

## **COVID-19 vaccine surveillance report** Week 23

8 June 2023

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## **Executive summary**

Rigorous clinical trials have been undertaken to understand the immune response, safety profile, and efficacy of all COVID-19 vaccines approved for use in the UK 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.

UK Health Security Agency (UKHSA), formerly 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 page <u>COVID-19</u>: vaccine surveillance strategy (<u>1</u>). As with all vaccines, the safety of COVID-19 vaccines is continuously <u>being monitored by the MHRA</u>. They conclude that overall, the benefits of COVID-19 vaccines outweigh any potential risks (<u>2</u>).

Please note that this report is published quarterly. The next report will be published on 31 August 2023.

This month's report contains updates on vaccine effectiveness, vaccine coverage, vaccination in pregnancy, vaccine impact on the proportion of the population with antibodies to COVID-19, and vaccination status by the time of admission through SARI-Watch.

## Vaccine effectiveness

Large clinical trials were undertaken for each of the COVID-19 vaccines first approved in the UK. These found that they were highly efficacious at preventing symptomatic disease in the populations that were studied. The clinical trials were able to assess the efficacy of the vaccines against laboratory confirmed symptomatic disease caused by the strains circulating in 2020 and early 2021. They also assessed with lower power efficacy against more severe disease.

Post implementation real world vaccine effectiveness (VE) studies are needed to understand vaccine effectiveness against different outcomes (such as severe disease and onward transmission), effectiveness in different subgroups of the population and against different variants as well as to understand the duration of protection. Vaccine effectiveness is estimated by comparing rates of disease (or positivity among those tested) in vaccinated individuals to rates in unvaccinated individuals. Below we outline the latest real-world evidence on vaccine effectiveness from studies in UK populations. Where available we focus on data related to the Omicron variant which is currently dominant in the UK. Previous reports demonstrated high levels of effectiveness against Alpha and Delta variants, particularly for severe disease and death.

Please note that vaccine effectiveness data will be updated in this report as it becomes available.

## 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 National Immunisation Management System (NIMS), cohort studies such as the COVID-19 Infection Survey, and GP electronic health record data. After community testing for COVID-19 was reduced in April 2022, we were no longer able to assess vaccine effectiveness against symptomatic disease using this data but results from previous analyses are available in earlier iterations of this report. Evidence prior to April from community testing and in other studies since this time has shown that effectiveness of all COVID-19 vaccines, including after first booster doses, against symptomatic disease with Omicron BA.1 and subsequent sub-lineages of the Omicron variant is low (starting at about 50 to 60%) and wanes rapidly to near 0 by 6 months.

## Effectiveness against hospitalisation

Several studies have estimated vaccine effectiveness against hospitalisation, all of which indicate higher levels of protection against hospitalisation than symptomatic disease with all vaccines against the Alpha, Delta and Omicron variants ( $\underline{3}$  to  $\underline{8}$ ). Given that Omicron generally causes milder disease than previous variants ( $\underline{9}$ ), in particular among younger individuals, an increasing proportion of individuals hospitalised with a positive COVID-19 test are likely to have COVID-19 as an incidental finding rather than the primary reason for admission ( $\underline{8}$ ).

We have previously found that using broader definitions of hospitalisation has given lower vaccine effectiveness estimates, reflecting outcome misclassification where cases are likely coincidentally positive whilst in hospital, without this being the primary reason for admission (and therefore these cases cannot be prevented by vaccination). We, therefore, use stricter definitions to define a COVID-19 hospitalisation. Here, a COVID-19 hospitalisation is defined as requiring at least 2 days stay in the hospital and a respiratory code in the primary diagnostic field. To estimate effectiveness against more severe disease we also estimate effectiveness for those requiring oxygen, mechanical ventilation or ICU (Table 1). There are likely still some incidental admissions in our data, in particular among younger adults. Using data from mid-July when BA.5 sub-lineage of Omicron has been dominant shows that protection against hospitalisation is maintained to a reasonable degree long term; VE after a second dose was 40.2% after 15 or more months and 52.3% at 12 to 14 months after a first booster dose (Table 1).

Dose	Interval	VE against hospitalisation
	2 weeks to 2 months	70.3 (30.9 to 87.2)
	3 to 5 months	71.7 (50.5 to 83.8)
	6 to 8 months	57.6 (38.6 to 70.7)
	9 to 11 months	51.2 (31.2 to 65.4)
	12 to 14 months	35.5 (23.3 to 45.8)
	15+ months	40.2 (31.0 to 48.1)
Booster (third+ dose)	2 weeks to 2 months	78.1 (75.7 to 80.3)
	3 to 5 months	65.3 (61.7 to 68.6)
	6 to 8 months	53.6 (48.8 to 58.0)
	9 to 11 months	51.1 (45.7 to 56.0)
	12 to 14 months	52.3 (43.7 to 59.6)
	15+ months	Insufficient data

Table 1. Vaccine effectiveness against hospitalisation in those aged 65 years and over. Hospitalisation is defined as a stay of at least 2 days with a respiratory code in the primary diagnosis field

## Effectiveness of the autumn 2022 bivalent booster

Bivalent boosters with either Pfizer BioNTech (Original/Omicron BA.1 Comirnaty®) or a Moderna bivalent (Spikevax® bivalent Original/Omicron vaccine) targeting both the ancestral strain and Omicron BA.1 were offered to those in clinical risk groups and those aged 50 years and older from September 2022. VE of the bivalent boosters was estimated against hospitalisation in the period following 5 September 2022 against all Omicron sub-lineages in circulation at the time. Only individuals who had received at least 2 COVID-19 vaccines before 5 September 2022 and with the last of these doses at least 6 months prior to sample date were included in analysis.

The effectiveness measured is therefore incremental effectiveness on top of at least 6 months waned protection. The incremental protection conferred by the bivalent vaccines estimated relative to those with waned immunity was 47% for Pfizer after 2 to 4 weeks, and 58% for Moderna. After 25 or more weeks after vaccination; VE was 21% for the Pfizer booster and 20% for the Moderna booster (Table 2).

Vaccine manufacturer	Interval after booster dose	Controls	Cases	VE (95% CI)
Pfizer	2 to 4 weeks	1,849	315	47.4 (40.1 to 53.7)
	5 to 9 weeks	5,395	884	44.9 (40 to 49.4)
	10 to 14 weeks	4,925	1128	34.5 (29 to 39.5)
	15 to 19 weeks	2,636	1008	28 (21.1 to 34.4)
	20 to 24 weeks	1,297	605	25.2 (15.6 to 33.6)
	25+ weeks	41	18	20.8 (-41.6 to 55.7)
Moderna	2 to 4 weeks	1,130	313	58.1 (51.9 to 63.5)
	5 to 9 weeks	3,807	672	48.1 (42.8 to 53)
	10 to 14 weeks	8,597	1649	36.9 (32.2 to 41.3)
	15 to 19 weeks	4,899	1273	31.1 (25.3 to 36.4)
	20 to 24 weeks	3,385	1580	23.6 (16.9 to 29.8)
	25+ weeks	1,158	594	19.6 (8.7 to 29.2)

## Table 2. Vaccine effectiveness of the bivalent booster vaccines against hospitalisation in those aged 50 years and older (VE = vaccine effectiveness, CI = confidence intervals)

## Effectiveness against mortality (vaccines given prior to the autumn 2022 bivalent boosters)

Vaccine effectiveness against mortality with the Omicron variant (all sublineages using tests taken until 5 September 2022) has been estimated for those aged 65 years and older using a test-negative case control study design (all vaccines combined) (Table 3). At 40-plus weeks following the second dose, vaccine effectiveness was around 50%. At 2 or more weeks following third and fourth dose vaccination, effectiveness was boosted to 85.0% and 80.9%, respectively. At 40 or more weeks after a third dose VE waned to 56.9% while at 20 or more weeks after a fourth dose (spring 2022 booster) VE waned to 68.2%. This analysis is also likely to include some incidental deaths of individuals who died with COVID-19 as opposed to from COVID-19, and we suspect the true VE against mortality is likely higher than the estimates presented here.

Dose	Interval after dose (weeks)	VE (95% CI)
2	40+	49.7 (41.5 to 56.7)
3	2 to 4	85.0 (80.8 to 88.2)
	5 to 9	83.1 (80.3 to 85.5)
	10 to 14	79.5 (76.6 to 82.0)
	15 to 19	75.6 (72.3 to 78.6)
	20 to 24	68.8 (64.3 to 72.7)
	25 to 39	62.6 (57.4 to 67.2)
	40+	56.9 (43.1 to 67.4)
4	2 to 4	80.9 (76.8 to 84.3)
	5 to 9	79.5 (75.8 to 82.7)
	10 to 14	71.2 (66.2 to 75.5)
	15 to 19	68.2 (61.2 to 73.9)
	20+	68.2 (58.4 to 75.7)

Table 3. Vaccine effectiveness against mortality in those aged 65 years and older (all
vaccine brands combined) (VE = vaccine effectiveness, CI = confidence intervals)

## 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 it 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 Alpha and Delta variants (10 to 13).

