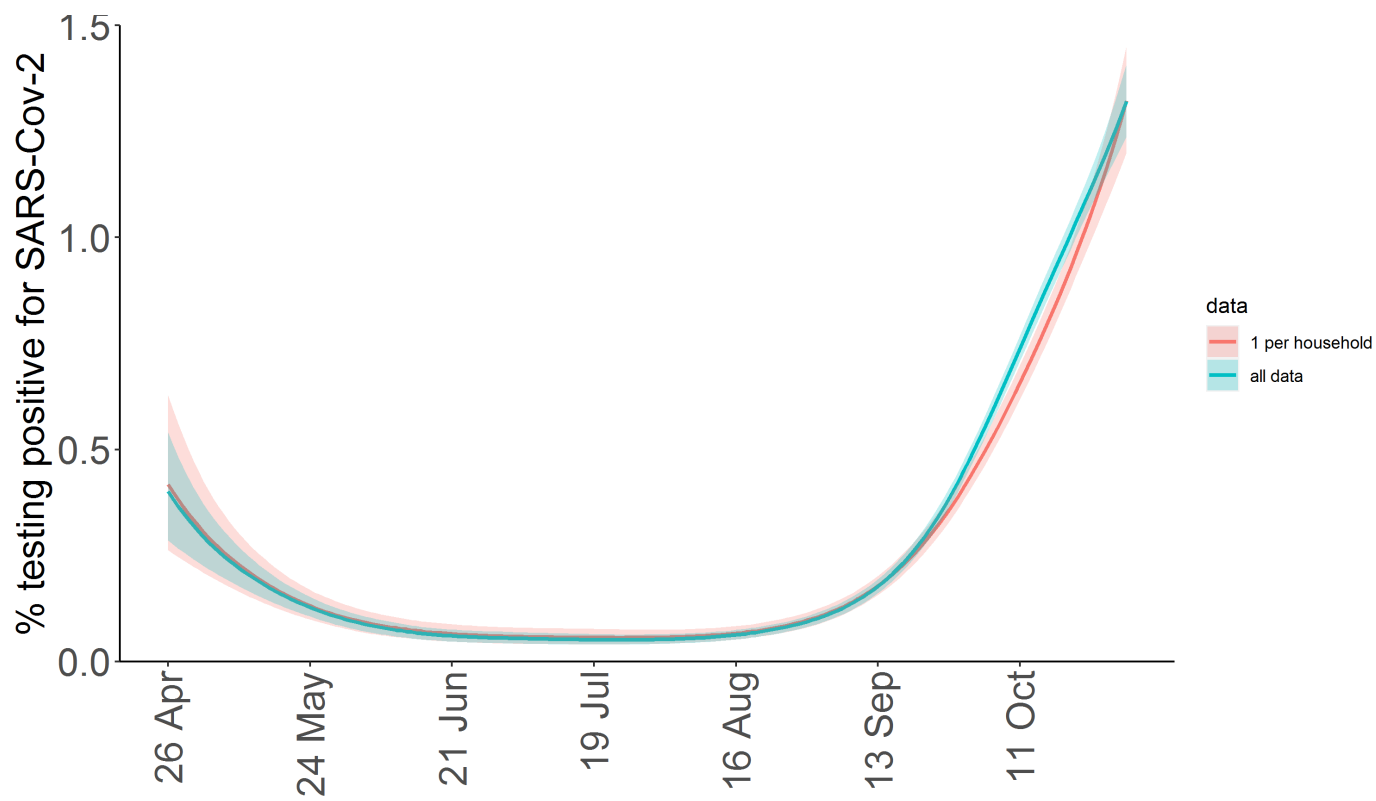


Figure S2. Comparison of using all data for the multi-level regression with post-stratification for estimation of the percentage of inhabitants testing positive over time versus use one randomly selected person per household. The shaded area falls within the 95% credible intervals.

