

Literature screening report

COVID-19 vaccines in the WHO's Emergency Use Listing (EUL) Report (10)

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Abstract

This report provides an in-depth review of the **seven¹** World Health Organization's (WHO) Emergency Use Listing (EUL) authorized vaccines: BNT162b2/ COMIRNATY (Pfizer-BioNTech, USA), Spikevax/ Moderna COVID-19 Vaccine/ mRNA-1273 (Moderna, USA), Vaxzevria/ ChAdOx1 nCoV-19/ AZD1222/ Covishield (AstraZeneca/Oxford, UK, India), Janssen Covid-19 vaccine/ Ad26CoV2.S/ Johnson & Johnson (Janssen, USA), Sinopharm/ BBIBP-CorV (China), Sinovac/ CoronaVac (China), and COVAXIN/ BBV152 (Bharat Biotech, India)]. The current report summarises the latest data on COVID-19 vaccine-related literature as of 13 December 2021, and presents the information in the form of a synoptic table. This report covers vaccine effectiveness, protection against variants, transmissibility, breakthrough infections, booster doses, COVID-19 vaccines for children, and further important information for each vaccine. The latest changes and additions to the synoptic table are highlighted in yellow.

Since the Covishield vaccine uses the same formulation and platform as Vaxzevria (AstraZeneca's COVID-19 vaccines), we combined both vaccines into one column in the synoptic table. Henceforth, seven vaccines will be referenced as WHO EUL approved (including Covishield)





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Preamble

A large number of scientific publications become available on a daily basis, reflecting the rapid development of knowledge and progress of science on COVID-19 related issues. Leading authorities should base decisions or policies on this knowledge; hence they need to master the actual state of this knowledge. Due to the large number of publications shared daily, decision makers heavily depend on accurate summaries of these publications, in the different public health domains. Therefore, the authors of this report were mandated by the Swiss School of Public Health plus (SSPH+), upon request of the Federal Office of Public Health (FOPH), to inform the FOPH on recent findings from the literature.

Background

According to the current global data on vaccinations, 56.1% of the world populations, of which only 7.3% of people in low-income countries, have received at least one dose of a marketed COVID-19 vaccine as of 15 December 2021². Currently, seven vaccines [namely, Comirnaty/BNT162b2 (Pfizer-BioNTech, USA), Spikevax/Moderna COVID-Vaccine/mRNA-1273 (Moderna, USA). Vaxzevria/ChAdOx1 19/AZD1222/Covishield (AstraZeneca/Oxford, UK, India), Janssen Covid-19 vaccine/ Ad26CoV2.S/Johnson & Johnson (Janssen, USA), Sinopharm/BBIBP-CorV (China), Sinovac/CoronaVac (China), and COVAXIN/BBV152 (Bharat Biotech, India)] were assessed and granted an authorization by WHO as of 15 December 2021³. **Articles** regarding the latest data on vaccine effectiveness, particularly against the omicron variant, vaccine induced immune response, breakthrough infections and transmission, and booster doses were prioritized during the literature search and are the latest additions to the table. Data from clinical trials and observational studies for the seven EUL-accepted vaccines and the vaccine candidate Novavax regarding these highlighted topics were summarized and can be found in the synoptic table below.

³ Status of COVID-19 vaccines within WHO EUL/ PQ evaluation process. World Health Organization. https://extranet.who.int/pqweb/sites/default/files/documents/Status COVID VAX 11Nov2021.pdf [Last updated 11 November 2021; Accessed 15 December 2021]



² https://ourworldindata.org/covid-vaccinations (accessed on 15.12.2021).



Methodology

We screened the data for the EUL-accepted vaccines and the vaccine candidate Novavax as of 13 December 2021 from PubMed, Embase, medRxiv, bioRxiv, Cochrane, and clinical trials databases such as ClinicalTrials and WHO Trial Registry. The methods used were reported previously and can be found in prior reports⁴.

Results

As phase III COVID-19 vaccine trials confirmed vaccine efficacy and safety for all seven WHO EUL authorized vaccines, and as the share of fully vaccinated people begin to increase across countries, it is important to assess vaccine effectiveness in real-world conditions, especially in relation to evolving variants of concern (VOC).

The Newest Variant of Concern: Omicron (B.1.1.529)

On 25 November 2021, a new variant of concern, later named Omicron (B.1.1.529), was firstly reported. The Omicron variant was identified to contain several mutations on the spike glycoprotein leading to a higher affinity to the ACE-2 domain of the SARS-CoV-2 virus^{5,6}. Although not confirmed, this increased affinity could impact the behavior of the virus in terms of its ability to spread, escape existing immunity, and the severity of illnesses it causes. Even though information on the transmissibility and the severity of disease are not yet clear, few studies and preliminary analyses on the effect of the new variant on the efficacy, effectiveness, and immunogenicity of vaccines and booster doses have been released. Based on results examining the effects of the Omicron variant on the immunogenicity induced by the two-dose COVID-19 vaccines,

⁶ Structural insights of SARS-CoV-2 spike protein from Delta and Omicron variants. bioRxiv. https://www.biorxiv.org/content/10.1101/2021.12.08.471777v1



⁴ COVID-19 vaccines: efficacy and safety (Literature Review 1). Swiss School of Public Health. https://www.bag.admin.ch/dam/bag/de/dokumente/mt/k-und-i/aktuelle-ausbrueche-pandemien/2019-nCoV/Literaturrecherchen/literaturrecherchen_covid-19-impfstoffe_20210209.pdf.download.pdf/20210209_Literaturrecherchen_Covid-19-Impfstoffe_EN.pdf

⁵ The Omicron variant increases the interaction of SARS-CoV-2 spike glycoprotein with ACE2. *bioRXiv*. https://www.biorxiv.org/content/10.1101/2021.12.06.471377v2



a 11.4 to 41-fold decrease in mean neutralization titers was reported in comparison to either the wild-type or the Delta variant^{7,8,9}. Regarding the effects of the new variant on the effectiveness of vaccines against symptomatic infection, one study estimated the effectiveness of the BNT162b2 vaccine to be 88.0% (95% CI, 65.9-95.8) 2-9 weeks following the second dose, 48.5% (95% CI, 24.3-65.0) 10-14 weeks following the second dose, and 34-37% from 15 weeks following the second dose¹⁰. Nevertheless, results based on preliminary laboratory studies demonstrate that three doses of the Pfizer-BioNTech COVID-19 vaccine neutralize the Omicron variant and increase the antibody titers by 25-fold compared to two doses8 and to have an effectiveness against symptomatic infection of **75.5%** (95% CI, 56.1-86.3)⁹. Although the booster dose has demonstrated to greatly increase the overall immune response and maintain a high effectiveness, a study reported that, after 3 months, a homologous BNT162b2 booster dose had a 24.5-fold decrease in neutralization compared to the Delta variant⁶. Similar results can be applied to the administration of heterologous booster doses. Overall, the vaccine-induced immunity of the current COVID-19 vaccines have shown to decrease in comparison to the wild-type and even Delta variant, while the administration of a booster dose has shown to provide a better protection against the new variant.

Latest Data on Vaccine Effectiveness, Duration of Protection & Transmissibility

A study that analysed global vaccine data across 187 countries between 20 December 2020 and 25 April 2021, analysed the correlation between the administration of vaccines and daily COVID-19 cases and deaths. The study concluded that COVID-19

Effectiveness of COVID-19 vaccines against the Omicron (b.1.1529) variant of concern. https://khub.net/documents/135939561/430986542/Effectiveness+of+COVID-19+vaccines+against+Omicron+variant+of+concern.pdf/f423c9f4-91cb-0274-c8c5-70e8fad50074



Reduced neutralization of SARS-CoV Omicron B.1.1.529 variant by post-immunization serum. *medRxiv*. https://www.medrxiv.org/content/10.1101/2021.12.10.21267534v1

⁸ SARS-CoV-2 Omicron has extensive but incomplete escape of Pfizer BNT162b2 elicited neutralization and requires ACE2 for infection. medRxiv. https://www.medrxiv.org/content/10.1101/2021.12.08.21267417v2

⁹ Pfizer and BioNTech Provide Update on Omicron Variant. [press release] *Pfizer and BioNTech.*

https://www.pfizer.com/news/press-release/press-release-detail/pfizer-and-biontech-provide-update-omicron-variant



cases and deaths would reduce by **24.43%** (**95% CI**, **18.89-29.59**) and **13.32%** (**95% CI**, **3.81-21.89**) if 10,000 persons become fully vaccinated per day, respectively¹¹. However, the study stopped collecting and analysing data in April 2021, a period of mostly Alpha variant predominance, and multiple new variants with mutations able to escape vaccine immunity have emerged since (*see above*). Given the emergence of VOCs such as the Delta (B.1.617.2) and now the Omicron (B.1.1.529) strain, mutations on the SARS-CoV-2 spike protein partially evades vaccine immunity¹², the original vaccine-induced herd immunity threshold must be updated¹³.

In line with previous data on vaccine duration of protection, the effectiveness of BNT162b2, mRNA-1273, and Ad26.COV2.S against COVID-19 infection declined with the rise of the more transmissible Delta variant in New York. By the week of 28 August vaccine effectiveness against SARS-COV-2 infection had declined to **72.3% (range: 63.7-77.5)**, **77.8% (range: 70.1-86.8)**, and **69.4% (range: 63.4-77.3)** for recipients of BNT162b2, mRNA-1273, and Ad26.COV2.S, respectively¹⁴. All three vaccines declined by similar percentage points: **20.7**, **19.5**, and **19.0** percentage points, respectively¹⁵. Likewise, a study in the UK demonstrated that Pfizer-BioNTech's BNT162b2 declined from **81% (95% CI, 68-89)** in the first two months after the administration of the second dose to **46% (95% CI, 22-63)** after six months¹⁶. Nevertheless, protection remained high for fully vaccinated (two-dose schedule) SARS-CoV-2 recovered persons; adjusted vaccine effectiveness remained consistently over the **90%** effectiveness mark "even in those infected over 15-months

¹⁶ Effectiveness and durability of protection against future SARS-CoV-2 infection conferred by COVID-19 vaccination and previous infection; findings from the UK SIREN prospective cohort study of healthcare workers March 2020 to September 2021. *medRxiv*. https://www.medrxiv.org/content/10.1101/2021.11.29.21267006v1



6/118

¹¹ The effect of the COVID-19 vaccine on daily cases and deaths based on global vaccine data. Vaccines. https://www.mdpi.com/2076-393X/9/11/1328/htm

¹² Mutations of SARS-CoV-2 spike protein: Implications on immune evasion and vaccine induced-immunity. *Seminars in Immunology*. https://www.sciencedirect.com/science/article/pii/S1044532321000646?via%3Dihub

¹³ The herd immunity threshold must be updated for multi-vaccine strategies and multiple variants. Scientific Reports. https://www.nature.com/articles/s41598-021-00083-2

¹⁴ COVID-19 Vaccine effectiveness in New York State. New England Journal of Medicine. https://www.nejm.org/doi/full/10.1056/NEJMoa2116063?query=featured_home

 ¹⁵ COVID-19 Vaccine effectiveness in New York State. New England Journal of Medicine. https://www.nejm.org/doi/full/10.1056/NEJMoa2116063?query=featured_home



ago"¹⁷, however further studies would need to confirm whether past COVID-19 infection + two-dose vaccination retains high protection over time against the new Omicron variant. In regards to AstraZeneca/Oxford's ChAdOx1 nCoV-19 vaccine, a large-scale test-negative, case-control study demonstrated that the vaccine remained effective against moderate to severe SARS-CoV-2 infection during the highly transmissible Delta variant surge in India¹⁸. After a full two-dose vaccination schedule, vaccine effectiveness was estimated to be 63.1% (95% CI, 51.5-72.1) against SARS-CoV-2 infection and 81.5% (95% CI, 9.9-99.0) against moderate-to-severe COVID-19 infection¹⁹.

Despite inactivated viral vaccines (Sinopharm's BBIBP-CorV and Sinovac's CoronaVac) being approved and administered in 72 and 47 countries worldwide, real world evidence on both vaccines continues to remain scare and vaccine effectiveness data are published at a slower rate compared to its other vaccine counterparts. A Brazilian study analysed CoronaVac's effectiveness over the Gamma (P.1) variant predominant months (study period: February to April) and determined the vaccine was effectiveness against symptomatic COVID-19 cases (80.5%; 95% CI, 75.1-84.7), severe disease (95.0%; 95% CI, 86.9-98.1), and death (94.9%; 95% CI, 76.4-98.9)²⁰. Additionally, the authors observed an indirect protective effect in unvaccinated persons among higher vaccinated communities (≥52% of the adult population over the age of 18). Given reports of rapidly waning immunity, particularly for the

²⁰ Projeto S: A Stepped-Wedge Randomized Trial to Assess CoronaVac Effectiveness in Serrana, Brazil. SSRN – Preprint. https://papers.ssrn.com/sol3/papers.cfm?abstract_id=3973422&download=yes



¹⁷ Effectiveness and durability of protection against future SARS-CoV-2 infection conferred by COVID-19 vaccination and previous infection; findings from the UK SIREN prospective cohort study of healthcare workers March 2020 to September 2021. *medRxiv*. https://www.medrxiv.org/content/10.1101/2021.11.29.21267006v1

¹⁸ Effectiveness of ChAdOx1 nCoV-19 vaccine against SARS-CoV-2 infection during the delta (B.1.617.2) variant surge in India: a test-negative, case-control study and a mechanistic study of post-vaccination immune responses. The Lancet Infectious Diseases. https://www.thelancet.com/journals/laninf/article/PIIS1473-3099(21)00680-0/fulltext

¹⁹ Effectiveness of ChAdOx1 nCoV-19 vaccine against SARS-CoV-2 infection during the delta (B.1.617.2) variant surge in India: a test-negative, case-control study and a mechanistic study of post-vaccination immune responses. *The Lancet Infectious Diseases*. https://www.thelancet.com/journals/laninf/article/PIIS1473-3099(21)00680-0/fulltext



CoronaVac vaccine^{21,22,23}, these results are quite outdated and may not reflect current vaccine effectiveness. Vaccine effectiveness data covering from April to September 2021 confirm this, as vaccine effectiveness was measured to be **72.0%** (**95% CI**, **69.9-73.9**) against ICU admission and **82.4%** (**95% CI**, **81.0-83.7**) against death²⁴, demonstrating that CoronaVac is effective at preventing severe COVID-19 related outcomes. Nevertheless, CoronaVac's estimated effectiveness was lower than BNT162b2 (**90.3**; **95% CI**, **88.8-91.6**) and AZD1222 (**95.6%**; **95% CI**, **88.3-98.4**) against ICU admission and death (**92.7%** and **95.3%**, respectively)²⁵. Likewise, Pfizer-BioNTech's mRNA vaccine demonstrates a higher quantitative efficiency compared to Sinopharm's BBIBP-CorV vaccine; immunoglobulin G (IgG) antibody titres were significantly higher for BNT162b2 (**515.5 BAU/mL**; **99.3% of participants had positive IgG titres**) when compared to BBIBP-CorV (**170.0 BAU/mL**; **85.7% had positive IgG titres**)²⁶.

A study that analysed the effect of staff vaccination rates on COVID-19 infections in nursing homes observed that nursing homes with low staff vaccination coverage (those in the lowest vaccination quartile among 12,364 selected nursing homes across the U.S) within counties with high COVID-19 prevalence (highest quartile) had **1.56** (**95% CI, 1.05-2.07**) and **0.19** (**95% CI, 0.08-0.30**) COVID-19 cases and COVID-19-related deaths of residents per 100 beds when compared to nursing homes in the same county but had higher vaccination coverage²⁷. Among staff, an additional **1.50** (**95% CI, 1.06-1.94**) COVID-19 cases per 100 beds were observed. Among nursing

²⁷ Nursing home staff vaccination and COVID-19 outcomes. The New England Journal of Medicine. https://www.nejm.org/doi/10.1056/NEJMc2115674



²¹ Comparison of an inactivated COVID-19 vaccine-induced antibody response with concurrent natural COVID-19 infection. International Journal of Infectious Diseases. https://www.ijidonline.com/article/S1201-9712(21)00768-2/fulltext

²² China's COVID vaccines have been crucial – now immunity is waning. *Nature News*. https://www.nature.com/articles/d41586-021-02796-w

²³ Limited and short-lasting virus neutralizing titers induced by inactivated SARS-CoV-2 vaccine. *Emerging Infectious Diseases*. https://wwwnc.cdc.gov/eid/article/27/12/21-1772_article

²⁴ PICK-ing Malaysia's epidemic apart: Effectiveness of a diverse COVID-19 vaccine portfolio. *Vaccines*. https://www.mdpi.com/2076-393X/9/12/1381

²⁵ PICK-ing Malaysia's epidemic apart: Effectiveness of a diverse COVID-19 vaccine portfolio. *Vaccines*. https://www.mdpi.com/2076-393X/9/12/1381

²⁶ Pfizer-BioNTech and Sinopharm: A Comparative Study on Post-Vaccination Antibody Titers. *Vaccines*. https://www.mdpi.com/2076-393X/9/11/1223/htm



homes in counties with low COVID-19 prevalence, there was only a small difference in COVID-19 outcomes between nursing homes with low and high vaccination rates²⁸.

Booster Dose

The characterisation of the durability of protective immunity continues to be one of the toughest challenges many immunologists and vaccinologists are facing. By now, multiple studies and a great deal of evidence have demonstrated that the immunogenicity of fully vaccinated individuals wanes over time, making the implementation of booster vaccination programs crucial for the restoration of immune protection and the reduction of the burden of disease. This issue has been highlighted after the increasing number of breakthrough infections with the Delta variant and the identification of the worrisome new variant of concern Omicron. Multiple countries have approved and started implementing booster vaccination programs to prioritize risk-groups and older individuals, all while expanding the program to include all individuals. Based on recent studies, homologous booster doses for the BNT162b2 vaccine have shown to be highly effective against infection and hospitalization, while increasing the neutralizing antibody count to 9.34 times higher than the second dose and increasing the IgG antibodies to 33-fold higher than the second dose in adults older than 60 years²⁹. Additionally, a homologous booster dose of BNT162b2 demonstrated to protect against COVID-19 across all age groups by decreasing the rate of confirmed infections by 17.2 (95% CI, 15.4-19.2) in individuals aged 16-29, by 9.0 (95% CI, 8.4-9.7) in individuals aged 30-39, by 9.7 (95% CI, 9.2-10.3) in individuals aged 40-49, by 12.2 (95% CI, 11.4-13.0) in individuals 50-59, and by 12.3 (95% CI, **11.8-12.8)** in individuals aged 60 and over³⁰. Heterologous booster doses have also demonstrated to be safe and effective in preventing infections, as demonstrated in a

³⁰ Protection against Covid-19 by BNT162b2 Booster across Age Groups. NEJM. https://www.nejm.org/doi/full/10.1056/NEJMoa2115926



²⁸ Nursing home staff vaccination and COVID-19 outcomes. The New England Journal of Medicine. https://www.nejm.org/doi/10.1056/NEJMc2115674

²⁹ Early Immunogenicity and safety of third dose of BNT162b2 mRNA Covid-19 vaccine among adults older than 60years; real world experience. *The Journal of Infectious Diseases*. https://academic.oup.com/jid/advance-article/doi/10.1093/infdis/jiab584/6446235



study examining seven different COVID-19 vaccines used as booster doses in people previously vaccinated with AstraZeneca/Oxford or Pfizer-BioNTech vaccines³¹. Based on the study's results, the administration of the Moderna vaccine as a heterologous booster produced the biggest antibody response, followed by the Pfizer-BioNTech vaccine. Although booster doses are becoming more and more crucial in the ongoing battle against COVID-19, the consideration of the substantial geopolitical and ethical implications of booster vaccination programs remain as many individuals in low-income countries have yet to receive the first jabs of COVID-19 vaccines.