Generally, estimates are similar to or slightly lower than vaccine effectiveness estimates against symptomatic disease and there is evidence of a significant waning in protection against infection over time. Effectiveness against infection with the Omicron variant is low and wanes rapidly (<u>Table 4</u>).

## Effectiveness against transmission

As described above, several studies have provided evidence that vaccines provide some protection against infection. Uninfected individuals cannot transmit. Therefore, the vaccines also provide some protection against transmission. There may be an 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). Several studies have provided evidence of reduced risk of household transmission from vaccinated cases compared to unvaccinated cases (<u>14</u> to <u>18</u>) with past variants.

### Consensus vaccine effectiveness estimates

Table 4. Consensus estimates of relative vaccine effectiveness against BA.4, BA.5, BQ.1 and CH1.1 Omicron for a booster dose of COVID-19 vaccine compared to 6+ months since the last dose (at least 2 doses)

Vaccine product of booster dose [Note 1]	Outcome	0 to 1 months	2 to 3 months	4 to 6 months	6+ months	Consensus narrative
Monovalent	All Infection [Note 2]	30% (20 to 40%)	20% (10 to 30%)	10% (0 to 20%)	0% (0 to 5%)	Post fourth dose estimates appear similar same time period, restoring VE to similar I booster dose (pre waning). VE estimates I individuals have had a prior infection. Son may be slightly higher against Omicron BA BA.4/BA.5.
Monovalent	Symptomatic [Note 2]	40% (30 to 50%)	40% (30 to 50%)	10% (0 to 20%)	Insufficient data	Post fourth dose estimates appear similar same time period, restoring VE to similar booster dose (pre waning). VE estimates individuals have had a prior infection.
Monovalent	Hospitalisation	60% (55 to 65%)	40% (30 to 50%)	20% (15 to 25%)	0% (0 to 5%)	These estimates are based on UKHSA test SUS data on hospitalisations in age 75+ w coded. Note: As absolute VE had waned t dose 3, a relative VE of 50% would increat of 85% compared to unvaccinated. Some against hospitalisation are similar for BA.2
Monovalent	Mortality	Insufficient data	Insufficient data	Insufficient data	Insufficient data	
Bivalent	All Infection [Note 2]	30% (20 to 40%)	20% (10 to 30%)	10% (0 to 20%)	0% (0 to 5%)	Estimates for the BA.1/BA.2 bivalent boos what is being seen with BA.4/BA.5 bivaler
Bivalent	Symptomatic [Note 2]	40% (30 to 50%)	40% (30 to 50%)	10% (0 to 20%)	Insufficient data	
Bivalent	Hospitalisation	55% (40 to 65%)	50% (40 to 65%)	Insufficient data	Insufficient data	Estimates for the BA.1/BA.2 bivalent boos what is being seen with BA.4/BA.5 bivaler evidence that VE against hospitalisation is compared to BA.4/BA.5.
Bivalent	Mortality	70% (65 to 80%)	70% (65 to 80%)	Insufficient data	Insufficient data	Current evidence based on age 65+.

The table presents estimates of VE compared to 6+ months since last dose (at least 2 doses) (estimates agreed by the vaccine expert panel).

Note 1. Refers to either Pfizer or Moderna.

Note 2. Estimates were not stratified according to monovalent or bivalent.

High confidence	Evidence from multiple studies which is consistent and comprehensive.		
Medium confidence	Evidence is emerging from a limited number of studies or with a moderate level of uncertainty.		
Low confidence	Little evidence is available, and results are inconclusive.		

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test negative case control using with 2+ days stay and respiratory to about 70% at 6+ months post ease this back to an absolute VE e studies suggest VE estimates .2 versus BA.4/BA.5.

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## Effectiveness against BQ.1

VE against hospitalisation for BQ.1 and BA.5 was estimated during a period of co-circulation (Table 5). This included the period from 5 September 2022 to 25 December 2022. Cases were classified as BA.5 (VOC-22APR-04) or BQ.1 (V-22OCT-01) based on sequencing information. VE was estimated for those who had received a bivalent booster vaccine as part of the autumn programme, as well as at least 2 previous doses at least 6 months previously, relative to those who were not boosted in the autumn but had at least 2 previous doses at least 6 months previously. Estimates are for those who received the booster 2 or more weeks ago. All vaccine manufacturers were combined in the analysis.

The effectiveness of the bivalent booster (manufacturers combined) against hospitalisation with BQ.1 was 52.1% as compared to 63.6% with BA.5, at 2 or more weeks after receiving the booster (<u>Table 5</u>). Although the effectiveness point estimate is lower for BQ.1 the confidence interval is fairly wide and overlaps the estimate for BA.5. Currently the number of BQ.1 cases in the analysis is too small to confidently assess differences in vaccine effectiveness between the 2 variants.

Variant	Vaccine	Controls	Cases	VE (95% CI)
BQ.1	Bivalent (any)	8,617	175	52.1 (40.2 to 61.6)
	Pfizer	2,730	49	54.1 (36.0 to 67.1)
	Moderna	5,887	126	51.0 (37.7 to 61.4)
BA.5	Bivalent (any)	8,617	100	63.6 (53.8 to 71.3)
	Pfizer	2,730	25	53.7 (28.5 to 70)
	Moderna	5,887	75	65.8 (55.5 to 73.7)

Table 5. VE estimates against hospitalisation with BQ.1 and BA.5 for the bivalent boosters in those aged 50 years and older

Estimates of earlier circulating Omicron sub-lineages (BA.4, BA.4.6 and BA.5) as compared to BA.2 are available in previous iterations of this report.

## Effectiveness against XBB.1.5, CH.1.1 and BQ.1

Sub-lineages CH.1.1 and XBB.1.5 increased in prevalent in England in December 2022 and January 2023, respectively. VE against hospitalisation for XBB.1.5, CH.1.1. and BQ.1 was estimated during a period of co-circulation between 5 December and 19 February (<u>Table 6</u>). During this period, BA.5 was not a prominent sub-lineage. Cases were classified based on sequencing information. Cases were classified as XBB.1.5 (V-23JAN-01), CH.1.1 (V-22DEC-01) or BQ.1 (V-22OCT-01) based on sequencing information, and VE was estimated as described above.

There was some evidence that VE against hospitalisation for CH.1.1 was lower than that of BQ.1, and that VE against XBB.1.5 was lower than that of CH.1.1 and BQ.1, but confidence intervals were wide and overlapped. Currently the number of cases in the analysis is too small to confidently assess differences in vaccine effectiveness between these variants.

Variant	Bivalent booster	Controls	Cases	VE
XBB.1.5	None	34,802	108	Baseline
	2 to 4 weeks	3,215	2	56.9 (-75.9 to 89.5)
	5 to 9 weeks	19,986	15	47.3 (7.8 to 69.9)
	10 to 14 weeks	42,149	80	25.7 (-0.9 to 45.3)
	15+ weeks	30,465	220	12.3 (-12.9 to 31.9)
CH.1.1.	None	34,802	164	Baseline
	2 to 4 weeks	3,215	11	31.4 (-27.4 to 63)
	5 to 9 weeks	19,986	71	35.1 (13 to 51.6)
	10 to 14 weeks	42,149	199	28.4 (10.7 to 42.5)
	15+ weeks	30,465	164	26.3 (5.4 to 42.5)
BQ.1	None	34,802	450	Baseline
	2 to 4 weeks	3,215	23	62.6 (42.9 to 75.5)
	5 to 9 weeks	19,986	214	46.5 (36.5 to 54.9)
	10 to 14 weeks	42,149	476	40.5 (31.8 to 48.1)
	15+ weeks	30465	258	31.3 (17.7 to 42.6)

Table 6. VE estimates against hospitalisation with XBB.1.5, CH.1.1 and BQ.1 for the bivalent boosters in those aged 50 years and older

## Vaccine effectiveness publications

UKHSA and collaborators have published a significant amount of <u>research into vaccine</u> <u>effectiveness</u>, which is summarised on pages 4 to 11. The publications listed in Table 7 provide further results and details on the methods used.