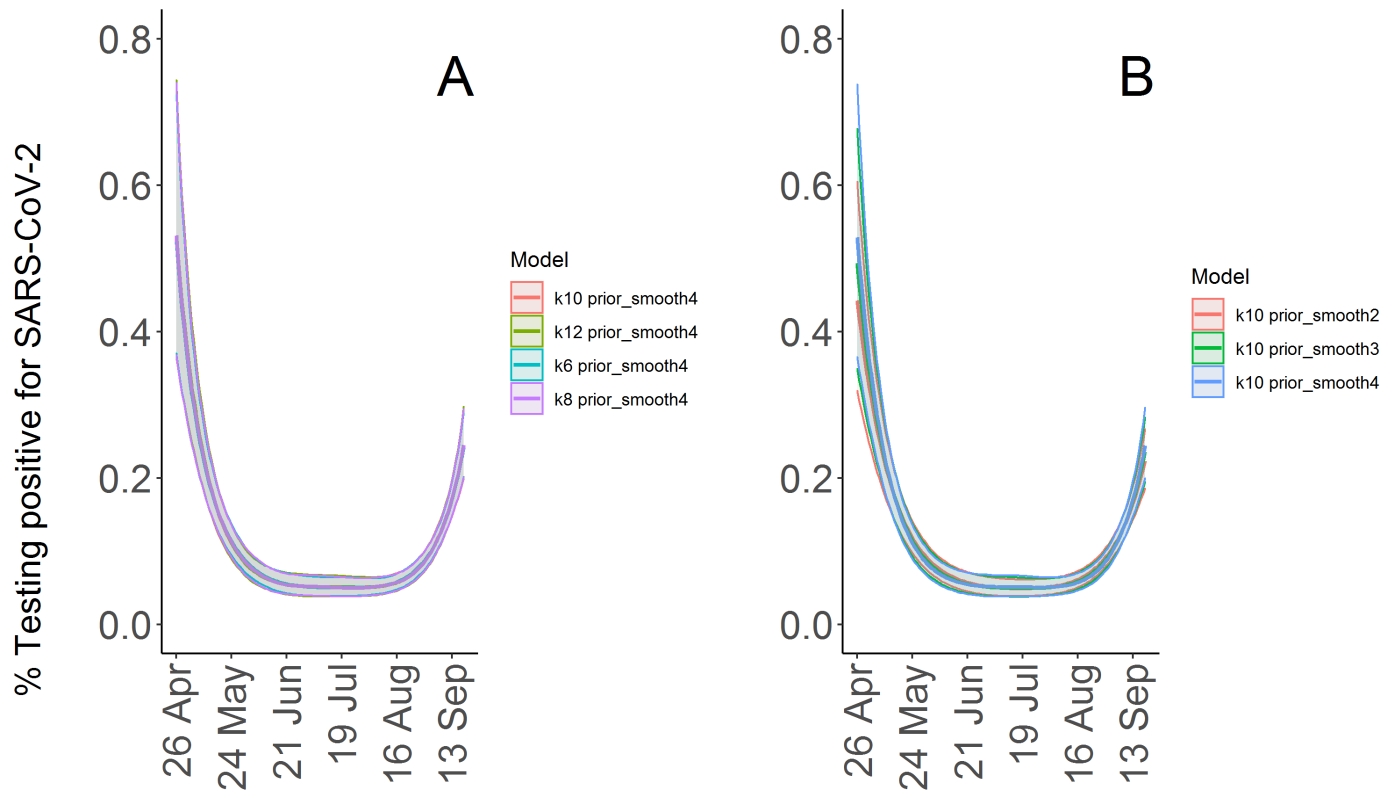


Figure S3. Comparison of models using different values for k (6, 8, 10, and 12, higher values resulted in divergent transitions) (A) and for the prior of the standard deviation of the smooth (normal prior with location 2, 3, and 4) for estimating the percentage of the population living in private households testing positive for SARS-CoV-2 with and without reporting symptoms.

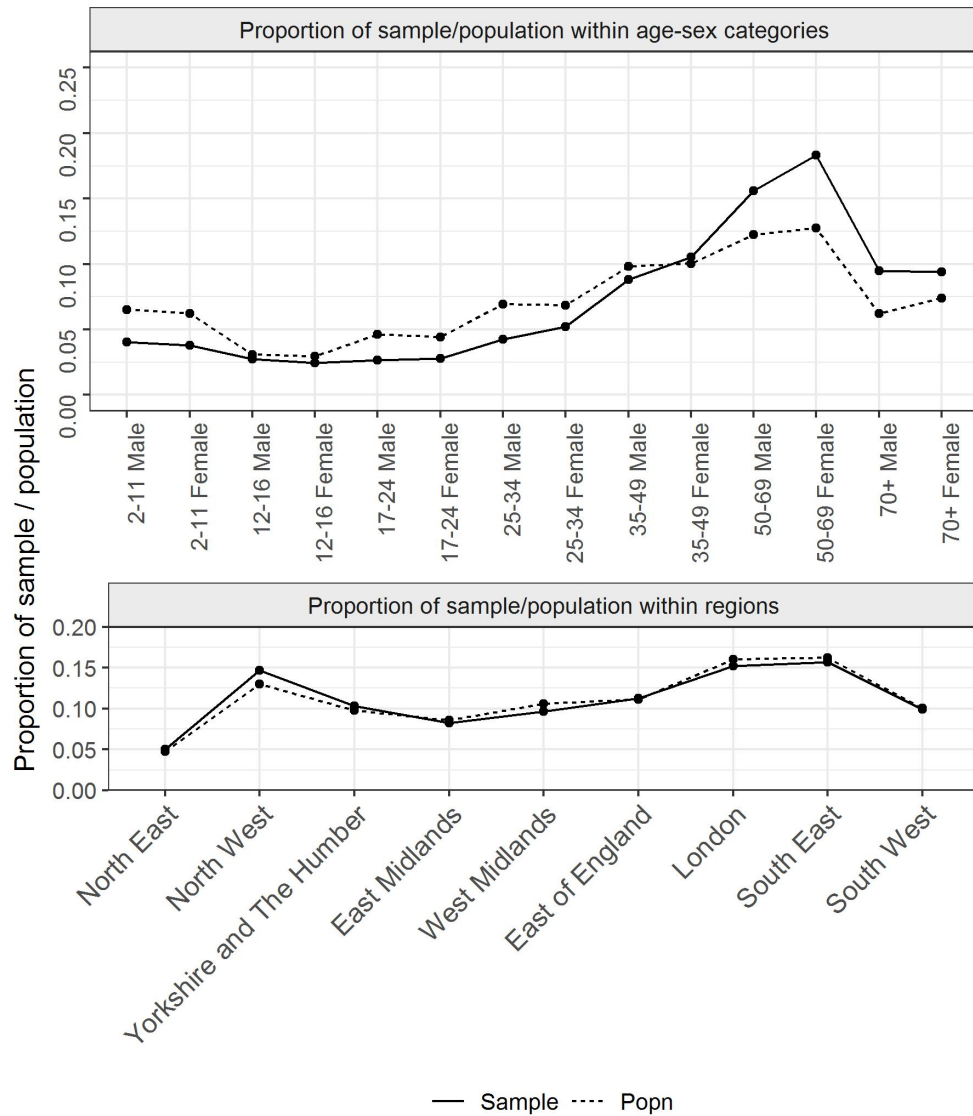


Figure S4. Representativeness of sample in terms of age and sex. Proportion of sample (solid line) and population in England (dashed line) within age- and sex-categories and regions.

