

Protecting and improving the nation's health

COVID-19 vaccine surveillance report Week 26

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Summary

Four coronavirus (COVID-19) vaccines have now been approved for use in the UK. Rigorous clinical trials have been undertaken to understand the immune response, safety profile and efficacy of these vaccines as part of the regulatory process. Ongoing monitoring of the vaccines as they are rolled out in the population is important to continually ensure that clinical and public health guidance on the vaccination programme is built upon the best available evidence.

Public Health England (PHE) works closely with the Medicines and Healthcare Regulatory Agency (MHRA), NHS England, and other government, devolved administration and academic partners to monitor the COVID-19 vaccination programme. Details of the vaccine surveillance strategy are set on the Public Health England page COVID-19: vaccine surveillance strategy (1). As with all vaccines, the safety of COVID-19 vaccines is continuously being monitored by the MHRA. They conclude that overall, the benefits of COVID-19 vaccines outweigh any potential risks (2).

Vaccine effectiveness

Several studies of vaccine effectiveness have been conducted in the UK which indicate that a single dose of either vaccine is between 55 and 70% effective against symptomatic disease, with higher levels of protection against severe disease including hospitalisation and death. Additional protection is seen after a second dose. There is now also evidence from a number of studies that the vaccines are effective at protecting against infection and transmission.

Population impact

The impact of the vaccination programme on the population is assessed by taking into account vaccine coverage, evidence on vaccine effectiveness and the latest COVID-19 disease surveillance indicators. Vaccine coverage tells us about the proportion of the population that have received 1 and 2 doses of COVID-19 vaccines. By 27 June 2021, the overall vaccine uptake in England for dose 1 was 59.9% and 44.1% for dose 2. In line with the programme rollout, coverage is highest in the oldest age groups.

Based on antibody testing of blood donors, 84.2% of the adult population now have antibodies to COVID-19 from either infection or vaccination compared to 14.9% that have antibodies from infection alone. Over 94% of adults aged 40 or older have antibodies from either infection or vaccination. Seropositivity among those aged 30 to 39 has begun to rise over the last few weeks. The latest estimates indicate that the vaccination programme has directly averted over 44,500 hospitalisations. Analysis on the direct and indirect impact of the vaccination programme on infections and mortality,

suggests the vaccination programme has prevented between 6.4 and 7.9 million infections and between 26,000 and 28,000 deaths.

Vaccine effectiveness

Large clinical trials have been undertaken for each of the COVID-19 vaccines approved in the UK which found that they are highly efficacious at preventing symptomatic disease in the populations that were studied. It is important to continue to evaluate the effectiveness of vaccines in the 'real world', as this may differ to clinical trial efficacy. The clinical trials are also performed in order to be able to assess the efficacy of the vaccine against laboratory confirmed symptomatic disease with a relatively short follow up period so that effective vaccines can be introduced as rapidly as possible. Nevertheless, understanding the effectiveness against different outcomes (such as severe disease and onwards transmission), effectiveness in different subgroups of the population and understanding the duration of protection are equally important in decision making around which vaccines should be implemented as the programme evolves, who they should be offered to and whether booster doses are required.

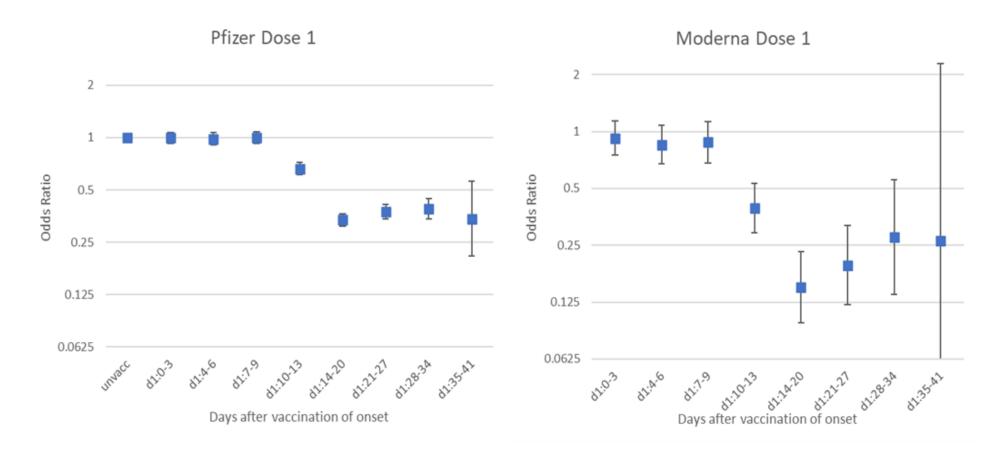
Vaccine effectiveness is estimated by comparing rates of disease in vaccinated individuals to rates in unvaccinated individuals. Below we outline the latest real-world evidence on vaccine effectiveness from studies in UK populations. The majority of this data relates to a period when the main circulating virus was the Alpha variant, emerging data on effectiveness against symptomatic disease with the Delta variant is also summarised below. The findings are also summarised in Tables 3 to 5.

