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Letter to the Editor

Observations of SARS-CoV-2 variant of concern B.1.1.7 at the UK's largest hospital trust


To the editor,

The outbreak of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) has placed unprecedented strain on health-care services, leading to 134,125,854 cases and more than 2,905,412 deaths worldwide, as of 09/04/21.¹ Globally it has been observed that more transmissible variants develop.² Since October, one such variant of concern (VOC) (SARS-CoV-2 20B/GR clade; VOC-202012/01; lineage B.1.1.7) has been described in the UK.³ B.1.1.7 appears to have substantially increased transmissibility compared to other variants and has quickly become the dominant variant in much of the UK.⁴ A report by the Office for National Statistics suggested that people infected with B.1.1.7 are more likely to have a cough, sore throat, fatigue, or myalgia than those infected with other variants.⁵ Furthermore, preliminary data indicate that infection with VOC B.1.1.7 is associated with an increased risk of death compared with other variants.⁴ At University Hospitals Birmingham (UHB) NHS Foundation Trust, we undertake whole genome sequencing on a substantial proportion of our SARS-CoV-2 positive samples. Here, we report our observations and outcomes of VOC-202012/01 B.1.1.7 infected patients admitted to UHB during December 2020.

UHB is one of the largest Trusts in the UK, covering 4 NHS hospital sites, treating over 2.2 million patients per year and housing one of the largest single critical care units in the world. UHB's patient testing programme detects SARS-CoV-2 RNA from nasopharyngeal swabs/sputum using molecular platforms including Panther, Hologic, M2000, Abbott and Cepheid, GeneXpert., Positive samples were collected from new admissions from the 15th to 31st December 2020 and then subjected to whole-genome sequencing at the University of Birmingham. Lineages were assigned with Pangolin.⁶ To understand the impact of the VOCs: VOC-202012/01 B.1.1.7 and B.1.351, clinical details for patients whose samples resulted in an assignment of lineage were collected. This included: hospital length of stay, ethnicity, age, sex, critical care admission, critical care length of stay, treatment given for COVID-19 (dexamethasone or remdesivir), oxygen and ventilatory support, death, comorbidities and presenting symptoms. Multiple logistic, linear and quasipoisson regression models were used to examine the role of variants and patient characteristics such as age and sex and clinical outcomes. Statistical modelling was performed using R.⁷

SARS-2 positive samples from 152 patients were assigned a lineage following whole genome sequencing - 79 belonging to the B.1.1.7 lineage, 1 to the South African VOC B.1.351 and 72 to other variants (Fig. 1). During the study period, 52.3% of positive samples were the UK 'Kent' variant (B.1.1.7), 40.2% the 'Spanish' variant (B.1.177), and one South African variant (B.1.351) (Fig. 1). During December, there was a steady increase in the proportion of

B.1.1.7 detected at UHB (18% and 63% in weeks 50 and 52 respectively) (Fig. 1). A quasipoisson regression model fitted to the number of patient isolates with COVID-19 demonstrated a significant interaction between collection date and B.1.1.7, leading to a significant increase in the recovery of B.1.1.7 (but not other variants) as time progressed ($p=0.0077$). The following non-significant observations were made between the other variants vs B.1.1.7 and B.1.351 (Table 1); for the former, total length of stay was longer, length of stay on critical care was shorter, the proportion of patients on critical care was lower, the proportion of patients who died was higher and patients were older. A multiple logistic regression suggests that VOCs are significantly associated with the probability of developing a cough when compared to a similar patient with one of the other variants, with an odds ratio of 2.19 ($p=0.0443$).