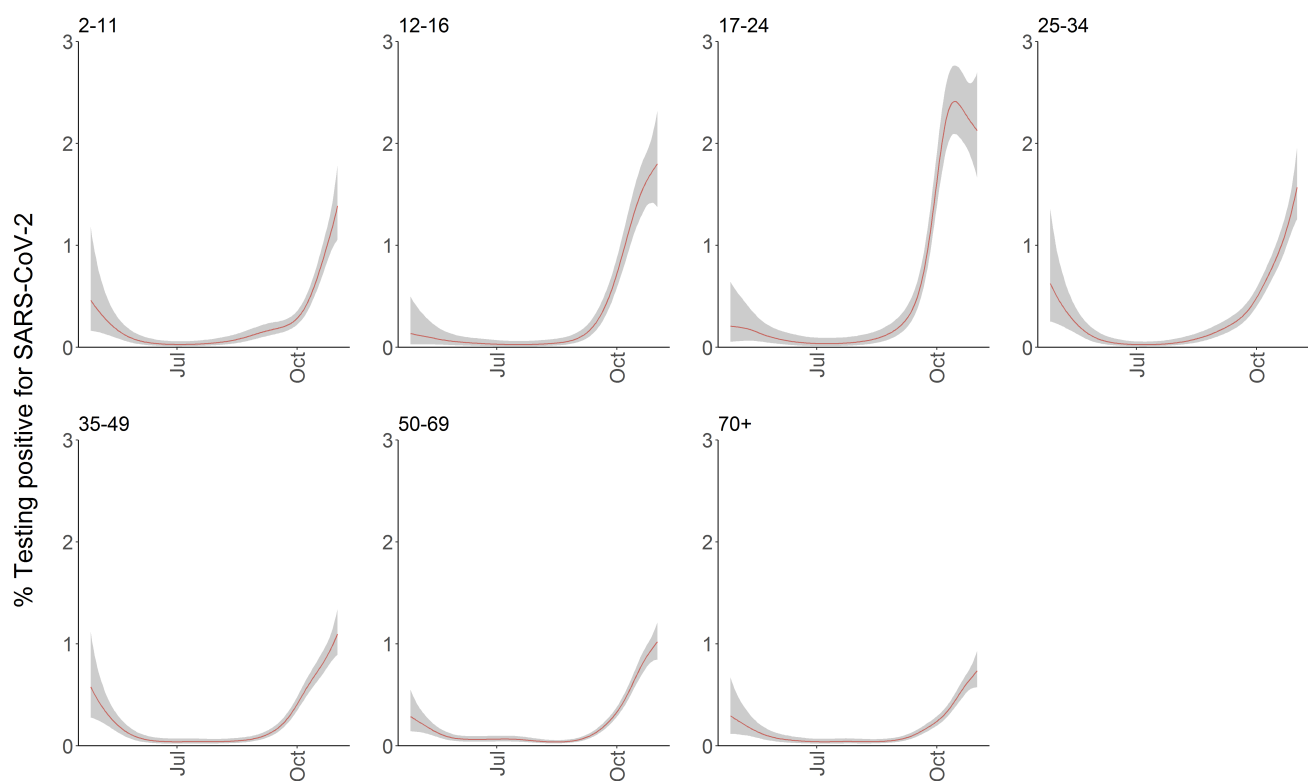


Figure S5. Percentage of population within age (in years) subgroups testing positive for SARS-CoV-2. Estimates are from a multilevel Bayesian GAM without post-stratification. The shaded area falls within the 95% credible intervals.