Publication	Subject
Effectiveness of the COVID-19 vaccines against severe disease with Omicron sub- lineages BA.4 and BA.5 in England	The latest evidence shows that vaccine effectiveness against hospitalisation is similar for the BA.4/5 variants as it is for BA.2. In somebody who received their second dose around 6 months previously, a booster dose increases protection against hospitalisation by 50 to 60%. This is the most comprehensive analysis of vaccine effectiveness against hospitalisation for BA.4/5 undertaken to date.
Effectiveness of AstraZeneca COVID-19 booster vaccination against the Omicron and Delta variants	This study estimates the effectiveness of booster vaccination with AstraZeneca against symptomatic disease and hospitalisation in individuals who were not able to receive mRNA vaccines in the UK.
<u>COVID-19 vaccine effectiveness against the</u> <u>Omicron BA.2 variant in England</u>	This study estimates the effectiveness of booster vaccination against symptomatic disease caused by the BA.2 sub-lineage of the Omicron (B.1.1.529) variant.
Vaccine effectiveness against hospitalisation with the Omicron variant	This study estimates vaccine effectiveness against hospitalisation with the Omicron variant and investigates the impact of using different hospitalisation outcome definitions
Effectiveness of COVID-19 vaccines against hospitalisation with the Omicron variant in adults aged 75 years and older	This study reports on vaccine effectiveness against hospitalisation with the Omicron variant in adults aged 75 years and older.
Effectiveness of BNT162b2 and ChAdOx1 against SARS-CoV-2 household transmission: a prospective cohort study in England	This study reports on vaccine effectiveness against transmission of COVID-19 with the Alpha and Delta variants.
Effectiveness of 3 doses of COVID-19 vaccines against symptomatic COVID-19 and	Updated analysis on the effectiveness of 3 doses of COVID-19 vaccines against

Table 7. UKHSA publications on the effectiveness of COVID-19 vaccination

Publication	Subject
hospitalisation in adults aged 65 years and older	symptomatic COVID-19 and hospitalisation in adults aged 65 years and older.
Effectiveness of BNT162b2 COVID-19 booster vaccine against COVID-19 related symptoms and hospitalisation in England	This study provides real world evidence of significantly increased protection from the booster vaccine dose against symptomatic disease and hospitalisation irrespective of the primary course.
Effectiveness of COVID-19 vaccines against the Omicron (B.1.1.529) variant of concern	This study reports on the vaccine effectiveness against symptomatic disease with 2 dose courses of BNT1622 and ChAdOx1-S as well as booster doses of BNT162b2 following a primary course of either BNT1622 or ChAdOx1-S.
Effectiveness of BNT162b2 (Comirnaty, Pfizer- BioNTech) COVID-19 booster vaccine against COVID-19 related symptoms in England: test negative case-control study	Results from the first UK real-world study by UKHSA show significantly increased protection against symptomatic disease from a booster dose of the Pfizer-BioNTech vaccine in those aged 50 years and older.
Duration of protection against mild and severe disease by COVID-19 vaccines	This study reports on the vaccine effectiveness and duration of protection of Comirnaty, Vaxzevria and Spikevax against mild and severe COVID-19 in the UK.
Serological responses and vaccine effectiveness for extended COVID-19 vaccine schedules in England	This study investigates the impact of different dosing schedules on immune response and vaccine effectiveness.
Pfizer-BioNTech and Oxford AstraZeneca COVID-19 vaccine effectiveness and immune response among individuals in clinical risk groups	This study reports on the immune response and clinical effectiveness of COVID-19 vaccine among individuals in clinical risk groups. A <u>supplementary appendix</u> is also available to download.
Effectiveness of COVID-19 vaccines against hospital admission with the Delta (B.1.617.2) variant	This study reports on the effectiveness of COVID-19 vaccines on hospitalisation disease with the Delta variant. A supplementary appendix is also available to download.
Effectiveness of COVID-19 vaccines against the B.1.617.2 (Delta) Variant	This study reports on the effectiveness of COVID-19 vaccines on symptomatic disease with the Delta variant.

Publication	Subject
Effectiveness of BNT162b2 mRNA and ChAdOx1 adenovirus vector COVID-19 vaccines on risk of hospitalisation among older adults in England: an observational study using surveillance data	A study using the SARI watch surveillance system of COVID-19 hospitalisations found high levels of protection against hospitalisation after both a single dose and 2 doses of COVID-19 vaccines.
Effectiveness of BNT162b2 mRNA vaccine and ChAdOx1 adenovirus vector vaccine on mortality following COVID-19	A study on deaths with COVID-19 indicates that COVID-19 vaccines offer high levels of protection against mortality.
Effect of Vaccination on Household Transmission of SARS-CoV-2 in England	Impact of vaccination on household transmission of SARS-COV-2 in England is an analysis to determine whether individuals who have received the vaccine, but still become infected with SARS-COV-2 up to 60 days after the first dose, are less likely than unvaccinated cases to transmit to their unvaccinated household contacts.
Vaccine effectiveness of the first dose of ChAdOx1 nCoV-19 and BNT162b2 against SARS-CoV-2 infection in residents of Long- Term Care Facilities (VIVALDI study)	The VIVALDI study found evidence that COVID-19 vaccines were associated with a substantially reduced risk of infection in care home residents.
Effectiveness of BNT162b2 and ChAdOx1 nCoV-19 COVID-19 vaccination at preventing hospitalisations in people aged at least 80 years: a test-negative, case-control study	The Avon CAP study, conducted in 2 hospitals in Bristol, found evidence of high levels of protection against hospitalisation in 80+ year olds with a single dose of either vaccine.
<u>COVID-19 vaccine coverage in health-care</u> workers in England and effectiveness of <u>BNT162b2 mRNA vaccine against infection</u> (SIREN): a prospective, multicentre, cohort study	Early data from UKHSA's SIREN study shows a promising impact on infection in healthcare workers aged under 65. Healthcare workers in the study are tested for COVID-19 every 2 weeks – whether or not they have symptoms.
Effectiveness of the Pfizer-BioNTech and Oxford-AstraZeneca vaccines on COVID-19 related symptoms, hospital admissions, and mortality in older adults in England: test negative case-control study	Early data from routine COVID-19 testing in older adults shows that vaccines are effective at preventing COVID-19 disease and severe outcomes.
Impact of COVID-19 vaccination programme on seroprevalence in blood donors in England, 2021	Report on the impact of COVID-19 vaccination programme on seroprevalence in blood donors in England, 2021.

## **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.

UKHSA and other government and academic partners monitor the impact 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

By the end of week 20 2023 (week ending 21 May 2023), 52.9% (2,853,424 out of 5,398,425) of all people aged over 75 years old who are living and resident in England who had been vaccinated with a Spring 2023 booster dose since 3 April 2023. For further detail on the vaccine uptake and the current booster campaign, please the flu and COVID-19 weekly surveillance report weekly national influenza and COVID-19 surveillance report.

## Vaccination in pregnancy

Vaccination of pregnant women alongside their peers is recommended in the UK and other countries as an important way to protect pregnant women and their unborn children against COVID-19 disease. Vaccination of pregnant women is strongly recommended by the <u>Royal</u> <u>College of Obstetricians and Gynaecologists and the Royal College of Midwives</u>.

Increased severity of COVID-19 disease in pregnant and recently pregnant women was reported after the first SARS-CoV-2 wave in England (<u>19</u>, <u>20</u>) and Scotland (<u>21</u>, <u>22</u>). The disease is generally reported to be milder during the Omicron period with reduced risk of complications in pregnant women when compared to the Delta period (<u>23</u>, <u>24</u>). Omicron has, however, also been associated with higher rates of infection in pregnant women when compared to Delta (<u>24</u>). Pregnant women who develop the severe disease have increased rates of admission to the ICU, need for invasive ventilation and pre-term delivery.

Data from the US Centers for Disease Control and Prevention (CDC) found that pregnant women were around 3 times more likely to be admitted to ICU and nearly 3 times more likely to require invasive ventilation compared to non-pregnant women with COVID-19 disease and 25% more likely to die ( $\underline{25}$ ).

From 16 April 2021, the Joint Committee on Vaccination and Immunisation (JCVI) advised that pregnant women be offered COVID-19 vaccines at the same time as people of the same age or risk group (<u>26</u>). Therefore, any pregnant women not in a high-risk group would likely have received their first dose from mid-April 2021 as part of the general adult population programme in those aged under 50 years. This was offered by decreasing age group (<u>26</u>). As part of the ongoing review of the programme, the JCVI met on 2 December 2021 and considered further data on severity of SARS-CoV-2 infection in pregnant women and their pregnancies together with data on vaccine safety. As a result, pregnant women were added to the UK's priority COVID-19 vaccine list (<u>27</u>). The booster dose, made available to all individuals with severe immunosuppression from September 2021 and then extended to all eligible adults in England from 30 November 2021, is important to confer high levels of protection against Omicron strains (see report section vaccine effectiveness). Pregnant women are included as one of the priority groups to be offered the autumn 2022 COVID-19 booster dose.

Prior to 16 April 2021, COVID-19 vaccine was delivered to priority groups, based on clinical risk and risk of exposure, and delivered in order of priority. On 22 December 2020, JCVI advised that vaccine could be offered to pregnant and breast-feeding women who were in these risk categories. The Pfizer vaccine was rolled out from early December 2020, AstraZeneca vaccine was used from 4 January 2021 and the Moderna vaccine became available from April 2021. From 17 April 2021 pregnant women have been offered the Pfizer-BioNTech or Moderna (mRNA) vaccines where available for their first dose due to reassuring global safety data (28).