Effectiveness against symptomatic disease

Vaccine effectiveness against symptomatic COVID-19 has been assessed in England based on community testing data linked to vaccination data from the NIMS and from the COVID Infection Survey. Current evidence is primarily from older adults, who were among the earliest group vaccinated. Estimates of vaccine effectiveness range from around 55 to 70% after 1 dose, with little evidence of variation by vaccine or age group (3, 4, 5). Data on 2 doses is indicates effectiveness of around 65 to 90% (3, 6).

Offer of the Pfizer and Moderna mRNA vaccines to adults aged under 40 years began on 10 May 2021. Early estimates of effectiveness of a single dose of either vaccine indicate a vaccine effectiveness of 61% (95% CI: 56 to 66%) after 1 dose of the Pfizer vaccine and 72% (95% CI: 46 to 86%) after 1 dose of the Moderna vaccine. Effectiveness is particularly high 14 to 27 days after vaccination with the Moderna vaccine, however, confidence intervals overlap with the later periods (Figure 1). Both the Alpha and Delta variant were circulating during this period and the changing distribution of variants could affect the results by period after vaccination.

Figure 1: Odds ratios for becoming a case by days after vaccination with the first dose (age 15 to 39 years)



Effectiveness against hospitalisation

Several studies have estimated the effectiveness against hospitalisation in older adults, all of which indicate higher levels of protection against hospitalisation after a single dose than that seen against symptomatic disease, around 75 to 85% after 1 dose of the Pfizer-BioNTech or Oxford-AstraZeneca vaccine (3, 8, 9, 10). Data on VE against hospitalisation with 2 doses for all ages with the Alpha variant is shown in Table 1.

Table 1. Vaccine effectiveness against hospitalisation Alpha variant

Vaccination Status	OR v symptomatic disease	HR vs hospitalisation	VE vs hospitalisation
Any vaccine			
Dose 1	0.5 (0.47-0.52)	0.39 (0.34-0.45)	81% (78-83)
Dose 2	0.11 (0.1-0.13)	0.3 (0.16-0.59)	97% (93-98)
Pfizer			
Dose 1	0.49 (0.44-0.54)	0.38 (0.32-0.45)	82% (78-85)
Dose 2	0.06 (0.05-0.08)	0.3 (0.13-0.67)	98% (96-99)
AstraZeneca			
Dose 1	0.5 (0.47-0.54)	0.42 (0.35-0.51)	79% (74-82)
Dose 2	0.18 (0.15-0.22)	0.32 (0.1-1.02)	94% (81-98)

Effectiveness against mortality

Data is also emerging which suggests high levels of protection against mortality. Studies linking community COVID-19 testing data, vaccination data and mortality data indicate that both the Pfizer-BioNTech and Oxford-AstraZeneca vaccines are around 75 to 80% effective at preventing death with COVID-19 after a single dose (3, 11). Data on 2 doses is shown in Table 2.

Table 2. Vaccine effectiveness against mortality

Vaccination Status	Age group	OR v symptomatic disease	HR vs mortality	VE vs mortality
Pfizer				
D 4	40-64	0.52 (0.48-0.57)	0.53 (0.44- 0.62)	73% (67-77)
Dose 1	65+	0.44 (0.4-0.48)		77% (72-81)
Dose 2	40-64	0.1 (0.09-0.11)	0.25 (0.11-	98% (94-99)
	65+	0.1 (0.08-0.13)	0.56)	98% (94-99)
AstraZeneca				
Dose 1	40-64	0.53 (0.51-0.56)	0.4 (0.32-	79% (73-83)
	65+	0.43 (0.39-0.47)	0.51)	83% (78-86)
Dose 2	40-64	0.26 (0.25-0.28)	0.29 (0.09- 0.91)	92% (76-98)
	65+	0.22 (0.18-0.26)		94% (80-98)

Effectiveness against infection

Although individuals may not develop symptoms of COVID-19 after vaccination, it is possible that they could still be infected with the virus and could transmit to others. Understanding how effective vaccines are at preventing infection is therefore important to predict the likely impact of the vaccination programme on the wider population. In order to estimate vaccine effectiveness against infection, repeat asymptomatic testing of a defined cohort of individuals is required. Studies have now reported on vaccine effectiveness against infection in healthcare workers, care home residents and the general population. With the Pfizer-BioNTech, estimates of effectiveness against infection range from around 55 to 70%, with the Oxford-AstraZeneca vaccine they range from around 60 to 70% (5, 12, 13, 14). Estimates for 2 doses are currently only available for the Pfizer-BioNTech vaccine and indicate effectiveness against infection of 70 to 90% (5,12).