Safety and adverse events

In order to build population trust in the newly introduced COVID-19 vaccines, it is important to collect data on and report on Adverse Events Following Immunization (AEFI). Self-reports of adverse events for all 7 WHO EUL approved vaccines confirm that while systemic vaccine adverse events (VAE) can occur (common symptoms include pain at injection site, headache, fatigue, and muscle pain), they are minor, short lived, and self-resolving^{32,33,34,35}. Typically, local and systemic VAE occur at higher rates in women^{36,37}, younger individuals^{38,39,40}, and persons with allergy

⁴⁰ Evaluation of adverse effects with COVID-19 vaccination in Pakistan. Pakistan Journal of Medical Sciences. https://pims.org.pk/index.php/pjms/article/view/4522



³¹ Safety and immunogenicity of seven COVID-19 vaccines as third dose (booster) following two doses of ChAdOx1 nCov-19 or BNT162b2 in the UK (COV-BOOST): a blinded, multicentre, randomized, controlled, phase 2 trial. The Lancet. https://www.thelancet.com/journals/lancet/article/PIIS0140-6736(21)02717-3/fulltext

³² Adverse events following immunization against SARS-CoV-2 (COVID-19) in the state of Minas Gerais. *Revista de Saúde Pública*. https://www.revistas.usp.br/rsp/article/view/191771/176666

³³ Self-reported adverse events of COVID-19 vaccines in Polish Healthcare workers and medical students. Cross-sectional study and pooled analysis of CoVaST project results in Central Europe. *Journal of Clinical Medicine*. https://www.mdpi.com/2077-0383/10/22/5338

³⁴ Adverse events report of inactivated COVID-19 vaccine from 4040 healthcare workers. Postgraduate Medicine. https://www.tandfonline.com/doi/full/10.1080/00325481.2021.1999708

³⁵ Self-reported adverse events of COVID-19 vaccines in Polish Healthcare workers and medical students. Cross-sectional study and pooled analysis of CoVaST project results in Central Europe. *Journal of Clinical Medicine*. https://www.mdpi.com/2077-0383/10/22/5338

³⁶ Adverse events report of inactivated COVID-19 vaccine from 4040 healthcare workers. *Postgraduate Medicine*. https://www.tandfonline.com/doi/full/10.1080/00325481.2021.1999708

³⁷ Reactogenicity within 2 weeks after mRNA COVID-19 vaccines: Findings from the CDC v-safe surveillance system. *Vaccine*. https://pubmed.ncbi.nlm.nih.gov/34763946/

³⁸ Adverse events report of inactivated COVID-19 vaccine from 4040 healthcare workers. Postgraduate Medicine. https://www.tandfonline.com/doi/full/10.1080/00325481.2021.1999708

³⁹ Reactogenicity within 2 weeks after mRNA COVID-19 vaccines: Findings from the CDC v-safe surveillance system. *Vaccine*. https://pubmed.ncbi.nlm.nih.gov/34763946/



histories⁴¹. mRNA vaccines were observed to lead to higher rates of short-term adverse events (i.e. pain at injection site), while viral vector based vaccines lead to higher incidences of mild systemic side effects (i.e. fatigue and fever)⁴²; a study that compared the adverse events of four COVID-19 vaccines (Pfizer-BioNTech, AstraZeneca, Sinopharm, and Sputnik) in Bahrain reported that the Sinopharm's BBIBP-CorV vaccine recipients reported the mildest side effects⁴³. Nevertheless, severe adverse events (SAE) post-COVID-19 vaccination can sometimes occur and it is important to properly report on SAE for their timely recognition, diagnosis, and management⁴⁴. Comparisons across the BNT162b2, ChAdOx1 nCoV-19 and Ad26CoV2.S vaccines demonstrated that recipients of the viral-vector-based vaccines had higher SAE frequencies related to coagulation disorders and arterial, cardiac, and nervous system events than Pfizer-BioNTech's mRNA vaccine⁴⁵. The study is corroborated by an Australian study that demonstrated the possible association between immune thrombocytopenia (ITP) and the AstraZeneca vaccine⁴⁶. When comparing across the two viral vector vaccines, higher rates of SAE were reported in younger individuals who had received ChAdOx1 nCov-19 and among older individuals who had received the Janssen's Ad26CoV2.S⁴⁷. Lastly, we would like to highlight that a severe headache and/ or vaccine-induced thrombocytopenia (VIT) 48, in addition to blurred vision, shortness of breath, chest pain, leg swelling and abdominal pain⁴⁹,

⁴¹ Adverse events report of inactivated COVID-19 vaccine from 4040 healthcare workers. *Postgraduate Medicine*. https://www.tandfonline.com/doi/full/10.1080/00325481.2021.1999708

⁴⁹ CDC recommends use of Johnson & Johnson's Janssen COVID-19 vaccine resume. *Centers for Disease Control and Prevention*. https://www.cdc.gov/coronavirus/2019-ncov/vaccines/safety/JJUpdate.html#symptoms-list



⁴² Self-reported adverse events of COVID-19 vaccines in Polish Healthcare workers and medical students. Cross-sectional study and pooled analysis of CoVaST project results in Central Europe. *Journal of Clinical Medicine*. https://www.mdpi.com/2077-0383/10/22/5338

⁴³ Unfolding the mild to moderate short-term side effects of four COVID-19 vaccines used in Bahrain: A cross-sectional study. *Vaccines*. https://www.mdpi.com/2076-393X/9/11/1369/htm

⁴⁴ The Diagnostic Process. *Improving Diagnosis in Health Care*. https://www.ncbi.nlm.nih.gov/books/NBK338593/

⁴⁵ Cardiovascular, neurological, and pulmonary events following vaccination with the BNT162b2, ChAdOx1 nCoV-19, and Ad26.COV2.S vaccines: An analysis of European data. *Journal of Autoimmunity*. https://www.sciencedirect.com/science/article/pii/S0896841121001505?via%3Dihub

⁴⁶ Immune thrombocytopenia following immunization with Vaxzevria ChadOx1-S (AstraZeneca) vaccine, Victoria, Australia. *Vaccine*. https://www.sciencedirect.com/science/article/pii/S0264410X21013505?via%3Dihub

⁴⁷ Cardiovascular, neurological, and pulmonary events following vaccination with the BNT162b2, ChAdOx1 nCoV-19, and Ad26.COV2.S vaccines: An analysis of European data. *Journal of Autoimmunity*.
https://www.sciencedirect.com/science/article/pii/S0896841121001505?via%3Dihub

⁴⁸ Vaccine-induced thrombocytopenia with severe headache. *The New England Journal of Medicine*. https://www.nejm.org/doi/10.1056/NEJMc2112974



following either ChAdOx1 nCoV-19 or Ad26CoV2.S may be early signs of vaccineinduced immune thrombotic thrombocytopenia (VITT). Patients presenting with these severe headaches, blurred vision, shortness of breath, chest pain, leg swelling, and persistent abdominal pain should "undergo immediate testing for thrombocytopenia and D-dimer levels and, if available, anti-PF4-heparin IgG antibodies"50. Despite incidences of SAEs, we would like to emphasize that the benefits of COVID-19 vaccination far outweigh the risks of developing SAEs following vaccination and adverse events related to SARS-CoV-2 infection⁵¹.

Further (biweekly) updated data on the seven WHO EUL vaccines and the vaccine candidate Novavax are synthesized in the synoptic table and new data has been highlighted in yellow

⁵¹ Adverse effects after BNT162b2 vaccine and SARS-CoV-2 infection, according to age and sex. The New England Journal of Medicine. https://www.nejm.org/doi/full/10.1056/NEJMc2115045



⁵⁰ Vaccine-induced thrombocytopenia with severe headache. The New England Journal of Medicine. https://www.nejm.org/doi/10.1056/NEJMc2112974



Synoptic Table

Synoptic table about SARS-CoV-2 vaccines accepted in the WHO's Emergency Use Listing and Novavax Vaccine (as of 15 December 2021)

	BNT162b2/ COMIRNATY (Pfizer-BioNTech, USA)	Spikevax/ Moderna COVID- 19 Vaccine/ mRNA-1273 (Moderna, USA)	Vaxzevria/ ChAdOx1 nCoV- 19/ AZD1222/ Covishield (AstraZeneca/Oxf ord, UK, India)	Janssen COVID- 19 vaccine/Johnson & Johnson (Janssen, USA)	BBIBP-CorV, (Sinopharm, China)	CoronaVac (Sinovac, China)	COVAXIN / BBV152 (Bharat Biotech, India)	Novavax/ NVX- CoV2373	
	GENERAL VACCINE INFORMATION								
Platform	mRNA-based vaccine	mRNA-based vaccine	Non-replicating vector-based vaccine	Non-replicating vector-based vaccine	Inactivated virus (Vero cell)	Inactivated virus (Vero cell)	Whole-virion inactivated Vero cell	Recombinant protein (nanoparticle) vaccine with Matrix-M adjuvant	
Dose and frequency	2 doses, 21 days apart	2 doses, 28 days apart	2 doses, 4-12 weeks apart	1 dose, once [Phase III trials currently testing 2- dose regime, 56 days apart]i	2 doses, 21 days apart	2 doses, 14 days apart	2 doses, 28 days apart	2 doses, 21 days apart	
Target population	12 years old and over	12 years old and over	18 years old and over	18 years old and over	18 years old and over	18 years old and over	18 years old and over	18 years old and over	

ⁱ Johnson & Johnson Announces Real-World Evidence and Phase 3 Data Confirming Strong and Long-Lasting Protection of Single-Shot COVID-19 Vaccine in the U.S. *Johnson & Johnson*. https://www.jnj.com/johnson-johnson-announces-real-world-evidence-and-phase-3-data-confirming-strong-and-long-lasting-protection-of-single-shot-covid-19-vaccine-in-the-u-s





Storage conditions	2°C to 8 °C (for 1 month)	2°C to 8 °C (for 1 month)	2°C until 8 °C	2°C to 8 °C (for 3 months)	2°C until 8 °C	2°C until 8 °C	2°C until 8 °C	2°C to 8 °C
Approving authorities	FDA (11.12.20) ⁱⁱ ; EMA (21.12.20); WHO EUL (31.12.20); and list of 112 countries (including Switzerland – approved on 20.12.20)	FDA (18.12.20); EMA (06.01.21); WHO EUL (30.04.21); and list of 79 countries (including Switzerland – approved 12.01.21)	FDA (awaiting on approval); EMA (29.01.21); WHO EUL (15.02.21); and list of 127 countries (Switzerland awaiting on approval)	FDA (27.02.21); EMA (11.03.21), WHO EUL (12.03.21), and list of 85 countries (including Switzerland – approved 22.03.21)	WHO EUL (07.05.21); and list of 72 countries (including Argentina, Bahrain, Brazil, China, Indonesia, United Arab Emirates, Zimbabwe)	WHO EUL (01.06.21), and list of 47 countries (including Albania, Chile, Egypt, Hong Kong, Malaysia, Tunisia, Turkey, Ukraine)	WHO EUL (03.11.21) and list of 12 countries (including Guyana, Inidia, Iran, Mauritius, Mexico, Nepal, Paraguay, Philippines & Zimbabwe)	Waiting on approval (Not-yet- approved by countries or WHO for emergency use)
Booster shot approving authorities	EMA approved booster for those aged 18 and above, 6 months after the 2 nd dose ¹ FDA approved booster for those ages 16 and above, 6 months after the 2 nd dose ⁱⁱⁱ Swissmedic approves booster	EMA authorised booster dose for immunocompromi sed individuals FDA approved third booster dose for individuals >65 and high-risk individuals, 6 months after the 2 nd dose ^{vi} Swissmedic approves booster	-	-	-	-	-	-

ii Pfizer-BioNTech's Comirnaty Vaccine received full FDA approval on 23 August 2021 for people age 16 and above, moving it beyond emergency use status. https://www.fda.gov/news-events/press-announcements/fda-approves-first-covid-19-vaccine

FDA authorizes booster dose of Pfizer-BioNTech COVID-19 vaccine for certain populations. FDA News Release. https://www.fda.gov/news-events/press-announcements/fda-authorizes-booster-dose-pfizer-biontech-covid-19-vaccine-certain-populations

^v Comirnaty and Spikevax: EMA recommendations on extra doses and boosters. *European Medicines Agency*. https://www.ema.europa.eu/en/news/comirnaty-spikevax-ema-recommendations-extra-doses-boosters

^{*} F.D.A. Panel recommends booster for many Moderna vaccine recipients. The New York Times. https://www.nytimes.com/2021/10/14/us/politics/fda-moderna-vaccine-boosters.html



	dose for everyone aged 16 and overiv	dose for adults aged 18 and over ^{vii}						
			EFFECTIVENESS	AGAINST ANY SAR	S-COV-2 INFECTIO	N		
	BNT162b2/ COMIRNATY	Spikevax/ Moderna COVID- 19 Vaccine/ mRNA-1273	Vaxzevria/ ChAdOx1 nCoV- 19/ AZD1222/ Covishield	Janssen COVID- 19 vaccine/Johnson & Johnson	BBIBP-CorV,	CoronaVac	COVAXIN / BBV152	Novavax/ NVX- CoV2373 (Awaiting approval from WHO EUL)
Effectiveness single dose	Against any SARS-CoV-2 infection: 70% ² . 77.6% (95% CI, 70.9-82.7) ³ 36.8% (95% CI, 33.2-40.2) [3 weeks after first dose] ⁴ 57% (95% CI, 52-61; Spain) [Apr-Aug] ⁵ 72% (pooled meta-analysis) ⁶ 64% (95% CI, 59%-68%; United	Against SARS-COV-2 infection: 60% (95% CI, 57-64; >2 weeks after dose) ^{10, ix} 88.9% (95% CI, 78.7-94.2) ³ 66% (95% CI, 56-73; Spain) [Apr-Aug] ⁵ 69% (pooled meta-analysis) ⁶ 64% (95% CI, 59%-68%; United States) [May to July 2021] ^{7x} 39.6% (95% CI, 36.3-42.8;	Against SARS- CoV-2 infection: 31.4% (95% CI, 25.7-36.7; Norway) [Jan- Sep] ⁸ Symptomatic disease: 67% ¹¹ 49% (95% CI, 32.0-62.0; India) [Apr-Jun] ¹² 41% (95% CI, 34- 48; Spain) [Apr- Aug] ⁵ 51% (pooled meta-analysis) ⁶	Against SARS-CoV-2 infection: 50.6% (95% CI, 14.0-74.0) [<2 weeks after dose]; 76.7% (95% CI, 30.3-95.3) [>2 weeks after dose] ¹³ ; 79% (95% CI, 77-80) (when corrected for under-recording, VE was estimated to be 69% (95% CI, 67-71) ¹⁴ .	Partial protection ²³ .xiii	15.5% for preventing COVID-19; 37.4% for preventing hospitalization; 44.7% for preventing admission to the ICU; and 45.7% for preventing of COVID-19 related death ²⁴ . 18.6% (95% CI, 17.6-19.6) against SARS-CoV-2 infection, 28.1% (95% CI, 26.3-	Against symptomatic disease: 45% (95% CI,6.0- 68.0; India) [Apr- Jun] ¹² 40% (95% CI, -21- 71; India) less than 7 days after first dose [April- May] ²⁶ 1% (95% CI, -30- 25); India) at least 7 days after first dose [April-May] ²⁶	Ongoing studies in South Africa ²⁷ and the United Kingdom ²⁸

^{iv} COVID-19 vaccine from Pfizer/BioNTech: Swissmedic approves the extension of the booster dose to everyone aged 16 years and over. *Swissmedic*. https://www.swissmedic.ch/swissmedic/en/home/news/coronavirus-covid-19/covid-19-impfstoff-pfizer-biontech-boosterdosis.html

xiii Study did not report numerical data on vaccine effectiveness. Further studies are required to validate results.



vii Swissmedic approves booster dose of the Moderna COVID-19 vaccine for adults aged 18 and over. Swissmedic. https://www.swissmedic.ch/swissmedic.ch

ix mRNA-aggregated data (results do not disaggregate between BNT162b2 and mRNA-1273).

^{*} Results do not disaggregate between mRNA vaccines, BNT162b2 and mRNA-1273.



July 2 19.6% 17.3-2 Norwa Sep] ⁸ Again symp disea 66% (71; Sp Aug] ⁵ Indivin	2021] ^{7√iii} % (95% CI, .21.9; yay) [Jan- 3 7 nst otomatic ase: (95% CI, 60- Spain) [Apr- 5 iduals ≥ 70:	Against Sep] ⁸ Against Symptomatic disease: 71% (95% CI, 61-79; Spain) [Apr-Aug] ⁵ Andividuals ≥ 70: Symptomatic disease: 64% (95% CI, 46-78; >2 weeks after dose) ¹⁰ .xi	46% (95% CI, 37- 54; Spain) [Apr- Aug] ⁵ Individuals ≥70: Symptomatic disease: 58% ⁹ .	71% (95% CI, 56-81) [11 March – 15 August] ¹⁵ . 61% (95% CI, 29-84) [January-June] ¹⁶ 50.9% (95% CI, 35.1-63.0) [June-September; Brazil] ¹⁷ 50.0% (95% CI, 42.0-57.0; Spain) [Apr-Aug] ⁵ 73.6% (95% CI, 65.9-79.9; US) [Feb-Jul] ¹⁸ 82.3% (95% CI, 75.1-87.4%; USA) [16 Dec 2020 to 30 Sep 2021] ^{19xii} Symptomatic disease: 54% (95% CI, 45-62; Spain) [Apr-Aug] ⁵ 81% (95% CI, 79-84) for preventing hospitalization when corrected for under-recording,		29.9) against hospitalization, 28.5% (95% CI, 25.4-31.4) against ICU admission, and 29.4% (95% CI, 26.7.3-31.9) against death [January-April] ²⁵	-1% (95% CI, -51-33; India) at least 21 days after first dose [April-May] ²⁶		
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viii Results do not disaggregate between mRNA vaccines, BNT162b2 and mRNA-1273.

xi mRNA-aggregated data (results do not disaggregate between BNT162b2 and mRNA-1273).

xii Study does not differentiate between Pfizer, Moderna, and Janssen.



				VE was estimated to be 73% (95% CI, 69-76) ¹⁴ . 75% (95% CI, 65-82) against severe critical COVID-19 ²⁰ 66.1% against moderate to severe-critical COVID-19 cases after 28 days [ENSEMBLE study; Sep 2020-Nov 2021) ²¹ 85.4% against severe COVID-19 cases after 28 days [ENSEMBLE study; Sep 2020-Nov 2021) ²¹ 85.4% against severe COVID-19 cases after 28 days [ENSEMBLE study; Sep 2020-Nov 2021) ²¹ <i>Individuals</i> ≥50: 68% (95% CI, 50-79) ²² .				
Effectiveness of two doses	SARS-Cov-2 infection: 85% ² .	SARS-Cov-2 infection: 100% ²⁹ .	Asymptomatic efficacy: 61.9% ³⁹	Not Applicable (one dose schedule)	Partial protection ²³ .xxiii	65.9% for preventing COVID-19; 87.5%	Against symptomatic disease:	Ongoing studies in South Africa ²⁷

xxiii Study did not report numerical data on vaccine effectiveness. Further studies are required to validate results. Death reports on fully vaccinated doctors (10 cases during June 2021 in Indonesia). It may be related to new variants [media report]. Indonesian Covid deaths add to questions over Sinovac vaccine. *The Guardian* [press release]. https://www.theguardian.com/world/2021/jun/28/indonesian-covid-deaths-add-to-questions-over-sinovac-vaccine





94.6% ²⁹ . 94.5% ³⁰ . 76% (95% CI, 69-81) [Jan-Jul] ³¹ . 88.8% (95% CI, 84.6-91.8) [Dec 2020-May] ³ 74% (95% CI, 72-76) [Jan-Jun] ¹⁶ 77.5% (95% CI, 76.4-78.6) [first month after second dose] ⁴ 47% (95% CI, 43-51) [5 months after second dose] ³² 56% (95% CI, 53-59) [4 months after second dose] ³³ 69% (95% CI, 66-72; Spain) [Apr-Aug] ⁵	86% (95% CI, 81-90.6) [January-July] ³¹ . 96.3% (95% CI, 91.3-98.4) [December-May] ³ 85% (95% CI, 80-90) [January-June] ¹⁶ 71% (95% CI, 68-74) [4 months after second dose] ³³ 63% (95% CI, 44-76) [June-August] ³⁸ 82% (95% CI, 78-86; Spain) [Apre-	SARS-CoV-2 infection: 53% (95% CI, 12-84) [January-June] ¹⁶ 27% (95% CI, 17-37) [4 months after second dose] ³³ 88% (95% CI, 79.0-94.0; India) [Apr-Jun] ¹² 54.0% (95% CI, 48-60; Spain) [Apr-Aug] ⁵ 43.4% (95% CI, 4.4-66.5; Norway) [Jan-Sep] ⁸		for preventing hospitalization; 90.3% for preventing ICU admission; and 86.3% for preventing COVID-19 related death ²⁴ . 52.7% (95% CI, 52.1-53.4) against SARS-CoV-2 infection, 72.8% (95% CI, 71.8-73.7) against hospitalization, 73.8% (95% CI, 72.2-75.2) against ICU admission, and 73.7% (95% CI, 72.3-75.0) against death I January Aprill ²⁵	71% (95% CI, 41-85; India) [Apr-Jun] ¹² Effectiveness of full vaccination: 69% (95% CI; 54-79; India) [May -July 2021] ⁴⁰ 50% (95% CI, 33-62; India) 14 days after second dose [April-May] ²⁶ 47% (95% CI, 29-61; India) 14 days after second dose – excluding participants with previous SARS-CoV-2 infections [April-May] ²⁶	and the United Kingdom ²⁸ 89.7% protection against SARS- CoV-2 infection (95% CI, 80.2- 94.6; United Kingdom) ⁴²
_	-	•		SARS-CoV-2	after second dose	
					[April-May] ²⁶	
, <u>-</u>		[Apr-Jun] ¹²		•	470/ (050/ CL 00	
		54.0% (05% CI				
-	uosoj			· · · · · · · · · · · · · · · · · · ·		
	63% (95% CI, 44-	• • •		•		
				•		
-	August] ³⁸	•				
	000/ (050/ 01 70			•		
72; Spain) [Apr- Aug] ⁵	82% (95% CI, 78- 86; Spain) [Apr-	[Jan-Sep]°		against death [January-April] ²⁵	[Aprii-May] ²⁰	
88% (pooled	Aug] ⁵	80% (95% CI; 73-		[January April]	46% (95% CI, 22-	
meta-analysis)6	- 01	86; India) [May -			62; India) 28 days	
84% (95% CI, 40-	80% (pooled	July 2021] ⁴⁰		<u>In pregnant</u>	after second dose	
96; Italy) [27 Dec	meta-analysis)6	60% (95% CI, 50-		<u>women</u> :	[April-May] ²⁶	
2020 – 24 Mar 2021] 14-21 days	95% (95% CI,	67; Sweden) [27 Dec 2020-2 Nov		41% (95% CI, 27.1-52.2%;	57% (95% CI, 21-	
from the first dose	93%-96%; United	2021] ³⁵		Brazil) against	76; India) 42 days	
and 95% (95% CI,	States) [May to			symptomatic	after second dose	
62-99; Italy) [27	July 2021] ^{7xviii}	<u>Symptomatic</u>		COVID-19, 85%	[April-May] ²⁶	
Dec 2020 – 24		<u>disease</u> : 90% ¹¹ .		(95% CI, 59.5-		

xviii Results do not disaggregate between mRNA vaccines, BNT162b2 and mRNA-1273.





Mar 2021 at least 7 days from the second dose 4 Span							TODETC HEAL
	7 99 99 99 99 99 99 99 99 99 99 99 99 99	days from the econd dose ³⁴ 5% (95% CI, 3%-96%; United states) [May to uly 2021] ^{7xiv} 9.7% (95% CI, 8.6-70.8; lorway) [Jansep] ⁸ 2.3% (95% CI, 5.1-87.4%; USA) 16 Dec 2020 to 0 Sep 2021] ^{19xv} 5% (95% CI, 73-7; Sweden) [27 Dec 2020-2 Nov 021] ³⁵ Symptomatic lisease: 2% (95% CI, 69-5; Spain) [Apraug] ⁵ Asymptomatic SARS-CoV-2	76.7-79.6; Norway) [Jan-Sep] ⁸ 82.3% (95% CI, 75.1-87.4%; USA) [16 Dec 2020 to 30 Sep 2021] ^{19xix} 85% (95% CI, 82-87; Sweden) [27 Dec 2020-2 Nov 2021] ³⁵ Symptomatic disease: 91% (95% CI, 89-93; >2 weeks after dose) ¹⁰ .xx 85% (95% CI, 80-89; Spain) [Apr-Aug] ⁵ Asymptomatic SARS-CoV-2 infection:	63; Spain) [Apr-		against severe COVID-19, and 75% (95% CI 27.9-91.2;	

xiv Results do not disaggregate between mRNA vaccines, BNT162b2 and mRNA-1273.

xv Study does not differentiate between Pfizer, Moderna, and Janssen.

