## In brief

Omicron (B.1.1.529)

updated 27 January 2022

## Summary

- Omicron (B.1.1.529) was designated a variant of concern by the World Health Organization on 26 November 2021. It was first reported to WHO from South Africa on 24 November 2021. It has 50 mutations, including 26-32 mutations on its spike protein.<sup>1,2</sup>
- Omicron has been separated into two lineages: BA.1 and BA.2. <u>BA.2 does not contain the spike</u> <u>deletion</u> and therefore is S-gene positive (SGTP). BA.1 was previously dominant, however <u>BA.2 is</u> <u>increasing</u> in India, South Africa, Denmark and the UK. The UK designated BA.2 as a 'variant under investigation' on 21 January 2022.<sup>3, 4</sup>
- The World Health Organization notes that based on current evidence, Omicron has a substantial growth advantage over Delta.<sup>2</sup> Likely factors contributing to the growth rate include <u>immune</u> evasion and potential intrinsic increased transmissibility.<sup>5</sup> One study estimates that <u>Omicron is</u> 36.5% more transmissible than Delta and that Omicron erodes 63.7% of the population immunity accumulated from prior infection and vaccination.<sup>6</sup>
- Omicron infections feature lower peak viral RNA and a shorter clearance phase than Delta.
   Preliminary data suggests that the amount of viral RNA is <u>highest three to six days after diagnosis</u> or symptom onset.<sup>7,8</sup>
- <u>Household secondary attack rate</u> estimates range from 15.8% to 31% for Omicron compared to 10.3% to 21% for Delta. One study suggests <u>increased transmission</u> for unvaccinated individuals, and reduced transmission for booster-vaccinated individuals, compared to fully vaccinated individuals.<sup>9, 10</sup>
- Epidemiological data in the Gauteng Province, South Africa, showed SARS-CoV-2 infection rates increased more rapidly than in previous waves but have now plateaued.<sup>11</sup>
- The risk of reinfection is estimated to be <u>16 times higher</u> than Delta. Unvaccinated individuals are twice as likely to be reinfected than people who had their second vaccine 14 to 89 days ago. Individuals are more likely to be reinfected if they had lower viral loads at their first infection. <u>Reinfection has been reported in several countries</u> including South Africa, Denmark, Israel and the United Kingdom.<sup>12, 13</sup>
- Preliminary data from South Africa, England, Scotland and Denmark show that people infected with the Omicron variant are less likely to require hospitalisation compared with Delta. This ranges from 40-45%, up to <u>90%</u> less likely.<sup>3, 14-17</sup> <u>Length of stay</u> is significantly shorter for Omicron compared to Delta.<sup>18</sup>
- International data shows an <u>increase in hospital admissions</u> for children under five years old.
   Expert comment suggests few children admitted to hospital are needing <u>intensive care</u>.<sup>3, 19</sup>
- <u>Laboratory studies</u> suggest Omicron does not infect cells deep in the lungs as readily as it does those in the upper airways.<sup>20</sup>
- Therapeutic interventions for the management of severe or critical COVID-19 that target host responses (such as corticosteroids, interleukin-6 receptor blockers and prophylaxis with anticoagulation) are expected to remain effective.<sup>2</sup> Evidence for other interventions include:



- Antiviral medications will still likely be effective for managing COVID-19, including Paxlovid (nirmatrelvir plus ritonavir), molnupiravir and remdesivir.<sup>21-23</sup>
- Treatments that target the spike protein of the virus, such as monoclonal antibodies, may be less effective, but this will require assessment.<sup>9, 23, 24</sup> Early findings suggest Omicron will likely compromise the binding of many monoclonal antibodies.<sup>25, 26</sup>
- Preprint results for monoclonal antibody sotrovimab show it retains neutralising activity against all tested individual Omicron substitutions in laboratory tests, while early tests show Regen-Cov (casirivimab and imdevimab) is not as effective against Omicron.<sup>22, 27, 28</sup>
- Studies are underway to understand the effectiveness of vaccines. Early estimates of vaccine
  effectiveness against symptomatic infection indicate significantly lower effectiveness against
  Omicron compared with Delta. However, moderate to high vaccine effectiveness of up to 70%
  against symptomatic infection is seen in the early period after a booster dose.<sup>29</sup> Early results have
  found:
  - The T-cell immune response in previously infected, and most likely vaccinated individuals, should still be effective against Omicron.<sup>30</sup>
  - Neutralising activity of sera from individuals who are vaccinated plus infected, or infected plus vaccinated (also called hybrid immunity), holds well against Omicron.<sup>31</sup>
  - Studies on specific vaccines include:
    - Comirnaty
      - A 20- to 40-fold reduction in neutralising activity by two doses of Comirnaty compared with other strains.<sup>32</sup>
      - A booster dose of Comirnaty resulted in an increase in neutralising activity irrespective of primary vaccination type (approximately 71% for those who received Vaxzevria as the primary course and approximately 76% for those who received Comirnaty).<sup>32</sup>
      - Two doses of Comirnaty offers 70% protection against hospitalisation <sup>33</sup> and up to 92% following three doses. <sup>3</sup>
    - Spikevax
      - preliminary data suggests an unadjusted vaccine effectiveness of <u>16.5% for</u> <u>two doses of Spikevac and 100% for three doses</u>.<sup>34</sup>
      - a booster dose of Spikevax at 50 microgram (ug) level increases Omicron neutralising antibody levels approximately 37-fold compared to pre-boost level. A booster dose at 100ug increased the Omicron neutralising antibody levels approximately 83-fold.<sup>35</sup>
    - Vaxzevria
      - Preliminary studies suggest a significant reduction in vaccine effectiveness at 15 weeks after the second dose of Vaxzevria.<sup>36</sup>
    - Johnson & Johnson
      - Preliminary data suggests vaccine effectiveness of two doses of Janssen-<u>Ad26.COV2.S against hospitalisation with Omicron</u> 14-27 days post booster, as compared to unvaccinated healthcare workers, was 84% (67 to 92%), which was maintained 1-2 months after a booster. <sup>37</sup>





- <u>News reports from Israel</u> on four doses of COVID-19 vaccine suggest threefold protection against serious illness and twofold protection against infection in the current wave driven by Omicron. <sup>38</sup>
- The diagnostic accuracy of routinely used polymerase chain reaction (PCR) and antigen-based rapid diagnostic test (Ag-RDT) assays does not appear to be influenced by Omicron.<sup>2</sup>
- Most Omicron variant sequences reported include a deletion in the S gene, causing some S gene targeting PCR assays to appear negative, and so S gene target failure can be used as a useful proxy marker of Omicron for surveillance.<sup>2</sup>

To inform this brief, PubMed and Google searches were conducted using terms related to Omicron on 9 December 2021 and updated on 24 January 2022. Wording of the summary was updated on 9 March 2022 but an updated search was not carried out. The Critical Intelligence Unit maintains a living evidence table on <u>SARS-CoV-2 variants</u>. <sup>39</sup>