Effectiveness against transmission

As described above, several studies have provided evidence that vaccines are effective at preventing infection. Uninfected individuals cannot transmit; therefore, the vaccines are also effective at preventing transmission. Data from Scotland has also shown that household contacts of vaccinated healthcare workers are at reduced risk of becoming a case, which is in line with the studies on infection (15). There may be additional benefit,

beyond that due to prevention of infection, if some of those individuals who become infected despite vaccination are also at a reduced risk of transmitting (for example, because of reduced duration or level of viral shedding). A household transmission study in England found that household contacts of cases vaccinated with a single dose had approximately 35 to 50% reduced risk of becoming a confirmed case of COVID-19. This study used routine testing data so would only include household contacts that developed symptoms and went on to request a test via pillar 2. It cannot exclude asymptomatic secondary cases or mildly symptomatic cases who chose not to request a COVID-19 test (16).

Table 3. Summary of evidence on vaccine effectiveness against different outcomes (data relate to period when the Alpha variant dominated)

	Vaccine effectiveness			
Outcome	Pfizer-BioNTech		Oxford-AstraZeneca	
	1 dose	2 doses	1 dose	2 doses
Symptomatic disease	55-70%	85-95%	55-70%	70-85%
Hospitalisation	75-85%	90-99%	75-85%	80-99%
Mortality	70-80%	95-99%	75-85%	75-99%
Infection	55-70%	70-90%	60-70%	No data
Transmission (secondary cases)*	45-50%	No data	35-50%	No data

High	Evidence from multiple studies which is consistent
Confidence	and comprehensive
Medium Confidence	Evidence is emerging from a limited number of studies or with a moderately level of uncertainty
Low	Little evidence is available at present and results
Confidence	are inconclusive

^{*} effectiveness in reducing symptomatic secondary cases in households of a symptomatic index case

Vaccine effectiveness against the Delta variant

Analysis of routine testing data up to 13 June 2021, linked to sequencing and S-gene target status has been used to estimate vaccine effectiveness against symptomatic disease using a test negative case control design. Methods and detailed results are available in Effectiveness of COVID-19 vaccines against the Delta variant. After a single dose there was an 14% absolute reduction in vaccine effectiveness against symptomatic disease with Delta compared to Alpha, and a smaller 10% reduction in effectiveness after 2 doses (Table 4).

Table 4. Vaccine effectiveness against symptomatic disease for Alpha and Delta variants

Vaccine Status	Vaccine Effectiveness		
	Alpha	Delta	
Dose 1	49 (46 to 52)	35 (32 to 38)	
Dose 2	89 (87 to 90)	79 (78 to 80)	

Vaccine effectiveness against hospitalisation was estimated by evaluating hospitalisation rates via emergency care among symptomatic confirmed cases using survival analysis. This analysis used available data from linkage of symptomatic cases, 12 April to the 10 June 2021 (updated from the previous analysis to 4 June 2021). Hazard ratios for hospitalisation are combined with odds ratios against symptomatic disease from the test negative case control analysis described above to estimate vaccine effectiveness against hospitalisation. Methods and detailed results are available here (17). Similar vaccine effectiveness against hospitalisation was seen with the Alpha and Delta variants (Table 5).

Table 5. Vaccine effectiveness against hospitalisation for Alpha and Delta variants

Vaccine Status	Vaccine Effectiveness		
	Alpha	Delta	
Dose 1	78 (64 to 87)	80 (69 to 88)	
Dose 2	93 (80 to 97)	96 (91 to 98)	

Population impact

Vaccines typically have both direct effects on those who are vaccinated and indirect effects on the wider population due to a reduced probability that people will come into contact with an infected individual. The overall impact of the vaccination programme may therefore extend beyond that estimated through vaccine effectiveness analysis.

Estimating the impact of a vaccination programme is challenging as there is no completely unaffected control group. Furthermore, the effects of the vaccination programme need to be differentiated from that of other interventions (for example, lockdowns or outbreak control measures), changes in behaviour and any seasonal variation in COVID-19 activity.