 $^{^{\}mbox{\scriptsize xix}}$ Study does not differentiate between Pfizer, Moderna, and Janssen.

 $^{^{\}mbox{\tiny XX}}$ Results do not disaggregate between BNT162b2 and mRNA-1273.

xxi Results do not disaggregate between BNT162b2 and mRNA-1273



90.6% ³⁶ 73.1 (95 70.3-75.	% CI, 78) [January-			
93) [Jan July] ³¹ .	% CI, 73- uary- "% CI, 85- "91.6% (95% CI, 81-97) [January- July] ³¹ .			
15 Augu 89% (95	95) [11 March – 15 August) ¹⁵ .			
≥50 yea January June ²² . ³ 90% (95	.22 ≥50 years [1 √ii January-22 % CI, 89- June ²² . ^{xxii}			
92) [Dec Aug 202 <i>Individu</i> a	1] ³² a <u>ls ≥65:</u>			
65) agai CoV-2 ir and 86% 82-88) a	(95% CI,			
<i>Individua</i> VE of 68				

xvi Results do not disaggregate between BNT162b2 and mRNA-1273

xvii mRNA-aggregated data (results do not disaggregate between BNT162b2 and mRNA-1273).

xxii mRNA-aggregated data (results do not disaggregate between BNT162b2 and mRNA-1273).



United Kingdom)
against non-

								TOBLIC HEAL
	infections, 73.2% (95% CI, 65.3-79.3) for hospitalization, 85.1% (95% CI, 80.0-89.0) for mortality [Germany, 09 Jan – 11 Apr 2021] ³⁷							
	_		EFFECTI	/ENESS AGAINST V	ARIANTS**iv			
	BNT162b2/ COMIRNATY	Spikevax/ Moderna COVID- 19 Vaccine/ mRNA-1273	Vaxzevria/ ChAdOx1 nCoV- 19/ AZD1222/ Covishield	Janssen COVID- 19 vaccine/Johnson & Johnson	BBIBP-CorV,	CoronaVac	COVAXIN / BBV152	Novavax/ NVX- CoV2373 (Awaiting approval from WHO EUL)
Alpha (B.1.1.7)	Single dose: 48.7% (95% CI, 45.5 to 51.7) ⁴³ 66% (95% CI,64-68) ⁴⁴ . 54.5% (95 CI, 50.4-58.3) ⁴⁵ Two doses: 93.7% (95% CI, 91.6 to 95.3) ⁴³ 92% (95% CI, 90-93) ⁴⁶ . 89% (95% CI, 86-91) ⁴⁴ .	Single dose: 88.1% (95% CI, 83.7 to 91.5) ⁴⁸ 83% (95% CI, 80- 86) ⁴⁴ . Two doses: 100% (95% CI, 91.8 to 100) ⁴⁸ 92% (95% CI, 86- 96) ⁴⁴ . 98.4% (95% CI, 96.9-99.1) ⁴⁹	Single dose: 48.7% (95% CI 45.5 to 51.7) ⁴³ 64% (95% CI, 60-68) ⁴⁴ . Two doses: 74.5% (95% CI, 66-78) ⁴⁶ . 79% (95% CI, 56-90) ⁴⁷ .	-	No published data	Two doses: Equally effective (~76%) in neutralizing D614G, B.1.1.7 and B.1.429 as the wild-type strain.	No available data	Ongoing studies in South Africa ²⁷ and the United Kingdom ²⁸ Post hoc analysis showed efficacy of 86.3% (95% CI, 71.3-93.5; United Kingdom) against B.1.1.7 variants and 96.4% (95% CI, 73.8-99.5; United Kingdom)

78% (95% CI, 68-84)⁴⁷



xxiv Effectiveness data against the latest variant of interest (Mu) will be included in upcoming reports based on data availability.



	84.4% (95 CI, 81.8-86.5) ⁴⁵							B.1.1.7 variants. ⁴²
Beta (1.351)	Against SARS-CoV-2 infection: Single dose: 60% (95% CI, 52-67) ⁴⁴ . Two doses: 84% (95% CI, 69-92) ⁴⁴ . 72% (95% CI, -5-97; Israel) [Dec 2020-Mar 2021] ⁵⁰ Against symptomatic infection: 100% (95% CI, 19-100; Israel) [Dec 2020-Mar 2021] ⁵⁰	Single dose: 61.3% (95% CI, 56.5 to 65.5) ⁴⁸ 77% (95% CI, 69- 92) ⁴⁴ . Two doses: 96.4% (95% CI, 91.9 to 98.7) ⁴⁸	<u>Single dose:</u> 48% (95% CI, 28-63) ⁴⁴ .	-	No published data	Neutralization capacity was decreased by factor 5.27 ⁵¹ .	No available data	No available data
Gamma (P.1)	Neutralization activity reduced by 3.3-fold ⁵² .	No available data	No available data	No available data	No published data	Demonstrated 42% vaccine effectiveness in a setting with high P.1 transmission, in individuals aged 70 and above ⁵³ . 50.2% against P.1 (>14 days after 2 nd dose) ⁵⁴ .	No available data	No available data





















						Neutralization was decreased by factor 3.92 ⁵¹ . Against symptomatic COVID-19: 80.5% (95% CI, 75.1-84.7) ⁵⁵		
Delta (1.617.2)	Single dose: 30.7% (95% CI, 25.2 to 35.7) ⁴³ ; 57% (95% CI, 50-63) ⁴⁷ 22.5% (95 CI, 17.0-27.4) ⁴⁵ 22% (95% CI, 10-32; France) [May-August 2021] ⁵⁶ Two doses: 88.0% (95% CI, 85.3 to 90.1) ⁴³ ; 80% (95% CI, 77-83) ⁴⁷ 79% (95% CI, 75-82) ⁴⁶ . 80% (95% CI, 77-83) ⁴⁷ 40.5% (95% CI, 8.7-61.2) ⁵⁷ .	Single dose: 72% effective against symptomatic SARS-Cov-2 infection ⁶¹ . ≥ 14 days after second dose: 76% (95% CI, 58- 87) ³¹ . 94.5% (95% CI, 94.1-95) [2-9 weeks after second dose] ⁵⁸ . 50.6% (95% CI, 45.0-55.7) [among nursing home residents] ⁵⁹ . 86.7% (95% CI, 84.3-88.7) ⁴⁹	Single dose: 30.7% (95% CI 25.2 to 35.7) ⁴³ 73% (95% CI, 64- 80; India) [May – July 2021] ⁴⁰ Two doses: 67.0% (95% CI, 61.3 to 71.8) ⁴³ 67% (95% CI, 62- 71) ⁴⁷ . 60% (95% CI, 53- 66) ⁴⁶ . 66.7% (95% CI, 45-49.6) [2-9 weeks after second dose] ⁵⁸ . 47.3% (95% CI, 66.3-67.0) [≥20 weeks after second dose] ⁵⁸ .	78% (95% CI, 73-82) against SARS-CoV-2 infection ¹⁴ . 3% (95% CI, -7-12) [August] ⁶⁰ 76.5% (95% CI, 40.9-90.6; USA) [01 Jul 2021 to 30 Sep 2021] ^{19xxvii} Individuals ≥50: 83% (95% CI, 81-85) ¹⁴	No available data	Single dose: 13.8% (95% CI, -60.2-54.8) ⁶⁴ . Two doses: 59% (95% CI, 16-81.6) against SARS-CoV-2 infection and 70.2% (95% CI, 29.6-89.3) against moderate COVID-19 infection ⁶⁴ .	Single dose: 44% (95% CI, 0-71; India) [May – July 2021] ⁴⁰ Two doses: 64% (95% CI, 40-79; India) [May – July 2021] ⁴⁰	No available data

xxvii Study does not differentiate between Pfizer, Moderna, and Janssen.





76.5% (95% CI, 40.9-90.6; USA) [01 Jul 2021 to 30 Sep 2021] ^{19xv} 67% (95% CI, 63- 71; France) [May- August 2021] ⁵⁶		40.9-90.6; USA) [01 Jul 2021 to 30 Sep 2021] ^{19xxv} 67% (95% CI, 63- 71; France) [May-	56.6% (95% CI, 42.0-67.5) against infection ⁶² 84.2% (95% CI, 56.4-94.3) against symptomatic infection ⁶² 64% (95% CI, 62-66) [August; elderly Veteran population] ⁶⁰ 76.5% (95% CI, 40.9-90.6; USA) [01 Jul 2021 to 30 Sep 2021] ^{19xxvi} 10-14 weeks after second dose: 90.3% (95% CI, 67.2-97.1) ⁵⁸ .	81% (95% CI, 71-88; India) [May – July 2021] ⁴⁰ Odds ratio of 5.45 (95% CI, 1.39-21.4) to become infected with B.1.167.2 compared to non-B.1.167.2 ⁶³ .					
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xxv Study does not differentiate between Pfizer, Moderna, and Janssen.

xxvi Study does not differentiate between Pfizer, Moderna, and Janssen.



	Against severe <u>COVID-19:</u> 91.4% (95% CI, 82.5-95.7) ⁵⁷ .	T						
Mu (B.1.621)	Mu variant is 9.1 times more resistant than the wild type strain when vaccinated with BNT162b2 ⁶⁵	Two doses: 90.4% (95% CI, 73.9-96.5) ⁴⁹ (demonstrated similar protective measures as against the Alpha variant)	No available data	No available data	No available data	No available data	No available data	No available data
Omicron (B.1.1.529)	88.0% (95% CI, 65.9-95.8) after 2-9 weeks following second dose, 48.5% (95% CI, 24.3-65.0) after 10-14 weeks following second dose, 34-37% from 15 weeks after second dose ⁶⁶ If assuming a 25-fold decrease in pseudovirus neutralization 66% (95% CI, 42-86) ⁶⁷	No available data	No protective effect of vaccination against symptomatic disease with Omicron from 15 weeks after the second dose66					
			EFFECTIVE	NESS AGAINST HOS	SPITALIZATION			
	BNT162b2/ COMIRNATY	Spikevax/ Moderna COVID-	Vaxzevria/ ChAdOx1 nCoV-	Janssen COVID- 19	BBIBP-CorV,	CoronaVac	COVAXIN / BBV152	Novavax/ NVX- CoV2373



		19 Vaccine/ mRNA-1273	19/ AZD1222/ Covishield	vaccine/Johnson & Johnson				(Awaiting approval from WHO EUL)
Any SARS-CoV- 2 infection	Single dose: 85% (pooled meta-analysis) ⁶ Hospitalization risk reduced by 35-45% ⁹ . Risk of death reduced by 54% ⁹ . Individuals ≥50: ≥14 days after first dose: 54% (95% CI, 47-61) [1 Jan-22 Jun ²² . xxviii Two doses: 91% (pooled meta-analysis) ⁶ 91% (95% CI, 93%-96%; United States) [May to July 2021] ^{7xxix} 89% (95% CI, 84-93; Sweden) [27 Dec 2020-2 Nov 2021] ³⁵	Single dose: 73% (pooled meta-analysis) ⁶ Individuals ≥50: ≥14 days after first dose: 54% (95% CI, 47-61) [1 Jan-22 Jun ^{22,xxx} Two doses: 88% (pooled meta-analysis) ⁶ 91% (95% CI, 93%-96%; United States) [May to July 2021] ^{7xxxi} 79% (95% CI, 60-89; Sweden) [27 Dec 2020-2 Nov 2021] ³⁵	Single dose: 56% (pooled meta-analysis) ⁶ Hospitalization risk reduced by 35-45% ⁹ . Two doses: 91% (pooled meta-analysis) ⁶ 92% (95% CI, 80-97; Sweden) [27 Dec 2020-2 Nov 2021] ³⁵ Against ICU admission: 95.6% (95% CI, 88.3-98.4; Malaysia) [Apr-Sep 2021] ⁶⁸ Against death: 95.3% (95% CI, 91.3-97.4;	No available data	No available data	Against ICU admission: 72.0% (95% CI, 69.9-73.9; Malaysia) [Apr-Sep 2021]68 Against death: 82.4% (95% CI, 81.0-83.7; Malaysia) [Apr-Sep 2021]68	No available data	No available data

xxviii mRNA-aggregated data (results do not disaggregate between BNT162b2 and mRNA-1273).

xxix Results do not disaggregate between mRNA vaccines, BNT162b2 and mRNA-1273.

xxx mRNA-aggregated data (results do not disaggregate between BNT162b2 and mRNA-1273).

 $^{^{\}text{xxxi}}$ Results do not disaggregate between mRNA vaccines, BNT162b2 and mRNA-1273.



	Against ICU admission: 90.3% (95% CI, 88.8-91.6; Malaysia) [Apr-Sep 2021] ⁶⁸ Against death: 92.7% (95% CI, 91.7-93.6; Malaysia) [Apr-Sep 2021] ⁶⁸		Malaysia) [Apr-Sep 2021] ⁶⁸					
Alpha	Single dose: 83% (95% CI, 62-93) 53% (95% CI, 7-83; England) [Feb-Sep 2021] ⁶⁹ Two doses: 95% (95% CI, 78-99) ⁷⁰ . 71% (95% CI, 12-95; England) [Feb-Sep 2021] ⁶⁹ <i>Against death:</i> 98.2% (95% CI, 95.9-99.2) [2-9 weeks] ⁵⁸ . 90.4% (95% CI, 85.1-93.8) [≥20 weeks] ⁵⁸ .	No available data	Single dose: 76% (95% CI, 61-85) 3% (95% CI, -38 – 39; England) [Feb-Sep 2021] ⁶⁹ Two doses: 86% (95% CI, 53-96) ⁷⁰ . 26% (95% CI, -39 – 73; England) [Feb-Sep 2021] ⁶⁹ <i>Against death:</i> 94.1% (95% CI, 91.8-95.8) [2-9 weeks] ⁵⁸ . 78.7% (95% CI, 52.1-90.4) [≥20 weeks] ⁵⁸ .	Beta 67% effective at preventing hospitalizations ⁷¹ . Against death: 96% effective at preventing death ⁷¹ .	No available data	No available data	No available data	No available data
Gamma	No available data	No available data	No available data	72.9% (95% CI, 35.1-91.1) ¹⁷	No available data	Against hospitalization:	No available data	No available data





















								PUBLIC HEA
				Against ICU admission: 92.5% (95% CI, 54.9-99.6) ¹⁷ Against death: 90.5% (95% CI, 31.5-99.6) ¹⁷		95% (95% CI, 86.9-98.1) ⁵⁵ Against death: 94.9% (95% CI, 76.4-98.9) ⁵⁵		
Delta	Single dose: 94% (95% CI, 46- 99) ⁷⁰ . 91% (95% CI, 90- 93) ⁷² 4% (95% CI, -21 – 44; England) [Feb- Sep 2021] ⁶⁹ Two doses: 96% (95% CI, 86- 99) ⁷⁰ . 88% (95% CI, 78.9-93.2) ⁵⁷ . 75% (95% CI, 24- 93.9) ³¹ . 84% (95% CI, 79- 89) ⁷³ . 98.4% (95% CI, 97.9-98.8) [2-9 weeks] ⁵⁸ .	Single dose: 81% (95% CI, 81- 90.6) ³¹ . Two doses: 84% (95% CI, 80- 87) ⁷² 95% (95% CI, 92- 97) [Jun-Aug 2021] ⁷⁴ 96.7% (95% CI, 93.9-98.2) ⁸ 97.3% (95% CI, 95.9-98.4; New York) [Aug 2021] ⁷⁶ Individuals ≥65: 93.7% (95% CI, 92.9-94.4; New York) [Aug 2021] ⁷⁶	Single dose: 71% (95% CI, 51-83) ⁷⁰ 88% (95% CI, 83-91) ⁷² 2% (95% CI, -19 – 31; England) [Feb-Sep 2021] ⁶⁹ Two doses: 92% (95% CI, 75-97) ⁷⁰ . 95.2% (95% CI, 75-97) ⁷⁰ . 95.2% (95% CI, 75-97) ⁷⁰ . 95.2% (95% CI, 70.3-82.3) [≥20 weeks] ⁵⁸ . 77.0% (95% CI, 70.3-82.3) [≥20 weeks] ⁵⁸ . 94% (95% CI, 92-95) ⁷²	71% ⁷¹ 85% (95% CI, 73-91) ¹⁴ . 91% (95% CI, 88-94) ⁷² 93.5% (95% CI, 89.6-96.1; New York) [Aug 2021] ⁷⁶ 85% effective at preventing severe disease and hospitalization ⁸¹ . Individuals ≥ 50: 84% (95% CI, 81-85) ¹⁴	Single dose: Does not offer clinically meaningful protection against severe illness 82,xxxiii Two doses: 88% (95% CI, 55-98) adjusted risk reduction in developing severe illness. 82,xxxiii	Single dose: Does not offer clinically meaningful protection against severe illness 82,xxxiv Two doses: 88% (95% CI, 55-98) adjusted risk reduction in developing severe illness. 82,xxxv	No available data	No available data

 $^{^{\}text{\tiny XXXII}}$ Study does not differentiate between the two inactivated vaccines, BBIBP-CorV and CoronaVac.

 $^{^{\}mbox{\tiny XXXV}}$ Study does not differentiate between the two inactivated vaccines, BBIBP-CorV and CoronaVac.



 $^{^{\}text{xxxiii}}$ Study does not differentiate between the two inactivated vaccines, BBIBP-CorV and CoronaVac.

xxxiv Study does not differentiate between the two inactivated vaccines, BBIBP-CorV and CoronaVac.



92.7% (95% CI, 90.3-94.6) [≥20 weeks] ⁵⁸ . 96% (95% CI, 95-96) ⁷² 80% (95% CI, 73-85) [June-August] ⁷⁴ 93% (95% CI, 84-96) ⁷⁵ 96.8% (95% CI, 84-96) ⁷⁵ 96.8% (95% CI, 84-96) ³² 91.5% (95% CI, 83-94) ⁷⁶ Against death: 90% (95% CI, 83-94) ⁷⁶ Against death: 90% (95% CI, 83-94) ⁷⁸	Against ICU admission: 86% (95% CI, 79-90) ⁷² 96% against severe COVID-19 infection ⁶¹ . Estimated risk of SARS-CoV-2 infection is 4.52 events per 1000 persons (95% CI, 4.17-4.84) ⁷⁹	14% (95% CI, -5 – 46; England) [Feb-Sep 2021] ⁶⁹ 63.1% (95% CI, 51.5-72.1; India) (Apr – May 2021) ⁸⁰ Against moderate to severe disease: 81.5% (95% CI, 9.9-99.0; India) (Apr – May 2021) ⁸⁰ Against ICU admission: Single dose: 92% (95% CI, 84-96) ⁷² Two doses: 96% (95% CI, 94-98) ⁷² Against death: 91% (95% CI, 86-94) [≥2 weeks after second dose] ⁷⁷ All ages: 91% (95% CI, 86-94) ⁷⁸ 40-59: 88% (95% CI, 76-93) ⁷⁸ 60+: 90% (95% CI, 84-94) ⁷⁸	Individuals ≥65: 81.8% (95% CI, 77.8-85.3; New York) [Aug 2021] ⁷⁶ Against ICU admission: 94% (95% CI, 88- 98) ⁷²				
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	40-59: 95% (95% CI, 79-99) ⁷⁸ 60+: 87% (95% CI, 77-93) ⁷⁸ Estimated risk of SARS-CoV-2 infection is 5.75 events per 1000 persons (95% CI, 5.39-6.23) ⁷⁹								
Omicron	Estimated VE against hospitalization 4 to 5-fold increased compared to Delta ⁸³ * 84.9% (95% CI, 83.0-86.6) against Omicron variant for recently vaccinated Pfizer ⁸³ *No differention between mRNA vaccines	Estimated VE against hospitalization 4 to 5-fold increased compared to Delta ⁸³ * *No differention between mRNA vaccines							
DURATION OF PROTECTION, TRANSMISSION & BREAKTHROUGH INFECTIONS									
	BNT162b2/ COMIRNATY	Spikevax/ Moderna COVID- 19 Vaccine/ mRNA-1273	Vaxzevria/ ChAdOx1 nCoV- 19/ AZD1222/ Covishield	Janssen COVID- 19 vaccine/Johnson & Johnson	BBIBP-CorV,	CoronaVac	COVAXIN / BBV152	Novavax/ NVX- CoV2373 (Awaiting approval from WHO EUL)	