## References

- 1. World Health Organization. Classification of Omicron (B.1.1.529): SARS-CoV-2 Variant of Concern [Internet]. Switzerland: WHO; 26 Nov 2021 [cited 3 Dec 2021]. Available from: <u>https://www.who.int/news/item/26-11-2021-classification-of-omicron-(b.1.1.529)-sars-cov-2-variant-of-concern</u>.
- 2. World Health Organization. Enhancing Readiness for Omicron (B.1.1.529): Technical Brief and Priority Actions for Member States [Internet]. Switzerland: WHO; 17 Dec 2021 [cited 22 Dec 2021]. Available from: <u>https://www.who.int/publications/m/item/enhancing-readiness-for-omicron-(b.1.1.529)-technical-brief-and-priority-actions-for-member-states</u>.
- UK Health Security Agency. SARS-CoV-2 variants of concern and variants under investigation in England, Technical briefing 34 [Internet]. England: UK Health Security Agency; 14 Jan 2022 [cited 19 Jan 2022]. Available from: <u>https://assets.publishing.service.gov.uk/government/uploads/system/uploads/attachment\_data/fi</u> <u>le/1046853/technical-briefing-34-14-january-2022.pdf</u>.
- 4. World Health Organization. Enhancing Readiness for Omicron (B.1.1.529): Technical Brief and Priority Actions for Member States [Internet]. Switzerland: WHO; 21 Jan 2022 [cited 27 Jan 2022]. Available from: <u>https://www.who.int/publications/m/item/enhancing-readiness-for-</u>omicron-(b.1.1.529)-technical-brief-and-priority-actions-for-member-states.
- 5. Pearson CAB, Silal SP, Li MWZ, et al. Bounding the levels of transmissibility & immune evasion of the Omicron variant in South Africa. medRxiv. 2021:2021.12.19.21268038. DOI: 10.1101/2021.12.19.21268038
- Yang W, Shaman J. SARS-CoV-2 transmission dynamics in South Africa and epidemiological characteristics of the Omicron variant. medRxiv. 2021:2021.12.19.21268073. DOI: 10.1101/2021.12.19.21268073
- Hay JA, Kissler SM, Fauver JR, et al. Viral dynamics and duration of PCR positivity of the SARS-CoV-2 Omicron variant. medRxiv. 2022:2022.01.13.22269257. DOI: 10.1101/2022.01.13.22269257
- 8. National Institute of Infectious Diseases. Active epidemiological investigation on SARS-CoV-2 infection caused by Omicron variant (Pango lineage B.1.1.529) in Japan: preliminary report on infectious period [Internet] Japan: National Institute of Infectious Diseases; 5 Jan 2022 [cited 20 Jan 2022] Available from: https://www.niid.go.jp/niid/en/2019-ncov-e/10884-covid19-66-en.html.
- UK Health Security Agency. SARS-CoV-2 variants of concern and variants under investigation in England, Technical briefing 32 [Internet]. England: UK Health Security Agency; 17 Dec 2021 [cited 22 Dec 2021]. Available from: https://assets.publishing.service.gov.uk/government/uploads/system/uploads/attachment\_data/fi

https://assets.publishing.service.gov.uk/government/uploads/system/uploads/attachment\_data/fi le/1042688/RA Technical Briefing 32 DRAFT 17 December 2021 2021 12 17.pdf.



