Predictors of adverse outcome in the first and second waves of the COVID-19 pandemic: results from a UK centre

Christopher A. Martin*, Daniel Pan†, George Hills†, Deborah Modha, Prashanth Patel, Laura J. Gray, David R. Jenkins, Linda Barton, William Jones, Nigel J. Brunskill, Pranab Haldar, Kamlesh Khunti† and Manish Pareek†

Abstract

Background/Aims: Data concerning differences in demographics/disease severity between the first and second waves of COVID-19 are limited. We aimed to examine prognosis in patients presenting to hospital with COVID-19 amongst different ethnic groups between the first and second waves in the UK.

Methods: In this retrospective cohort study, we included 1763 patients presenting to a regional hospital centre in Leicester (UK) and compared those in the first (n = 956) and second (n = 807) waves. Admission National Early Warning Scores, mechanical ventilation and mortality rate were lower in the second wave compared with the first.

Results: Thirty-day mortality risk in second wave patients was approximately half that of first wave patients (adjusted hazard ratio [aHR] 0.55, 95% confidence interval [CI] 0.40–0.75). In the second wave, Black patients were at higher risk of 30-day mortality than White patients (4.73, 1.56–14.3).

Conclusion: We found that disproportionately higher risks of death in patients from ethnic minority groups were not equivalent across consecutive waves of the pandemic. This suggests that risk factors for death in those from ethnic minority groups are malleable and potentially reversible. Our findings need urgent investigation in larger studies.

Keywords: COVID-19, ethnicity, mortality, SARS-CoV-2

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Introduction

The COVID-19 pandemic has caused significant global morbidity and mortality and has disproportionately affected ethnic minority groups. In the absence of widespread immunity, many countries will continue to suffer surges and/or waves of infection, in association with the implementation and release of lockdown measures. Ethnic minority groups have been disproportionately affected by COVID-19 – suffering increased infection, hospitalisation and death during the first wave of the pandemic. Multiple reasons exist for this, but it is thought that the disproportionate risk is mainly attributable to a higher risk of infection, from living in deprived areas, working in high-exposure or frontline occupations and living in larger, multigenerational households compared with White groups. Limited evidence suggests that mortality in the second wave may be lower than that of the first, but it is not known whether this is true for all ethnic groups.

To address this, we compared the demographics and clinical outcomes of patients with COVID-19 admitted to a large UK centre serving a multiethnic population during the first and second waves of the current pandemic. We aimed to estimate the effect of ethnicity as an important determinant of mortality in both waves – and hypothesised that ethnicity was related to mortality in at least one wave of the pandemic.
Molecular testing was made available for a nationwide policy of universal testing for all presence of symptoms during the first wave; however, Testing for SARS-CoV-2 was based on the presence of symptoms during the first wave; however, a nationwide policy of universal testing for all emergency admissions was adopted from 27 April 2020. Molecular testing was made available for hospitalised patients since the start of the first cases of COVID-19 in the United Kingdom, on 27 January 2020.

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We extracted data from the electronic hospital record concerning age, sex, self-reported ethnicity, postcode, comorbidities (see Supplemental Table 1), NEWS 2 on admission, admission/discharge date, highest supplementary oxygen flow rate during admission, dexamethasone use, requirement for mechanical ventilation and date of death. Ethnicity was categorised as White, South Asian, Black and Other (see Supplemental Table 2). Postcode was used to derive Index of Multiple Deprivation (IMD).

Continuous variables were summarised as median [interquartile range (IQR)] and categorical variables as count (%). Comparison was by Wilcoxon rank-sum test for continuous variables and chi-square test for categorical variables.

We established survival time in days by calculating the difference between date of positive swab and date of data extraction or death and used univariable and multivariable Cox regression to establish factors associated with all-cause mortality within 30 days of COVID-19 diagnosis. Multiple imputation was used to replace missing data in all models fitted, and the multiple imputation model included all variables for those being imputed (for further details, see Supplementary information). To investigate the effects of multiple imputation on the results, we conducted a sensitivity analysis using only complete cases. All analyses were conducted using Stata (StataCorp. 2019). Values of p < 0.05 were considered statistically significant.

Table 1 shows a description of the cohort stratified by pandemic wave. A total of 1763 patients were included in the final analysis, 956 (54.2%) from the first wave and 807 (45.8%) from the second wave. There were no significant demographic/comorbidity differences between the first and second wave cohorts.