Duration of protection (antibodies)	Median time between second dose and infection: 146 days (IQR, 121-167)84 Anti-SARS-CoV-2 Antibodies: 1 month after 2nd dose: 1762 KU/L (IQR: 933-3761) 3 months after 2nd dose: 1086 KU/L (IQR: 629-2155) 6 months after 2nd dose: 802 KU/L (IQR, 447-1487)85 No health worker had antibodies BELOW method-dependent cut-off (0.8 KU/L) Neutralizing antibodies: At peak immunity, NAb titre was 1,789, after 8 months titre was 5386 Pseudovirus neutralizing antibodies:	Preliminary phase I results: Antibody activity remained high in all age groups at day 209 (approximately 6 months) GMT were lower in ≥56 years old89 Neutralizing antibodies: At peak immunity, NAb titre was 5,848, after 8 months titre was 13386 Pseudovirus neutralizing antibodies: At peak immunity, pseudovirus NAb titre was 1,569, after 8 months titre was 27386 Anti-spike Protein RBD IgG Antibodies: At peak immunity, RBD titre was 25,677, after 8 months titre was 1,54686	Antibody Response: After single dose, antibody response declined within one year, but remained above baseline levels. Antibody levels after day 180: 0.54 GMR (CI, 0.47-0.61). Antibody levels after day 320: 0.30 GMR (CI, 0.24-0.39)90 Cellular Immune Response: Day 182 after first dose: median of 237 SFUx106 PBMC (IQR, 109- 520)90 6 months after second dose: (median 1240, IQR 432-2002) in groups with 15-25 week interval between doses90 Anti-spike Protein RBD IgG Antibodies:	Neutralizing antibodies: Remained largely stable for 8-9 months91 Remained stable for 8 months; At 4 weeks after immunization NAb titre was 146, after 8 months titre was 62986 Pseudovirus neutralizing antibodies: Remained stable for 8 months; At 4 weeks after immunization pseudovirus NAb titre was 391, after 8 months titre was 18586 Binding antibodies: Remained stable 6 months irrespective of age group91 Humoral & Cellular Immune Response:	Antibody Response: Unexposed subjects: After 1st dose: 43.6 IU/mL (95% CI, 30.3-62.8) After 2nd dose: 377.0 IU/mL (95% CI: 324.3-438.3) 3 months after 2nd dose: 125.4 IU/mL (95% CI: 88.2- 178.4)93 Exposed subjects: Before 1st dose: 203.2 UI/mL (95% CI: 42.9-962.4) After 1st dose: 761.7 UI/mL (95% CI: 381.1-1522) After 2nd dose: 719.9 UI/mL (95% CI: 264.6-1959) 3 months after 2nd dose: 484.4 IU/mL (95% CI: 147.3- 1593)93 Anti-RBD IgG: Decreased up to 41.8% 2 months after second dose and dropped to	A phase I/II clinical trial found that NAbs titres dropped below the seropositive cutoff of 8, 6 months after the administration of the first dose ⁹⁵ . 80-90% of anti-S IgG and Nab titers against wild type waned 6 months after second vaccination ⁹⁶ Anti-spike Protein RBD IgG Antibodies: Younger age groups (<60): 1 month after 2 nd dose: 97% seropositivity, 11.3 (IQR, 6.2-20.7) 3 months after 2 nd dose: 76% seropositivity, 2.4 (IQR, 1.0-5.0) ⁸⁷ Older age groups (≥60): 1 month after 2 nd dose: 88% seropositivity, 6.4 (IQR, 2.5-13.6)	No available data	No available data
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At peak immunity, pseudovirus NAb titre was 700, after 8 months titre was 16086 Anti-spike Protein RBD IgG Antibodies: At peak immunity, RBD titre was 21,564, after 8 months titre was 75586 Younger age groups (<60): 1 month after 2nd dose: 100% seropositivity, 35.3 (IQR, 27.6-40.0) 3 months after 2nd dose: 100% seropositivity, 19.2 (IQR, 8.2-23.1)87 Older age groups (<60): 1 month after 2nd dose: 100% seropositivity, 19.2 (IQR, 8.2-23.1)87	Humoral & Cellular Immune Response: CD8+ T cell response was 0.017% 8 months after full vaccination86	Younger age groups (<60): 1 month after 2 nd dose: 100% seropositivity, 17.1 (IQR, 9.9-23.6) 3 months after 2 nd dose: 97% seropositivity, 6.5 (IQR, 3.5-9.3) ⁸⁷ Older age groups (≥60): 1 month after 2 nd dose: 96% seropositivity, 13.3 (IQR, 6.9-27.7) 3 months after 2 nd dose: 90% seropositivity, 3.9 (IQR, 1.9-8.4) ⁸⁷	Antibody responses were detected in all vaccine recipients on day 239 (stable response for at least 8 months) ⁹² CD8+ T cell response was 0.12% 8 months after vaccination ⁸⁶ Anti-spike Protein RBD IgG Antibodies: Remained stable for 8 months; At 4 weeks after immunization titre was 1,361, after 8 months titre was 843 ⁸⁶	42.9% decrease after 7 months ⁹⁴ Binding Antibodies: Decreased 82.1% 7 months after second dose ⁹⁴	3 months after 2 nd dose: 60% seropositivity, 1.3 (IQR, 0.5-3.3) ⁸⁷	
(IQR, 22.5-33.3) 3 months after 2 nd						



dose: 100%

seropositivity, **14.8** (IQR, 7.4-18.7)⁸⁷



















			100210 112112
Sub-populations: Older age (≥65): 38% to 42% decrease of humoral antibodies compared to 18- to 45-year-old ⁸⁸			
Older age (≥65) AND men: 37% to 46% decrease compared to 18- to 45-year-old women ⁸⁸			
Immunosuppress ion: 65% to 70% decrease compared to non-immunosuppresse d88			
Obesity (BMI ≥30): 31% increase in neutralizing antibody compared with nonobese ⁸⁸			
Humoral & Cellular Immune Response:			























	CD8+ T cell							
	response was							
	0.016% 8 months							
	after full							
	vaccination86							
	Against any	36.4 (95% CI,	VE reduced by 7%	A study observed				
	SARS-CoV-2	17.1-51.5)	(95% CI, -18 - 2)	sustained and				
	<u>Infection:</u>	reduction of	for every 30 days	stable vaccine				
	After reaching	observed	from the second	effectiveness				
	peak VE (77.5%) 1 month after 2 nd	incidence rate (SARS-CoV-2	dose for those aged 18 to 64	starting 14 days post vaccination to				
	dose, VE dropped	infection) if	years ⁴⁷ .	a maximum of 152				
	to 20% in months	vaccinated from	years .	days after				
	5-7 after 2 nd	Dec 2020 – Apr	VE reduced from	vaccination ¹⁴ .				
	dose ⁹⁷	2021 than Jul	58% (95% CI, 51-					
		2021 – Dec	65) to 27% (95%	VE decreased				
	VE reduced from	2020.103	CI, 17-37) after 4	from 89.4% in				
	87% (95% CI, 85-		months.33	May to 51.7% in				
Duration of	89) to 56% (95%	46.0 (95% CI, -	\ <i>(</i> = 1 1 <i>(</i>	July ³⁸				
protection (vaccine	CI, 53-59) after 4 months. ³³	52.4-83.2) reduction of	VE reduced from	VE decreased	No available data	No available data	No available data	No available data
effectiveness)	1110111115.99	observed	88% (95% CI, 87-89) in March to	from 86.4% (95%				uala
	VE reduced from	incidence rate	3% (95% CI, -7-	CI, 85.2-87.6) in				
	91% (95% CI, 91-	(severe SARS-	12) in August ⁶⁰	March 2021 to				
	92) in March to	CoV-2 infection) if	, 25:21	13.1% (95% CI,				
	50% (95% CI, 47-	vaccinated from	VE decreased by	9.2-16.8) in				
	52) in August ⁶⁰	Dec 2020 – Apr	18.5% points	September				
		2021 than Jul	(95% CI 8.4-33.4)	2021 101				
	VE reduced from	2021 – Dec	among all ages	\/ =				
	89.0% (95% CI,	2020.103	and 19.9% points	VE decreased by 18.5% points				
	84.6-92.1; United States) [May to	VE against the	among older individuals (95%	(95% CI 8.4-33.4)				
	August] to 62.7%	Delta variant	CI; 9.2-36.7)	among all ages				
	(95% CI, 62.4-	declined from	[Overall average	and 19.9% points				
	63.1; United	94.1% (95% CI,	from Systematic	among older				





















90.5-96.3) 14-60	Review and Meta-	individuals (95%
days after		CI; 9.2-36.7)
vaccination to	,	Overall average
80.0% (95% CI,	VE reduced from	from Systematic
70.2-86.6) 151-	96.9% (range,	Review and Meta-
180 days after	93.7-98.0) for the	Regression]99xlix
vaccination.49	week of 1 May	
	2021 to 77.8%	VE reduced from
91% [January-	(range, 70.1-86.8)	86.6% (range,
March]	by the week of	77.8-89.7) for the
71% (95% CI, 53-	August 28 2021 ⁷⁶	week of 1 May
83) [April-May]		2021 to 69.4%
63% (95% CI, 44-	<u>Against</u>	(range, 63.4-77.3)
76) ³⁸	<u>symptomatic</u>	by the week of
		August 28 2021 ⁷⁶ .
VE reduced from		
•	,	<u>Against</u>
	,	<u>symptomatic</u>
		<u>COVID-19:</u>
months ³³		VE decreased by
	,	25.4% (95% CI,
		13.7-42.5) among
•	-	all ages and
,	•	32.0% (95% CI,
		11.0-69.0) among
	Regression	older individuals
	E00/ (050/ CL 4C	[Overall average
	to the same of the	from Systematic
Julie-August**	11 11 11 11 11 11 11 11 11 11 11 11 11	
	vaccination to 80.0% (95% CI, 70.2-86.6) 151-180 days after vaccination. ⁴⁹ 91% [January-March] 71% (95% CI, 53-83) [April-May] 63% (95% CI, 44-76) ³⁸	days after vaccination to 80.0% (95% CI, 70.2-86.6) 151-180 days after vaccination. 49

xxxvi Study does not differentiate between the two mRNA vaccines, Pfizer and Moderna.

xxxviii Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.

xlvi Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.

xivii Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.

xlix Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.



13.7-42.5) among	VE reduced from	Effectiveness did	Review and Meta-		
all ages and	92% (95% CI, 92-	not fall	Regression]991		
32.0% (95% CI,	93) in March to	significantly after			
11.0-69.0) among	64% (95% CI, 62-	longer intervals,	Against severe		
older individuals	66) in August ⁶⁰	however this could	<u>COVID-19:</u>		
[Overall average	\/ / = ==:==t	be influenced by	VE decreased by		
from Systematic	VE against	the study's small	8.0% (95% CI,		
Review and Meta-	infection was 82%	number of	3.6-15.20) among		
Regression ^{99xxxviii}	(95% CI, 79-85)	participants ¹⁰²	all ages and 9.7%		
VE reduced by	14-90 days after the second dose		(95% CI; 5.9-14.7) among older		
22% (95% CI, 6-	and appeared to	Against severe	individuals		
41) for every 30	wane over time	COVID-19:	[Overall average		
days from the	and was 63%	VE decreased by	from Systematic		
second dose for	(95% CI, 55-68)	8.0% (95% CI,	Review and Meta-		
those aged 18 to	91-180 days after	3.6-15.20) among	Regression] ^{99li}		
64 years ⁴⁷ .	the second dose	all ages and 9.7%			
,	[27 Dec 2020 – 26	(95% CI; 5.9-14.7)			
VE against	Oct 2021;	among older			
infection was 82%	Finland]100xli	individuals			
(95% CI, 79-85)	-	[Overall average			
14-90 days after	VE decreased	from Systematic			
the second dose	from 89.2% (95%	Review and Meta-			
and appeared to	CI, 88.8-89.6) in	Regression]99xlviii			
wane over time	March 2021 to				
and was 63%	58.0% (95% CI,				
(95% CI, 55-68)	56.9-59.1) in				
91-180 days after	September				
the second dose	2021 ¹⁰¹				
[27 Dec 2020 – 26					

xxxviiii Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.

xli Study does not differentiate between COMIRNATY/BNT162b2 and SPIKEVAX/ mRNA-1273.

xiviii Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.

¹ Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.

^{II} Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.



Oct 2021;	VE reduced from			
Finland] ^{100xxxix}	89.0% (95% CI,			
_	84.6-92.1; United			
VE decreased	States) [May to			
from 86.9% (95%	August] to 62.7%			
CI, 86.5-87.3) in	(95% CI, 62.4-			
March 2021 to	63.1; United			
43.3% (95% CI,	States) [May to			
41.9-44.6) in	August]98xlii			
September				
2021 ¹⁰¹	VE decreased by			
	18.5% points			
VE declined from	(95% CI 8.4-33.4)			
<mark>81%</mark> (95% CI, 68-	among all ages			
89) 14-73 days	and 19.9% points			
<mark>after second dose.</mark>	among older			
4-6 months after	individuals (95%			
second dose, VE	CI; 9.2-36.7)			
remained at 70%	[Overall average			
<mark>(95% CI, 62-76)</mark>	from Systematic			
and declined to	Review and Meta-			
46% (95% CI, 22-	Regression]99xliii			
63) after six	V/= 1 1 (
months. [second	VE reduced from			
dose was	96.9% (range,			
administered ≥6	93.7-98.0) for the			
weeks after first dose]. ¹⁰²	week of 1 May 2021 to 77.8%			
uusej.'**	(range, 70.1-86.8)			
VE declined from	by the week of			
86% (95% CI, 73-	August 28 2021 ⁷⁶ .			
00 /0 (90% CI, / 3-	August Zo Zuz I			

xxxix Study does not differentiate between COMIRNATY/BNT162b2 and SPIKEVAX/ mRNA-1273.

xlii Study does not differentiate between the two mRNA vaccines, Pfizer and Moderna.

xliii Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.



after second d				
6 months after	<u>symptomatic</u>			
second dose, '				
declined to 61°	VE decreased by			
(95% CI, 45-73	3). 25.4% (95% CI,			
[second dose	was 13.7-42.5) among			
administered <	<mark>≤6</mark> all ages and			
weeks after firs	st 32.0% (95% CI,			
dose] ¹⁰²	11.0-69.0) among			
	older individuals			
Against severe				
<u>COVID-19:</u>	from Systematic			
VE decreased				
8.0% (95% CI,				
3.6-15.20) amo				
all ages and 9.				
(95% CI; 5.9-1				
among older	<u>disease:</u>			
individuals	VE decreased by			
[Overall average				
from Systemat				
Review and M	<u> </u>			
Regression]99x				
	among older			
<u>Against</u>	individuals			
Hospitalization				
and Death:	from Systematic			
After reaching				
peak VE (96.8				
2 months after				
dose, VE did r	not			
decline over				

xl Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.

xliv Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.

xlv Study does not differentiate between Pfizer/BioNTech-Comirnaty, Moderna-mRNA-1273, Janssen-Ad26.COV2.S and AstraZeneca-Vaxrevria.



Transmission prevention	time, except for 7th months (VE 55.6%) with very few cases97 Prior Delta Variant: Vaccine effectiveness against infectiousness given infections 41.3%104 VE against transmission 88.5%104 VE against onwards transmission of Alpha 57% (95% CI, 5-85)69 During Delta Variant: Similar Ct values	VE against onwards transmission: 52% (95% CI, 33-69) ¹⁶ VE against transmission from vaccinated index case to unvaccinated contact is 63% (95% CI, 46-75) and 40% (95% CI, 20-54) to a vaccinated contact. ¹⁰⁸	48% (limited data) May not be able to block the transmission of the alpha variant as efficiently as the wild type ¹⁰⁹ . VE against transmission from vaccinated index case to unvaccinated contact is 63% (95% CI, 46-75) and 40% (95% CI, 20-54) to a vaccinated contact. ^{108liv} Evidence of fully	Limited data	Unknown	Unknown	No available data	No available data
	(<25) were found in both vaccinated and unvaccinated groups ¹⁰⁵		vaccinated individuals infecting other fully vaccinated individuals ¹¹⁰					

liii Study does not differentiate between Comirnaty/BNT162b2, Spikevax/ mRNA-1273, and Vaxzevria/ ChAdOz1 nCOV-19.

 $^{^{\}text{liv}} \ Study\ does\ not\ differentiate\ between\ Comirnaty/BNT162b2,\ Spikevax/\ mRNA-1273,\ and\ Vaxzevria/\ ChAdOz1\ nCOV-19.$



Studies from Scotland and England demonstrated reductions in secondary infections among	81 breakthrough infections among 1100 HCWs; 32 breakthrough infections among 4000 HCWs ¹¹⁰		
families of vaccinated individuals compared to families of unvaccinated individuals ^{106,107} .	VE against onwards transmission of Alpha 35% (95% CI, -26 – 74) ⁶⁹		
VE against onwards transmission: 62% (95% CI, 57-67) ¹⁶	VE against onwards transmission of Delta 42% (95% CI, 14-69) ⁶⁹		
VE against transmission from vaccinated index case to unvaccinated contact is 63% (95% CI, 46-75) and 40% (95% CI, 20-54) to a vaccinated contact. ^{108lii}			
VE against onwards			

iii Study does not differentiate between Comirnaty/BNT162b2, Spikevax/ mRNA-1273, and Vaxzevria/ ChAdOz1 nCOV-19.





	transmission of Delta 31% (95% CI, -3 – 61) ⁶⁹ VE against infection [within a ten-day window] when having a confirmed household exposure 80.4% (95% CI, 73.6-85.5) ⁵⁶							
	Additional infections occurred in 49.8% (95% CI, 48-51.6) of homogenously unvaccinated household members and 12.5% (95% CI, 9.1-17) of homogenously vaccinated household members [within a ten-day window] ⁵⁶							
Breakthrough infections	From 6,161 patients with a positive nasopharyngeal SARS-CoV-2 PCR, 1,120 (18%) were breakthrough	From 6,161 patients with a positive nasopharyngeal SARS-CoV-2 PCR, 1,120 (18%) were	As of 10 June, 1.5 million individuals have been fully vaccinated with Covishield in Odisha Province, India. Between 1	From 6,161 patients with a positive nasopharyngeal SARS-CoV-2 PCR, 1,120 (18%) were	No available data	No available data	As of 10 June, 380,000 individuals have been fully vaccinated with Covaxin in Odisha Province, India.	No available data