There is evidence of high levels of protection against SARS-CoV-2 infection in pregnant women after COVID-19 vaccination (<u>29</u> to <u>31</u>) and evidence that vaccination induces higher antibody levels than after disease (<u>31</u>). There is also evidence from a recent US study that 2-doses of mRNA COVID-19 vaccination during pregnancy might help prevent COVID-19 hospitalisations in young infants under 6 months of age (<u>32</u>). Between February and September 2021, 0.4% of 1,714 pregnant women with COVID-19 symptoms who required hospital treatment in the UK had received 2 doses of COVID-19 vaccine and, of 235 pregnant women who were admitted to intensive care with COVID-19 disease in that period, none had received 2 doses of vaccine (<u>33</u>). Similar findings have been reported from Scotland with the report that 90.9% (748 out of 823; 95% CI 88.7 to 92.7) of SARS-CoV-2 associated with hospital admission, 98% (102 out of 104; 95% CI 92.5 to 99.7) of SARS-CoV-2 associated with critical care admission and all baby deaths, occurred in pregnant women who were unvaccinated at the time of their COVID-19 diagnosis (<u>22</u>, <u>34</u>). The researchers also found a higher extended perinatal mortality rate for women who gave birth within 28 days of a COVID-19 diagnosis compared to rates across the pandemic period and in women vaccinated and going on to give birth within 28 days.

COVID-19 vaccines used in the UK programme do not contain live SARS-CoV-2 virus and therefore cannot infect a pregnant woman or her unborn child with the virus. Whilst as is commonly the case in trials of medicinal products, pregnant women were excluded from the original COVID-19 vaccine trials, there is accumulating experience and evidence of the safe and effective use of mRNA vaccines (such as the Pfizer-BioNTech or Moderna) in pregnant women. In Scotland, COVID-19 vaccine had been administered to more than 30,000 pregnant women to the end of March 2022 (21). In the USA data collected by the US Centre for Disease Control indicated that around 71% of pregnant people were fully vaccinated before or during pregnancy in week ending 20 August 2022 (35).

No safety concerns relating to COVID-19 vaccination of pregnant women have been found in published studies to date (35 to 39). The rate of vaccine side-effects appears to be similar in pregnant and non-pregnant populations (35). Studies from Norway, the USA and Scotland have found no association between COVID-19 vaccination and the risk of miscarriage (37 to 40).

This report presents data on vaccine coverage only for women delivering up to the end of February 2023 and updates the coverage data on COVID-19 vaccination in pregnant women published in the <u>COVID-19 vaccine surveillance report</u> – weeks 47 of 2021, 4, 8, 12, 16, 19 and 35 of 2022, week 5, week 9 and week 14 of 2023. Of note, these figures have previously excluded women who have given birth more than once in the period covered. All women who give birth, including those who have given birth more than once in the period covered, are now included in the figures.

Findings continue to be provisional and are not directly comparable between reports as data is updated through the complete time period under consideration. All previous reports to week 9 2023 additionally included pregnancy outcomes (stillbirth, low birth weight and preterm deliveries) but women delivering in the most recent months with maternity data available have overwhelmingly been vaccinated with doses 1 to 3 before the pregnancy for which they are giving birth (Figure 1). These pregnancy outcomes would therefore not be expected to substantively alter previously published data. Similarly, breakdown of coverage by ethnicity, index of multiple deprivation (IMD) and maternal age group for doses 1 to 3 will no longer be updated. Data will be collated and published on women who have delivered from September 2022, during the period of vaccination offered under the Autumn boost programme, once numbers have accumulated.

This report is the third to include data on coverage of the autumn boost that was recommended to those at higher risk of severe COVID-19 disease in the population who had not been boosted for at least 3 months, including pregnant women. It is the first to include women who had given birth previously in the whole period covered which has not changed the overall patterns of uptake. Autumn boost vaccination began in September 2022.

#### Vaccine coverage

COVID-19 vaccine coverage in women at any point before they gave birth increased as more women became eligible for vaccination. In November 2021, 48.2% of women giving birth had received at least one dose of vaccine before they delivered. This increased to 53.4% of women who gave birth in December 2021, to 72.9% of women delivering by June 2022 when it stabilised, peaking in October 2022 at 75.3%. (Table 8). Of women who gave birth in December 2021, 42.9% had received 2 doses of the vaccine before they gave birth increasing to 67.1% of women giving birth in June 2022 and peaking at 70.3% of women who gave birth in October 2022. By October 2022 most 1st, 2nd and 3rd doses of vaccine given to women who gave birth were administered before the start of their pregnancy (Figure 1).

A small proportion of women who gave birth in September (0.5%) 2022 had received an autumn booster dose prior to delivery, rising to 18.1% in January 2023 and 19.0% in February 2023. More women eligible for the booster will give birth over the coming months. A smaller proportion of women received the autumn boost after they gave birth, falling from 6.5% of women who gave birth in September 2022 to 0.1% of women who delivered in January and in February 2023.

In the 26-month period between January 2021 and February 2023, a total of 1,134,487 women gave birth with linked records on vaccination status for 1,130,448 (99.6%) of them. Of women with linked records who gave birth over this period, 529,098 (46.6%) were known to have received at least one dose of COVID-19 vaccine prior to giving birth, 459,311 (40.5%) received at least 2 doses, 234,383 (20.7%) women had received at least 3 doses. Through September 2022 to February 2023, 25.2% of women had not received a COVID-19 vaccine before they gave birth.

There were 378,104 women who received their first dose prior to pregnancy and went on to conceive and deliver by end February 2023. Of the 29,744 autumn boost doses administered to women before they gave birth between September 2022 and February 2023, 12,222 were the Moderna bivalent vaccine and 17,522 were the Pfizer bivalent vaccine.

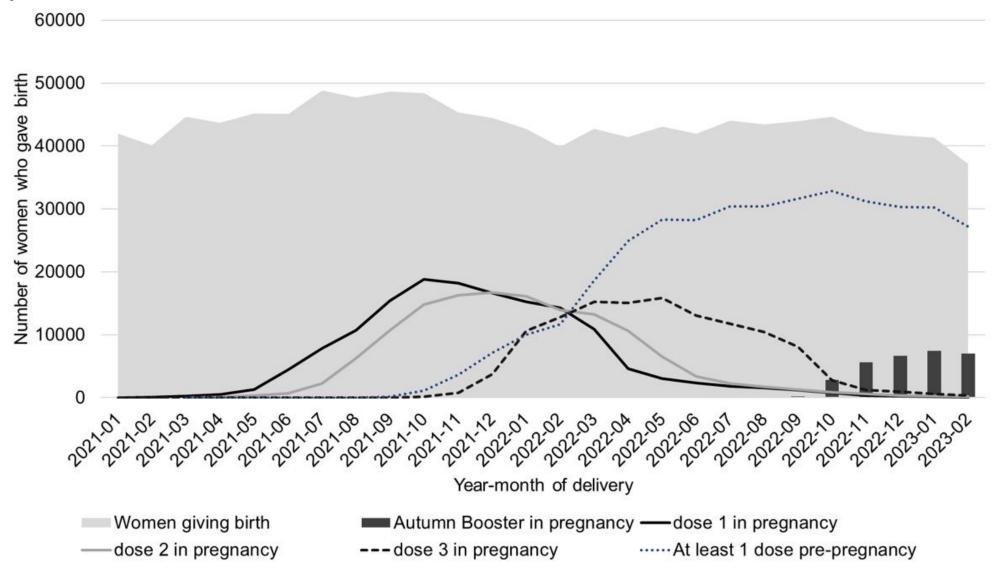


Figure 1. Women who gave birth and had received COVID-19 vaccine whilst pregnant or pre-pregnancy by month of delivery and by dose number administered

Table 8. Vaccine coverage in women giving birth from September 2022 (when the Autumn boost was first offered), by month of delivery<sup>1,2</sup>

Date	September 2022	October 2022	November 2022	December 2022	January 2023	February 2023
Women giving birth	43,973	44,658	42,285	41,680	41,324	37,125
1 or more doses by time of delivery	32,852 (74.7%)	33,617 (75.3%)	31,562 (74.6%)	30,608 (73.4%)	30,420 (73.6%)	27,324 (73.6%)
2 or more doses by time of delivery	30,601 (69.6%)	31,396 (70.3%)	29,461 (69.7%)	28,590 (68.6%)	28,437 (68.8%)	25,547 (68.8%)
3 or more doses by time of delivery	18,867 (42.9%)	20,119 (45.1%)	18,705 (44.2%)	17,964 (43.1%)	18,153 (43.9%)	16,276 (43.8%)
Autumn boost by time of delivery	202 (0.5%)	2,788 (6.2%)	5,581 (13.2%)	6,659 (16.0%)	7,466 (18.1%)	7,048 (19.0%)
Unvaccinated at delivery	10,933 (24.9%)	10,828 (24.2%)	10,511 (24.9%)	10,845 (26.0%)	10,671 (25.8%)	09,551 (25.7%)
Women who received autumn boost after delivery to 6 June 2023	2,853 (6.5%)	1,726 (3.9%)	526 (1.2%)	126 (0.3%)	053 (0.1%)	029 (0.1%)

#### Methods

Data on COVID-19 vaccination status together with details of each vaccine administered is recorded in a central data set called the NIMS<sup>3</sup>. In addition, NHS Digital manages the Hospital Episode Statistics (HES) data sets, containing information about hospital activity in England.