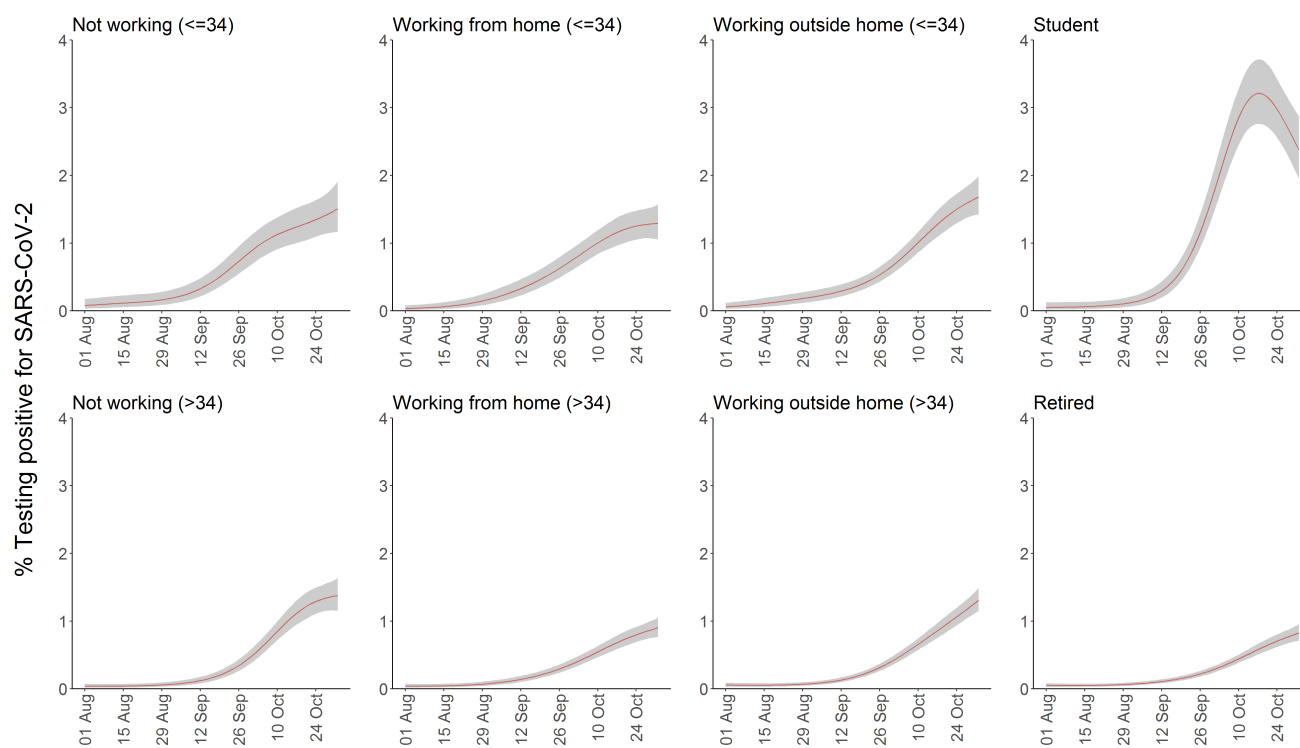


Figure S6. Percentage of population testing positive for SARS-CoV-2 by work location. Estimates are from a multilevel Bayesian GAM without post-stratification. The shaded area falls within the 95% credible intervals.

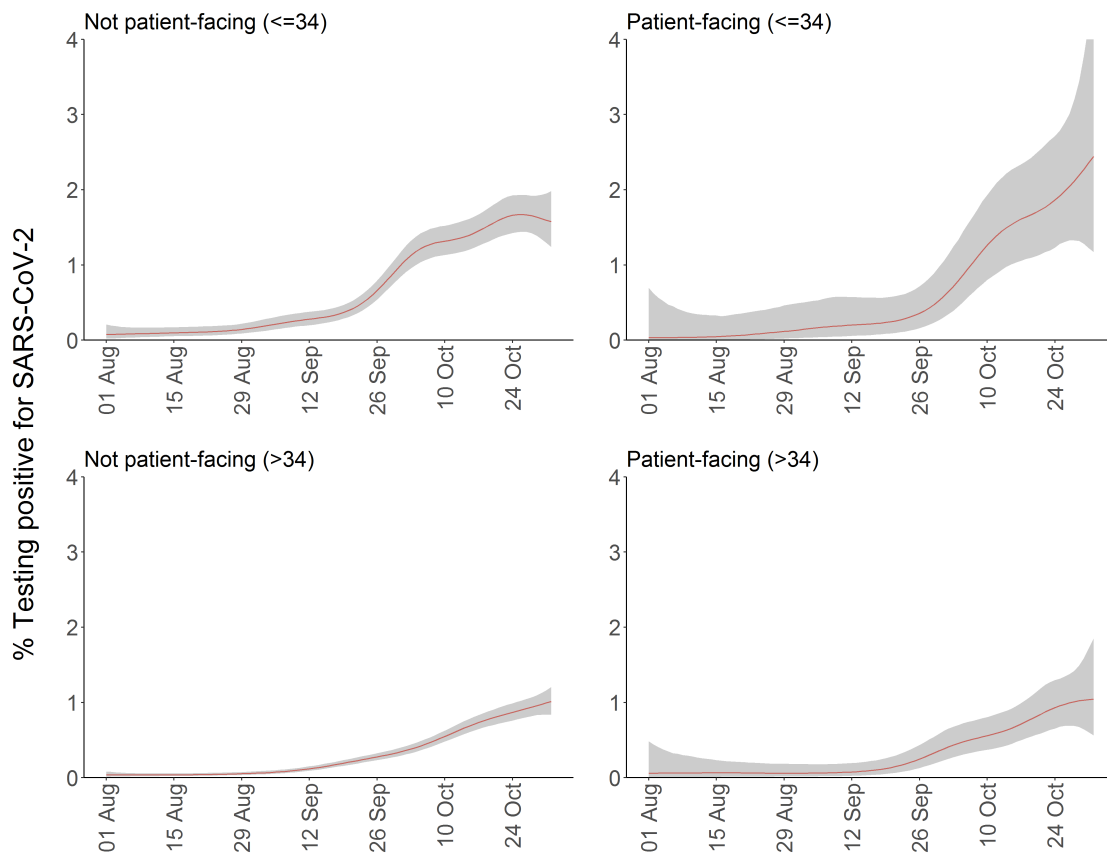


Figure S7. Percentage of population testing positive for SARS-CoV-2 by having a patient-facing role or not. Estimates are from a multilevel Bayesian GAM without post-stratification. The shaded area falls within the 95% credible intervals.