UHB has seen the largest number of COVID-19 cases in the country compared to other hospital Trusts: between 01/03/20–09/04/21, the Trust had 13,677 cases of COVID-19. In our small data set, B.1.1.7 was associated with younger age, an increased proportion of patients being admitted to critical care for longer periods but the association wasn't statistically significant. Whilst numbers of patients were relatively low, no increase in mortality was observed. Interestingly, significantly more patients infected with B.1.1.7 presented with a cough which could be a factor contributing to the increased transmission observed with this variant. This concurs with the report issued by the Office for National Statistics.⁵ The NERVTAG report on B.1.1.7 suggests an increase in transmission of up to 71% in comparison to other circulating variants of SARS-CoV-2.⁴ This is probably why the emergent new variant has contributed to UHB's record number of inpatients, which reached a peak of 1067 on 24th January, with 165 of those patients being on critical care. Gregson et al., (2020) showed that when a patient coughs, the average particle number is higher than that of someone talking, singing or shouting.⁸ This could partly explain why more transmission is seen with B.1.1.7. This is an important observation, as heightened transmissibility of this new variant may result in a more prolonged period of increased incidence within healthcare institutions due to nosocomial spread. Importantly, UHB have seen more outbreaks within the acute setting in January 2021 when the new variant became the dominant strain. Further work is warranted to observe the effect of the new variant on nosocomial outbreaks and the conversion rate of contacts exposed to patients infected with B.1.1.7.

Finally, it will be important to monitor the proportion of new variant cases in hospital settings, both within patients and health-care workers. Technologies such as whole genome sequencing are useful to pick up new variants quickly, especially with the emergence of other VOCs such as B.1.351 in South Africa and P1 in Brazil. In our dataset we were able to identify the South African variant B.1.351 in one of our patients, where normally this would