Records of women giving birth ('delivery records') in the months since 1 January 2021 were identified in HES. De-duplication of delivery records resulted in a data set of women who had given birth with one record per woman, identified by her NHS Number, and the latest 'delivery

<sup>&</sup>lt;sup>1</sup> In this period 4,039 women could not be matched with a NIMS record. Their vaccine status is therefore unknown and they are excluded from these coverage figures.

<sup>&</sup>lt;sup>2</sup> Unlike previous reports, women giving birth in this period includes women who have previously given birth in the period of the HES maternity data extracts used from January 2021.

<sup>&</sup>lt;sup>3</sup> NIMS Data controllers are NHSEI and NHSD. The NIMS IT software is commissioned by NHSEI via South Central West CSU and is provided by the System C and Graphnet Care Alliance.

episode' associated with her. An 'earliest' and 'latest' likely pregnancy start date were assigned to each woman's record, using the known delivery date and further information from her record, where available:

- Where a valid gestational age was recorded (GESTAT\_1 between 24 and 42), the woman's earliest pregnancy start date was calculated by taking the number of weeks away from the delivery date, and then calculating an additional earlier week, to account for GESTAT\_1 recording completed weeks of pregnancy. In a similar way, latest pregnancy start date was calculated by taking the number of weeks of GESTAT\_1 away from the delivery date.
- 2. Where no valid GESTAT\_1 was available, the first 12 diagnostic codes were examined to identify any with a code suggesting delivery at term (O60.2). In this case, the gestational age at delivery was assumed to be between 37 and 42 completed weeks of pregnancy, and a similar method was used to establish earliest and latest pregnancy start dates.
- 3. Where no valid GESTAT\_1 was available and there were no codes suggesting term delivery, the first 12 diagnoses codes were examined to identify any suggesting preterm delivery (O60.1 or O60.3). In this case, the gestational age at delivery was assumed to be between 24 and 36 completed weeks of pregnancy, and these values were used to establish earliest and latest pregnancy start dates.
- 4. In the absence of any additional information in the woman's record (or in conflicting cases where diagnostic codes suggesting both term and pre-term delivery appeared in the same record), the gestational age at delivery was assumed to be between 24 and 42 completed weeks of pregnancy, and these values were used to establish earliest and latest pregnancy start dates.

Earliest and latest dates for the start of each trimester were established in a similar way, using the windows of trimester 1: day 0 to day 97 (where day 0 is the earliest or latest pregnancy start date, as established using the method above), trimester 2: day 98 to day 195 and trimester 3: day 196 to delivery. Each woman's delivery record was linked to her record(s) in the NIMS using the NHS Number, establishing her vaccine status as either having had one or more doses before delivery (including any prior to becoming pregnant) or not having had any doses of the vaccine prior to delivery, using the NIMS vaccine records. For a women to be identified as having had the autumn booster dose bivalent vaccine was recorded in NIMS on or after 1 September 2022.

For each vaccine dose (this analysis considered doses 1 to 4) the woman was known to have received, the following information was ascertained:

Dose administered pre-pregnancy	Dose administered before the earliest pregnancy start date
Dose administered in pregnancy	Dose administered after the latest pregnancy start date and before the delivery date
Dose administered post- pregnancy	Dose administered on or after the delivery date based on NIMS records extracted on 6 June 2023
Dose in pregnancy: unknown	Dose administered around the start or pregnancy: after the earliest pregnancy start date and before the latest pregnancy start date
Unvaccinated	No vaccine records exist for the woman, based on NHS number

And the following information about trimester:

Dose administered pre-pregnancy	Dose administered before the earliest pregnancy start date
Dose administered in trimester 1	Dose administered after the latest pregnancy start date and before the earliest pregnancy start date +97 days
Dose administered in trimester 2	Dose administered after the latest pregnancy start date +98 days and before the earliest pregnancy start date +195 days
Dose administered in trimester 3	Dose administered after the latest pregnancy start date + 196 days and before the delivery date
Dose administered post-pregnancy	Dose administered on or after the delivery date based on NIMS records extracted on 28 February 2023
Dose in trimester unknown	Dose administered in the 'gap' between trimesters, because of inaccuracy in establishing pregnancy start date
Unvaccinated	No vaccine records exist for the woman, based on NHS number

The ethnicity, residence and age information when used is taken from the NIMS record. The analysis within this section was carried out on 6 June 2023. The latest HES data available was for February 2023, and all HES data since April 2022 is considered provisional.

#### Interpretation and limitations

Previously published findings have supported vaccine safety in pregnancy from COVID-19 vaccine surveillance reports from week 47 2021 onwards <u>COVID-19 vaccine weekly</u>

<u>surveillance reports (weeks 39 to 40, 2021 to 2022)</u> and from analyses published by Public Health Scotland. <u>Outputs and information for the public – the University of Edinburgh</u>.

More detailed statistical analyses are planned (see <u>COVID-19 vaccination in pregnancy</u> <u>surveillance protocol</u>). There are recognised limitations of the data sets including the level of completeness of the relevant fields. Adverse pregnancy outcomes considered in earlier reports are routinely reported as official statistics annually by ONS, (see <u>COVID-19 hospital admissions</u> <u>by vaccination and pregnancy status</u>, <u>England</u>), however HES data was used to monitor outcomes more quickly than ONS data allows.

#### Main findings

COVID-19 vaccination is the safest and most effective way for women to protect themselves and their pregnancies against severe COVID-19 disease. The JCVI has advised that women who are pregnant are in a clinical risk group within the COVID-19 vaccine programme. Unvaccinated women who become pregnant are strongly encouraged to come forward for vaccination. Women who are pregnant and have previously been vaccinated should have been offered a booster dose this autumn (Joint Committee on Vaccination and Immunisation (JCVI) updated statement on the COVID-19 vaccination programme for autumn 2022).

COVID-19 vaccine coverage in pregnant women at delivery increased as more women have become eligible for vaccination, stabilising from May 2022. 73.2% of women who gave birth in December 2022 had received one or more dose before their baby was born. In Scotland, 78% of the women delivering in July 2022 had received at least one dose of COVID-19 vaccine prior to delivery, with 73% having received at least 2 doses and 48% having received at least 3 doses (42). In Wales estimated coverage at time of delivery for 2,018 women with delivery dates during July 2022 was 81% for at least one dose, 77% for at least 2 doses and 45% for at least one booster dose (data provided by Public Health Wales). This is the third report that includes coverage data for women who received the COVID-19 autumn boost vaccination before giving birth in England between September and February 2023 with 19.0% who gave birth in February 2023 receiving this vaccine before they gave birth. In Wales the estimated coverage for women who were pregnant as of 1 February 2023 was 10.7% (95% CI 10.8 to 11.9%) for the autumn 2022 booster (data provided by Public Health Wales). Outcome data and further break down of coverage data will be generated for the autumn boost when further vaccinated pregnancies have accumulated.

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

### Seroprevalence

The results from testing samples provided by healthy adult blood donors aged 17 years and older, supplied by the NHS Blood and Transplant (NHS BT collection) between weeks 35 2020 and week 21 2023 are summarised. As of week 44 2020, approximately 250 samples from each geographic NHS region are tested each week.

The COVID-19 vaccination campaign began on 8 December 2020 (week 50) with a phased roll out by age and risk group. From the beginning of September 2021, a third dose was offered to individuals with severe immunosuppression. A booster dose was introduced from 16 September 2021 for individuals aged 50 years and over, frontline health and social care staff, individuals aged 16 to 49 with certain underlying health conditions and household contacts of immunosuppressed individuals.

Eligibility for booster doses was extended to individuals aged 40 years and over from 22 November and from December to those aged 18 to 39 in a phased rollout by age group. Booster doses are generally given at least 6 months after the second dose, although the minimum interval was reduced to at least 3 months from the second or third dose in an effort to accelerate the roll out with the emergence of the Omicron variant. A second booster (spring booster) was introduced from 21 March 2022 for individuals aged 75 years and older, older residents in care homes and individuals with severe immunosuppression. The spring 2022 booster was offered 3 to 6 months after the last dose. An autumn booster was introduced from 7 September 2022 for individuals aged 50 years and over, those in care homes, individuals aged 5 years and over in clinical risk groups, frontline health and social care staff, those who care for vulnerable individuals and families of individuals with weakened immune systems, with those at highest risk being vaccinated first. The autumn 2022 booster was offered at least 3 months after the last dose. A spring 2023 booster was introduced from 3 April 2023 for adults aged 75 years and older, older adults in care homes and immunosuppressed individuals aged 5 years and older. The offer of the spring 2023 booster is available until the end June 2023 and is offered at least 3 months after the last dose.