Markers of COVID-19 severity and duration of hospital stay stratified by ethnicity are shown in Table 2. Admission NEWS was significantly higher in the first wave compared with the second wave in those of White and South Asian ethnicity. The proportion of deaths after diagnosis with COVID-19 was higher in the first wave than in the second wave. This was observed in White and South Asian patients (26.1% versus 10.8%, p < 0.001 and 22.1% versus 10.9%, p = 0.001, respectively) but not in Black patients (16.3% versus 15.4%, p = 0.922). When those who died and those who were not discharged at the point of extraction were excluded, median duration of hospital stay was longer in the second wave compared with the first (4 days, IQR 1–10 versus 5 days IQR 1–9.5, p = 0.016).
versus 19 (9–27), p < 0.0001]. Factors associated with increased risk of 30-day mortality were increasing age, male sex, presence of diabetes and admission NEWS. Compared with those in the first wave, those in the second wave had around half the risk of 30-day mortality [adjusted hazard ratio (aHR) 0.55, 95% confidence interval (CI) 0.40–0.75].

| Variable                        | Total n = 1763 | First Wave (1 March–28 April) n = 956 | Second Wave (1 September–15 November) n = 807 |
|---------------------------------|---------------|--------------------------------------|-----------------------------------------------|
| Age in years, med (IQR)        | 66 [52–79]    | 66 [52–78.5]                         | 67 [51–79]                                    |
| Sex, n (%)                     |               |                                      |                                               |
| Female                         | 759 (43.0)    | 425 (44.5)                           | 334 (41.4)                                    |
| Male                           | 1004 (57.0)   | 531 (55.5)                           | 473 (58.6)                                    |
| Ethnicity, n (%)               |               |                                      |                                               |
| White                           | 1096 (62.2)   | 605 (63.3)                           | 491 (60.8)                                    |
| South Asian                    | 501 (28.4)    | 263 (27.5)                           | 238 (29.5)                                    |
| Black                           | 69 (3.9)      | 43 (4.5)                             | 26 (3.2)                                      |
| Other                           | 97 (5.5)      | 45 (4.7)                             | 52 (6.5)                                      |
| IMD Quintile, n (%)            |               |                                      |                                               |
| 1 (most deprived)              | 329 (18.6)    | 186 (19.5)                           | 143 (17.7)                                    |
| 2                               | 444 (25.2)    | 254 (26.6)                           | 190 (23.5)                                    |
| 3                               | 308 (17.5)    | 150 (15.7)                           | 158 (19.6)                                    |
| 4                               | 355 (20.1)    | 187 (19.6)                           | 168 (20.8)                                    |
| 5 (least deprived)             | 314 (17.8)    | 178 (18.5)                           | 136 (16.9)                                    |
| Missing                         | 13 (0.8)      | 1 (0.1)                              | 12 (1.5)                                      |
| Comorbidity type, n (%)        |               |                                      |                                               |
| Hypertension                    | 525 (29.8)    | 277 (29.0)                           | 248 (30.7)                                    |
| Other cardiovascular            | 194 (11.0)    | 94 (9.8)                             | 100 (12.4)                                    |
| Cerebrovascular                 | 93 (5.3)      | 52 (5.4)                             | 41 (5.1)                                      |
| Respiratory                     | 242 (13.7)    | 133 (13.9)                           | 109 (13.5)                                    |
| Diabetes                        | 301 (17.0)    | 167 (17.5)                           | 134 (16.6)                                    |
| Comorbidity number, n (%)      |               |                                      |                                               |
| 0                               | 881 (50.0)    | 466 (48.7)                           | 415 (51.4)                                    |
| 1                               | 325 (18.4)    | 193 (20.2)                           | 132 (16.4)                                    |
| ≥2                              | 557 (31.6)    | 297 (31.1)                           | 260 (32.2)                                    |

IMD, Index of Multiple Deprivation; IQR, interquartile range; med, median
Table 2. Markers of severity and outcome data stratified by pandemic wave and ethnicity.