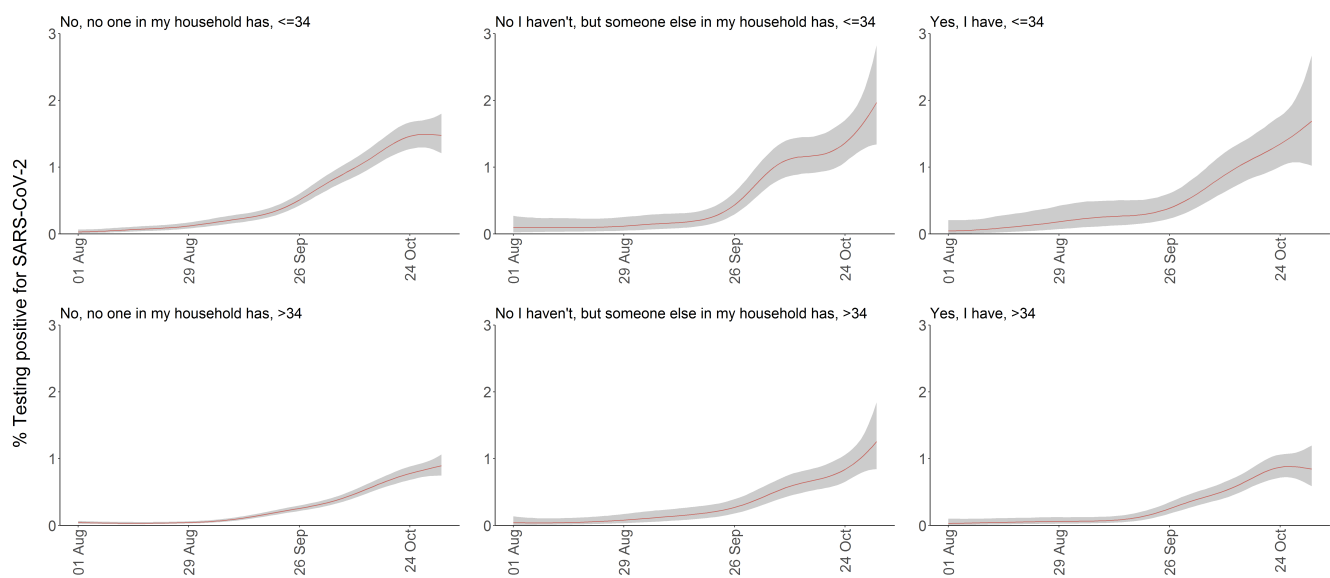


Figure S8. Percentage of population testing positive for SARS-CoV-2 by contact with hospital. Estimates are from a multilevel Bayesian GAM without post-stratification. The shaded area falls within the 95% credible intervals.

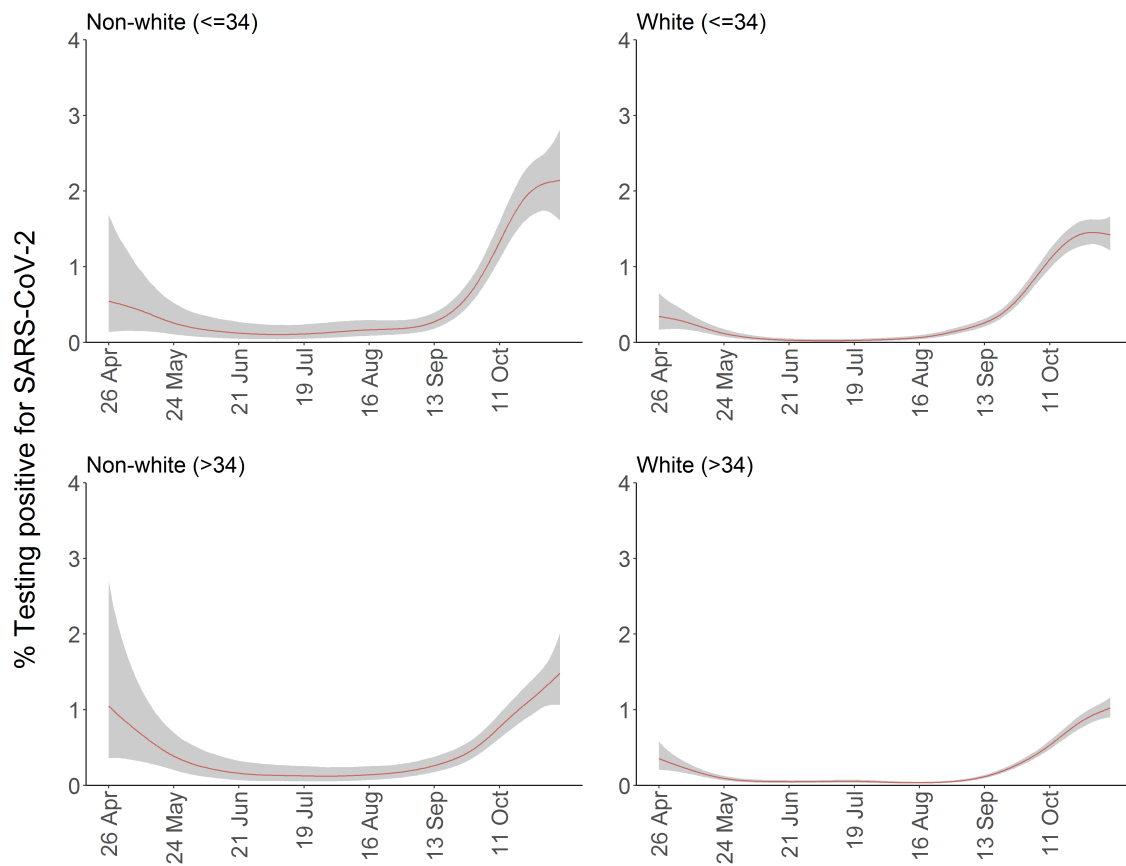


Figure S9. Percentage of population testing positive for SARS-CoV-2 by ethnicity (white / non-white). Estimates are from a multilevel Bayesian GAM without post-stratification. The shaded area falls within the 95% credible intervals.