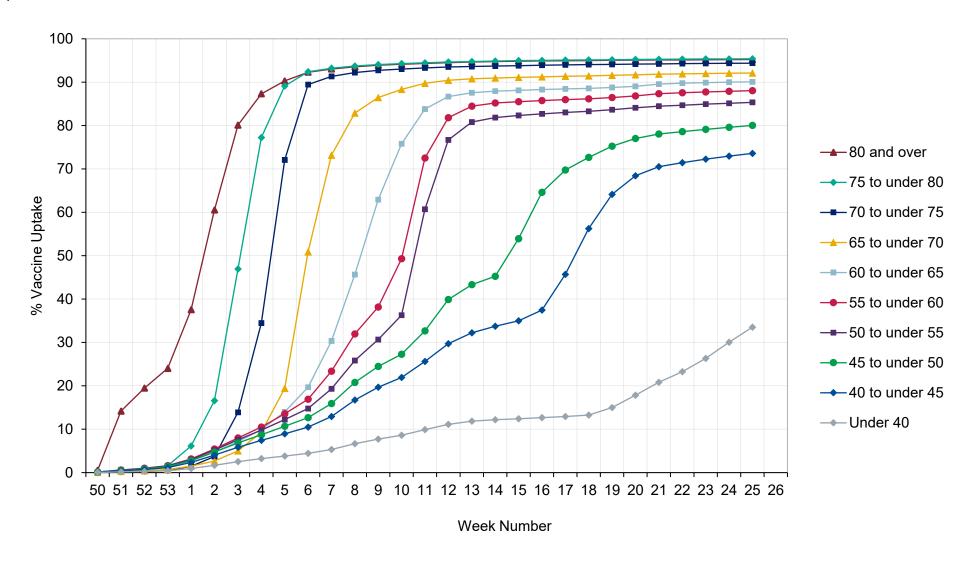
PHE and other government and academic partners monitor the impact of the of the vaccination programme on levels of COVID-19 antibodies in the population and different disease indicators, including hospitalisations and mortality. This is done through population-based testing and through modelling which combines vaccine coverage rates in different populations, estimates of vaccine effectiveness and disease surveillance indicators.

Vaccine Coverage

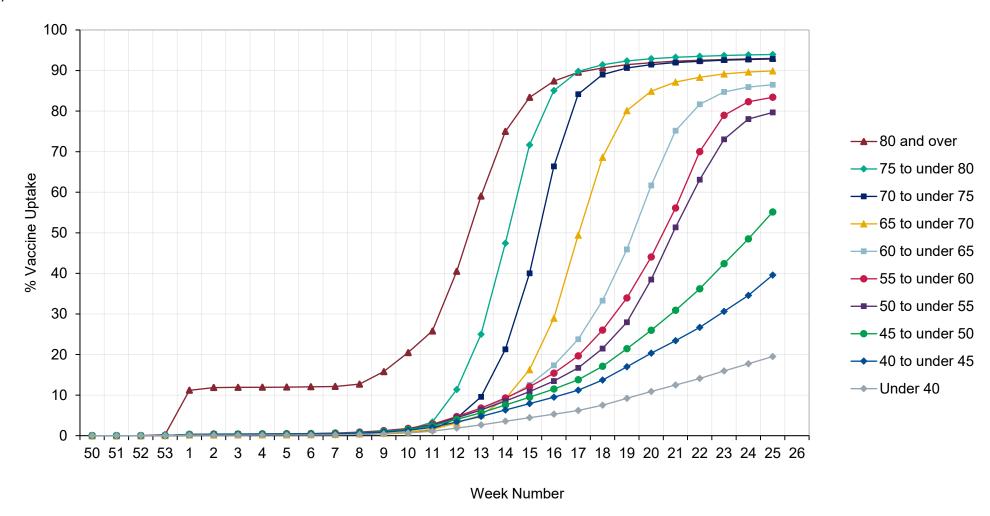
The data in this week's report covers the period from 8 December 2020 to 27 June 2021 (week 25) (Figure 2). It shows the provisional number and percentage of people in England who have had received 1 dose or 2 doses of a COVID-19 vaccination by age group and week since the start of the programme.