| Variable | First wave (n = 956) | Second wave (n = 807) |
|----------|----------------------|-----------------------|
|          | Total | White | South Asian | Black | Other | Total | White | South Asian | Black | Other |
| NEWS score on admission, med [IQR] | 3 [1–4.5]* | 2 [1–4]* | 3 [1–5]* | 3 [1–5] | 3 [1–7] | 2 [0–4]* | 2 [0–4]* | 2 [0–4]* | 2 [0–4] | 0 [1–4] |
| Missing, n (%) | 20 (2.1) | 10 (1.7) | 5 (1.9) | 1 (2.3) | 4 (8.9) | 4 (0.5) | 1 (0.2) | 3 (1.3) | 0 (0.0) | 0 (0.0) |
| Maximum oxygen flow rate received during admission [L/min], med [IQR] | 2 [0–10]* | 3 [0–10]* | 2 [0–8] | 2 [0–4] | 0 [0–4] | 2 [0–6]* | 2 [0–6]* | 2 [0–8] | 0 [0–8] | 0 [0–3] |
| Not mechanically ventilated | 886 (92.7%) | 571 (94.4%) | 238 (90.5%) | 39 (90.7%) | 38 (84.4%) | 778 (96.5%) | 479 (97.6%) | 226 (95.0%) | 25 (96.2%) | 48 (92.3%) |
| Mechanically ventilated | 70 (7.3%)† | 34 (5.6%)† | 25 (9.5%) | 4 (9.3%) | 7 (15.6%) | 29 (3.6%)† | 12 (2.4%)† | 12 (5.0%) | 1 (3.9%) | 4 (7.7%) |
| Survived | 728 (76.2%) | 447 (73.9%) | 205 (78.0%) | 36 (83.7%) | 40 (88.9%) | 723 (89.6%) | 438 (89.2%) | 212 (89.1%) | 22 (84.6%) | 51 (98.1%) |
| Died | 228 (23.9%)† | 158 (26.1%)† | 58 (22.1%)† | 7 (16.3%) | 5 (11.1%) | 84 (10.4%)† | 53 (10.8%)† | 26 (10.9%)† | 4 (15.4%) | 1 (1.9%) |

Excluding those not discharged or dead at the time of data extraction

| Variable | First wave (n = 576) | Second wave (n = 492) |
|----------|----------------------|-----------------------|
| Not mechanically ventilated | 760 (95.7%) | 482 (96.2%) | 211 (95.1%) | 36 (94.7%) | 31 (93.9%) | 562 (97.6%) | 332 (97.9%) | 180 (97.3%) | 19 (95.0%) | 31 (96.9%) |
| Mechanically ventilated | 34 (4.3%) | 19 (3.8%) | 11 (5.0%) | 2 (5.3%) | 2 (6.1%) | 14 (2.4%) | 7 (2.1%) | 5 (2.7%) | 1 (5.0%) | 1 (3.1%) |
| Survived | 566 (71.3%) | 343 (68.5%) | 164 (73.9%) | 31 (81.6%) | 28 (84.9%) | 492 (85.4%) | 286 (84.4%) | 159 (86.0%) | 16 (80.0%) | 31 (96.9%) |
| Died | 228 (28.7%)† | 158 (31.5%)† | 58 (26.1%)† | 7 (18.4%) | 5 (15.2%) | 84 (14.6%)† | 53 (15.6%)† | 26 (14.1%)† | 4 (20.0%) | 1 (3.1%) |

Excluding those who died and those not discharged at the time of data extraction

| Variable | First wave (n = 556) | Second wave (n = 492) |
|----------|----------------------|-----------------------|
| Length of stay in days, med [IQR] | 4 [1–10]* | 5 [1–10] | 3 [1–6]* | 3 [0–7] | 1 [0–6.5] | 5 [1–9.5]* | 5 [2–11] | 4 [1–9]* | 4 [1.5–6] | 3 [1–8] |

IQR, interquartile range; NEWS, National Early Warning Score.  
*Wilcoxon rank-sum p < 0.05.  
†Chi-square p < 0.05.
Table 3. Multivariable Cox proportional hazards model for survival at 30 days from COVID-19 diagnosis in the first and second pandemic waves.