- 10. Lyngse FP, Mortensen LH, Denwood MJ, et al. SARS-CoV-2 Omicron VOC Transmission in Danish Households. medRxiv. 2021:2021.12.27.21268278. DOI: 10.1101/2021.12.27.21268278
- Madhi S, Kwatra G, Myers JE, et al. South African Population Immunity and Severe Covid-19 with Omicron Variant. medRxiv. 2021:2021.12.20.21268096. DOI: 10.1101/2021.12.20.21268096
- Office for National Statistics. Coronavirus (COVID-19) Infection Survey, characteristics of people testing positive for COVID-19, UK: 19 January 2022 [Internet] United Kingdom: Office for National Statistics; 19 Jan 2022 [cited 20 Jan 2022] Available from: <u>https://www.ons.gov.uk/peoplepopulationandcommunity/healthandsocialcare/conditionsanddise</u> <u>ases/bulletins/coronaviruscovid19infectionsurveycharacteristicsofpeopletestingpositiveforcovid1</u> <u>9uk/19january2022</u>.
- World Health Organization. Enhancing response to Omicron SARS-CoV-2 variant [Internet]. Switzerland: WHO; 7 Jan 2022 [cited 14 Jan 2022]. Available from: <u>https://www.who.int/publications/m/item/enhancing-readiness-for-omicron-(b.1.1.529)-technical-brief-and-priority-actions-for-member-states</u>.
- 14. Ferguson N, Ghani A, Hinsle W, et al. Report 50 Hospitalisation risk for Omicron cases in England [Internet] United Kingdom: Imperial College London; 22 December 2021 [cited 23 December 2021] Available from: <u>https://www.imperial.ac.uk/mrc-global-infectious-disease-analysis/covid-19/report-50-severity-omicron/</u>.
- 15. Wolter N, Jassat W, Walaza S, et al. Early assessment of the clinical severity of the SARS-CoV-2 Omicron variant in South Africa. medRxiv. 2021:2021.12.21.21268116. DOI: 10.1101/2021.12.21.21268116
- 16. Sheikh A, Kerr S, Woolhouse M, et al. Severity of Omicron variant of concern and vaccine effectiveness against symptomatic disease: national cohort with nested test negative design study in Scotland. [Internet]. University of Edinburgh. 22 Dec 2021. Available from: <u>https://www.research.ed.ac.uk/en/publications/severity-of-omicron-variant-of-concern-and-vaccine-effectiveness-</u>.
- 17. Barnes O, Burn-Murdoch J, Milne R. Omicron cases less likely to require hosptial treatment, studies show [Internet] Financial times; 22 Dec 2021 [Accessed 23 Dec 2021] Available from: https://www.ft.com/content/19065fba-025c-43fd-bd76-37234af97953.
- 18. Peralta-Santos A, Rodrigues EF, Moreno J, et al. Omicron (BA.1) SARS-CoV-2 variant is associated with reduced risk of hospitalization and length of stay compared with Delta (B.1.617.2). medRxiv. 2022:2022.01.20.22269406. DOI: 10.1101/2022.01.20.22269406
- 19. Royal College of Paediatrics and Child Health. RCPCH comments on reports of increased admissions of under 5s in hospital with COVID-19 [internet] United Kingdom: RCPCH; 13 Jan 2022 [cited 20 Jan 2022] Available from: <u>https://www.rcpch.ac.uk/news-events/news/rcpch-comments-reports-increased-admissions-under-5s-hospital-covid-19</u>.
- 20. Diamond M, Halfmann P, Maemura T, et al. The SARS-CoV-2 B.1.1.529 Omicron virus causes attenuated infection and disease in mice and hamsters. Res Sq. 2021 Dec 29. DOI: 10.21203/rs.3.rs-1211792/v1
- 21. McGregor G. Why COVID pills are likely to maintain efficacy against the Omicron COVID variant—even if vaccines don't [Internet]. United States: Fortune; 1 Dec 2021 [cited 3 Dec 2021]. Available from: <u>https://fortune.com/2021/12/01/covid-omicron-new-variant-antiviral-pill-vaccine-pfizer-mutations/</u>.
- 22. Terry M. Early Tests Show Omicron Tougher for Current COVID Antibody Therapies to Tackle [Internet]. United States: BioSpace; 30 Nov 2021 [cited 3 Dec 2021]. Available from: <u>https://www.biospace.com/article/preliminary-tests-suggest-covid-19-antibody-therapies-may-not-be-as-effective-against-omicron/</u>.
- 23. Willis O, Smith B. Omicron: What we know about the newest COVID variant of concern [Internet]. Australia: ABC News; 29 Nov 2021 [cited 3 Dec 2021]. Available from: <u>https://www.abc.net.au/news/health/2021-11-29/omicron-covid-19-variant-of-concern-what-we-know/100658678</u>.