Figure 2. Cumulative weekly vaccine uptake by age

a) Dose 1



b) Dose 2



Vaccine impact on proportion of population with antibodies to COVID-19

PHE monitors the proportion of the population with antibodies to COVID-19 by testing samples provided by healthy adult blood donors aged 17 years and older, supplied by the NHS Blood and Transplant (NHS BT collection). This is important in helping to understand the extent of spread of COVID-19 infection (including asymptomatic infection) in the population and the impact of the vaccine programme. 250 samples from every geographic region in England are tested each week using 2 different laboratory tests, the Roche nucleoprotein (N) and Roche spike (S) antibody assays. This dual testing helps to distinguish between antibodies that are produced following natural COVID-19 infection and those that develop after vaccination. Nucleoprotein (Roche N) assays only detect post-infection antibodies, whereas spike (Roche S) assays will detect both post-infection antibodies and vaccine-induced antibodies. Thus, changes in the proportion of samples testing positive on the Roche N assay will reflect the effect of natural infection and spread of COVID-19 in the population. Increases in the proportion positive as measured by S antibody will reflect both infection and vaccination. Antibody responses reflect infection or vaccination occurring at least 2 to 3 weeks previously given the time taken to generate an antibody response.

In this report, we present the results using a 4-weekly average, of testing samples up to 20 June 2021, which takes account of the age and geographical distribution of the English population. Overall, the proportion of the population with antibodies using the Roche N and Roche S assays respectively were 14.9% and 84.2% for the period 24 May to 20 June (weeks 21 to 24) (Figure 3). This compares with 15.5% Roche N seropositivity and 75.5% Roche S seropositivity for the period of 26 April 2021 to 23 May 2021 (weeks 17 to 20). During this period seropositivity using the Roche N assay has remained stable suggesting there hasn't been significant ongoing spread of infection in the population and the continuing increase in seropositivity using the Roche S assay reflects the growing proportion of adults who have developed antibodies following vaccination.

Figure 4a and 4b show the proportion of the population with antibodies by age group. Roche N seropositivity has continued to plateau across most age groups and this was first observed in the 70 to 84 age group.

The pattern of increases in Roche S seropositivity which are observed follow the roll out of the vaccination programme with the oldest age groups offered vaccine first. (Figure 4b). Roche S seropositivity increased first in donors aged 70 to 84 and has plateaued since week 13, reaching 99.3% in weeks 21 to 24. Seropositivity has also plateaued since week 16 for those aged 60 to 69 reaching 98.8% in weeks 21 to 24. Plateauing in Roche S seropositivity has been observed since week 19 in those aged 50 to 59

reaching 98.5% in weeks 21 to 24 2021. A notable increase has been observed in the 40 to 49 year olds from 79.3% in weeks 17 to 20 to 94.5% in weeks 21 to 24. Currently the greatest increase observed is in the 30 to 39 year olds increasing from 47.9% in weeks 17 to 20 to 76.5% in weeks 21 to 24.

The impact of the vaccination programme is clearly evident from the increases in the proportion of the adult population with antibodies based on Roche S testing. This is particularly evident amongst individuals aged 50 years and above who have been prioritised for vaccination as part of the phase 1 programme and since week 15 in those aged 40 to 49 and 30 to 39 years old as part of phase 2 of the vaccination programme. This is further supported by the earlier plateauing in the proportion testing positive using the Roche N assay, in older age groups and likely reflects the additional role of vaccination is having in reducing infection ahead of reductions seen from national restrictions alone in younger age groups.

Figure 3: Overall population weighted 4-weekly rolling SARS-CoV-2 antibody seroprevalence (% seropositive) in blood donors from the Roche S and Roche N assays.