of these occurred after 2 May (emergence of Delta variant). Of the 1,120 cases, hospitalized. Of the 1,120 cases, hospitalized. Of the 1,120 cases, hospitalized of								
after 2 May (emergence of Delta variant). Of the 1,120 cases, 126 (12%) were hospitalized. Of the 126 breakthrough admissions, 59 were vaccinated with BNT162b21***. Individuals vaccinated in January and February had a S19 (88) increased risk for breakthrough infections compared to i	infec	ctions – 97%		•				
(emergence of Delta variant). Of the 1,120 cases, 126 (12%) were the 126 coverable damissions, 59 were vaccinated with BNT162b2111. January and February had a 51% (95% Cl., 40-68) increased risk for breakthrough infections compared to individuals not compared to individuals vaccinated in March and Aprill¹²2 Breakthrough infections remained under 1% for fully vaccinated 1% March and Aprill¹²2 Breakthrough infections remained under 1% for fully vaccinated 1	of th	ese occurred	infections – 97%		infections – 97%			
Delta variant). Of the 1,120 cases, 126 (12%) were hospitalized. Of the 1,120 cases, 126 (12%) were hospitalized. Of breakthrough admissions, 59 were vaccinated with BNT162b2¹¹¹¹. Individuals vaccinated in January and February had a pffwarb for breakthrough infections compared to individuals vaccinated in March and April¹¹² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹² Breakthrough infections remained under 1% for fully vaccinated 10 ads measured at 116 and 50 comparable to vaccinated 10 ads measured at 16 comparable to vaccinated 10 comparable to vaccinated 116 comparable to vaccina	after	· 2 May	of these occurred	infections (SARS-	of these occurred		breakthrough	
the 1,120 cases, 126 (12%) were hospitalized. Of the 126 hospitalized.	(eme	ergence of	after 2 May	CoV-2 positive	after 2 May		infections (SARS-	
126 (12%) were hospitalized. Of the 126 preakthrough admissions, 59 were vaccinated with BNT162b2 ¹¹¹ . Individuals vaccinated in January and February had a 51% (95% CI, 40-68) increased risk for breakthrough infections remained under 1% vaccinated in March and April ¹¹² Breakthrough infections remained under 1% for fully vaccinated in March and April ¹¹² Breakthrough infections remained under 1% for fully vaccinated in March and April ¹¹² Breakthrough infections remained under 1% for fully vaccinated in March and April ¹¹² Breakthrough infections remained under 1% for fully vaccinated in March and April ¹¹³ Breakthrough infections remained under 1% for fully vaccinated in March and April ¹¹⁴ breakthrough infections remained under 1% for fully vaccinated in March and April ¹¹⁵ Breakthrough infections remained under 1% for fully vaccinated in March and April ¹¹⁶ breakthrough infections remained under 1% for fully vaccinated under 1% for fully vaccinated in March and April ¹¹⁶ breakthrough infections remained under 1% for fully vaccinated under 1% for fully vaccinated in Dark Prize and National Residual	Delta	a variant). Of	(emergence of	after having	(emergence of		CoV-2 positive	
126 (12%) were hospitalized. Of the 126 preakthrough admissions, 59 were vaccinated with BNT162b2 ¹¹¹ . Individuals vaccinated in January and February had a 51% (95% CI, 40-68) increased risk for breakthrough infections remained under Narch and Aprill ¹¹² between Pfizer or individuals vaccinated in March and Aprill ¹¹² Breakthrough infections remained under 1% for fully vaccinated in March and Aprill ¹¹² Breakthrough infections remained under 1% for fully vaccinated in March and Aprill ¹¹³ Breakthrough infections remained under 1% for fully vaccinated in March and Aprill ¹¹⁴ between Pfizer or bospitalized and sease of Covishield) were identified. Of the 126 breakthrough infections remained under 1% for fully vaccinated in March and Aprill ¹¹⁴ between Pfizer or bospitalized and sease of Covishield) were doese of the 126 breakthrough infections remained under 1% for fully vaccinated in letters and sease of covishield) were doese of the 126 breakthrough infections and the 126 breakthrough infections remained under 1 hospitalized of the 126 breakthrough infections and the 126 breakthrough infections remained under 1 hospitalized of the 126 breakthrough infections and the 126 breakthrough infections and symptomatic but waccinated in letters and any symptomatic but breakthrough infections remained under 1 hospitalization letters and any symptomatic but waccinated in letters and any symptomatic but breakthrough infections and any symptomatic but waccinated in letters and any symptomatic but breakthrough infections and any symptomatic but waccinated in letters and any symptomatic but any symptomatic but waccinated in letters and any symptomatic but any symptom	the 1	1,120 cases,	Delta variant). Of	received two	Delta variant). Of		after having	
hospitalized. Of the 126 breakthrough admissions, 59 were vaccinated with BNT162b2¹¹¹. Individuals vaccinated in January and February had a 51% (95% CI, 40-68) increased risk for breakthrough infections compared to individuals (no difference between Pfizer or March and April¹¹²² breakthrough infections remained under 1 March and April¹²² breakthrough infections remained under 1 March and April²² breakthrough infections remain	126	(12%) were		doses of	the 1,120 cases,		received two	
the 126 breakthrough admissions, 59 were vaccinated with BNT162b2 ¹¹¹ . Individuals vaccinated in January and February had a 51% (95% CI, 40-68) increased risk for breakthrough infections of individuals (no infections of individuals (no infections of individuals (no infections of infections of infections of infections remained under hard and Aprill ¹¹² Breakthrough infections remained under of infections remained under infections of infections of infections of infections remained under hard and Aprill ¹¹² Breakthrough infections hard and aprill ¹¹² Breakthrough infections remained under hard and aprill ¹¹² Breakthrough infections hard and aprill ¹¹³ Breakthrough infections hard and aprill ¹¹⁴ Breakthrough infections hard and aprill ¹¹⁵ Breakthrough infections hard and aprill ¹¹⁶ Breakthrough infections hard and aprill ¹¹⁸ Breakthrough i	hosp	oitalized. Of	126 (12%) were	Covishield) were	126 (12%) were		doses of	
breakthrough admissions, 59 were vaccinated with BNT162b2¹¹¹¹. Individuals vaccinated in January and February had a 51% (95% Cl, 40-68) increased risk for breakthrough infections compared to individuals vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹²² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹² Breakthrough infections remained under 1% for fully vaccinated in March and April¹¹² Breakthrough infections remained under 1% for fully vaccinated in March and Pril¹² Breakthrough infections remained under 1% for fully vaccinated in Priem			hospitalized. Of	identified. Of			Covishield) were	
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were vaccinated with BNT162b2 ¹¹¹ . were vaccinated with mRNA-1273. Individuals vaccinated in January and February had a remained under of individuals (no morbidities) February had a remained under of individuals (no		- U	breakthrough	•	breakthrough		these, 29 (82.9%)	
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January and February had a February had a 51% (95% CI, 40-68) increased risk for breakthrough infections compared to individuals vaccinated in March and April ¹¹² between May and August 2021. ⁹⁸ Breakthrough infections remained under 1% for fully vaccinated in floor f	vacc	inated in	Breakthrough	comorbidities114			comorbidities ¹¹⁴	
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51% (95% CI, 40-68) increased risk for breakthrough infections compared to individuals March and Aprill 12 Breakthrough infections remained under 1% for fully vaccinated 1% for fully vaccinated individuals (no difference between Pfizer or individuals were observed 2-3 to infections are cases were as the case were observed 2-3 to infections are cases were as the case were as th		•	remained under	Median antibody			Median antibody	
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infections compared to between Pfizer or individuals vaccinated in March and April 112 Breakthrough infections remained under 1 % for fully vaccinated 1% for fully vaccinated infections compared to between Pfizer or Moderna recipients between May and August 2021.98 Breakthrough infections remained under 1 % for fully vaccinated infections vaccinated Vietnamese study: Cases were symptomatic but mild, only one case required hospitalization 10x 116 kate of breakthrough infections was case required hospitalization wild, only one cases (out of 62). Their peak viral loads measured at loads were observed 2-3 days before case required hospitalization loads were observed 2-3 days before symptomatic but mild, only one cases were symptomatic but mild, only one cases were symptomatic but mild, only one cases required hospitalization loads measured at loads measured at loads were observed 2-3 days before case required hospitalization loads mild, only one cases were symptomatic but mild, only one cases (out of 62). Their peak viral loads were observed 2-3 days before save required hospitalization loads measured at loads measured at loads measured at loads were observed 2-3 days before case required hospitalization loads measured at loads were observed 2-3 days before case required hospitalization loads measured at loads were observed 2-3 days before case required breakthrough infections was case required hospitalization loads measured at loads were observed 2-3 days before case required breakthrough infections was case required hospitalization loads measured at loads were observed 2-3 days before case required breakthrough infections was case required hospitalization loads measured at loads were observed 2-3 days before case required breakthrough loads measured loads were observed 2-3 days before case required breakthrough loads measured loads were observed 2-3 days before case required loads were observed 2-3 days before case re	,		individuals (no					
compared to individuals Moderna Moderna vaccinated in March and April ¹¹² Breakthrough infections remained under 1% for fully vaccinated Setween Pfizer or Moderna were observed 2-3 mild, only one case required hospitalization were observed 2-3 mild, only one case required hospitalization were observed 2-3 mild, only one case required hospitalization were observed 2-3 mild, only one case required hospitalization were observed 2-3 mild, only one case required hospitalization were observed 2-3 mild, only one case required hospitalization were observed 2-3 mild, only one case required hospitalization wild infections — all cases were symptomatic but breakthrough mild, only one case required hospitalization wild infections was case required hospitalization wild loads measured at prize and were observed 2-3 mild, only one case required hospitalization wild mild, only one case required hospitalization wild loads measured at prize and water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required hospitalization water observed 2-3 mild, only one case required and a mild observed 2-3 mild				Vietnamese study:	cases were		4.2% of fully	
individuals vaccinated in March and April ¹¹² Breakthrough infections remained under 1% for fully vaccinated Moderna recipients days before symptom onset among 49 symptom onset breakthrough breakthrough breakthrough infections was cases (out of 62). Their peak viral loads measured at Moderna recipients days before symptom onset among 49 symptom onset hospitalization ^{lv} 116 hospitalization ^{lv} 116 symptomatic breakthrough infections was case required hospitalization mild, only one cases (out of 62). Their peak viral loads measured at Pfizer and	com	pared to	between Pfizer or		symptomatic but			
vaccinated in March and April ¹¹² between May and August 2021. ⁹⁸ days before symptom onset among 49 Breakthrough infections remained under 1% for fully vaccinated and an approximate to the symptomatic but loads measured at the symptom onset among 49 symptom onset among 49 symptomatic but breakthrough breakthrough breakthrough infections was case required to the symptomatic but mild, only one cases (out of 62). Their peak viral loads measured at Pfizer and 116		•	Moderna	U			developed	
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Breakthrough symptomatic Rate of breakthrough breakthrough cases (out of 62). 1% for fully vaccinated symptomatic but breakthrough breakthrough cases (out of 62). Their peak viral comparable to loads measured at Pfizer and symptomatic but mild, only one case required hospitalization loads measured at loads measured measured measured					•		cases were	
infections remained under 1% for fully vaccinated breakthrough cases (out of 62). Their peak viral loads measured at breakthrough infections was case required hospitalization loads Pfizer and mild, only one case required hospitalization loads	Brea	akthrough	ŭ		Rate of		symptomatic but	
remained under 1% for fully vaccinated cases (out of 62). infections was comparable to loads measured at Pfizer and case (out of 62). infections was comparable to loads measured at Pfizer and case required hospitalization loads loads measured at loads measured		_		•	breakthrough			
1% for fully Their peak viral comparable to loads measured at Pfizer and hospitalization loads measured at Pfizer and hospitalization loads measured at Pfizer and loads measured at Pfizer an	rema	ained under		<u> </u>				
vaccinated loads measured at Pfizer and	1% f	for fully						
		•		•	•			
				any point in time				

^{IV} Study does not differentiate between Covishield (*n*=62.4%) and Covaxin (*n*=37%).

[™] Study does not differentiate between Covishield (*n*=62.4%) and Covaxin (*n*=37%).





	difference between Pfizer or Moderna recipients between May and August 2021.98 Omicron (B.1.1529): Breakthrough cases described symptoms as mild or moderate, had viral loads ranging from 15,011.2 to over 40,000 AU.mL ¹¹³		were higher than that of asymptomatic cases (IQR: 16.5 log10/mL vs 30.8 log10/mL, respectively). NAbs were measured for 10 breakthrough cases, all 10 cases had lower NAbs at day 14 and 90 post second vaccination compared to controls ¹¹⁵	recipients during the initial stages of the study, but increased to 1.96% (2 times the breakthrough rate of mRNA vaccines).98				
			SAFE	TY AND ADVERSE E	VENTS			
	BNT162b2/ COMIRNATY	Spikevax/ Moderna COVID- 19 Vaccine/ mRNA-1273	Vaxzevria/ ChAdOx1 nCoV- 19/ AZD1222/ Covishield	Janssen COVID- 19 vaccine/Johnson & Johnson	BBIBP-CorV,	CoronaVac	COVAXIN / BBV152	Novavax/ NVX- CoV2373 (Awaiting approval from WHO EUL)
Common side effects	Pain at the injection site, fatigue, headache, myalgia, chills and fever ¹¹⁷ , arthralgia ¹¹⁸ Optimal safety for asthma patients ¹¹⁹ .	Pain at injection site, headache, fatigue, myalgia, arthralgia ¹²¹ , Covid arm (cutaneous hypersensitivity) ¹²² .	Fatigue, myalgia, arthralgia, headache ¹²³ , lethargy, fever, & nausea ¹²⁴ .	Headache, fever, chills, fatigue, myalgia, and nausea ¹²⁵ .	Pain at the injection site, dizziness, fever, headache, fatigue, nausea, vomiting, & allergic dermatitis 124,126.	Pain at injection site, headache, fatigue, tremors, & flushing ¹²⁷ , inflammatory reaction, urticaria ¹²⁸ , myalgia ¹²⁹	Pain at injection site, headache, pyrexia, fatigue, myalgia ¹³⁰	Pain at injectionsite, headache, muscle pain, fatigue ⁴²



















								PODEIC HEALT
	The vaccine is considered safe for cancer patients undergoing treatments ¹²⁰ .	The vaccine is considered safe for cancer patients undergoing treatments 120.						
Rare adverse events	Myocarditis & myopericarditis ¹³¹⁻¹³³ , anaphylaxis and swelling of the lips, face, and tongue related to anaphylaxis ¹³⁴ (11 anaphylaxis cases per million doses administered) ¹³⁵ , axillary aadenopathy, paroxysmal ventricular arrhythmia, leg paresthesia ¹³⁶ , pityriasis rosea ¹³⁷ (lesions improved completely after ~8 weeks) ¹³⁸ , lymphocytic vasculitis ¹³⁹ , varicella-zoster reactivation ¹⁴⁰⁻¹⁴² , Kikuchi-Fujimoto disease ¹⁴³ ,	Myocarditis & myopericarditis ¹³¹⁻¹³³ , orofacial swelling & anaphylaxis ¹³⁴ . Potential risk factor for Bell's palsy ¹⁵⁴ (most improve upon follow-up) ¹⁸³ , herpes zoster reactivation ¹⁴¹ , varicella zoster reactivation ¹⁴¹ , herpes zoster ophtalmicus ¹⁸⁴ , eczema & urticaria ¹⁸⁵ , transverse myelitis ¹⁸⁶ , Guillain-Barré syndrome ^{187,188} , acute generalized exanthematous pustulosis ¹⁸⁹ , rhabdomyolysis ^{190,}	Transverse myelitis, high fever 123,198, cutaneous hypersensitivity 198, vasculitis 199, thromboembolism 200, vaccine induced immune thrombotic thrombocytopenia 101, 202-204, intracerebral haemorrhage 205, small vessel vasculitis 202-204, psoriasis 206, rosacea, raynaud's phenomenon 185, Ischaemic stroke 207, anaphylaxis 208, recurrent herpes zoster 209, lvii, generalized	Thrombosis, thrombocytopenia ² 30, increased risk of developing Guillain-Barré syndrome post vaccination ²³¹ , herpes zoster ophtalmicus ¹⁸⁴ , pseudothrombocyt openia ²³² , vaccine induced thrombocytopic thrombosis ²³³ , cutaneous reactions ¹⁷³ , optic neuritis ²³⁴ , subacute thyroiditis ²³⁵ 97% of reported reactions after vaccine administration were nonserious ¹²⁵ .	Cutaneous reactions ¹⁷³ Rare adverse events were similar among the vaccine groups and control group within 7 days ²³⁶ . Pityriasis rosea ²³⁷ , uveitis ²³⁸	Myalgia, fever ¹²⁷ , pityriasis rosea (lesions improved completely after ~8 weeks) ¹³⁸ , reactivation of herpes zoster and herpes simplex ¹²⁸ . Most reactions improved without treatment within a few weeks ¹²⁸ , Guillain-Barré syndrome ²³⁹ , subacute thyroiditis ²⁴⁰ , erythema multiforme ²⁴¹ , uveitis ²³⁸ , vaccine induced thrombotic thrombocytopenia ² ⁴² , serum sickness-like reaction ²⁴³ , cutaneous	Subacute thyroiditis ²⁴⁶	Cutaneous reactions ¹⁷³ Myocarditis was reported in one vaccine recipient, occurring 3 days after second dose ⁴²

Wii All cases occurred in patients with chronic urticaria and were being treated with cyclosporine.





thrombotic thrombocytopenic purpura ^{144,145} , IgA nephropathy flare- up ¹⁴⁶ , Guillain- Barré syndrome ^{147,148} , pustural psoriasis ¹⁴⁹ , immunoglobulin A vasculitis ¹⁵⁰ , immune complex vasculitis ¹⁵¹ , Rhabdomyolysis ¹⁵ ² , subacute thyroiditis ¹⁵³ , Bell's Palsy ¹⁵⁴ , erythema multiforme ¹⁵⁵ , vaccine induced interstitial lung disease ¹⁵⁶ , macular neuroretinopathy ¹⁵ ⁷ , brachial neuritis ¹⁵⁸ , thyroid eye disease ¹⁵⁹ , exacerbation of subclinical hyperthyroidism ¹⁶⁰ , rhabdomyolysis ¹⁶¹ , internal jugular vein thrombosis ¹⁶² , herpes simplex virus keratitis ¹⁶³ , cervical lymphadenopathy ¹	ophtalmicus ¹⁸⁴ , eczema & urticaria ¹⁸⁵ , transverse myelitis ¹⁸⁶ , Guillain-Barré syndrome ^{187,188} , acute generalized exanthematous pustulosis ¹⁸⁹ , rhabdomyolysis ¹⁹⁰ , oervical lymphadenopathy ¹⁹² , glomerulonephritis ¹⁶⁵ , Behçet's disease ¹⁹³ , neurological autoimmune disease ¹⁶⁸ , axillary adenopathy ¹⁶⁹ , multiple sclerosis ¹⁷⁰ , cutaneous reactions ¹⁷³ , Löfgren's syndrome ¹⁹⁴ , erythema multiforme major ¹⁹⁵ . Pemphigus vulgaris ¹⁹⁶ , graft rejection (corneal)	bullous fixed drug eruption ²¹⁰ , Guillain-Barré syndrome ^{148,211} , pityriasis rosea ^{212,213} . Vaccination in individuals with adrenal insufficiency can lead to adrenal crises ^{148,211} , Dariers disease ^{212,213} , vaccine induced acute localized exanthematous pustulosis ²¹⁴ , Henoch-Schönlein Purpura ²¹⁵ , rhabdomyolysis ²¹⁶ , Grave's disease ²¹⁷ , acute demyelinating polyradiculoneuro pathy ²¹⁸ , erythema nodosum ²¹⁹ , polyarthralgia ²²⁰ , recurrence of cutaneous T-cell lymphoma ²²¹ , neurological autoimmune disease ¹⁶⁸ , multiple sclerosis ¹⁷⁰ , sudden			reactions ¹⁷³ , neuromyelitis optica spectrum disorders (transverse myelitis or optic neuritis) ²⁴⁴ , bullous pemphigoid ²⁴⁵		
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⁶⁴ ,glomerulonephri	sensorineural	
tis ¹⁶⁵ , Ramsay-	hearing loss ²²² ,	
Hunt syndrome ¹⁶⁶ ,	acute-onset	
Sweet's	polyradiculoneuro	
syndrome ¹⁶⁷ ,	pathy ²²³ ,	
neurological	cutaneous	
autoimmune	reactions ¹⁷³ ,	
disease ¹⁶⁸ , axillary	<mark>leukocytoclastic</mark>	
adenopathy ¹⁶⁹ ,	<mark>vasculitis</mark> ²²⁴ ,	
multiple	Löfgren's	
sclerosis ¹⁷⁰ ,	<mark>syndrome¹⁹⁴,</mark>	
meningoencephali	<mark>acute eosinophilic</mark>	
tis ¹⁷¹ , intracerebral	<mark>pneumonia²²⁵,</mark>	
haemorrhage due	bullous sweet	
to vasculitis ¹⁷² ,	<mark>syndrome²²⁶,</mark>	
cutaneous	neuralgic	
reactions ¹⁷³ ,	amyotrophy of the	
pigmented	lumbosacral	
purpuric	<mark>plexus²²⁷, sudden</mark>	
dermatosis ¹⁷⁴	sensorineural	
Customia allavaia	hearing loss ²²⁸ ,	
Systemic allergic	graft rejection	
symptoms were	(corneal) ¹⁹⁷ ,	
more common in	erythema envitore	
BNT162b2 than	annulare	
mRNA-1273,	centrfugum ²²⁹	
however, anaphylaxis rates		
were similar for		
both mRNA		
vaccines ¹⁷⁵ , could		
potentially worsen		
migraines in		
people who		
already suffer from		
migraines ¹⁷⁶ , graft		
rejection		
rojection		





















								PUBLIC HEA
	(corneal) ¹⁷⁷ , flexural exanthema ¹⁷⁸ , severe non-anaphylatic allergic reaction ¹⁷⁹ , uveitis ¹⁸⁰ , erythroderma ¹⁸¹ Having adverse reactions is associated with enhanced SARS-CoV-2 IgG antibody response ¹⁸²							
Potential associated adverse events (causal links not yet proven)	Cerebral venous sinus thrombosis and intracranial haemorrhage ²⁴⁷ , aseptic meningitis ²⁴⁸ , autoimmune hepatitis ^{249,250} , multiple sclerosis relapse ²⁵¹ , myeloperoxidase anti-neutrophil cytoplasmic antibody-positive optic perineuritis ²⁵² , central retinal vein occlusion ²⁵³ , paracentral acute	Cerebral venous sinus ²⁷¹ , Autoimmune hepatitis ²⁴⁹ , myocardial infarction ²⁷² , autoimmune haemolytic anaemia ²⁷³ , hypophysitis & panhypopituitaris m ²⁷⁴ , erythema nodosum-like rash ²⁷⁴ , pulmonary embolism ²⁷⁵ , minimal change disease ²⁷⁶ , encephalomyelitis ²	Autoimmune hepatitis ^{249,281,282} , Acute hyperglycaemic crisis ²⁸³ , Facial nerve palsy, cervical myelitis ²⁰⁷ , alopecia areata ²⁸⁴ , takotsubo (stress) cardiomyopathy ²⁸⁵ , acute disseminated encephalomyelitis ² ⁸⁶ , cerebral venous sinus thrombosis ^{287,271} (higher risk for women) ²⁰¹ , ophthalmic vein	Facial Diplegia ²⁹³ , acute macular neurotinopathy ²⁹⁴ , cerebral venous sinus thrombosis ^{271,295} , oral lichen planus ²⁹⁶	Longitudinally extensive transverse myelitis ²⁹⁷	Likely vaccine associated disease enhancement (VADE) ²⁹⁸ , autoimmune hepatitis ²⁹⁹	No available data	No available data





















middle maculopathy & acute macular neurotinopathy ²⁵⁴ , Stevens-Johnson syndrome/ toxic epidermal necrolysis ^{255,256} , lichenoid cutaneous skin eruption ²⁵⁷ , acute mania and psychotic features ²⁵⁸ , acute psychosis due to anti-N-methyl-D- aspartate receptor (anti-NMDAR) encephalitis ²⁵⁹ , alopecia areata ²⁶⁰ , rhombencephalitis inflammation and organ dysfunction ²⁶² , aplastic anaemia ²⁶³ , bullous pemphigoid ²⁶⁴ , minimal change disease ²⁶⁵ , miller fisher syndrome ²⁶⁶ ,	nephritis ²⁷⁸ , retinal vein occlusion ²⁷⁹ One case developed IgA Nephropathy after receiving the second dose of mRNA-1273 ²⁸⁰ .	thrombosis ²⁸⁸ , retinal vein occlusion ²⁸⁹ , Still's disease ²⁹⁰ , autoimmune encephalitis ²⁹¹ , acute abducens palsy ²⁹²			





















	multifocal placoid pigment epitheliopathy ²⁶⁹ , trigeminal neuralgia ²⁷⁰							
Myocarditis da	Mainly reported in young adults and adolescents 300 Israeli study: Estimated incidence within 42 days after receipt of first dose per 100,000 vaccinated persons was 2.13 cases (95% CI, 1.56-2.7)301 Male patients Incidence of 4.12 (95% CI, 2.99-5.26) per 100,000 vaccinated301 3.19 cases (95% CI, 2.37-4.02) per 100,000 vaccinated302 Female patients Incidence of 0.23 (95% CI, 0-0.49)	Mainly reported in young adults and adolescents ³⁰⁰ 5.8 cases per 1 million second dose administrations ³⁰³	No available data	Myocarditis was reported as viral myocarditis. Participant fully recovered after 2 days of hospitalisation. No episode of anaphylaxis or vaccine-associated enhanced COVID-19 was reported ⁴²				























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per 100,000 vaccinated ³⁰¹			
0.39 cases (95% CI, 0.10-0.68) per 100,000 vaccinated ³⁰²			
≥30 years Incidence of 1.13 (95% CI, 0.66- 1.60) per 100,00 vaccinated ³⁰¹			
5.8 cases per 1 million second dose administrations ³⁰³			
4.8 cases			
5.07 cases per 100,000 ³⁰⁴			
Disease severity Mild: 1.62 (95% CI, 1.12-2.11) Intermediate: 0.47 (95% CI, 0.21- 0.74) Fulminant: 0.04 (95% CI, 0- 0.12) ³⁰¹			
Risk per 100,000 persons			





















	1st dose (male): 0.64 2nd dose (male); 3.83 1st dose (female): 0.07 2nd dose (female): 0.46 1st dose (male 16- 19): 1.34 2nd dose (male 16- 19): 15.07 ³⁰²			HILDREN VACCINAT	TON.			
	BNT162b2/ COMIRNATY	Spikevax/ Moderna COVID- 19 Vaccine/ mRNA-1273	Vaxzevria/ ChAdOx1 nCoV- 19/ AZD1222/ Covishield	Janssen COVID- 19 vaccine/Johnson & Johnson	BBIBP-CorV,	CoronaVac	COVAXIN / BBV152	Novavax/ NVX- CoV2373 (Awaiting approval from WHO EUL)
Efficacy	Adolescents (12-15): After one dose had efficacy of 75% (CI, 7.6-95.5) After second dose efficacy of 100% (CI, 78.1-100) ³⁰⁵ . Children (5-11):	Adolescents (12-17): 14 days after one dose had efficacy of 92.7% (CI, 67.8-99.2) After second dose efficacy of 93.3% (CI, 47.9-99.9) ³⁰⁸ Against SARS-CoV-2 Infection:	No available data Paused ongoing trials in children aged 6-17 due to concerns over rare blood clots reported in adult population ³¹⁰ .	No available data Announced at begging of April ongoing study in adolescents but paused to investigate blood clots in adult population ³¹⁰ .	Children (3-17): Unknown. Ongoing clinical trial only looked at safety, tolerability, and immunogenicity viii * * The study design administered three	Children (3-17): Unknown. Clinical trial only looked at safety, tolerability and immunogenicity ³¹¹ .	No available data	Adolescents (16-17): PREVENT-19 clinical trial ^{lix} expanded to assess efficacy, safety, and immunogenicity in 12–17-year- old adolescents ³¹²