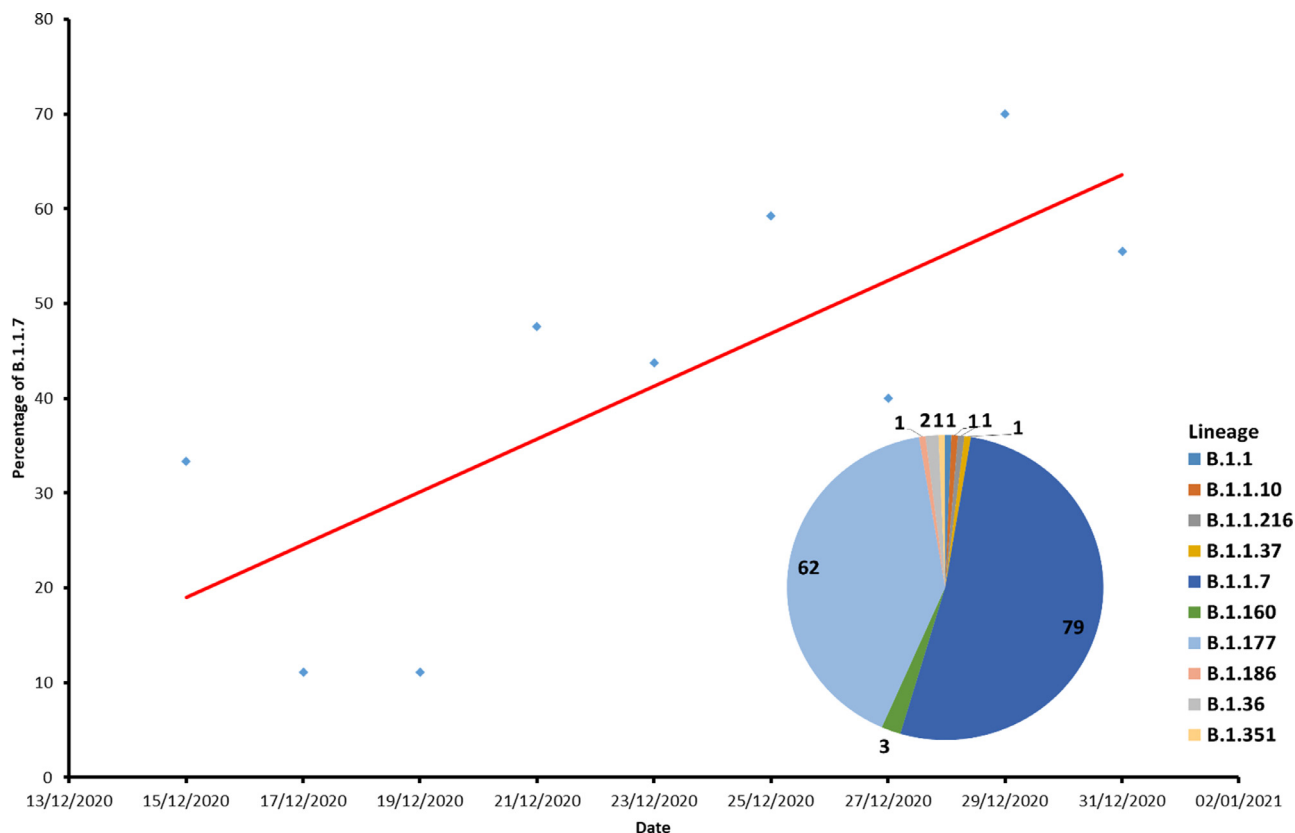


Fig. 1. The pie chart within the graph shows proportion of variants seen at UHB and the graph itself shows the increasing proportion (as a percentage) of B.1.1.7 at UHB during December 2020.

Note: The pie chart within the Figure shows the proportion of variants seen at UHB during December. Numbers in the pie chart represent actual numbers. The line chart shows the percentage increase of B.1.1.7 from 15th – 31st December. The red line in the chart represents a trend line. The blue diamonds represent percent of B.1.1.7 vs other variants.

Table 1
Data of other variants vs VOCs B.1.1.7 & B.1.351.

| Variant | Number | Mean LoS days | Mean critical care LoS days | Proportion patients admitted to critical care | Proportion patients died | Mean age | Proportion of patients with a cough |
|---------|--------|---------------|-----------------------------|---|--------------------------|----------|-------------------------------------|
| Other | 72 | 11.51 | 15.25 | 0.111 | 0.21 | 64.39 | 0.38 |
| VOCs | 80 | 9.56 | 18.93 | 0.19 | 0.18 | 58.2 | 0.59* |

Key:.
LoS – length of stay, * P <0.05.

not have been detected. This has enabled Public Health England (PHE) to undertake contact tracing on this patient and implement surge testing in the community.

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Not applicable.

Consent for publication

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Authors information

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Declaration of Competing Interests

None.

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Mark I. Garvey*

University Hospitals Birmingham NHS Foundation Trust, Queen Elizabeth Hospital Birmingham, Edgbaston, Birmingham, B15 2WB, England

Institute of Microbiology and Infection, The University of Birmingham, Edgbaston, Birmingham, England

Claire McMurray

Institute of Microbiology and Infection, The University of Birmingham, Edgbaston, Birmingham, England

Anna L Casey, Liz Ratcliffe
University Hospitals Birmingham NHS Foundation Trust, Queen Elizabeth Hospital Birmingham, Edgbaston, Birmingham, B15 2WB, England

Joanne Stockton
Institute of Microbiology and Infection, The University of Birmingham, Edgbaston, Birmingham, England

Martyn A C Wilkinson, Elisabeth Holden, Husam Osman
University Hospitals Birmingham NHS Foundation Trust, Queen Elizabeth Hospital Birmingham, Edgbaston, Birmingham, B15 2WB, England

Nick J Loman
Institute of Microbiology and Infection, The University of Birmingham, Edgbaston, Birmingham, England

*Corresponding author.

E-mail addresses: mark.garvey@uhb.nhs.uk, m.i.garvey@bham.ac.uk (M.I. Garvey), C.L.McMurray@bham.ac.uk (C. McMurray), anna.casey@uhb.nhs.uk (A.L. Casey), liz.Ratcliffe@uhb.nhs.uk (L. Ratcliffe), J.D.Stockton@bham.ac.uk (J. Stockton), martyn.wilkinson@uhb.nhs.uk (M.A.C. Wilkinson), elisabeth.holden@uhb.nhs.uk (E. Holden), Husam.osman@uhb.nhs.uk (H. Osman), n.j.loman@bham.ac.uk (N.J. Loman)