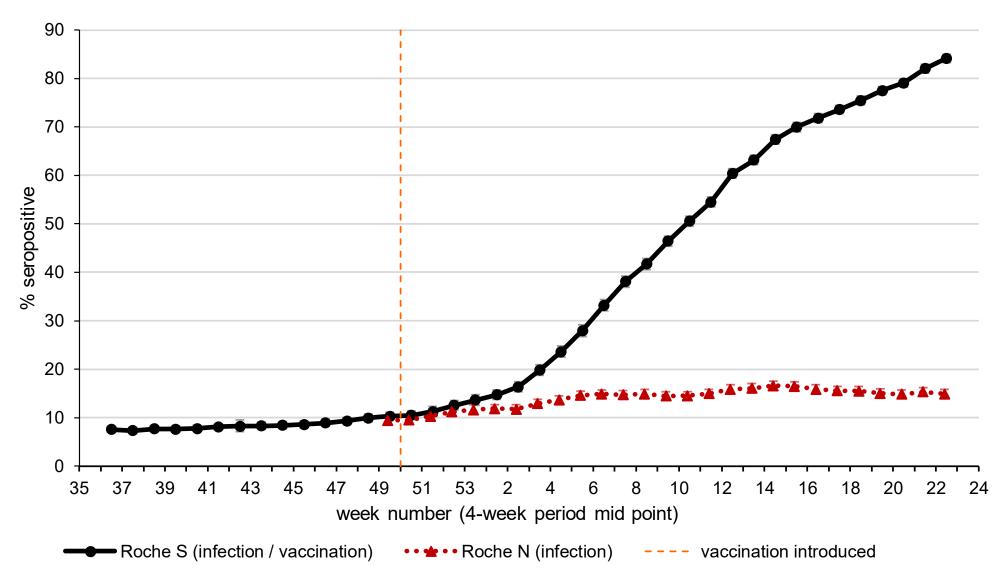
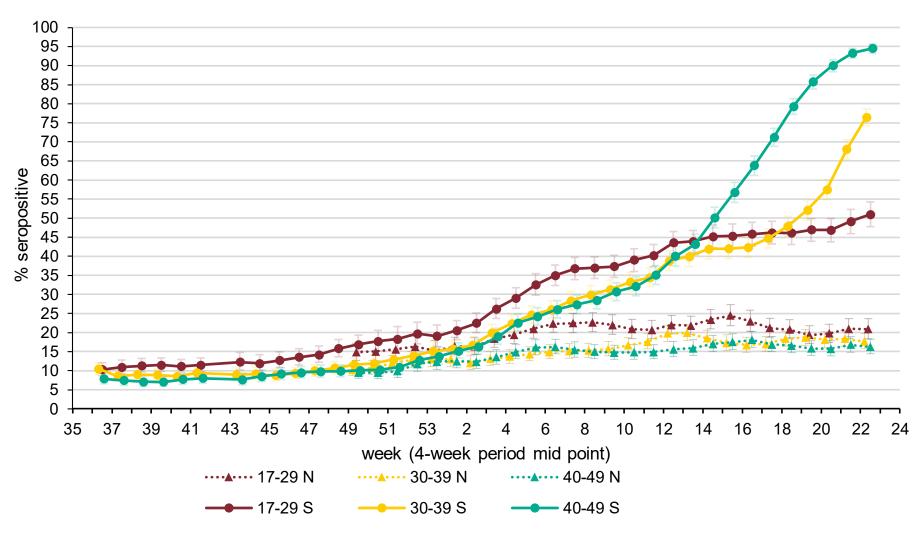
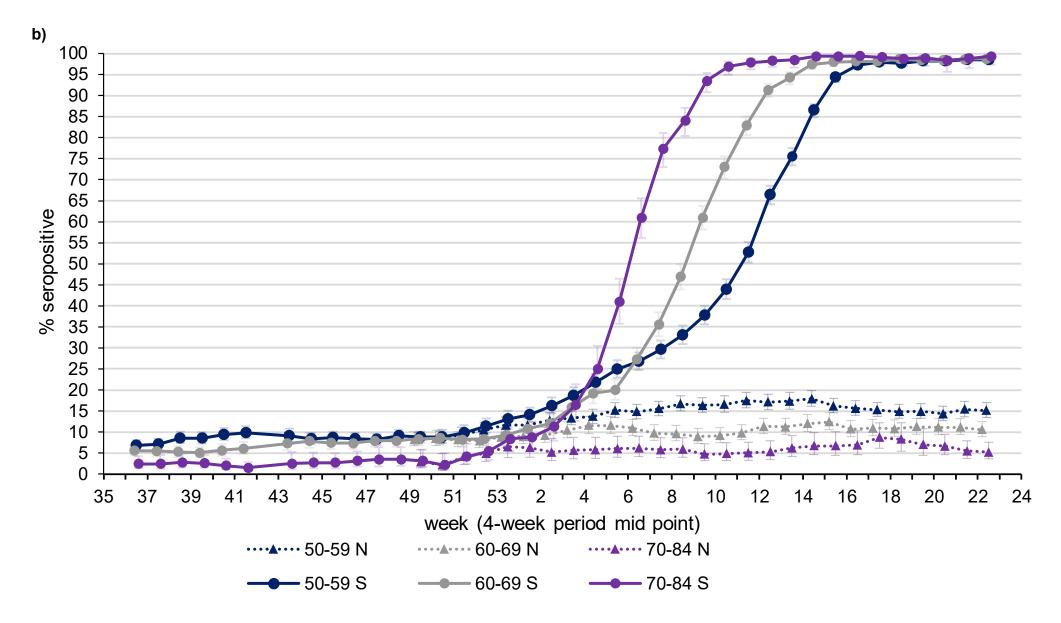