Safety and immunogenicity of an inactivated COVID-19 vaccine, BBIBP-CorV, in people younger than 18 years: a randomised, double-blind, controlled, phase 1/2 trial. *The Lancet Infectious Diseases*. https://www.thelancet.com/journals/laninf/article/PIIS1473-3099(21)00462-X/fulltext

ix A Study to Evaluate the Efficacy, Immune Response, and Safety of a COVID-19 Vaccine in Adults ≥18 Years With a Pediatric Expansion in Adolescents (12 to <18 Years) at Risk for SARS-CoV-2. ClinicalTrials.gov. ClinicalTrials.gov Identifier: NCT04611802. https://clinicaltrials.gov/ct2/show/NCT04611802?term=Novavax&cond=Covid19&draw=2





	After second dose efficacy of 90.7% (CI, 67.7-98.3) ³⁰⁶ Children (Under 5 years): Ongoing trials ³⁰⁷	14 days after first dose efficacy of 68.9% (95% CI, 49.9-82.1) 14 days after second dose efficacy of 55.7% (95% CI, 16.8,82.1) ³⁰⁸			doses of 2 μg, 4 μg, or 8 μg of vaccine			
		Against asymptomatic: 14 days after first dose efficacy of 59.5% (95% CI, 28.4-77.3) 14 days after second dose efficacy of 39.2 (95% CI, -24.7- 69.7) ³⁰⁸ Children (6month-						
		11): Ongoing trials ³⁰⁹						
Effectiveness	Adolescents Against SARS- CoV-2 infection: 91.5% (95% CI, 88.2-93.9) ³¹³ 91% (95% CI, 88- 93) ³¹⁴ Adolescents Against hospitalisation:	No available data	No available data	No available data	No available data	No available data	No available data	No available data



















84% (95% CI, -55- 98) ³¹⁻¹⁴ 933 (95% CI, 83- 97) ³¹⁵ Adolescents (12- 15) Serum- neutralizing itier: 1 month after 2nd dose had 1283.0 GMNs ₀ (CI, 1095.5-1402.5) ³⁰⁵ . Adolescents/youn g adult (16-25) serum-neutralizing antibodies after 28 days after 2 nd dose ranged from 105.3-180.2 GMT in 6-12 years cohort, 84.1-168.6 GMT in 6-12 years cohort, 34.1-168.6 GMT in 6-12 years cohort, add 88.0 155.7 GMT in 13- 1537.4) No available data
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Safety and Adverse events	Adolescents (12-15): Local and systemic events were generally mild to moderate Severe injectionsite pain (1.5%) Fever (20%) High Fever (0.1%) Adverse events (6%) Severe adverse events (0.6%) ³⁰⁵ . Adolescent/young adults (16-25): Local and systemic events were generally mild to moderate Severe injectionsite pain (3.4%) Fever (17%) Adverse events (6%) Severe adverse events (1.7%) ³⁰⁵ .	Adolescents (12-17): Solicited local reactions after 2nd dose (93.4%) Most common solicited adverse reactions were Injection-site pain (92.7%) Headache (70.2%) Fatigue (67.8%) Grade 3 adverse events (6.8%))322 Most common solicited local reaction: injection-site pain after first injection (93.1%) and second injection (92.4%) Most common systemic reactions: fatigue, myalgia, and chills308	No available data	No available data	Children (3-17): Most common adverse reaction was pain at injection site in 3–5-year group (4%), 6-12-year group (1.2%), and 13-17-year group (7.9%) Most common systemic reactions in all three age cohorts were mild to moderate fever and cough Adverse events were mostly mild to moderate in severity ³¹⁷	Children (3-17): Adverse reactions in 12–17 year group (35%), 3-5 year group (26%), and 6-11 year group (18%) Reported at least one adverse event (27%) Most reported events were mild and moderate and only (<1%) grade 3 events Injection-site pain (13%) Fever (25%) ³¹¹	Ongoing clinical trial ³¹⁸	Ongoing clinical trial ³¹⁹
	Children (5-11): Pain at injection site, fatigue, headache, chills were reported. Overall, the	Children (6-11): Vaccine was generally well tolerated ³¹⁶ Children (6month- 11):						





















	vaccine is safe and tolerable ³⁰⁶	Ongoing trials ³⁰⁹						
	<u>Children (Under</u> <u>5):</u> Ongoing trials ³⁰⁷							
	Multisystem inflammatory syndrome (causal link not yet proven) ³²⁰							
	Adverse events cases: 15-year old boy developed nephrotic syndrome ³²¹							
Managed Military Protection	Few reported cases of acute myocarditis and pericarditis in 16-25 year olds (mainly in males) ³²²	Few reported cases of acute myocarditis and pericarditis (mainly in males) ³²²	Na cuellable dete	Na sveilakla data	No queilable dete	No sucitable data		
Myocarditis Data	16-29 years Incidence of 5.49 (95% CI, 3.59- 7.39) per 100,00 vaccinated ³⁰¹	16-17 year old boys in US: Second dose: 31.2 cases per million doses administered ³²³	No available data					
	Male patients (16- 29 years)							





















			Tobero nene
Incidence of 10.69 (95% CI, 6.93-14.46) per 100,000 vaccinated ³⁰¹			
Incidence of 13.6 cases (95% CI, 9.30-19.20) per 100,000 vaccinated ³⁰²			
12-15 year old boys in US: First dose: 4.8 cases per million doses administered ³²³			
Second dose: 42.6 cases per million doses administered ³²³			
girls in US: First dose: 0.5 cases per million doses administered ³²³ Second dose: 4.3			
cases per million doses administered ³²³ 16-17 year old boys in US:			





















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	First dose: 5.2 cases per million doses administered ³²³ Second dose: 71.5 cases per million doses administered ³²³ 16-17 year old girls in US: First dose: 0.0 cases per million doses administered ³²³ Second dose: 8.1 cases per million doses administered ³²³							
			HETE	ROLOGOUS VACCI	NATION			
Vaccine Schedule	BNT162b2/ChAd Ox1 Administration of ChAdOx1 as second/booster dose	ChAdOx1/mRNA- 1273 Administration of mRNA-1273 as second/booster dose	ChAdOx1/BNT16 2b2 Administration of BNT162b2 as second/booster dose	Not Applicable (one dose schedule) For more information refer to booster section	BBIBP/BNT162b2	CoronaVac/ChAd Ox1 Press releases have confirmed that Thailand will use the AstraZeneca vaccine as the second dose for individuals whose	ChAdOx1/BBV15 2 Administration of Covaxin as second/booster dose	Ongoing trial ³²⁴ (Com-Cov2) ^{lxi}

lxi Comparing COVID-19 Vaccine Schedule Combinations. *University of Oxford*. https://comcovstudy.org.uk/about-com-cov2





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						first dose was Sinovac ^{lx} CoronaVac/Conv idecia		
Immunogenici	GMCs of SARS-CoV-2 anti-spike IgG at 28 days post booster: Heterologous (7133 ELU/mL, CI 6415-7932) vs. Homologous (14080 ELU/mL, CI 12491-15871) ³²⁵ . SFC frequency (TOcell ELISpot): Heterologous (99 SFC/10 ⁶ PBMCs) vs. Homologous (80 SFC/10 ⁶ PBMCs) ³²⁵ . Heterologous (80 SFC/10 ⁶ PBMCs) ³²⁵ . Heterologous (80 SFC/10 ⁶ PBMCs) ³²⁵ .	*Spike-specific IgG antibodies: Heterologous (3602 BAU/mL) vs. Homologous (4189 BAU/mL) ⁴⁸ *Neutralizing antibodies: Heterologous (100%) vs. Homologous (100%) ³²⁶ . Heterologous mRNA: 84.7% effectiveness (95% CI, 83.1-86.1) ⁸ *Results based on immunosuppressed population	RBD antibody titres: Heterologous (7756.68 BAU/mL, CI 7371.53-8161.96) vs. Homologous (99.84 BAU/mL, CI 76.93-129.59) at day 14 ³²⁷ . IgG antibody titres: Heterologous (3684 BAU/mL) vs. Homologous (101.2 BAU/mL) at day 14 ³²⁷ . Neutralizing antibodies: Heterologous (100%) at day 14 vs.	Not Applicable (one dose schedule) For more information refer to booster section	Unknown (on- going clinical trial) ⁴⁹	CoronaVac/ChAd Ox1: Anti-S Antibodies: Heterologous (797 U/mL; 95% CI, 598.7-1062) vs. Homologous CoronaVac (94.4 U/mL; 95% CI: 76.1-122.1) vs. Homolougous ChAdOx1 (818 U/mL; 95% CI: 662.5-1010) CoronaVac/Conv idecia Neutralizing antibodies: Heterologous 54.4 GMT (95% CI, 37.9-78) vs. Homologous CoronaVac	RBD antibody titres: Heterologous (1866 GMT; 95% CI, 1003-3472) vs. Homologous Covishield (2260 GMT; 95% CI, 1881-2716) vs. Homologous Covaxin (710 GMT, 95% CI, 461-1092) ³³¹ N-protein IgG: Heterologous (1145 GMT; 95% CI, 520.7-2520) vs. Homologous Covishield (353.7 GMT; 95% CI, 219.9-568.9) vs. Homologous Covaxin (742.4	No available data Ongoing trial ³²⁴

^{lx} Malaysia to stop using Sinovac vaccine after supply ends - minister. *Reuters* [press release]. https://www.reuters.com/world/asia-pacific/malaysia-stop-using-sinovac-vaccine-after-supply-ends-minister-2021-07-15/



			Homologous (30%) at day 14 ³²⁷ . Heterologous (median 99%) vs. Homologous (BNT162b2/BNT1 62b2) (median 62%) ³²⁸			12.8 GMT (95% CI, 9.3-17.5) ³³⁰	GMT; 95% CI, 485.8-1134) ³³¹ Neutralizing antibody titres: Heterologous (171.4 GMT; 95% CI, 121.3-242.3) vs. Homologous Covishield (111 GMT; 95% CI, 98.59-124.9) vs. Homologous Covaxin (86 GMT; 95% CI, 138.2- 252.0) ³³¹	
Immunogenicity against variants	No available data	No available data	Neutralizing Antibodies for Alpha, Beta, Gamma, and Delta: Heterologous 2.3-fold to 3.6- fold higher neutralizing antibodies than homologous ³²⁸ Omicron (B.1.1.529): 13/20 seropositive against Omicron ³³²	No available data	No available data	No available data	Neutralizing antibody titres B.1: 539.4 GMT (95% CI, 263.9-1103) ³³¹ Neutralizing antibody titres Alpha: 396.1 GMT (95% CI, 199.1-788) ³³¹ Neutralizing antibody titres Beta: 151 GMT (95% CI, 80.21-284.3) ³³¹	No available data



















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							Neutralizing <u>antibody titres</u> <u>Delta:</u> 241.2 GMT (95% CI, 74.99-775.9) ³³¹	
Reactogenicity	Observed increase in systemic reactogenicity after boost in heterologous schedules in comparison with homologous schedules ³²⁵ Adverse events in heterologous: Adverse events (90) Grade 1 (54.4%) Grade 2 (37.8%) Grade 3 (7.8%) Grade 4 (0%) Arthralgia, Migraine, Back Pain ³²⁵ . Adverse events in homologous: Adverse events (81) Grade 1 (59.3%) Grade 2 (39.5%) Grade 3 (1.2%)	*Adverse events in heterologous and homologous vaccination groups were very similar ³²⁶ . *Majority of adverse events self-reported were Pain at injection site, Swelling at injection site, Fever, Headaches, Fatigue, Chills, GI effects, Myalgia, Arthralgia ³²⁶ . *Results based on immunosuppressed population	Adverse events in heterologous: Headache (44%), Myalgia (43%), Malaise (42%), Fever (2%), Injection site pain (88%), Induration (35%), Erythema (31%) ³²⁷ . Severity of adverse events in heterologous: Mild (68%), Moderate (30%), Severe (2%) ³²⁷ .	Not Applicable (one dose schedule) For more information refer to booster section	Unknown (on- going clinical trial) ³³³	CoronaVac/ChAd Ox1: Unknown CoronaVac/Conv idecia: Convidecia recipients reported more adverse reactions and reported higher occurrence of solicited injection- site pain)330	Most common local adverse events: Pain at injection site (11.1%) ³³¹ Most common systemic adverse events: Pyrexia (27.77%, 11.1%) after 1st and 2nd dose Malaise (33.3%, 5.5%) after 1st and 2nd dose ³³¹	No available data Ongoing trial ³²⁴





















	Grade 4 (0%) ³²⁵ .							
				BOOSTER DOSES				
Vaccine Schedule	BNT162b2/BNT16 2b2	mRNA- 1273/mRNA-1273	ChAdOx1/ChAdO X1	Ad26.CoV.2.S/ Ad26.CoV.2.S	SinoPharm/Sino Pharm	CoronaVac/Coro naVac	Covaxin/Covaxin	NVX- CoV2373/NVX- CoV2373
Approved Administration	Israel: 12-year-old and over can received homologous booster shot 5 months after full jab ^{xii} United States: Starting September, adults who received mRNA vaccine 8 months ago are eligible for booster	Phase II booster trial of three booster doses are ongoing ³³⁴ Moderna sought FDA approval of its COVID-19 vaccine booster ^{lxiv} <u>United States:</u> Starting September, adults who received mRNA vaccine 8 months ago are	Preliminary results on tolerability and immunogenicity of third dose of ChAdOx1 vaccines showed strong boost to the immune response ³³⁵	Johnson & Johnson has said it will submit all of their new data to the FDA for potential consideration for adding a booster dose and consideration to authorize two- dose regimen ^{IXV}	UAE: Offering booster doses of Pfizer and Sinopharm to people who received full Sinopharm jab ≥6 months ago	Turkey and the United Arab Emirates began homologous booster shots Indonesia and Thailand are considering giving homologous booster shot to HCW ^{lxvi}	Ongoing clinical trials ^{lxvii}	Ongoing phase II trials ³³⁶ Results below are based on ongoing phase II trial

kii Israel offers COVID-19 booster to all vaccinated people. Reuters [press release]. https://www.reuters.com/world/middle-east/israel-offers-covid-19-booster-shots-all-vaccinated-people-2021-08-29/

waccine-booster-us-fda-2021-09-01/

Two dose version of Johnson & Johnson shot 94% effective against Covid-19, study finds. CNN. https://edition.cnn.com/2021/09/21/health/johnson-vaccine-two-doses-booster/index.html

Indonesia and Thailand consider booster shots amid doubts over Sinovac vaccine. Reuters [press release]. https://www.reuters.com/world/china/indonesia-thailand-consider-booster-shots-amid-doubts-over-sinovac-vaccine-2021-07-08/

bharat Biotech to initiate trials of booster dose of Covid-19 vaccine. Clinical Trials Arena. https://www.clinicaltrialsarena.com/news/bharat-biotech-booster-dose/



	Europe: Starting in fall, most European countries are planning on rolling out booster shots to immunocompromi sed and elder populations with some countries administering to overall population xiii	eligible for booster.						
Time-to-booster dose	6 months to 8 months after initial two-dose regimen Israel offers up to 5 months after initial two-dose regimen UK has shortened time interval up to 3 months after initial two-dose regimen due to new Omicron variantlxviii	6 months to 8 months after initial two-dose regimen	6-9 months after initial two-dose regimen	6 months after one dose regimen ⁹¹	6 months after initial two-dose regimen	6 months to 12 months After primary vaccination 8 months after the primary vaccination to healthy adults ≥60 years	Ongoing clinical trials ^{xxxvii}	6 months after initial two-dose regimen (189 days) ³³⁶

kxiii A country-by-country guide to coronavirus vaccine booster plans. POLITICO [press reléase]. https://www.politico.eu/article/vaccine-booster-coronavirus-covid-19-europe-delta-varian-who/lxviii UK's minimum gap for Covid-19 booster jabs to be halved to three months. The Guardian [press release]. Accessed on 12 December 2021.

https://www.theguardian.com/world/2021/nov/29/covid-booster-jabs-to-be-offered-to-all-uk-adults-after-three-month-gap



Efficacy	Symptomatic COVID-19: 95.6% during Delta prevalent period ³³⁷ 95.3% (95% CI, 89.5-98.3) ³³⁸ 96.5% (95% CI, 89.3-99.3) in 16- 55 year old ³³⁸ 93.1% (95% CI, 78.4-98.6) in ≥55 year old ³³⁸	No available data	No available data	No available data	No available data	No available data	Ongoing clinical trials ^{xxxvii}	No available data
Effectiveness	Effectiveness against testing positive: 12% (95% CI, 8- 17) in first 7 days after booster 58% (95% CI, 56- 61) 14 days after booster 85% (95% CI, 83- 86) 28 days after booster ³³⁹ Effectiveness against infection: 92% (95% CI, 91- 92) ³⁴⁰ Effectiveness in ≥50:	Effectiveness against infection: 94% (95% CI, 91- 95) ³⁴⁰	No available data	No available data				





















	84.4% (95% CI, 82.8-85.8) against symptomatic COVID-19 ³⁴¹ 94.0% (93.4-94.6) against symptomatic COVID-19 compared with unvaccinated ³⁴¹				
	Effectiveness against hospitalization: 87% 0-6 days after receiving booster dose 92% to 97% lower than those who received 2 doses ³³⁹				
Effectiveness against Variants	Omicron (B.1.1.529): 75.5% (95% CI, 56.1-86.3) effectiveness against symptomatic infection ⁶⁶ If assuming 25- fold decrease compared to wild- type, 81% (95% CI, 59-95) ⁶⁷				























Immunogenicity	Neutralizing titers: Elicits >5-8 more for wild type after 6 months after 2nd dose³42 ≥ 60 years: Neutralizing antibody: 9.34 times higher than second dose³43 IgG Antibodies in 97% seroconversion with increase in IgG antibody titers³44 33-fold increase in IgG after booster dose³43	Booster doses (mRNA1273 or mRNA1273.351) increased neutralizing antibody titers against wild- type ³⁴⁵	Antibody Levels: Higher levels after third dose (tlgG EU 3746; IQR: 2047-6420) 335 Spike Cellular Immune Response: Increased from 200 SFUx10 ⁶ PBMC (IQR, 127-389) after the second dose to 399 SFUx10 ⁶ PBMC (IQR, 314-662) after the third one ³³⁵	5X10 ¹⁰ vp booster dose elicited 9-fold increase at day 7 compared to first dose after 29 days in 18-55-year-olds ⁹¹ 1.25X10 ¹⁰ vp booster dose elicited 6-7.7-fold increase at day 28 compared to first dose after 29 days in 18-55 and ≥65-year-old ⁹¹	Ongoing trial ³³³ IgG Seroconversion: 175/176 vaccinees were seropositive for IgG 14 days after receiving third dose ⁹⁴ Mean IgG value increased 8.00-fold compared to before third vaccination ⁹⁴ Anti-RBD IgG: Increased by 8.14-fold higher than before third vaccine ⁹⁴ Memory B cells: Third dose increased the percentage of RBD-specific memory B cells (0.96%) ⁹⁴	Neutralizing Antibodies: 60% higher NAbs activity against wild-type compared to 2- doses ⁹⁶ Anti-S IgG and NAbs: 20-fold increase 4 weeks post booster vaccination NAbs were maintained 60 to 180 days post booster ⁹⁶	Ongoing clinical trials******ii	Anti-spike IgG: Increase of 4.6- fold compared to peak response after 2nd dose (Day 217 GMEU = 200408; 95% CI: 159796- 251342)336 Wild-type Neutralizing Response: Increase of 4.3- fold compared to peak response after 2nd dose (IC50 = 6231; 95% CI: 4738-8195) 336 Older Participants (60- 84): 5.4-fold increase in antibody response ³³⁶ Younger Participants (18- 59):
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								3.7-fold increase in antibody response ³³⁶
Immunogenicity against variants	Beta (B.1.351): Elicits 15-21 more neutralizing titers for Beta variant after 6 months after 2 nd dose ³⁴² Delta (B.1.671.2): >5-fold increase in neutralizing titers against Delta compared to dose 2 titers in 18–55-year-olds >11-fold increase in neutralizing titers against Delta compared to dose 2 titers in 65–85-year-olds ³⁴² Omicron (B.1.1.529): 37.0-fold decrease in neutralization compared to Delta after 0.5 months after booster 24.5-fold decrease in neutralization compared to Delta compared to Delta after 0.5 months after booster	Preliminary results of booster doses of mRNA-1273 vaccine show robust antibody response against Delta variant ³³⁴	Third dose provided higher antibody titters against Alpha, Beta, and Delta variants ³³⁵	No available data	Ongoing trial ³³³ Beta (B.1.351): 71.6% plasma inhibitions against Beta variant ⁹⁴ Delta (B.1.671.2): 83.4%% plasma inhibitions against Delta variant ⁹⁴ Lambda: 89.0% plasma inhibitions against Lambda variant ⁹⁴	Beta (B.1.351): 3.0-fold decrease in neutralizing antibodies compared to wild type ⁹⁶ Gamma (P.1): 3.1-fold decrease in neutralizing antibodies compared to wild type ⁹⁶ Delta (B.1.671.2): 2.3-fold decrease in neutralizing antibodies compared to wild type ⁹⁶ Delta (B.1.671.2): 2.5-fold decrease in neutralizing antibodies compared to wild type 2.5-fold higher neutralizing potency than 2-dose vaccination ⁹⁶	Ongoing clinical trials XXXXVII	High levels of functional antibodies against Alpha (B.1.1.7), Beta (B.1.351), and Delta (B.1.671.2): Increase of 6.6-fold in antibody response compared to Delta response observed with primary vaccination ³³⁶





















	after 3 months after booster ³⁴⁶	Similar safety and						
Reactogenicity	Preliminary results show consistent tolerability ³⁴² 25% reported at least one adverse event ³³⁸ Common solicited AE: Injection site pain, injection site redness, injection site swelling, fatigure, muscle pain, fever ³³⁸ ≥Grade 3 AE: 6.6% reported grade 3 or higher reactogenicity with 0.7% being local reactions and 5.9% systemic events ³³⁸	tolerability compared to second dose ³³⁴ Common solicited local adverse events: Injection-site pain (68.4% for mRNA-1273.351, 90% for mRNA-1273) fatigue (36.8% for mRNA-1273) headache (36.8% for mRNA-1273) myalgia (31.6% for mRNA-1273) myalgia (31.6% for mRNA-1273) arthralgia (21.1% for mRNA-1273, 50.0% for mRNA-1273)	Lower reactogenicity after third dose compared to first dose ⁹⁰	No available data	Ongoing trial ³³³	The third shot is considered to be safe ⁹⁵ . Common side effects: Pain at the injection site. Adverse events: Unrelated to the vaccination	Ongoing clinical trials ^{xxxvii}	Booster dose was well tolerated Local and systemic reactogenicity increased between Dose 1, Dose 2, and Dose 3 90% of symptoms were rated as mild or moderate ³³⁶



















against COVID-19 Dooster group ³⁴⁷ information info	Protection against COVID-19	12.2 (95% CI, 11.4-13.0) lower rate in booster group ³⁴⁷ Oldest age group (≥60): 12.3 (95% CI, 10.4-12.3) lower rate in booster group ³⁴⁸ 12.3 (95% CI, 11.8-12.8) lower	No available information	Ongoing clinical trials************************************	No available information				
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Severe Illness: 40-59 age group: 21.7 (95% CI, 10.6-44.2) lower rate in booster group ³⁴⁷			
Older population (≥60): 19.5 (95% CI, 12.9-29.5) lower rate in booster group ³⁴⁸ 17.9 (95% CI, 15.1-21.2) lower rate in booster group ³⁴⁷			
Mortality: ≥60 years old: 14.7 (95% CI, 10.0-21.4) lower rate in booster group ³⁴⁷			
≥50 years old: Adjusted hazard ratio for death due to COVID-19 in booster compared to non-booster was 0.10 (95% CI, 0.07 to 0.14) or			