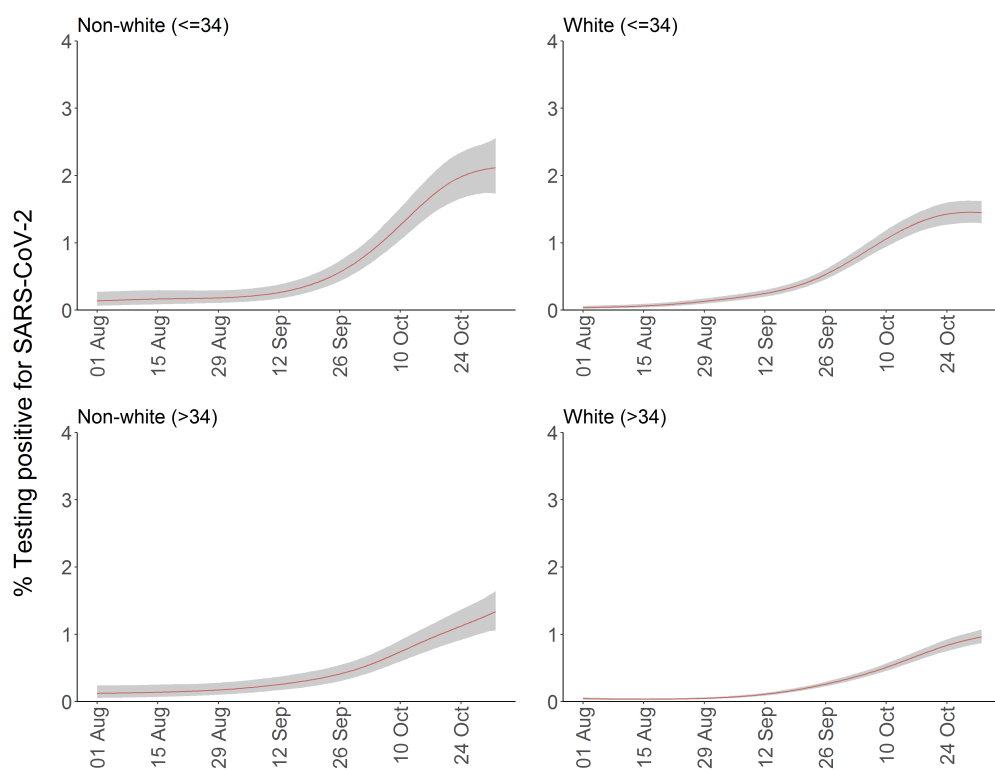


Figure S10. Percentage of population living in private households testing positive for SARS-CoV-2 stratified by ethnicity and age (≤ 34 and > 34 years of age). Shaded areas are 95% credible intervals.