Please note that this section will be updated quarterly. This update was published on 8 June 2023.

#### Seroprevalence in blood donors aged 17 years and older

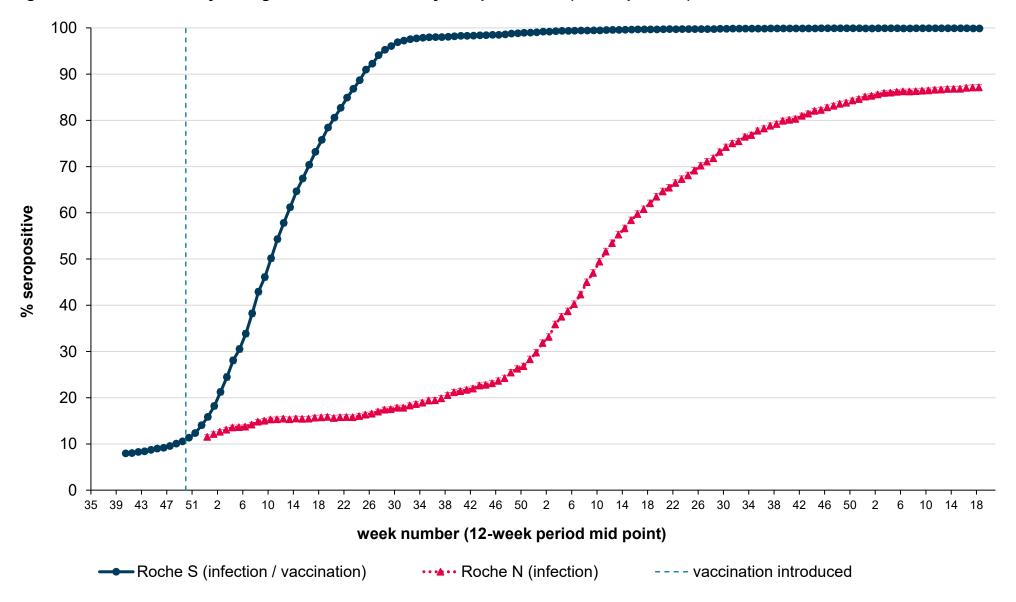
The results presented here are based on testing samples with Roche nucleoprotein (N) and Roche spike (S) antibody assays.

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 seropositivity for the Roche N assay reflect the effect of natural infection. Increases in seropositivity as measured by S antibody reflect both infection and vaccination. Antibody responses to both targets reflect infection or vaccination occurring at least 2 to 3 weeks previously given the time taken to generate a COVID-19 antibody response. Currently donors are asked to defer donations for at least 48 hours post vaccination (previously 7 days), and for at least 10 full days after a positive COVID-19 test as well as 7 days following resolution of any symptoms (previously 28 days, changes were implemented during January 2022).

This report presents Roche N and Roche S seropositivity estimates on the same set of samples, using a 12-week rolling prevalence for national, age group and regional estimates. Seropositivity estimates are plotted using the mid-point of a 12-weekly rolling period that reduces to 8 weeks in the most recent weeks to allow for a more representative current estimate of seropositivity. However, this also means the data will reflect seroprevalence several weeks previously. Seroprevalence estimates reported are based on seropositivity which are unadjusted for the sensitivity and specificity of the assays used.

#### National prevalence

Overall population weighted (by age group, sex and NHS region) antibody prevalence among blood donors aged 17 years and older in England was 87.1 % (95% CI 86.5% to 87.8%) using the Roche N assay and 99.9% (95% CI 99.8% to 99.9%) using the Roche S assay for the period 3 April to 28 May 2023 (week 14 to week 21 2023). 11,533 out of 13,224 were Roche N positive and 13,205 out of 13,224 samples were Roche S positive. This compares with 86.2% (95% CI 85.7% to 86.8%) Roche N seropositivity and 99.9% (95% CI 99.8% to 99.9%) Roche S seropositivity for the period of 12 January to 31 March 2023 (week 2 to week 13 2023). Seropositivity (weighted by region, age group and sex) varies over time. Figure 2 shows the overall 12-weekly rolling proportion seropositive over time for the Roche N and Roche S assays. Seropositivity estimates are plotted weekly using the mid-point of a rolling 12-weekly period.

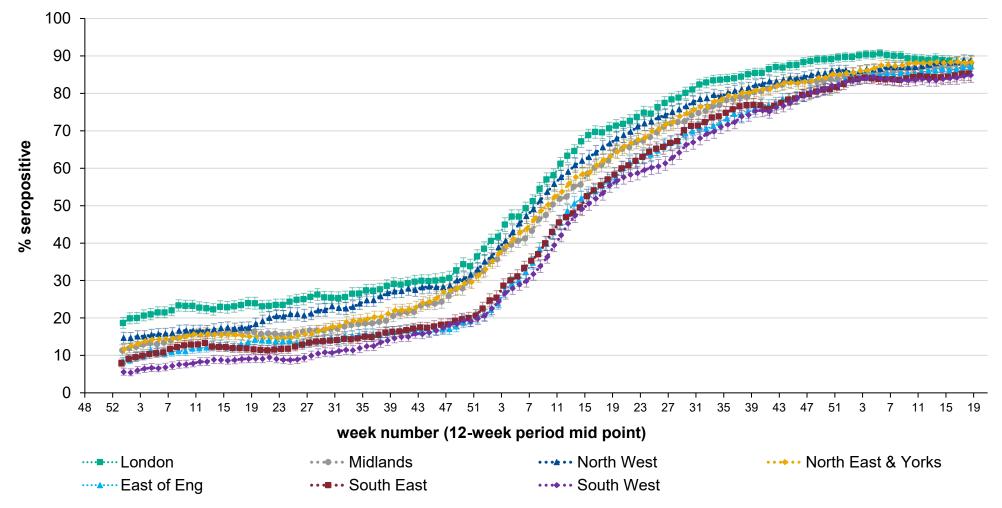




#### Regional prevalence of infection over time

Seropositivity (weighted by age group and sex) using the Roche N assay which detects infection only, varies by region (Figure 3).

## Figure 3. 12-weekly rolling SARS-CoV-2 antibody seroprevalence (% seropositive) in blood donors by region, using Roche N test; error bars show 95% confidence intervals



NHS region	Weeks 2 to 13 2023	Weeks 14 to 21 2023		
East of England	85.3% (83.9% to 86.6%)	86.9% (85.1% to 88.6%)		
London	90.2% (89.0% to 91.3%)	87.7% (85.8% to 89.4%)		
Midlands	85.3% (83.8% to 86.7%)	87.5% (85.9% to 89.0%)		
North East and Yorkshire	87.5% (86.2% to 88.8%)	88.3% (86.6% to 89.8%)		
North West	87.0% (85.6% to 88.3%)	88.6% (86.8% to 90.2%)		
South East	83.7% (82.2% to 85.1%)	85.3% (83.4% to 87.0%)		
South West	83.8% (82.3% to 85.3%)	84.9% (83.0% to 86.6%)		

Table 9. Roche N seropositivity (95%CI) estimates by NHS region

Increases in Roche N seropositivity have recently been observed in most regions with the largest increase seen in the Midlands (Table 9) compared to the previous 12-week period whilst a decrease was observed in London. Decreases in prevalence estimates are likely to reflect waning immunity and potential changes in the locations of sampling over time.

The difference in seropositivity by region has narrowed over time with the lowest seropositivity observed in the South West and the South East. The highest seropositivity has consistently been observed in London, closely followed by the North West, but in recent weeks seropositivity in the North West and in the North East and Yorkshire have surpassed London.

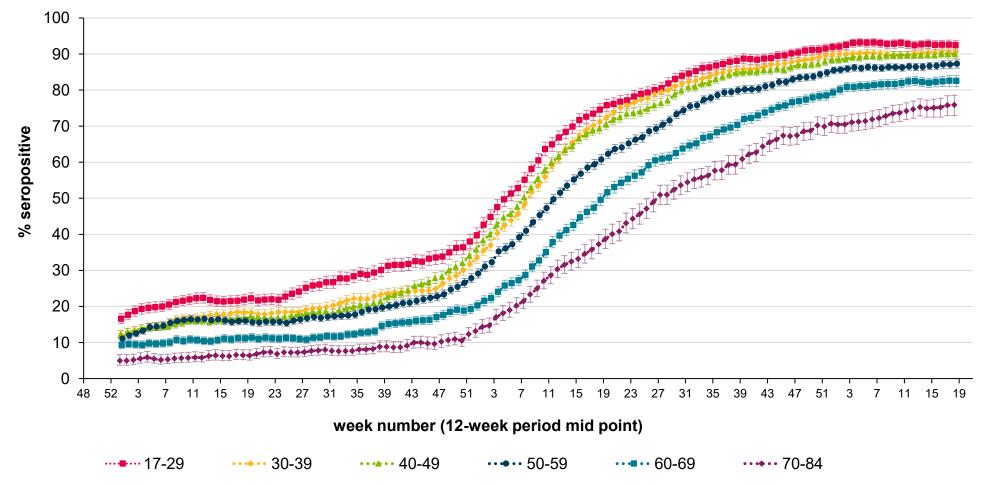
Overall COVID-19 case rates through Pillar 1, for week 20 2023, have decreased across all regions and overall positivity through Respiratory Datamart decreased compared to the previous week (Weekly national Influenza and COVID-19 surveillance report week 21 2023).