Figure 4: Population weighted 4-weekly rolling SARS-CoV-2 antibody seroprevalence (% seropositive) in blood donors from the Roche S and Roche N assays by a) age groups 17 to 29, 30 to 39 and 40 to 49, b) age group 50 to 59, 60 to 69 and 70 to 84.

a)





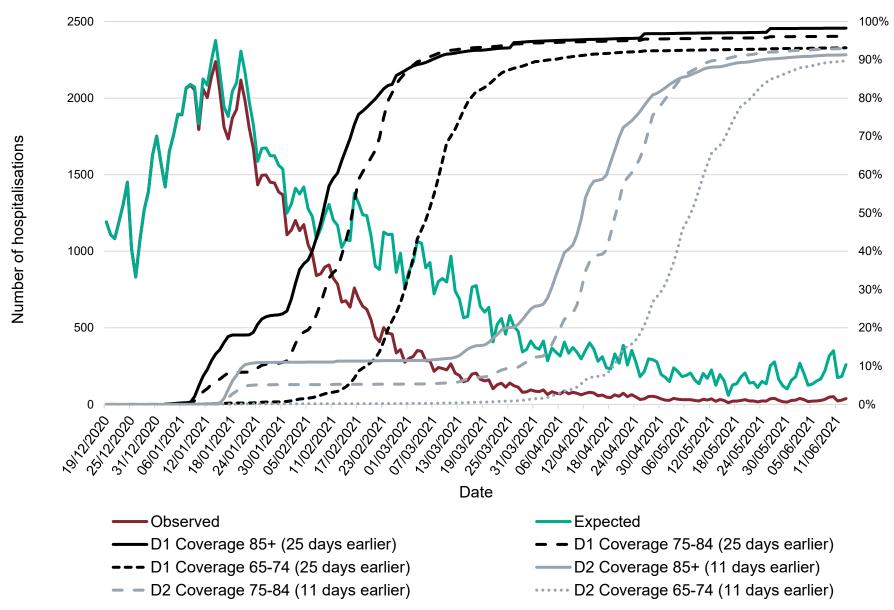
Direct impact on hospitalisations

The number of hospitalisations averted by vaccination, can be estimated by considering vaccine effectiveness against hospitalisation, vaccine coverage and observed hospitalisations and through modelling using a range of parameters.

PHE estimates to 13 June 2021 based on the direct effect of vaccination and vaccine coverage rates, are that around 44,500 hospitalisations have been prevented in those aged 65 years and over in England (approximately 6,000 admissions in those aged 65 to 74, 17,200 in those aged 75 to 84, and 21,300 in those aged 85 and over) as a result of the vaccination programme (Figure 5). There is increasing evidence that vaccines prevent infection and transmission. The indirect effects of the vaccination programme will not be incorporated in this analysis, therefore the figure of 44,500 hospitalisations averted is likely to be an underestimate.

Please note this analysis will be updated every 2 weeks, the next update will be in week 27.

Figure 5. Plot of daily observed and expected COVID-19 hospitalisations in adults aged 65 and over



Direct and indirect impact on infection and mortality

This analysis shows the direct and indirect impact of the vaccination programme on COVID-19 infections and mortality, and was first published here on 28 June 2021. This section will replace the section examining the direct impact of the vaccination on mortality, which was previously included in the weekly vaccine surveillance report.

The PHE and Cambridge real-time model has been used to track the COVID-19 infection throughout the pandemic, providing key epidemic insights, including estimation of the reproduction number, R, to the Scientific Pandemic Influenza subgroup on Modelling (SPI-M) and to the Scientific Advisory Group on Emergencies (SAGE). The application to data from the first wave has been published in Real-time nowcasting and forecasting of COVID-19 dynamics in England: the first wave (18). Since the first wave, the model has been constantly improved to capture the pandemic activity as it develops, in particular to account for the impacts, both direct and indirect, of the vaccination programme. The direct impact of vaccination is the number of deaths saved in those that get infected, whereas the indirect effect incorporates the additional prevention of infections. The history of real-time modelling outputs can be found at Nowcasting and Forecasting of the COVID-19 Pandemic (19), with the most recent results on which the figures here are based is currently available at COVID-19: nowcast and forecast (20).

Vaccination rates in the model are based on the actual number of doses administered, and the vaccine is assumed to reduce susceptibility to COVID-19 as well as mortality once infected. Estimates for vaccine efficacy are based on the best available published results (21). To infer the impact of vaccination, the model was fitted to both ONS prevalence and daily COVID-19 mortality data in England, resulting in posterior samples for a range of epidemiological parameters. The posterior samples were then used to simulate the number of infections and deaths that would have occurred without vaccination (Figure 6). Finally, the total impact was calculated by comparing the infection and mortality estimates with vaccination versus the simulated outcomes without vaccination (Figure 7; Table 6). The figures in Table 6 are expected to continue to grow rapidly as the number of deaths in the no-vaccination scenario is still showing exponential growth.