- 24. World Health Organization. Update on Omicron [Internet]. Switzerland: WHO; 28 Nov 2021 [cited 3 Dec 2021]. Available from: <u>https://www.who.int/news/item/28-11-2021-update-on-omicron</u>.
- 25. Zahradnik J, Tuekprakhon A, Ginn HM, et al. Receptor binding and escape from Beta antibody responses drive Omicron-B.1.1.529 evolution. bioRxiv. 2021:2021.12.03.471045. DOI: 10.1101/2021.12.03.471045
- 26. Woo HG, Shah M. Omicron: A heavily mutated SARS-CoV-2 variant exhibits stronger binding to ACE2 and potently escape approved COVID-19 therapeutic antibodies. bioRxiv. 2021:2021.12.04.471200. DOI: 10.1101/2021.12.04.471200
- 27. Wilhelm A, Widera M, Grikscheit K, et al. Reduced Neutralization of SARS-CoV-2 Omicron Variant by Vaccine Sera and monoclonal antibodies. medRxiv. 2021:2021.12.07.21267432. DOI: 10.1101/2021.12.07.21267432
- 28. Cathcart AL, Havenar-Daughton C, Lempp FA, et al. The dual function monoclonal antibodies VIR-7831 and VIR-7832 demonstrate potent in vitro and in vivo activity against SARS-CoV-2. bioRxiv. 2021:2021.03.09.434607. DOI: 10.1101/2021.03.09.434607
- 29. UK Health Security Agency. SARS-CoV-2 variants of concern and variants under investigation in England, Technical briefing 35 [Internet]. England: UK Health Security Agency; 28 Jan 2022 [cited 31 Jan 2022]. Available from: https://assets.publishing.service.gov.uk/government/uploads/system/uploads/attachment\_data/fi le/1050999/Technical-Briefing-35-28January2022.pdf.
- 30. Redd AD, Nardin A, Kared H, et al. Minimal cross-over between mutations associated with Omicron variant of SARS-CoV-2 and CD8+ T cell epitopes identified in COVID-19 convalescent individuals. bioRxiv. 2021:2021.12.06.471446. DOI: 10.1101/2021.12.06.471446
- 31. Roessler A, Riepler L, Bante D, et al. SARS-CoV-2 B.1.1.529 variant (Omicron) evades neutralization by sera from vaccinated and convalescent individuals. medRxiv. 2021:2021.12.08.21267491. DOI: 10.1101/2021.12.08.21267491
- 32. UK Health Security Agency. SARS-CoV-2 variants of concern and variants under investigation in England, Technical briefing 31 [Internet]. England: UK Health Security Agency; 10 Dec 2021 [cited 14 Dec 2021]. Available from: <u>https://assets.publishing.service.gov.uk/government/uploads/system/uploads/attachment\_data/fi</u> le/1040076/Technical Briefing 31.pdf.
- 33. Winning A, Roelf W. Pfizer shot less effective in South Africa after Omicron emerges, study shows [Internet]. United Kingdom: Reuters; 15 Dec 2021 [cited 15 Dec 2021]. Available from: <u>https://www.reuters.com/business/healthcare-pharmaceuticals/pfizer-vaccine-protecting-against-hospitalisation-during-omicron-wave-study-2021-12-14/.</u>
- 34. World Health Organization. Weekly epidemiological update on COVID-19 11 January 2022 [internet] Switzerland: WHO; 11 Jan 2022 [cited 20 Jan 2022] Available from: <u>https://www.who.int/publications/m/item/weekly-epidemiological-update-on-covid-19---11-january-2022</u>.
- 35. Moderna. Moderna announces preliminary booster data and updates strategy to address omicron variant [Internet]. United States: Moderna; 20 Dec 2021 [cited 22 Dec 2021 Dec]. Available from: <a href="https://investors.modernatx.com/news/news-details/2021/Moderna-Announces-Preliminary-Booster-Data-and-Updates-Strategy-to-Address-Omicron-Variant/default.aspx">https://investors.modernatx.com/news/news-details/2021/Moderna-Announces-Preliminary-Booster-Data-and-Updates-Strategy-to-Address-Omicron-Variant/default.aspx</a>.
- 36. Schubert M, Bertoglio F, Steinke S, et al. Human serum from SARS-CoV-2 vaccinated and COVID-19 patients shows reduced binding to the RBD of SARS-CoV-2 Omicron variant in comparison to the original Wuhan strain and the Beta and Delta variants. medRxiv. 2021:2021.12.10.21267523. DOI: 10.1101/2021.12.10.21267523
- 37. Gray GE, Collie S, Garrett N, et al. Vaccine effectiveness against hospital admission in South African health care workers who received a homologous booster of Ad26.COV2 during an Omicron COVID19 wave: Preliminary Results of the Sisonke 2 Study. medRxiv. 2021:2021.12.28.21268436. DOI: 10.1101/2021.12.28.21268436
- 38. Staff T. Health Ministry: 4th dose triples protection from serious illness for over-60s [internet] Israel: The times of Israel; 23 Jan 2022 [cited 27 Jan 2022] Available from:



https://www.timesofisrael.com/health-ministry-4th-dose-triples-protection-from-serious-illness-for-over-60s/

 COVID-19 Critical Intelligence Unit. Living Evidence - SARS-CoV-2 variants [internet] Australia: Agency for Clinical Innovation; 24 Jan 2022 [cited 27 Jan 2022] Available from: <u>https://aci.health.nsw.gov.au/covid-19/critical-intelligence-unit/sars-cov-2-variants</u>.

SHPN: (ACI) 220161 | TRIM: ACI/D21/695-62 | Edition 5