| Variable                  | First wave (n = 956) | Second wave (n = 807) |
|---------------------------|----------------------|-----------------------|
|                           | aHR (95% CI)         | p value               | aHR (95% CI)         | p value               |
| Age                       | 1.05 (1.04–1.07)     | <0.001                | 1.07 (1.05–1.10)     | <0.001                |
| Sex                       |                      |                       |                      |                       |
| Female                    | Ref                  | –                     | Ref                  | –                     |
| Male                      | 1.50 (1.13–2.00)     | 0.005                 | 2.22 (1.29–3.81)     | 0.004                 |
| Ethnicity                 |                      |                       |                      |                       |
| White                     | Ref                  | –                     | Ref                  | –                     |
| South Asian               | 1.04 (0.73–1.49)     | 0.81                  | 0.90 (0.52–1.56)     | 0.71                  |
| Black                     | 0.88 (0.38–2.04)     | 0.76                  | 4.73 (1.56–14.34)    | 0.006                 |
| Other                     | 0.71 (0.28–1.78)     | 0.47                  | 0.50 (0.06–3.87)     | 0.51                  |
| IMD quintile              |                      |                       |                      |                       |
| 1 (least deprived)        | Ref                  | –                     | Ref                  | –                     |
| 2                         | 1.20 (0.79–1.83)     | 0.38                  | 1.00 (0.49–2.05)     | 1.00                  |
| 3                         | 0.98 (0.60–1.59)     | 0.92                  | 1.20 (0.58–2.49)     | 0.62                  |
| 4                         | 0.89 (0.56–1.41)     | 0.62                  | 0.59 (0.27–1.29)     | 0.19                  |
| 5 (most deprived)         | 0.91 (0.58–1.43)     | 0.69                  | 0.81 (0.35–1.86)     | 0.62                  |
| Comorbidity type          |                      |                       |                      |                       |
| Hypertension              | 0.84 (0.58–1.22)     | 0.36                  | 1.96 (1.00–3.82)     | 0.05                  |
| Other cardiovascular      | 0.78 (0.51–1.21)     | 0.26                  | 1.19 (0.68–2.09)     | 0.55                  |
| Stroke                    | 1.25 (0.76–2.06)     | 0.38                  | 0.71 (0.24–2.06)     | 0.53                  |
| Respiratory disease       | 1.00 (0.64–1.58)     | 0.99                  | 1.49 (0.80–2.75)     | 0.21                  |
| Diabetes                  | 1.43 (0.99–2.08)     | 0.06                  | 2.03 (1.15–3.59)     | 0.02                  |
| Comorbidity number        |                      |                       |                      |                       |
| 0                         | Ref                  | –                     | Ref                  | –                     |
| 1                         | 1.46 (0.95–2.24)     | 0.08                  | 2.58 (1.07–6.23)     | 0.03                  |
| ≥2                        | 1.24 (0.69–2.22)     | 0.48                  | 1.23 (0.41–3.68)     | 0.71                  |
| NEWS score on admission   | 1.28 (1.23–1.33)     | <0.001                | 1.34 (1.23–1.46)     | <0.001                |
| Treatment                 |                      |                       |                      |                       |
| Did not receive dexamethasone | Ref                  | –                     | Ref                  | –                     |
| Received dexamethasone    | 0.14 (0.02–0.99)     | 0.049                 | 1.55 (0.95–2.51)     | 0.08                  |

aHR, adjusted hazard ratio; CI, confidence interval; IMD, Index of Multiple Deprivation; NEWS, National Early Warning Score.
Separate multivariable Cox regression models for survival to 30 days in the first and second waves are shown in Table 3. In both pandemic waves, age, male sex and admission NEWS score are significantly associated with a higher risk of 30-day mortality. In the first wave, ethnicity did not impact upon 30-day mortality; however, in the second wave, as compared with White patients, those of Black ethnicity were almost 5 times as likely to die within 30 days (aHR 4.72, 95% CI 1.56–14.43). When the waves were analysed together, the interaction between pandemic wave and Black ethnicity was not significant (aHR 3.25, 95% CI 0.87–12.07, \( p = 0.079 \)). When only complete cases are analysed, significant findings do not change.

**Discussion**

In this study within an ethnically diverse cohort, we found a number of novel observations. First, there were no significant differences in demographic characteristics between patients in the first and second COVID-19 pandemic waves and that markers of severe COVID-19 (NEWS, requirement for mechanical ventilation and COVID-19-associated mortality) were lower in the second wave of the pandemic compared with the first. We found those of Black ethnicity were at higher risk of 30-day mortality than those of White ethnicity in the second wave.