Pillar 1 testing is undertaken by NHS hospitals and UKHSA labs for those with a clinical need and some health and social care workers. <u>Testing recommendations</u> have been updated and routine asymptomatic testing through NHS settings has been paused since the end of August 2022, which will have an impact on Pillar 1 case rates and positivity rates. Changes in testing practices is likely to influence a range of surveillance indicators highlighting the importance of maintaining the serosurveillance programme to provide consistent data on exposure to infection and vaccine impact in the population over time.

### Prevalence by age group

Seropositivity estimates by age group using the Roche N assay are presented below.

Figure 4. Population weighted 12-weekly rolling SARS-CoV-2 antibody seroprevalence (% seropositive) in blood donors from the Roche N assay by age group



Based on testing samples using the Roche N assay (<u>Figure 4</u>) as a marker of infection, the highest seropositivity continues to be observed in those aged 17 to 29 and the lowest in those aged 70 to 84.

Age group	Weeks 2 to 13 2023	Weeks 14 to 21 2023
17 to 29	93.2% (92.0% to 94.2%)	92.5% (90.9% to 93.8%)
30 to 39	90.3% (89.3% to 91.2%)	90.4% (89.2% to 91.6%)
40 to 49	89.4% (88.4% to 90.4%)	90.0% (88.8% to 91.2%)
50 to 59	86.2% (85.2% to 87.1%)	87.3% (86.1% to 88.4%)
60 to 69	81.4% (80.1% to 82.6%)	82.6% (81.0% to 84.0%)
70 to 84	71.9% (69.4% to 74.2%)	75.9% (72.9% to 78.6%)

Table 10. Roche N seropositivity (95%CI) estimates by age group

Increases in N seropositivity have recently been observed across most age groups (Table 10) compared to the previous 12-week period. In the most recent period, the largest increase in seropositivity was observed in individuals aged 70 to 84 years. A small decrease in seropositivity was observed in individuals aged 17 to 29 years.

In England, Pillar 1 COVID-19 case rates for week 20 2023, decreased across all age groups with the highest rates currently seen in individuals aged 70 years and older (<u>Weekly national</u> <u>Influenza and COVIDto19 surveillance report week 21 2023</u>).

Roche S seropositivity in blood donors has plateaued and is now over 99% across all age groups. Seropositivity estimates for S antibody in blood donors are likely to be higher than would be expected in the general population and this probably reflects the fact that donors are more likely to be vaccinated. Seropositivity estimates for N-antibody will underestimate the proportion of the population previously infected due to (i) waning of the N-antibody response over time and (ii) observations from UKHSA surveillance data that N-antibody levels are lower in individuals who acquire infection following vaccination. These lower N antibody responses in individuals with breakthrough infections (post-vaccination) compared to primary infection likely reflect the shorter and milder infections in these patients. Patients with breakthrough infections do have significant increases in S antibody levels consistent with boosting of their antibody levels.

Vaccination has made an important contribution to the overall Roche S increases observed since the roll out of the vaccination programme. The impact of the booster vaccination programmes can be assessed by monitoring Roche S antibody levels across the population over time.

## Roche S levels by age group and month

The Roche S assay that the UKHSA uses for serological surveillance is fully quantitative, meaning that it measures the level of antibodies in a blood sample; an antibody level above 0.8

au/ml (approximately 1 IU/ml using the WHO standard) is deemed positive. The UKHSA surveillance since Autumn 2021 has found that over 99% of the population of blood donors test positive for S-antibodies, which may have resulted from either COVID-19 infection or vaccination. With such high seropositivity, it is important to look at population antibody levels in order to assess the impact of the vaccination booster programmes.

Figure 5 shows monthly categorised Roche S levels in N-antibody negative individuals by age group over the past year. After high antibody levels following the rollout of the accelerated booster programme due to the emergence of the Omicron variant in late 2021, the decreasing profile of antibody levels among 17 to 69 year olds from June through to August shows signs of waning. From April 2022 the proportion of donors, aged 70 to 84 years, with very high antibody levels of 25,000+ au/ml increased following the introduction of the spring booster for ages 75 and older. Antibody levels in this older age group have remained consistent from May to September 2022. A further increase in highest antibody levels can be seen in October following the Autumn booster in those aged 40 to 84, with the highest increase seen in those aged 70 to 84. In November 2022 the proportion of donors aged 70 to 84 with very high antibody levels of 25,000+ au/ml remained high and the proportion of donors aged 50 to 69 years with very high antibody levels increased. From January 2023 the proportion of donors, aged 50 to 84 years, with very high antibody levels of 25,000+ au/ml remained high and the proportion of donors, aged 50 to 69 years with very high antibody levels of 25,000+ au/ml remained high and the proportion of donors.

By 21 May 2023, 52.9% of all people aged 75 years and older, living in England, had been vaccinated with a Spring 2023 booster dose (<u>Weekly national Influenza and COVIDto19</u> <u>surveillance report week 21 2023</u>).

Figure 6 shows categorised Roche S levels in N-antibody positive individuals, those likely to have experienced past infection. Pre-vaccination antibody levels will be influenced by time since infection, variant and severity of infection, as well as individual factors such as underlying health conditions and age. From June to September 2022 the proportion of donors with very high antibody levels of 25,000+ au/ml decreased or remained stable across all age groups. Increases of the highest antibody levels were seen across those aged 60 to 84 in October following rollout of the Autumn booster. In those aged 17 to 59 very little change has been observed between June and October 2022. In November 2022 increases in the proportion of donors aged 50 to 84 years. Since January 2023 decreases were seen in the proportion of donors, aged between 50 to 84 years, with very high antibody levels of 25,000+ au/ml.

Comparing Figure 5 with Figure 6, the overall higher profile of antibody levels in those who have experienced past infection is evident; both vaccination post infection and breakthrough infection following vaccination are expected to boost existing antibody levels. Whilst it is thought that there is no threshold antibody level that offers complete protection against infection, higher antibody levels are likely to be associated with lower probability of infection.



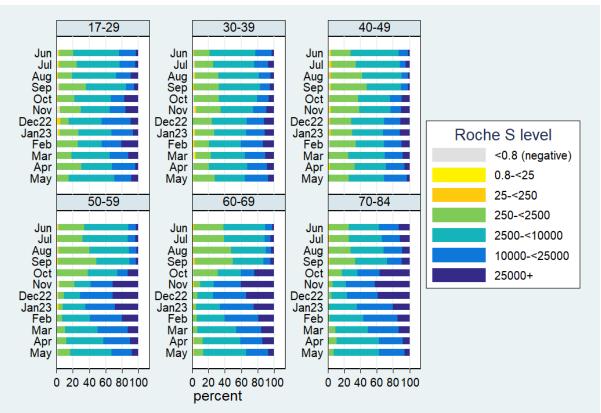
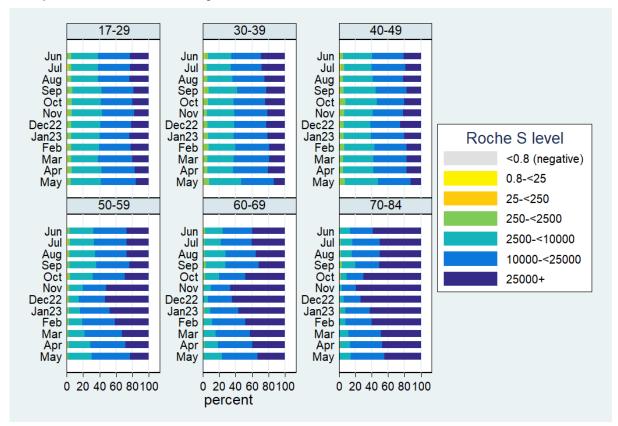


Figure 6. Categorised Roche S antibody levels by age group and month in N positive samples, June 2022 to May 2023



## **SARI-Watch surveillance data**

SARI-Watch is a national surveillance system in England that captures aggregate and individual data on laboratory confirmed COVID-19 hospitalisations (to all level of care) and ICU or HDU admissions. The individual level data is based on hospitalisations for laboratory confirmed COVID-19 submitted by a sentinel network of acute NHS trusts.