					TOBLIC HEA
	90% lower mortality rate ³⁴⁹				
	Detailed report from Pfizer				
	regarding booster doses can be				
	found here:				
	https://www.fda.go v/media/152161/d				
	<u>ownload</u>				
	14-20 days after				
	booster, marginal effectiveness				
	increases to 70-			For more detailed information	
Other	84% ³⁵⁰			regarding immunogenicity of	
	Incidence Rate:			third dose refer to	
	Infection in			study ^{lxix}	
	individuals <60: 0.22 (95% CI,				
	0.22-0.23) incidence rate in				
	booster compared				
	to non-booster ³⁵¹				
	Infection in individuals ≥60:				
	0.16 (95% CI,				
	<mark>0.15-0.17)</mark>				

lxix A third dose of inactivated vaccine augments the potency, breadth, and duration of anamnestic responses against SARS-CoV-2. *medRxiv*. https://www.medrxiv.org/content/10.1101/2021.09.02.21261735v1





	incidence rate in booster compared to non-booster ³⁵¹ Severe illness in individuals <60: 0.33 (95% CI, 0.21-0.52) incidence rate in booster compared to non-booster ³⁵¹ Severe illness in individuals ≥60: 0.12 (95% CI, 0.10-0.14) incidence rate in booster compared to non-booster ³⁵¹							
	_		HETER	OLOGOUS BOOSTE	R DOSES			
Vaccine Schedule	Heterologous 1: mRNA1273/BNT1 62b2 Heterologous 2: Ad26.CoV.2.S/BN T162b2	Heterologous 1: BNT162b2/mRNA 1273 Heterologous 2: Ad26.CoV.2.S/m RNA1273	Heterologous 1: BNT162b2/ChAd Ox1*	Heterologous 1: BNT162b2/Ad26. CoV.2.S Heterologous 2: mRNA1273/Ad26. CoV.2.S	Heterologous: SinoPharm/BNT1 62b2	Heterologous 1: CoronaVac/ChAd Ox1 Heterologous 2: CoronaVac/BNT1 62b2	No available data	Heterologous 1: BNT162b2/NVX -CoV2373 Heterologous 2: ChAdOx1/NVX-CoV2373
	Heterologous 3: ChAdOx1/BNT16 2b2 *Received BNT162b2 as booster dose	Heterologous 3: ChAdOx1/mRNA 1273 *Received mRNA1273 as booster dose	*Received ChAdOx1 as booster dose	Heterologous 3: ChAdOx1/Ad26.C oV.2.S. *Received Ad26.CoV.2 as booster dose		Heterologous 3: CoronaVac/Sino Pharm *Received CoronaVac as initial regimen		*Received NVX- CoV2373 as booster dose























Time-to-booster dose	At least 3 months after receiving two dose regimen	At least 3 months after receiving two dose regimen	6 months after initial two-dose regimen	4 months after initial two-dose BNT162b2 regimen ³⁵² At least 3 months after receiving two dose regimen	6 months after initial two-dose regimen	Heterologous 1: 21 to 26 days after full jab of CoronaVac Heterologous 2: 6 months after primary vaccination of CoronaVac Heterologous 3: 6 months after primary vaccination of CoronaVac	No available data	6 months after initial two-dose regimen
Effectiveness	Heterologous 1: 94% (95% CI, 91- 96) effectiveness against infection³40 Heterologous 2 - Effectiveness in ≥50: 87.4% (95% CI, 84.9-89.4) against symptomatic COVID-19³41 93.1% (95% CI, 91.7-94.3) against symptomatic COVID-19 compared to unvaccinated³41	Heterologous 1: 92% (95% CI, 88- 95) effectiveness against infection ³⁴⁰ Heterologous 3: 91% (95% CI, 63- 98) effectiveness against infection ³⁴⁰						



















	Heterologous 3: 82% (95% CI, 68- 90) effectiveness against infection ³⁴⁰		<u>Omicron</u> (B.1.1.529):					
Effectiveness against Variants			Heterologous 1: 71.4% (95% CI, 41.8-86.0) against symptomatic					
	Binding Antibody Responses: 2-fold or greater rise in bAb noted in 98-100% of BNT162b2 recipients ³⁵³	Binding Antibody Responses: 2-fold or greater rise in bAb noted in 96-100% of mRNA1273 recipients ³⁵³	infection ⁶⁶ Heterologous 1: Anti-spike IgG: In individuals <70: 12440 ELU/mL (95% CI, 10420- 14852)	Heterologous 1: 14.8 to 32.4-fold increase in neutralization titers against wild-type virus ³⁵²		Heterologous 1: Heterologous vaccination had a 9-fold greater GMT (7,947 U/mL) than fully		Heterologous 1: Anti-spike IgG: In individuals <70: 14961 ELU/mL (95% CI, 12065- 18551)
Immunogenicity	Neutralizing Antibody Responses: 341.3-677.9 IU50/mL 15 days after booster with BNT162b2 ³⁵³	Neutralizing Antibody Responses: 676.1-901.8 IU50/mL 15 days after booster with mRNA1273 ³⁵³	In individuals ≥70: 14961 ELU/mL (95% CI, 12065- 18551) ³⁵⁴ Cellular Response: In individuals <70: 105 (95% CI, 67-	Binding Antibody Responses (bAb): 2-fold or greater rise in bAb noted in 98-100% of Ad26.COV2.S. recipients ³⁵³	No available data	patients fully vaccinated with AZD1222 and the highest antibody response, IgA, and neutralizing antibodies than other groups ³⁵⁵	No available data	In individuals ≥70: 9130 EUL/mL (95% CI, 6783- 12289) ³⁵⁴ <u>Cellular</u> <u>Response:</u>
	Participants who received mRNA-based booster vaccination had four-fold increase	Participants who received mRNA-based booster vaccination had four-fold increase	164) In individuals ≥70: 84 (95% CI, 45- 156) ³⁵⁴	Neutralizing Antibody Responses: 31.2-382.2 IU50/mL 15 days		Anti-RBD Antibody: 9865 U/mL 14- days after booster ³⁵⁶		In individuals <70: 69 (95% CI, 45-156) In individuals ≥70: 45 (95% CI, 22-92) ³⁵⁴





















compared to Ad26.COV2.S. ³⁵³	compared to Ad26.COV2.S. 353	after booster with Ad26.COV2.S. ³⁵³	Heterologous 2:	<u>Heterologous</u>
Heterologous 3:	Heterologous 1:	Anti-spike IgG:	Median values of IgG-S titers were	<u> 2:</u>
ricterologous s.	ricterologous 1.	In individuals >70:	higher in group	Anti-spike IgG:
Anti-spike IgG:	Anti-spike IgG:	17312 ELU/mL	that received	In individuals
In individuals <70:	In individuals <70:	(95% CI, 13678-	BNT162b2 as	<70: 8389
22479 ELU/mL	44547 ELU/mL	21911)	booster than	ELU/mL (95%
(95% CI, 18276-	(95% CI, 38424-	<mark>In individuals ≥70:</mark>	CoronaVac	CI, 6599-10665)
<mark>27648)</mark>	<mark>51645)</mark>	16855 ELU/mL	BNT162b2	In individuals
Individuals ≥70:	In individuals ≥70:	(95% CI, 13360-	boosted IgG-S	<mark>≥70: 5822</mark>
19091 EUL/mL	25118 ELU/mL	<mark>21264)³⁵⁴</mark>	median titers by	ELU/mL (95%
(95% CI, 15554-	<mark>(95% CI, 17698-</mark>		factor of 46.6 but	<mark>CI, 4495-</mark>
23432) ³⁵⁴	35650) ³⁵⁴	<u>Cellular</u>	IgG-N titers	<mark>7541)³⁵⁴</mark>
- · · ·	A	Response:	decreased by	
<u>Cellular</u>	<u>Cellular</u>	In individuals <70:	factor of 6.5 ³⁵⁷	<u>Cellular</u>
Response:	Response:	114 (95% CI, 55-	And and a DDD	Response:
In individuals <70:		<mark>236)</mark> In individuals ≥70:	Anti-spike RBD:	In individuals
119 (95% CI, 83-	<mark>143 (95% CI, 82-</mark> 250)	109 (95% CI, 64-	Single booster dose of	<70: 137 (95% CL 88 243)
169) sport forming cells per 106	In individuals ≥70:	187) ³⁵⁴	BNT162b2	CI, 88-213) In individuals
peripheral blood	88 (95% CI, 46-	107)**	induced higher	≥70: 55 (95%
mononuclear cells	168)	Heterologous 3 :	anti-spike RBD	CI, 35-89) ³⁵⁴
In individuals ≥70:	,	<u> </u>	IgG antibody	C1, CC CC)
113 (95% CI, 64-		Anti-spike IgG:	levels, compared	
200) sport forming	Heterologous 3:	In individuals <70:	to single booster	
cells per 10 ⁶		5582 ELU/mL	dose of	
peripheral blood	Anti-spike IgG:	(95% CI, 4415-	CoronaVac ⁸⁷	
mononuclear	In individuals <70:	<mark>7057)</mark>		
cells ³⁵⁴	35522 ELU/mL	<mark>In individuals ≥70:</mark>	<mark>20,787 U/mL</mark> 14	
	<mark>(95% CI, 29205-</mark>	5464 ELU/mL	<mark>days after</mark>	
	43204)	(95% CI, 4266-	booster ³⁵⁶	
	In individuals ≥70:	<mark>6998)</mark>		
	27702 ELU/mL	0.11.1.	<u>Heterologous 3:</u>	
	(95% CI, 21337-	<u>Cellular</u>	Anti onika DBD.	
	35966) ³⁵⁴	Response: In individuals <70:	<u>Anti-spike RBD:</u>	
		in mulviduals 0.</td <td></td> <td></td>		























		Cellular Response: In individuals <70: 228 (95% CI, 177- 294) In individuals ≥70: 101 (95% CI, 54- 187) ³⁵⁴		141 (95% CI, 100-200) In individuals ≥70: 82 (95% CI, 54-124)		1073 U/mL 14 days after booster ³⁵⁶		
Immunogenicity against variants	Binding Antibody Responses: Baseline bAb levels for Delta were 34-45% lower compared to Wa-1 strain ³⁵³ Following boost, bAB levels for Delta were 15- 36% lower compared to Wa-1 strain ³⁵³ Heterologous 1: Neutralizing Ab: 22.7-fold decrease in neutralization after 0.5 months after booster compared to Delta ³⁴⁶	Binding Antibody Responses: Baseline bAb levels for Delta were 34-45% lower compared to Wa-1 strain ³⁵³ Following boost, bAB levels for Delta were 15- 36% lower compared to Wa-1 strain ³⁵³ Neutralizing Antibody Responses: Delta and Beta variants were only available in those boosted with mRNA-1273 ³⁵³ Heterologous 1:	Pseudovirus neutralizing antibody NT ₅₀ : 260 GMT (95% CI, 217-313) against Delta ³⁵⁴	Heterologous 1: 10.9 to 21.2-fold increase in pseudo virus neutralization assay (one volunteer did not have any against B.1.351) 352 Binding Antibody Responses: Baseline bAb levels for Delta were 34-45% lower compared to Wa-1 strain 353 Following boost, bAB levels for Delta were 15-36% lower compared to Wa-1 strain 353	No available data	Heterologous 1: Neutralizing activity against the wild type and variant strains showed higher neutralizing activity in the following order: wild type > B.1.617.2 > B.1.1.7 > B.1.351 ³⁵⁵	No available data	Heterologous 1: Pseudotype neutralizing antibody NT50: 165 GMT (95% CI, 131-209) against Delta ³⁵⁴ Heterologous 2: Pseudotype neutralizing antibody NT50: 124 GMT (95% CI, 99-156) against Delta ³⁵⁴





















	Pseudotype virus neutralizing antibody NT ₅₀ : 315 GMT (95% CI, 1314–1998) against Delta ³⁵⁴	Pseudotype virus neutralizing antibody NT ₅₀ : 508.7 GMT (95% CI, 408.6-633.4) against Delta ³⁵⁴ Heterologous 3: Pseudotype virus neutralizing antibody NT ₅₀ : 559.7 GMT (95% CI, 441.3-709.9) against Delta ³⁵⁴		Pseudotype virus neutralizing antibody NT ₅₀ : 418 GMT (95% CI, 330-530) against Delta ³⁵⁴ Heterologous 3: Pseudotype virus antibody NT ₅₀ : 125 GMT (95% CI, 99-159) against Delta ³⁵⁴				
	Adverse Events: 72-92% participants reported local pain or tenderness ³⁵³	Adverse Events: 75-86% participants reported local pain or tenderness ³⁵³		Adverse Events: 71-84% participants reported local pain or tenderness ³⁵³				
Reactogenicity	Malaise, myalgias, and headaches were commonly reported ³⁵³	Malaise, myalgias, and headaches were commonly reported ³⁵³	No available data	Malaise, myalgias, and headaches were commonly reported ³⁵³	No available data	Similar results to homologous booster administration	No available data	No available data
	14.4% of the participants reported unsolicited adverse events ³⁵³	15.6% of participants reported unsolicited adverse events ³⁵³		12% of participants reported unsolicited adverse events ³⁵³				





















Other		Ongoing clinical trial examining immunogenicity and safety of third dose vaccination with ChAdOx1 or BNT162b2 vaccine among adults who received full jab of CoronaVacinical received full page full received full jab of CoronaVacinical received full received full page full received full page full received full page full received full received full page full received full rece	

Lixx Third Dose Vaccination with AstraZeneca or Pfizer COVID-19 Vaccine Among Adults Received Sinovac COVID-19 Vaccine. ClinicalTrials.gov. https://clinicaltrials.gov/ct2/show/NCT05049226



ANNEXES

	BNT162b2/ COMIRNATY (Pfizer- BioNTech, USA)	Spikevax/ Moderna COVID- 19 Vaccine/ mRNA-1273 (Moderna, USA)	Vaxzevria/ ChAdOx1 nCoV- 19/ AZD1222/ Covishield (AstraZeneca/Ox ford, UK, India)	Janssen COVID- 19 vaccine/Johnson & Johnson (Janssen, USA)	Sinopharm/BBIB P-CorV, China	Sinovac CoronaVac, China	COVAXIN/ BBV152 (Bharat Biotech, India)	Novavax/ NVX- CoV2373
				FURTHER INFORM	IATION			
Storage conditions	2°C to 8 °C (for 1 month)	2°C to 8 °C (for 1 month)	2°C until 8 °C	2°C to 8 °C (for 3 months)	2°C until 8 °C	2°C until 8 °C	2°C until 8 °C	2°C to 8 °C
Approving authorities	FDA (11.12.20) ^{lxxi} ; EMA (21.12.20); WHO EUL (31.12.20); and list of countries (including Switzerland – approved on 20.12.20)	FDA (18.12.20); EMA (06.01.21); WHO EUL (30.04.21); and list of 51 countries (including Switzerland – approved 12.01.21)	FDA (awaiting on approval); EMA (29.01.21); WHO EUL (15.02.21); and list of 121 countries (Switzerland awaiting on approval)	FDA (27.02.21); EMA (11.03.21), WHO EUL (12.03.21), and list of 59 countries (including Switzerland – approved 22.03.21)	WHO EUL (07.05.21); and list of 55 countries (e.g., Argentina, Bahrain, Brazil, China, Indonesia, United Arab Emirates, Zimbabwe)	WHO EUL (01.06.21), and list of 33 countries (e.g., Albania, Chile, Egypt, Hong Kong, Malaysia, Tunisia, Turkey, Ukraine)	WHO EUL (03.11.21) and list of 9 countries (Guyana, Inidia, Iran, Mauritius, Mexico, Nepal, Paraguay, Philippines & Zimbabwe)	Waiting on approval (Not-yet-approved by countries or WHO for emergency use)
				IMMUNOGENICI	TY			

Pfizer-BioNTech's Comirnaty Vaccine received full FDA approval on 23 August 2021 for people age 16 and above, moving it beyond emergency use status. https://www.fda.gov/news-events/press-announcements/fda-approves-first-covid-19-vaccine





Immunogenicity	7-14 days after second dose: 18-55 years: GMT ranged from 1.7 to 4.6 times the GMT of the convalescent serum ³⁵⁸ . 65-85 years: GMT ranged from 1.1 to 2.2 times the GMT of the convalescent serum ³⁵⁸ .	14 days after second dose: 18-55 years: PRNT ₈₀ GMT 654.3 (95% CI, 460.1-930.5) ³⁵⁹ . 56-70 years: PRNT ₈₀ GMT 878 (95% CI, 516-1494) ³⁶⁰ . ≥71 years: PRNT ₈₀ GMT 317 (95% CI, 181-557) ³⁶⁰ .	28 days after second dose median antibody titres: 18–55 years: 20,713 AU/mL [IQR 13,898 - 33,550] ³⁶¹ 56–69 years: 16,170 AU/mL [IQR 10,233 - 40,353] ³⁶¹ . ≥70 years: 17,561 AU/mL [IQR 9,705 - 37,796] ³⁶¹ .	29 days after vaccination: 18-55 years: GMC 586 (95% CI, 445-771); GMT 224 (95% CI, 168-298)³6². ≥65 years: GMC 312 (95% CI, 246-396); GMT 212 (95% CI, 163-266)³6². 57 days after vaccination: 18-55 years: 754 (95% CI, 592-961); GMT 288 (95% CI, 221-376)³6².	14 days after second dose: 18-55 years: GMT 211.2 (95% CI, 158.9-280.6) ³⁶³ . ≥60 years: GMT 131.5 (95% CI, 108.2-159.7) ³⁶³ .	Single dose (≥4 weeks): 37.7±57.08 IU/mI (min: 0, max: 317.25); 57.02% of participants did not develop sufficient antibody titres (<25.6 IU mI) Two doses (≥4 weeks): 194.61±174.88 IU/mI (min: 0, max: 677.82); 11.48% of participants did not develop sufficient antibody titres (<25.6 IU mI) 364. 2 weeks after second dose: 164.4 BAU/ mL365 4 weeks after second dose: 94.8 BAU/ mL365 8-12 weeks after second dose: 34.7 BAU/ mL365	Single dose (≥4 weeks: 43.8% seropositive for anti-spike antibody > 15 AU/mL³66 GMT 16.8 (95% CI, 15.80-17.88) for SARS-CoV-2 spike antibody titre³66 Two doses (≥4 weeks): 80.0% seropositive for anti-spike antibody > 15 AU/mL³66 GMT 48.3 (95% CI, 47.46-48.92) for SARS-CoV-2 spike antibody titre³66	
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Immunogenicity against the Mu variant	6.8-fold decrease in neutralizing titres when compared to convalescent sera ³⁶⁷	Neutralizing titre similar to that of BNT162b2 sera ³⁶⁷	Neutralizing titre similar to that of BNT162b2 sera ³⁶⁷	No available data	No available data	No available data	No available data	No available data
Immunogenicity against Omicron variant	29.8-fold decrease in mean neutralizing titres compared to wild- type, 10.3-fold decrease compared to Beta, 25.1-fold decrease compared to Delta ³⁴⁶ 11.4-fold decrease in neutralization 6 months after vaccination compared to Delta 25-fold decrease in neutralization titers against Omicron variant compared to wild- type ³⁶⁸ 41-fold decrease in neutralization level against Omicron ³⁶⁹	20-fold decrease in neutralization 6 months after vaccination compared to Delta ³⁴⁶ 1/10 seropositive against Omicron ³³²	Mean neutralizing titres drop to below the detectable threshold in all but one participant ³⁴⁶ 0/20 seropositive against Omicron ³³²					























								PUBLIC HEAL
	9/20 seropositive against Omicron 332							
				EFFICACY				
Single dose ^{lxxii}	52% (95% CI, 29.5 to 68.4; starting at 12 days) or 82.2% (75.1 to 87.3; starting at ≥14 days) ³⁷⁰ . 91% (95% CI, 85-94) ³⁷¹ . ≥80 years: 71.4% (95% CI, 46.5-90.6) vaccine efficacy for symptomatic disease 14 days after one dose [United Kingdom, 18 Dec 2020 – 26 Feb 2021] ³⁷² ≥65 years: 56% (95% CI 19-76) at 28-34 days and 62% (95% CI	95.2% (95% CI, 91.2.8 to 97.4; starting at >14 days) ¹²¹ .	72.8% (starting at 22 days up to 60 days) ³⁷³ . 88% (95% CI, 75-94) ³⁷¹ . Ixxiv ≥80 years: 80.4% (95% CI, 36.4-94.5) vaccine efficacy for symptomatic disease 14 days after one dose [United Kingdom, 18 Dec 2020 − 26 Feb 2021 ³⁷² ≥65 years: 56% (95% CI 19-76) at 28-34 days and 62% (95% CI 23-81) at 35-48 days postvaccination [United Kingdom,	Single dose vaccine	Unknown	35.1% (95% CI, -6.6 to -60.5) [conducted in a setting with high P.1 transmission] ³⁷⁴ .	No available data	83.4% (95% CI, 73.6-89.5) starting at ≥14 days ⁴²

lxxii Against SARS-COV-2 infection

bxxiv Conducted between 8 December 2020 and 8 February 2021. Study sample = ≤1 million participants.