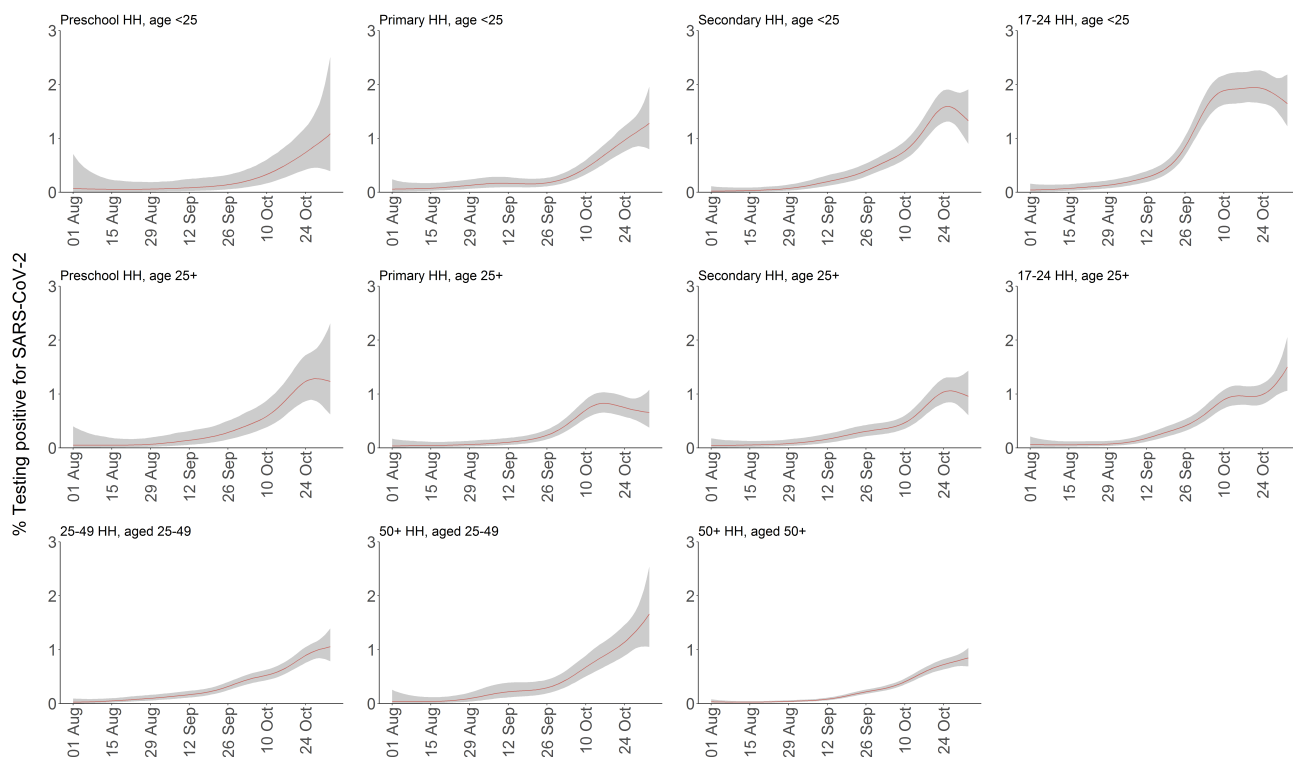


Figure S11. Percentage of population living in private households testing positive for SARS-CoV-2 stratified by their household composition. HH=household; Preschool HH = household with a preschool child (< 5 years of age); Primary HH = household with a primary school child ($\leq$ school year 6, $\leq$ 11/12 years of age); Secondary HH = household with a secondary school child ($\leq$ school year 11, $\leq$ 16/17 years of age). The first part of the title of each plot indicates the type of household (e.g. Preschool HH) and the second part indicates the age of the individual (e.g. age <25). Households were classified according to the following hierarchy: anyone aged 17-24 (17-24 HH); anyone in secondary school age (Secondary HH); anyone in primary school age (Primary HH); anyone in preschool age (Preschool HH); anyone aged 50+ (50+HH); all 25-49 (25-49 HH). Shaded areas are 95% credible intervals.

7 References

1. <https://cran.r-project.org/web/packages/rstanarm/vignettes/mrp.html>
2. [http://www.stat.columbia.edu/~gelman/research/unpublished/MRT\(1\).pdf](http://www.stat.columbia.edu/~gelman/research/unpublished/MRT(1).pdf)
3. <https://www.jstatsoft.org/article/view/v080i01>

8 Rstanarm code used for the regression model results from table S1 in the main paper and full details on all models considered

Model 1: Analysis from Table S1

Model 2: Model 1 + random intercept for household.

Model 3: Model 1 + hospital/care home contact added to the model. The relative exposure for contact with hospital and contact with care home reported in Table 1 come from this model. As these questions were only added since 8 May 2020, this regression is restricted to subset of data starting at 8 May.

Model 4: A reduced model with less covariates than model 1.