The analyses presented are based on individual level data from the sentinel collection in England. The analysis uses data from 3 April 2023 to 28 May 2023 inclusive submitted by 9 acute NHS trusts. 3 April 2023 was the official roll out of the Spring 2023 booster campaign (targeted at people aged 75 years and over, residents in care homes and those aged 5 years and over with immuno-suppression). The data analysed was up to 28 May 2023 as this was last date of ISO week 21 2023. To obtain vaccine history for admitted persons, the SARI-Watch data was linked to the NIMS on 2 June 2023 using key personal identifiers. NIMS captures data on vaccination history of people in England. Data on vaccination history was used to group admissions into 4 categories. Please note that vaccination status on admission does not represent VE against hospitalisation. Vaccination is highly protective against hospitalisation as VE studies show (see vaccine effectiveness section in this report) but does not completely eliminate the risk of being hospitalised. Hence those that get hospitalised even if vaccinated tend to be the vulnerable elderly population or those with significant underlying health issues. Other caveats are that London trusts are currently under-represented in sentinel data and the most recent data is typically subject to retrospective updates after submission.

## Vaccination status by time of admission by age group

Vaccination status on hospital admission was grouped into 4 categories:

- 'Unvaccinated' no evidence of previous vaccination at the time of admission
- 'D1' comprising only 1 dose of the primary course by the time of admission
- 'D2' comprising only 2 doses of the primary course by the time of admission
- 'D3 comprising 3 doses or more by the time of admission. This includes 3 doses of primary course only or 2 doses of the primary course plus any or all of the boosters (autumn 2021, spring 2022, autumn 2022 or spring 2023 boosters).

The unlinked group represents hospitalised cases that could not be matched to NIMS data either due to incorrect or missing personal identifiers. This accounted for 1% of data in the 3 April 2023 to 28 May 2023 period.

<u>Table 11</u> shows vaccination status by the time of admission and age group among admitted cases from 3 April 2023 to 28 May 2023 inclusive (n=797 admissions).

Those aged 75 years or more had the lowest proportion that were unvaccinated by the time of admission at 2.1% in this period. The highest proportion that were unvaccinated by time of admission was in <40 years at 61.8% (Table 11). Those in the category ' $\geq$ 3 doses' by time of admission accounted for 93.9% of admitted cases aged 75 years or more (Table 11). This compares with 17.1% in those aged <40 years.

Age group	Data type	Unvaccinated	1 dose (primary)	2 doses (primary)	≥3 doses	Unlinked	Total
Under 40	Number	47	5	9	13	2	76
	%	61.8	6.6	11.8	17.1	2.6	
40 to 49	Number	4	1	4	20	0	29
	%	13.8	3.4	13.8	69.0	0.0	
50 to 64	Number	8	2	4	73	1	88
	%	9.1	2,3	4.6	83.0	1.1	
65 to 74	Number	9	1	2	118	1	131
	%	6.9	0.8	1.5	90.1	0.8	
Over 75	Number	10	1	14	444	4	473
	%	2.1	0.2	3.0	93.9	0.8	

Table 11. Vaccination status at time of admission by age group for admissions from 3April 2023 to 28 May 2023, sentinel data, England

# Rate of hospitalisation for COVID-19 in vaccinated people by time since vaccination (any dose) and age group

Using linked sentinel data, admissions from 1 to 28 May 2023 inclusive were analysed as this represented 4 weeks mostly covering May 2023 (ISO weeks 18 to 21). This period will match the ISO week system used in the mandatory aggregate collection (ISO weeks 18 to 21 2023, used in second step of the calculation – see next paragraph). In this time period 9 sentinel acute NHS trusts contributed data. Time since last vaccination at the time of admission was calculated based on time in days between the last vaccination and hospital admission date. The last vaccination is irrespective of any dose whether from the primary or the booster schedule. The interval for each admitted case was grouped into <3 months, 3 to under 6 months, 6 to under 9 months, 9 to under 12 months and 12 months and over since last vaccination. Each month comprises 30 days. The proportion falling in each interval was calculated per age group.

The proportion by time since last vaccination and age group obtained from sentinel data was then applied to the corresponding age group for cases from the mandatory aggregate collection. Cases from the aggregate collection was used as this is a mandatory surveillance system based on wider national reporting. The mandatory surveillance is based on weekly data (based on ISO week system from Monday to Sunday). Four weeks in 2023 were used from the aggregate data (ISO weeks 18 to 21) covering 1 to 28 May inclusive. Due to mean 67% trust coverage in the period of study (1 to 28 May 2023 inclusive) in the aggregate collection, a corrective factor was applied to the second step of the calculation. The second step is where the proportions by age group and time since vaccination from sentinel data is applied to the cases from the aggregate collection in same age group. The corrective factor would approximate the true total cases if there was 100% trust coverage.

To estimate rates of hospitalisation among vaccinated people by time since last vaccination and age group, the NIMS denominator is required. Using NIMS data, the time since last vaccination in days was calculated from vaccination date (any dose) capped to 28 May 2023. The time in days was grouped in the same intervals described. The same age and/time since vaccination groups are used for SARI-Watch and NIMS, with the former being the numerator and latter being the denominator. The rate of hospitalisation is calculated for each age/time since last vaccination group, expressed as hospitalised cases in the study period (1 to 28 May 2023) per 100,000 vaccinated people in England.

It should be noted that the linked surveillance data did not yield a high overall number of cases for the period in question. This is likely due to a decrease in circulating Omicron variants observed since March 2023. A decrease in the hospitalisation rates for COVID-19 was also observed in the mandatory surveillance data separately. The smallest number of cases in the linked surveillance was for ages <40 and 40 to 49 years hence could not support meaningful stratification by time since last vaccination. Hence the rate per 100,000 vaccinated in the <40 and 40 to 49 year groups by time since last vaccination interval are not presented in this analysis.

In the linked surveillance data, 13.0% of hospitalised cases in May 2023 had the Spring 2023 booster. The majority (88.3%) of those who had this booster were aged 75 years and over.

Figure 7 shows that the highest hospitalisation rates were in those aged 75 years and over for most time since last vaccination intervals compared to corresponding intervals in other age groups. Usually, the rate in the shortest interval is highest in those aged 75 years compared to the corresponding interval in other age groups. However, in this analysis, 50 to 64 year age group had the highest rate for the <3 months interval.

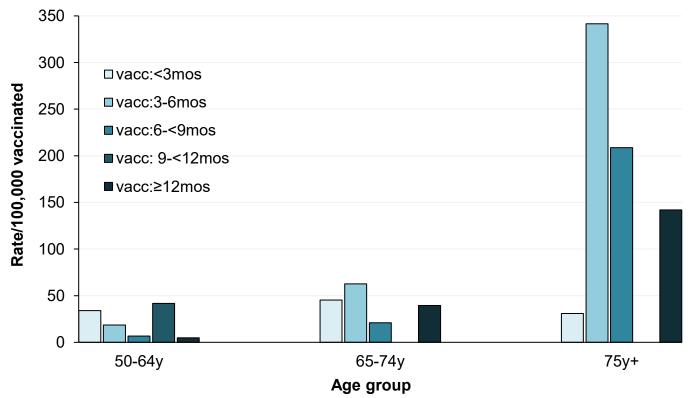
Among those aged 75 years or more, the hospitalisation rate was lowest in the shortest interval of <3 months (30.9 out of 100,000). All cases in this interval had the Spring 2023 booster based on linked data. The rate for this interval decreased since the previous report likely due to less circulating Omicron variants and the impact of recent vaccination by May 2023. The NIMS data showed that nationally 57.9% (3,169,030 out of 3,668,640) of people aged 75 years or more had their last vaccination within 3 months of 28 May 2023. This reflects the expansion of the Spring Booster 2023 vaccination programme since 3 April 2023. The rate in the next interval (3

to <6 months) increased to 341.5 out of 100,000 vaccinated, the increase coinciding with increases in XBB.1.5 and XBB Omicron sublineages. The rate per 100,000 vaccinated dropped in subsequent intervals although the rates in later intervals still exceeded the rate for the shortest interval. Note there was no data for the interval of 9 to <12 months since last vaccination due to 0 admitted cases for this interval in the individual level linked data.

The rates in younger age groups were much lower reflecting progressively lower risk of hospitalisation. However, the gradient is not clear, that is the general pattern of the rate increasing as time since last vaccination increases. In the 65 to 74 years age group, although the rate in the <3 months interval was lower than the 3 to <6 months interval (45.4 out of 100,000 and 62.6 out of 100,000 respectively), the subsequent intervals were lower than the shortest interval. The rate in the shortest interval in this age group was also the highest compared to corresponding interval in other age groups. Note there were no admitted cases reported for the 9 to <12 months interval based on the individual level linked data.

The rates in 50 to 64 years requires careful interpretation as the time since last vaccination percentages derived from linked data were based on small volumes for the period under study. It is possible that the clinical risk factors that make younger adults eligible for the spring 2023 booster vaccination may also make them more pre-disposed to hospital admissions than the general age-matched population. For example, patients with immunosuppression in whom the vaccine <u>may be less effective</u>. The combined effect of these factors may have elevated the risk in the shortest interval.

## Figure 7. Estimated rate of hospitalisation for COVID-19 per 100,000 vaccinated people by time since last vaccination (any dose) and age group, admissions from 1 May 2023 to 28 May 2023 inclusive (ISO weeks 18 to 21 2023), England



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