	23-81) at 35-48 days post- vaccination [United Kingdom, 8 Dec 2020 – 15 Mar 2021] ^{372 lxxiii}		8 Dec 2020 – 15 Mar 2021] ^{372 lxxv}					
Two doses ^{lxxvi}	95.0% (95% CI, 90.3-97.6) starting at ≥7 days in population without prior SARS-CoV-2 infection ¹³⁶ 94.6% (95% CI, 89.9-97.3) starting at ≥7 days in population with or without prior infection ¹³⁶	94.1% (95% CI, 89.3-96.8) after median follow-up of less than 63 days ¹²¹ 93.2% (95% CI, 91.0-94.8) ³⁷⁵ Against severe disease: 98.2% (95% CI, 92.8-99.6) ³⁷⁵ Prevention against COVID-19 illness: 93.2% (95% CI, 91.0-94.8; United States) ³⁷⁵ Prevention against severe disease:	63.1% (95% CI, 51.8-71.7) starting at ≥14 days for two standard doses ³⁷³ 80.7% (95% CI, 62.1-90.2) starting at ≥14 days for first low dose and standard second dose ³⁷³ 66.7% (95% CI, 57.4-74.0) starting at ≥14 days for pooled analysis efficacy ³⁷³ Against mild-to-moderate symptomatic COVID-19 >14 days after second injection:	66.9% (95% CI 59.0-73.4) after 14 days and 66.1% (95% CI 55.0-89.1) after 28 days for VE against moderate- severe-critical COVID-19 ³⁷⁷ 76.7% (95% CI 54.6 to 89.1) after 14 days and 85.4% (95% CI 54.2 to 96.9) after 28 days for VE against severe- critical COVID- 19 ³⁷⁷	After 14 days, efficacy against symptomatic cases was 72.8% (95% CI 58.1-82.4; in WIV04 vaccine) or 78.1% (95% CI 64.8 to 86.3; in HBO2 vaccine). ²³⁶	After 14 days, efficacy against symptomatic cases was 50.7% (95% CI 35.9 to 0-62.0). ¹²⁷ 99.17% of NAb titres were above or equal to the Nab positivity cutoff (20 units) against wild-type ³⁷⁸ .	Symptomatic SARS-CoV-2 infection: 77.8% (95% CI, 65.2-86.4) ³⁷⁹ Severe symptomatic SARS-CoV-2 infection: 93.4 (95% CI, 57.1-99.8) ³⁷⁹ Symptomatic COVID-19 in ≥60 years old: 67.8% (95% CI, 65.2-86.4) against symptomatic COVID-19 ³⁷⁹ Symptomatic COVID-19 in 18- 59 years old: 79.4% (95% CI, 66.0-88.2) against	89.7% (95% CI, 80.2-94.6) starting at ≥7 days ⁴² 90.4% (95% CI, 82.9-94.6) ³⁸⁰ 100% (95% CI, 87-100) against moderate-to-severe COVID-19 ³⁸⁰ 100% (95% CI, 34.6-100) against severe COVID-19 ³⁸⁰ 90% (95% CI, 80-95) (≥7 days after second dose) ³⁸¹

lxxiii Does not differentiate between BNT162b2 and ChAdOx1 nCoV-19.

lxxv Does not differentiate between BNT162b2 and ChAdOx1 nCoV-19.

lxxvi Against SARS-CoV-2 infection.



		98.2% (95% CI, 92.8-99.6; United States) 375 Prevention against asymptomatic infection starting 14 days after second infection: 63.0% (95% CI, 56.6-68.5; United States) 375	21.9% (95% CI, - 49.9 to 59.8; South Africa) [24 June – 09 November 2020] ³⁷⁶				symptomatic COVID-19 ³⁷⁹	
Against asymptomatic infection	90% (starting at 14 days) regardless of symptom status ³⁸²	63.0% (95% CI, 56.6-68.5) ³⁷⁵	Statistically non-significant reduction of 22.2% (95% CI - 9.9 to 45.0) for asymptomatic cases 61.9% efficacy ³⁹	At day 71, vaccine efficacy against asymptomatic infections was 65.5% (95% CI 39.9 to 81.1) ³⁷⁷ .	Efficacy against symptomatic and asymptomatic cases was 64% (95% CI 48.8 to 74.7; in WIV04 vaccine) or 73.5% (95% CI 60.6 to 82.2; in HBO2 vaccine) ²³⁶ .	Unknown	63.6 (95% CI, 29.0-82.4) efficacy against asymptomatic cases ³⁷⁹	Unknown
			EI	FICACY AGAINST	VARIANTS			
Alpha (B.1.1.7)	Two doses of the vaccine effectively neutralize the B.1.1.7 variant and the D614G substitution ³⁸³ .	NAbs remained high and consistent with titres of the wildtype for the B.1.1.7 variant ³⁸⁴ .	70.4% (95% CI, 43.6-84.5) against symptomatic infection with alpha variant (B.1.1.7); 28.9% (95% CI, -77.1 to 71.4) against asymptomatic	3.6-fold reduction in neutralization capacity when compared to wild-type.	Demonstrated reduced neutralizing capacity. However, there were no differences in the NAbs titres against B.1.351 in vaccinated	reduction in neutralization capacity when compared to natural infection sera ³⁷⁸ . 85.83% of NAb titres were above	PRNT ₅₀ 0.8 when compared with wild type against Alpha (no significant difference in neutralization capacity) ³⁸⁷	Two dose efficacy against the B.1.1.7 variant 86.3% (95% CI, 71.3-93.5) ⁴² 93.6% (95% CI, 81.7-97.8)



















			infection with B.1.1.7 ¹⁰⁹ .		individuals vs. those naturally infected, suggesting the vaccine has a similar level of protection against infection as natural infections ³⁸⁵ .	or equal to the Nab positivity cutoff (20 units) against wild-type ³⁷⁸ . Neutralization decreased by 4.1-fold when compared to wild-type ³⁸⁶ .		against the Alpha variant ³⁸⁰ Against non-B.1.1.7 variant 96% (95% CI, 74-99.5) (≥7 days after second dose) ³⁸¹ Against B.1.1.7 variant 86% (95% CI, 71-94) (≥7 days after second dose) ³⁸¹
Beta (B.1.351)	Neutralization was diminished by a factor of 5. Despite this, the BNT162b2 mRNA vaccine still provides some protection against B.1.351 ³⁸⁸ 100% (95% CI, 53.5-100) ³⁸⁹ .	NAbs were 6-fold lower. Nevertheless, NAbs were still found to be protective ³⁸⁴ .	Two doses of the vaccine had no efficacy against the B.1.351 (VE = 21.9%; 95% CI, -49.9 to 59.8) ³⁷⁶ . Against mild-to-moderate symptomatic COVID-19 associated with B.1.351 variant >14 days after second injection: 10.4% (95% -76.8 to 54.8; South Africa) [24 June – 09 November 2020] ³⁷⁶	Efficacy against moderate-severe-critical Covid-19 due to the variant was 52.0% (>14 days) and 64.0% (>28 days). Efficacy against severe-critical COVID-19 was 73.1% (>14 days) and 81.7% (>28 days) ³⁷⁷ . Demonstrated 3.6-fold reduction in neutralization sensitivity ³⁹⁰ . Neutralization titres were	No published data	NT _{GM} 35.03 (95% CI , 27.46-44.68); 8.75-fold reduction in neutralization capacity when compared to natural infection sera ³⁷⁸ . 82.5% of NAb titres were above or equal to the Nab positivity cutoff (20 units) against wild-type ³⁷⁸ .	GMT 61.57 (95% CI, 36.34-104.3) against Beta variant with significant reduction in neutralization titre ³⁹²	51.0% (95% CI, -0.6-76.2) efficacy against B.1.351 variant ³⁹³























								PUBLIC
				decreased by 6.7- fold ³⁹¹ .				
Gamma (P.1)	Single dose: ≥21 days: 83% against hospitalization and death ³⁹⁴ . Two doses: ≥14 days: 98% against hospitalization and death ³⁹⁴ .	3.2-fold reduction in neutralization capacity when compared to wild-type ³⁹⁵ .	Single dose: ≥21 days: 94% against hospitalization and death ³⁹⁴ . Two doses: 64% (95% CI, -2-87) [n=18] ³⁹⁶ Efficacy against Zeta (P.2) [2 doses]: 69% (95% CI, 55-78) ³⁹⁶	Demonstrated 3.4-fold reduction in neutralization sensitivity ³⁹⁰ .	No published data	49.6% against P.1 (>14 days after 1st dose) ³⁷⁴ . Neutralization decreased by 7.5-fold when compared to wild-type ³⁸⁶ .	No available data	No available data
Delta (B. 1.671.2)	Reduced NAb activity relative to B.1.1.7 strain ³⁹⁷ .	2.1-fold reduction in neutralization capacity when compared to wild-type ³⁹⁵ .	Single dose: ≥21 days: 90% against hospitalization and death ³⁹⁴ .	Demonstrated 1.6-fold reduction in neutralization sensitivity ³⁹⁰ . Neutralization titres were decreased by 5.4- fold ³⁹¹ .	Demonstrated reduced neutralizing capacity. However, there were no differences in the NAbs titres against B.1.617.2 in vaccinated individuals vs. those naturally infected, suggesting the vaccine has a similar level of protection against infection as	NT _{GM} 24.48 (95% CI,19.2-31.2) ³⁷⁸ . 69.17% of NAb titres were above or equal to the Nab positivity cutoff (20 units) against wild-type ³⁷⁸ .	65.2 (95% CI, 33.1-83.0) estimated efficacy ¹³⁰ GMT 68.97 (95% CI, 24.72-192.4) against Delta variant with significant reduction in neutralization titre ³⁹²	No available data



















Omicron (B.1.1.529)	22.5% (95% CI, 8.5-40.7) against symptomatic infection ³⁶⁹				natural infections ³⁸⁵ .			
			PH	ASE III TRIALS RES	SULTSIxxvii			
Number of participants (vaccine/placebo)	43,448 (21,720/ 21,728) ¹³⁶	30,420 (15,210/15,210) ¹²¹	17,178 (8597/8581) ³⁷³	39,321 (19,630/19,691) ³⁷⁷	26,917 (13,459/13458); or 26,914 (13,465/13,458) ²³⁶	9,823 (4,953/4,870) ¹²⁷	25,798 (12,899/12899) ¹³⁰	14,039 (7,020/7,019) ⁴²
Total COVID- 19 cases (vaccine/ control)	170(8/162) ¹³⁶	196 (11/185) ¹²¹	332 (84/248) ³⁷³	464 (116/348) ³⁷⁷	121(26/95) or 116(21/95) ²³⁶	253(85/168) ¹²⁷	130 (24/106) ¹³⁰	106(10/96)42
Efficacy estimates in Phase III trials	Starting from 7 days after 2nd dose: 95.0% (95% CI, 90.3 to 97.6) in population without prior SARS-CoV-2 infection. Efficacy of 94.6% (95% CI, 89.9 to 97.3) in	After a median follow-up of less than 63 days: Efficacy of 94.1% (95% CI, 89.3 to 96.8; P<0.001). 100% among adolescents (12 to <18 years old) ¹²¹ .	Two standard doses: efficacy was 63-1% (95% CI 51.8 to 71.7; ≥14 days) while those with first low dose and standard 2nd dose the efficacy	VE against moderate-severe- critical Covid-19 was 66.9% (95% CI 59.0 to 73.4) after 14 days post vaccine administration, and 66.1% (95%	After 14 days, efficacy against symptomatic cases was 72.8% (95% CI 58.1 to 82.4; in WIV04 vaccine) or 78.1% (95% CI 64.8 to	After 14 days, efficacy against symptomatic cases was 50.7% (95% CI 35.9 to 0-62.0). ¹²⁷	77.8% (95% CI, 65.2-86.4) against symptomatic COVID-19 starting at ≥14 days after second dose ¹³⁰	83.4% (95% CI, 73.6-89.5) starting at ≥14 days after first dose ⁴² 89.7% (95% CI, 80.2-94.6) starting at ≥7

Phase III trials were conducted between 27 July and 14 November 2020 for BNT162b2/ COMIRNATY, 27 July and 23 October 2020 for Spikevax/ Moderna, 23 April and 6 December 2020 for Vaxzevria/ ChAdOx1 nCoV-19/ AZD1222/ Covishield, 21 September 2020 and 22 January 2021 for Janssen Covid-19 vaccine/ Johnson & Johnson, 16 July and 20 December 2020 for Sinopharm/ BBIB-CorV, 21 July and 16 December 2020 for the Sinovac/ CoronaVac vaccine, 16 November 2020 and 7 January 2021 for the COVAXIN vaccine, and 28 September 2020 and 28 November 2020 for the Novavax vaccine. All trials were conducted prior to the transmission of the more contagious variant strains, particularly the delta variant (B.1.617.2). Studies are currently ongoing to determine the effectiveness of the vaccines against the delta variant.



	population with or without prior infection. 100% among adolescents (12-15 years old) ¹³⁶ .		was 80.7% (95% CI 62.1 to 90.2). Pooled analysis efficacy was 66.7% (95% CI 57.4 to 74.0). For any nucleic acid amplification test-positive swab: efficacy was 54.1% (95% CI 44.7 to 61.9) ³⁷³ .	CI 55.0 to 89.1) after 28 days. VE against severe-critical COVID-19 cases was 76.7% (95% CI 54.6 to 89.1) after 14 days and 85.4% (95% CI 54.2 to 96.9) after 28 days ³⁷⁷ . SII-ChAdOx1 nCoV-19 has a non-inferior immune response compared to AZD1222 and an acceptable safety/reactogenicity profile ³⁹⁸	86.3; in HBO2 vaccine) ²³⁶ .			days after second dose ⁴²
Efficacy against hospitalization and death	100% (after 7 days) ¹³⁶	100% (≥14 days) ¹²¹	100% (after 21 days) ³⁷³	76.7% (≥14 days) or 85.4% (≥28 days) ³⁷⁷	100% (>14 days) ²³⁶	100% (>14 days) ¹²⁷	93.4% (>14 days) against severe COVID-19 ¹³⁰	100% (after 7 days) ⁴² .
Phase III clinical trial serious adverse events	Serious adverse events were observed in a similar proportion of vaccine (0.6%) and placebo (0.5%) recipients. These events also occur at a similar frequency within the general population 117,399.	The frequency of grade 3 adverse events was similar in both the vaccine (1.5%) and placebo groups (1.3). Serious adverse events were observed in a similar proportion in both groups	Serious adverse events were balanced across the study arms. 79 cases occurred in the vaccine group and 89 cases in the placebo group – 3 cases were considered related to the experimental or	Serious adverse events were reported in 0.4% of vaccine recipients and 0.4% of placebo recipients. Seven of the serious adverse events were considered to be related to the vaccine:	A cross-sectional survey collected data on adverse events following vaccination in the UAE - none of the symptoms were of serious nature or required hospitalization ¹²⁶ .	Overall incidence of serious adverse events was 0.5% (31 in placebo group and 33 in vaccine group). All adverse events were determined to be unrelated to the vaccine ¹²⁷ .	Rates of local and systemic AEs reported in the BBV152 group as mild (11·2%), moderate (0·8%), or severe (0·3%) were comparable to the placebo group ¹³⁰	Phase II: Nine serious adverse events were reported, only one of which was assessed as related to the vaccine: acute colitis ⁴⁰⁰ .





















		(0.6%). 3 Bell's Palsy cases occurred in the vaccine group and one Bell's Palsy case occurred in the placebo group ¹²¹ .	control vaccine (out of 11 636 vaccine recipients): transverse myelitis, haemolytic anaemia and a case of fever higher than 40°C	Guillain-Barré syndrome (1), pericarditis (1), brachial radiculitis (1), hypersensitivity (1), Bell's Palsy (2), & severe generalized weakness, fever & headache (1) ³⁷⁷ .			15 deaths, none considered related to the vaccine or placebo ¹³⁰	
				PHASE III TF	RIAL OTHER			
Comments	Specific populations were excluded (HIV and immunocompromi sed patients, and pregnant women).	Calculation of efficacy were not based on the total number of confirmed Covid-19 cases.		2-DOSE EFFICACY Efficacy against symptomatic (moderate to severe/ critical) SARS-CoV-2 infection 94% (95% CI, 58- 100) in the US. 75% (95% CI, 55- 87) globally. ²⁰ Efficacy against severe/ critical SARS-CoV-2 infection 100% (95% CI, 33-100) ²⁰	Only 2 severe cases occurred in the control group and none in the vaccine group (very few cases to get a reliable estimate).	_	-	Novavax is currently awaiting FDA, EMA, and WHO EUL approval. Upcoming information regarding results of clinical trials or approval will be updated in upcoming reports





















		VACCINE PRODUCTION SITES								
	BNT162b2/ COMIRNATY (Pfizer- BioNTech, USA) ^{lxxviii}	Spikevax/ Moderna COVID- 19 Vaccine/ mRNA-1273 (Moderna, USA) ^{lxxix}	Vaxzevria/ ChAdOx1 nCoV- 19/ AZD1222/ Covishield (AstraZeneca/Oxf ord, UK, India) ^{lxxx}	Janssen COVID- 19 vaccine/Johnson & Johnson (Janssen, USA) ^{lxxxi}	Sinopharm/BBIB P-CorV, China ^{lxxxii}	Sinovac CoronaVac, China ^{lxxxiii}	COVAXIN / BBV152 (Bharat Biotech, India)	Novavax/ NVX- CoV2373		
EUL holder	BioNTech Manufacturing GmbH (Germany)	ModernaTX, Inc. (USA) ¹ Moderna Biotech (Spain) ²	AstraZeneca AB (Sweden)	Janssen-Cilag International NV (Belgium)	Beijing Institute of Biological Products Co., Ltd. (BIBP) (China)	Sinovac Life Sciences Co., Ltd. (China)	Bharat Biotech International Limited (India)	Novavax (USA)		
Production sites (Drug substance)	BioNTech Manufacturing GmbH (Mainz, Germany) BioNTech Manufacturing Marburg (Marburg, Germany)	Lonza Biologics, Inc., (USA) ¹ Moderna TX, Inc. (USA) ¹ Lonza AG (Switzerland) ²	Henogen S.A (Belgium) Catalent Maryland, Inc. (USA) Oxford Biomedica (UK) Ltd. (United Kingdom)	Janssen Vaccines & Prevention B.V. (The Netherlands) Janssen Biologics B.V. (The Netherlands) Emergent Manufacturing Operations Baltimore LLC	Beijing Institute of Biological Products Co., Ltd. (China)	Sinovac Life Sciences Co., Ltd. (China)	-	Novavax (Bohumil, Czech Republic)		

lxxviii WHO recommendation BioNTech Tozinameran - COVID-19 mRNA vaccine (nucleoside modified) - COMIRNATY. WHO. https://extranet.who.int/pqweb/vaccines/who-recommendation-covid-19-mrna-vaccine-nucleoside-modified-comirnaty

bxxiii WHO recommendation of Sinovac COVID-19 vaccine (Vero Cell [Inactivated]) - CoronaVac. WHO. https://extranet.who.int/pqweb/vaccines/who-recommendation-sinovac-covid-19-vaccine-vero-cell-inactivated-coronavac



^{1.} WHO recommendation ModernaTX, Inc/USFDA COVID-19 mRNA vaccine (nucleoside modified). WHO. https://extranet.who.int/pqweb/vaccines/who-recommendation-modernatx-incusfda-covid-19-mrna-vaccine-nucleoside-modified
2. WHO recommendation Moderna COVID-19 mRNA Vaccine (nucleoside modified). WHO. https://extranet.who.int/pqweb/vaccines/covid-19-mrna-vaccine-nucleoside-modified

WHO recommendation AstraZeneca/ EU approved sites COVID-19 vaccine (ChAdOx1-S) [recombinant]. WHO. https://extranet.who.int/pgweb/vaccines/covid-19-vaccine-chadox1-s-recombinant-0

kxxi WHO recommendation Janssen-Cilag International NV (Belgium) COVID-19 Vaccine (Ad26.COV2-S [recombinant]). WHO. https://extranet.who.int/pqweb/vaccines/who-recommendation-janssen-cilag-international-nv-belgium-covid-19-vaccine-ad26cov2-s

bxxii WHO recommendation COVID-19 vaccine BIBP/Sinopharm. WHO. https://extranet.who.int/pqweb/vaccines/who-recommendation-covid-19-vaccine-bibp



	Rentschler Biopharma SE (Laupheim, Germany) Wyeth BioPharma Division of Wyeth Pharmaceuticals (USA)		SK Bioscience (Republic of Korea) Halix B.V (Netherlands) WuXi Biologics (China)	(USA)			
Production sites (Drug product)	Baxter Oncology GmbH (Halle/ Westfallen, Germany) BioNTech Manufacturing GmbH (Mainz, Germany) Pfizer Manufacturing Belgium NV (Belgium) Novartis Pharma Stein AG (Switzerland) Mibe GmbH Arzneimittel (Brehna, Germany) Delpharm Saint- Remy (France)	Baxter Pharmaceutical Solutions, LLC. (USA) ¹ Catalent Indiana, LLC. (USA) ¹ Rovi Pharma Industrial Services, S.A. (Spain) ²	Catalent Anagni (Italy) CP Pharmaceuticals (United Kingdom) IDT Biologika (Germany) SK Bioscience (Republic of Korea) Universal Farma, S.L. ("Chemo") (Spain) Amylin Ohio LLC (USA)	Janssen Biologics B.V. (The Netherlands) Janssen Pharmaceutica NV (Belgium) Aspen SVP. (South Africa) Catalent Indiana LLC. (USA) Grand River Aseptic Manufacturing Inc. (USA) Catalent Anagni S.R.L. (Italy)	Beijing Institute of Biological Products Co., Ltd. (China)	Sinovac Life Sciences Co., Ltd. (China)	Novavax (Bohumil, Czech Republic)























	Sanofi-Aventis Deutschland GmbH (Germany)								
Diluent suppliers	Pfizer Perth, Australia Fresenius Kabi, USA	-	-	÷	-	-	-	-	





















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