COVID-19 case fatality rates have fallen in the second wave in 43 of the 53 countries with the highest total COVID-19-related deaths.\(^4\) Foremost among the explanations for this observation is the increased availability of testing in the second wave which may increase the number of paucisymptomatic/asymptomatic individuals who are tested. In our study, the change in SARS-CoV-2 testing policy from symptom-driven testing to universal testing would be expected to pick up bystander infection in those attending hospital for reasons unrelated to COVID-19 and could generate bias towards a less symptomatic sample in the second wave. Other suggested mechanisms include the changing demographic profile of COVID-19 cases over time, with a smaller proportion of elderly patients (a group at high risk of adverse outcome from COVID-19) in the second pandemic wave with more people shielding,\(^8,9\) although we found no such differences in our dataset. In addition, in contrast to the early part of the first wave, there is an effective pharmacological treatment for severe COVID-19 (dexamethasone).\(^10\) Clearly a reduction in COVID-19 mortality is desirable; however, increased survival from COVID-19 in the second wave may contribute to pressure on the healthcare system through increased hospital bed occupancy. Indeed our analysis shows that duration of hospital stay was longer in the second wave than in the first. Nonpharmaceutical interventions, such as mandating masks and lockdowns, may have also affected the number of patients infected and thus in turn, the number of deaths, but the city of Leicester remained in full lockdown (stay-at-home except for essential purchases, remote work, one exercise per day, cancellation of public gatherings and social events and no travel abroad) from 23 March 2020 to the end of the study.

We observed that mortality rates in the second pandemic wave may not be equal across ethnic groups. In our second wave cohort, those of Black ethnicity were at higher risk of 30-day mortality than White individuals (an effect not seen in the first wave cohort). These findings are opposite to a national UK study by Mathur et al.,\(^11\) which found that compared with the first wave, risks for death were attenuated for those from Black ethnic groups compared with White ethnic groups. Ethnicity is a social determinant of health. Therefore, it is likely that the differences in risks seen in our study reflect the number of infections that occurred in the local community and consequently, the number of people who present to hospital. A disproportionate number of infections in those of Black ethnicity may reflect socioeconomic factors, such as occupation, which prevent them from being able to stay at home on a second lockdown. Leicester is an especially ethnically diverse part of the United Kingdom, where approximately 30% of the population is born outside of Europe and North America, and individuals from the Indian subcontinent alone make up 15% of the population.\(^12\) However, we recommend cautious interpretation of this finding, due to the small number of Black patients in our sample and the nonsignificant interaction between ethnicity and pandemic wave. Nonetheless, given the higher risk of SARS-CoV-2 infection and adverse outcome from COVID-19 in ethnic minority groups compared with those of White ethnicity,\(^1,5,13,14\) this observation requires urgent follow-up by larger studies with a greater proportion of Black participants and emphasises the importance of targeted interventions such as tailored public health messaging in these groups.
This study has limitations. Data are from a single centre and our data do not allow us to control for the change in testing policy between waves. We found a shorter median follow-up time in second wave patients. This is likely due to the fact that while the first wave data capture peak COVID-19 hospital admissions in early April with data extraction a month later, COVID-19 admissions continued to increase after data extraction for the second wave, meaning that a greater proportion of the second wave cohort were admitted close to the point of data extraction when compared with the first wave. However, we conducted an exploratory analysis excluding those admitted after the first wave ‘peak’ and still demonstrated a significant protective effect of being admitted during the second wave compared with the first. Mortality rates and severity of COVID-19 may change in subsequent waves of the pandemic, particularly with the emergence of the new variants of SARS-CoV-2. We only collected routinely available variables that were recorded within our clinical systems and as such, we do not have granular information – such as occupation – which may have affected their risk of infection. No patients in the United Kingdom were vaccinated at the time of study as part of national policy – these findings may be different now that a significant proportion of the UK population has been vaccinated. The emergence of novel variants, especially during the period of the second wave, may have had an impact on our study outcomes – however, genomic sequencing is not routinely performed in all our hospitals. Despite these limitations, our findings, particularly those related to ethnicity, must be investigated by larger studies to ensure that the inequalities of the first wave are not allowed to widen.