The no-vaccination scenario assumes that no other interventions are implemented to reduce incidence and mortality. Therefore, the findings presented here should be interpreted as the impact of the vaccination programme on infection and mortality assuming no additional non-pharmaceutical interventions were implemented. In practice it is impossible to predict what interventions would have been implemented in the absence of vaccination, although it is reasonable to assume that lockdown measures would have remained in place for substantially longer.

The work presented in this section is joint work completed by PHE and Cambridge University's MRC Biostatistics Unit.

Figure 6. Inferred and predicted incidence, mortality and prevalence with and without vaccination in England

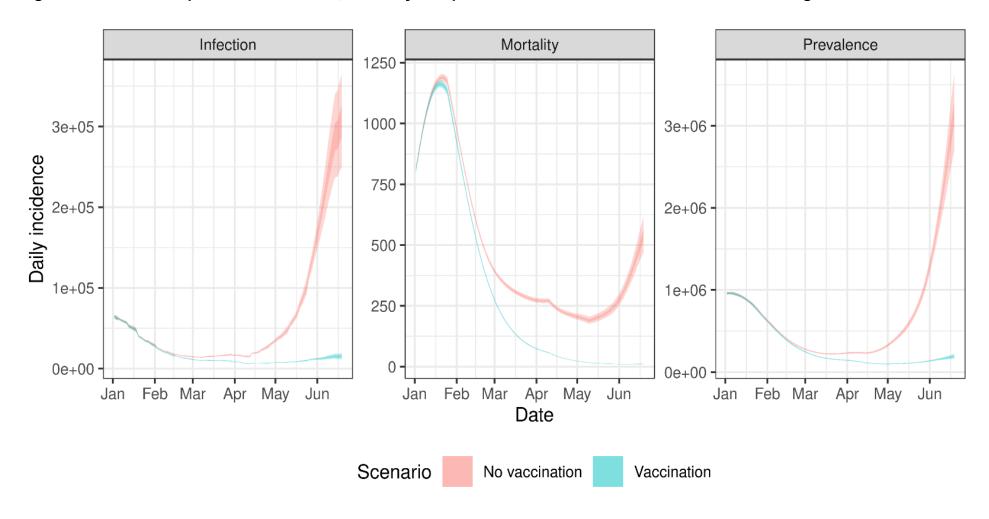


Figure 7. Averted number of infections (left) and deaths (right) due to vaccination (cumulatively)

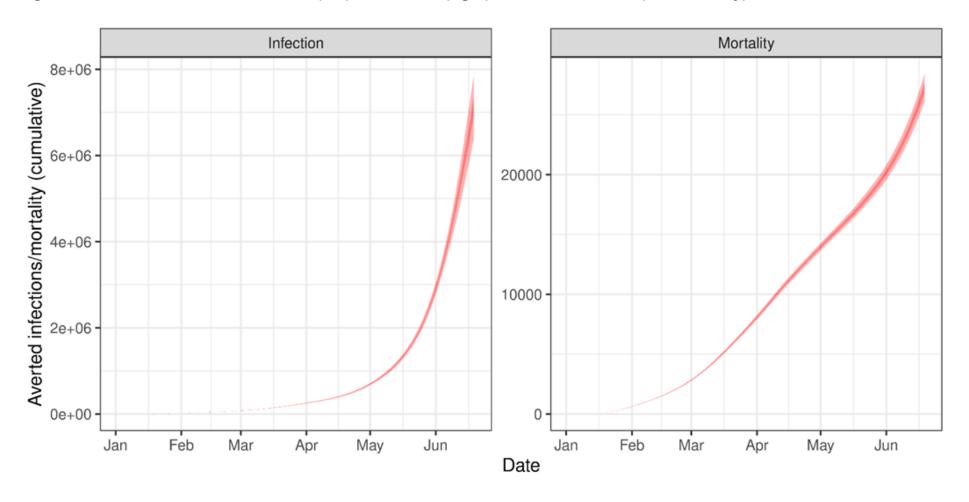


Table 6. Inferred reduction in infections and mortality as the result of vaccination up to 19 June 2021. (Infections are rounded to the nearest 1,000, deaths to the nearest 100.)

Model	Outcome	Reduction
ONS/Death	Infection	7151000 [6396000 , 7887000]
ONS/Death	Mortality	27200 [26100 , 28400]

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