	Model 1: Analysis from main text (table 1)	Model 2: model 1 + random intercept for household	Model 3: model 1 + hospital/carehome contact	Model 4: reduced model
<i>priors for covariables</i>	norm(0,1)	norm(0,1)	norm(0,1)	norm(0,1)
<i>prior for intercept</i>	norm(0,10)	norm(0,10)	norm(0,10)	norm(0,10)
<i>prior for covariance matrix (decov function in rstanarm)</i>	gamma(1,1)	gamma(1,1)	gamma(1,1)	gamma(1,1)
<i>regression formula</i>	stan_gamm4(result ~sex + s(study_day, by=region, k=5) + s(age, k=5) + work_location + patient_facing + resident_facing + ethnicity + householdsize + numchild, random= ~ (1 region), family=binomial(link="cloglog"), data=adults, iter=3000, cores=4, prior=normal(0,1), control=list(adapt_delta=0.95))	stan_gamm4(result ~sex + s(study_day, by=region, k=5) + s(age, k=5) + work_location + patient_facing + resident_facing + ethnicity + householdsize + numchild, random= ~ (1 region/household_id), family=binomial(link="cloglog"), data=adults, iter=3000, cores=4, prior=normal(0,1), control=list(adapt_delta=0.95))	stan_gamm4(result ~sex + s(study_day, by=region, k=5) + s(age, k=5) + work_location + patient_facing + resident_facing + ethnicity + householdsize + numchild + contact_hospital + contact_carehome, random= ~ (1 region), family=binomial(link="cloglog"), data=adults, iter=3000, cores=4, prior=normal(0,1), control=list(adapt_delta=0.95))	stan_gamm4(result ~sex + s(study_day, by=region, k=5) + s(age, k=5) + work_location + patient_facing + resident_facing, random= ~ (1 region), family=binomial(link="cloglog"), data=adults, iter=3000, cores=4, prior=normal(0,1), control=list(adapt_delta=0.95))
	Relative exposure (95% CrI)	Relative exposure (95% CrI)	Relative exposure (95% CrI)	Relative exposure (95% CrI)
Intercept	0.00045 (0.0002 to 0.00092)	0.00002 (0.000004 to 0.00005)	0.0005 (0.002 to 0.0011)	0.00043 (0.0002 to 0.00079)
Female	0.84 (0.57 to 1.25)	0.89 (0.59 to 1.35)	0.77 (0.49 to 1.19)	0.82(0.55 to 1.22)
<i>work location</i>				
Working outside of your home	2.47 (1.40 to 4.55)	2.34 (1.24 to 4.51)	2.11 (1.09 to 4.28)	2.73 (1.55 to 4.96)
Both (working from home and working outside of your home)	1.43 (0.53 - 3.54)	1.54 (0.55 to 4.05)	0.88 (0.24 to 2.60)	1.52 (0.54 to 3.76)
Not applicable	2.09 (1.20 to 3.75)	2.07 (1.14 to 4.03)	2.16 (1.17 to 4.11)	2.08 (1.18 to 3.78)
<i>Patient/resident facing</i>				
Patient-facing	4.06 (2.37 to 6.72)	4.73 (2.38 to 9.29)	3.76 (1.92 to 7.02)	4.24 (2.47 to 6.99)
Resident-facing	2.35 (0.85 to 5.27)	1.79 (0.57 to 5.06)	0.91 (0.19 to 3.17)	2.39 (0.88 to 5.52)
<i>Ethnicity</i>				
Asian	1.89 (0.87 to 3.64)	1.56 (0.50 to 4.47)	1.45 (0.56 to 3.30)	
Black	1.04 (0.28 to 3.07)	1.38 (0.32 to 5.46)	1.30 (0.31 to 3.97)	
Mixed	0.46 (0.09 to 1.84)	0.54 (0.09 to 2.46)	0.49 (0.09 to 1.98)	
Other	7.5 (2.86 to 16.50)	3.41 (0.74 to 13.09)	6.59 (2.14 to 16.07)	
<i>Household size</i>				
2	0.62 (0.35 to 1.09)	0.64 (0.32 to 1.30)	0.47 (0.24 to 0.91)	
3	1.51 (0.83 to 2.73)	1.28 (0.59 to 2.88)	1.61 (0.83 to 3.11)	
4	1.36 (0.68 to 2.63)	1.38 (0.59 to 3.36)	1.58 (0.76 to 3.21)	
5 or more	1.25 (0.51 to 2.88)	1.39 (0.47 to 4.10)	1.36 (0.49 to 3.52)	
<i>Number of children in household</i>				
1	1.01 (0.59 to 1.73)	1.12 (0.52 to 2.39)	0.80 (0.43 to 1.46)	
2	0.44 (0.20 to 0.94)	0.52 (0.19 to 1.38)	0.40 (0.16 to 0.89)	
3 or more	1.65 (0.64 to 4.17)	1.61 (0.46 to 5.19)	1.38 (0.47 to 3.83)	
<i>Contact with hospital</i>				
Yes, I have			2.18 (1.09 to 4.18)	
No I haven't, but someone else in my household has			1.99 (0.86 to 4.13)	
<i>Contact with care home</i>				
Yes, I have			0.77 (0.18 to 2.79)	
No I haven't, but someone else in my household has			0.48 (0.10 to 1.88)	

8 STROBE checklist

STROBE Statement—checklist of items that should be included in reports of observational studies

	Item No.	Recommendation	Page No.
Title and abstract	1	(a) Indicate the study’s design with a commonly used term in the title or the abstract	2
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	2
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	4
Objectives	3	State specific objectives, including any prespecified hypotheses	4
Methods			
Study design	4	Present key elements of study design early in the paper	4-5
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	4-5
Participants	6	(a) Cohort study—Give the eligibility criteria, and the sources and methods of selection of participants. Describe methods of follow-up	4-5
		Case-control study—Give the eligibility criteria, and the sources and methods of case ascertainment and control selection. Give the rationale for the choice of cases and controls	
		Cross-sectional study—Give the eligibility criteria, and the sources and methods of selection of participants	
		(b) Cohort study—For matched studies, give matching criteria and number of exposed and unexposed	
		Case-control study—For matched studies, give matching criteria and the number of controls per case	
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	5-7
Data sources/ measurement	8*	For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group	4-7
Bias	9	Describe any efforts to address potential sources of bias	6-7
Study size	10	Explain how the study size was arrived at	4-5

Continued on next page

Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why	5-6
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	5-7
		(b) Describe any methods used to examine subgroups and interactions	6-7
		(c) Explain how missing data were addressed	5-6
		(d) Cohort study—If applicable, explain how loss to follow-up was addressed Case-control study—If applicable, explain how matching of cases and controls was addressed Cross-sectional study—If applicable, describe analytical methods taking account of sampling strategy	NA
		(e) Describe any sensitivity analyses	6-7
Results			
Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed	7
		(b) Give reasons for non-participation at each stage	NA
		(c) Consider use of a flow diagram	NA
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders	Figure S3, Table S2
		(b) Indicate number of participants with missing data for each variable of interest	5-6
		(c) Cohort study—Summarise follow-up time (eg, average and total amount)	7
Outcome data	15*	Cohort study—Report numbers of outcome events or summary measures over time	7, Figure 1
		Case-control study—Report numbers in each exposure category, or summary measures of exposure	
		Cross-sectional study—Report numbers of outcome events or summary measures	7
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and why they were included	7-8, Table S3
		(b) Report category boundaries when continuous variables were categorized	Table S3
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period	NA

Continued on next page

Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses	7-8
Discussion			
Key results	18	Summarise key results with reference to study objectives	9
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias	10-11
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence	11
Generalisability	21	Discuss the generalisability (external validity) of the study results	9-11
Other information			
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based	

*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at <http://www.plosmedicine.org/>, Annals of Internal Medicine at <http://www.annals.org/>, and Epidemiology at <http://www.epidem.com/>). Information on the STROBE Initiative is available at www.strobe-statement.org.