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Author contributions
Christopher Martin: Conceptualisation; Data curation; Formal analysis; Investigation; Methodology; Project administration; Writing – original draft; Writing – review & editing
Daniel Pan: Conceptualisation; Data curation; Formal analysis; Investigation; Methodology; Writing – original draft; Writing – review & editing
George Hills: Conceptualisation; Data curation; Investigation; Writing – review & editing
Deborah Modha: Conceptualisation; Writing – review & editing
Prashanth Patel: Conceptualisation; Data curation; Investigation; Writing – review & editing
Laura Gray: Data curation; Formal analysis; Writing – review & editing
David Jenkins: Conceptualisation; Writing – review & editing
Linda Barton: Investigation; Writing – review & editing
William Jones: Data curation; Writing – review & editing
Nigel Brunskill: Methodology; Writing – review & editing
Pranab Haldar: Conceptualisation; Writing – review & editing
Kamlesh Khunti: Conceptualisation; Writing – review & editing
Manish Pareek: Conceptualisation; Formal analysis; Methodology; Project administration; Supervision; Writing – review & editing

Conflict of interest statement
The authors declared the following potential conflicts of interest with respect to the research, authorship and/or publication of this article: KK is the Director of the University of Leicester Centre for Black Minority Ethnic Health, Trustee of the South Asian Health Foundation and Chair of the Ethnicity Subgroup of SAGE. MP reports grants and personal fees from Gilead Sciences and personal fees from QIAGEN, outside the submitted work.

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Ethics statement
Ethical approval was not required after consultation with the National Health Service (NHS) Health Research Authority decision aid and the Caldicott Guardian and the study was approved as a service evaluation/audit (UHL10579).

ORCID iD
George Hills https://orcid.org/0000-0003-0172-6156

Supplemental material
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References
1. Sze S, Pan D, Nevill CR, et al. Ethnicity and clinical outcomes in COVID-19: a systematic review and meta-analysis. EClinicalMedicine 2020; 29: 100630.
2. Ferguson NM, Laydon D, Nedjati-Gilani G, et al. Report 9: impact of non-pharmaceutical interventions (NPIs) to reduce COVID-19 mortality and healthcare demand, https://www.imperial.ac.uk/media/imperial-college/medicine/sph/ide/gida-fellowships/Imperial-College-COVID19-NPI-modelling-16-03-2020.pdf (2020, accessed 21 December 2020).
3. Pan D, Martin CA, Nazareth J, et al. Ethnic disparities in COVID-19: increased risk of infection or severe disease? Lancet 2021; 398: 389–390.
4. Fan G, Yang Z, Lin Q, et al. Decreased case fatality rate of COVID-19 in the second wave: a study in 53 countries or regions. Transbound Emerg Dis 2020; 68: 213–215.
5. Martin CA, Jenkins DR, Minhas JS, et al. Socio-demographic heterogeneity in the prevalence of COVID-19 during lockdown is associated with ethnicity and household size: results from an observational cohort study. EClinicalMedicine 2020; 25: 100466.
6. Thomas R. All hospital emergency patients to be tested for coronavirus, https://www.hsj.co.uk/coronavirus/all-hospital-emergency-patients-to-be-tested-for-coronavirus/7027526.article (2020, accessed 6 January 2021).
7. Royal College of Physicians. National Early Warning Score (NEWS) 2: Standardising the assessment of acute-illness severity in the NHS. Updated report of a working party. London: RCP, 2017.
8. Burgess S, Smith D, Kenyon JC, et al. Lightening the viral load to lessen covid-19 severity. BMJ 2020; 371: m4763.
9. Venkatesan P. The changing demographics of COVID-19. Lancet Respir Med 2020; 8: e95.
10. Dexamethasone in Hospitalized Patients with Covid-19— preliminary report. N Engl J Med 2021; 384: 693–704.
11. Mathur R, Rentsh CT, Morton CE, et al. Ethnic differences in SARS-CoV-2 infection and COVID-19-related hospitalisation, intensive care unit admission, and death in 17 million adults in England: an observational cohort study using the OpenSAFELY platform. Lancet 2021; 397: 1711–1724.
12. Pareek M, Eborall HC, Wobi F, et al. Community-based testing of migrants for infectious diseases (COMBAT-ID): impact, acceptability and cost-effectiveness of identifying infectious diseases among migrants in primary care: protocol for an interrupted time-series, qualitative and health economic analysis. BMJ Open 2019; 9: e029188.
13. Martin CA, Patel P, Goss C, et al. Demographic and occupational determinants of anti-SARS-CoV-2 IgG seropositivity in hospital staff. J Public Health. Epub ahead of print 16 November 2020. DOI: 10.1093/pubmed/fdaa199.
14. Williamson EJ, Walker AJ, Bhaskaran K, et al. OpenSAFELY: factors associated with COVID-19 death in 17 million patients. Nature 2020; 584: 430–436.