



Unidad de Evidencia y Deliberación para la toma de decisiones UNED



COVID-19 Living Evidence <u>Synthesis # 8</u>

(Version 8.16: 13 Sep 2022)

Question

What is the effectiveness of available COVID-19 vaccines for children and adolescents, including variants of concern?

Findings

For vaccine effectiveness in variants of concern (VOC), we present a <u>visual summary</u> of evidence in Table 1 and Table 2.

Methods are presented in Box 1 and in the following appendices:

- 1) reference list
- 2) glossary
- 3) data-extraction template
- 4) process for assigning variant of concern to studies
- 5) research question and critical appraisal process
- 6) <u>detailed description of the narrative</u> summary statement.

Overall, 84 studies were appraised and 35 used to complete this summary. The <u>reasons for excluding</u> the remaining 49 studies are reported in the second section of Appendix 2.

One studies has been updated and three new studies had been added since the previous edition of this living evidence synthesis, which is signaled by a last updated date of 13 Sep 2022 (highlighted in yellow). The studies included results for VOC Delta (2) and VOC Omicron (4) – one reporting results by sublineage BA.2.

Studies examining effectiveness of vaccines in adults, including those covering periods beyond 120 days, are captured in COVID-END living evidence synthesis 6 and 10. The most recent version of all three syntheses (6,8,10) can always be found on the COVID-END website.

Box 1: Our approach

We retrieved candidate studies and updates to living evidence syntheses on vaccine effectiveness using the following mechanisms: 1) PubMed via COVID-19+ Evidence Alerts; 2) systematic scanning of pre-print servers; 3) updates to the COVID-END inventory of best evidence syntheses; and 4) crosscheck with updates from the VESPa team. We included studies and updates to living evidence syntheses identified up to two days before the version release date. We did not include press releases unless a preprint was available. A full list of included and excluded studies is provided in **Appendix 1**. A glossary is provided in **Appendix 2**.

Prioritized outcome measures: Infection, severe disease (as defined by the study investigators), death, and transmission.

Data extraction: We prioritized variant-confirmed and vaccine-specific data over total study population data (variant assumed and/or vaccine unspecified). We extracted data from each study in duplicate using the template provided in **Appendix 3**. Relevance to VOC is determined directly, when reported by study authors, or indirectly where reasonable assumptions can be made about the variant prevalent in the jurisdiction at the time of the study as described in **Appendix 4**.

Critical appraisal: We assessed risk of bias, direction of effect, and certainty of evidence. Risk of bias: assessed in duplicate for individual studies using an adapted version of ROBINS-I.

Direction of vaccine effect: "prevented" or "protects" was applied to mean estimates or range of mean estimates of effect that are greater than or equal to 70% (the lowest acceptable limit for vaccine effectiveness as determined by WHO). Certainty of evidence: assessed for the collection of studies for each vaccine according to variant of concern using a modified version of GRADE. Details of the research question for this synopsis and the critical appraisal process are provided in Appendix 5.

Summaries: We summarized the evidence by presenting narrative evidence profiles across studies, with or without pooling, as appropriate. A template for the summary statements used on page 1 under "Findings" and in Table 1 under each VOC is provided in **Appendix 6**.

We update this document Wednesday every two weeks and post it on the COVID-END website, but we are moving to every four weeks, with the next updates to be posted on 22 June, 20 July, 17 August, and 14 September

Highlights of changes this report

- New data on Pfizer [BNT162b2] against VOC Omicron has been added, with the data drawn from one study with a serious risk of bias (ref 35) and two studies with moderate risk of bias (ref 33, and ref 34). New data against VOC Omicron sub-lineage BA.2 has been added (ref 33).
- New data on Pfizer [BNT162b2] against VOC Delta has been added, with the data drawn from one study with a serious risk of bias (ref 35) and one study with moderate risk of bias (ref 34)
- New data on Sinovac [CoronaVac] against VOC Omicron has been added, with the data drawn from two studies with moderate risk of bias (ref 21 and ref 33). New data against VOC Omicron sub-lineage BA.2 has been added (ref 33).
- New data on Moderna [mRNA-1723] against VOC Delta has been added, with the data drawn from one study with a serious risk of bias (ref 35).
- New data on Moderna [mRNA-1723] against VOC Omicron has been added, with the data drawn from one study with a serious risk of bias (ref 35).

Pfizer/Comirnaty [BNT162b2]

• VOC Omicron

- We have low certainty evidence that <u>1 dose</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from infection from VOC **Omicron** (53.7% [95% CI, 43.3 to 62.2]- 1 Obs [<u>10</u>]) in adolescents age 12 to 17 years
- We have moderate certainty evidence that <u>1 dose</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from symptomatic infection from VOC **Omicron** (range of mean estimates: 25 to 53% 2 Obs [5][23]) in adolescents age 12 to 17 years
- We have low certainty evidence that <u>1 dose</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from severe disease from VOC **Omicron** (<u>56.3%</u> [95% CI, 45.9 to 64.6] 1 Obs [<u>23</u>]) in adolescents age 12 to 17 years
- We have low certainty evidence that 1 dose of BNT162b2 (Pfizer) did not reach threshold for protection from infection from VOC Omicron (range of mean estimates: 17 to 27% 2 Obs [25][27]) in children age 5 to 11 years
- We have low certainty evidence that <u>1 dose</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from infection from VOC **Omicron BA.2** (33.3% [95% CI, 3 to 53.3] 1 Obs [<u>29</u>]) in children age 3 to 11 years
- We have low certainty evidence that <u>1 dose</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from infection from VOC **Omicron BA.2** (26.1% [95% CI, -0.3 to 45.6] 1 Obs [<u>29</u>]) in adolescents age 12 to 18 years
- We have low certainty evidence that <u>1 dose</u> of BNT162b2 (Pfizer) did not reach threshold for protection from infection from VOC Omicron BA.2 (32.4% [95% CI, -29 to 64.6] 1 Obs [33]) in persons age 5 to 17 years
- We have low certainty evidence that <u>1 dose</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from symptomatic infection from VOC **Omicron** (range of mean estimates: 13 to 23% 2 Obs [<u>23</u>][<u>25</u>]) in children age 5 to 11 years
- We have low certainty evidence that <u>1 dose</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from severe disease from VOC **Omicron** (38.1% [95% CI, 20.9 to 51.5] 1 Obs [23]) in children age 5 to 11 years
- We have low certainty evidence that <u>2 doses</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from infection from VOC **Omicron** (range of mean estimates: 29 to 70% -4 Obs [25][27][28][35]) in children age 5 to 11 years

- We have low certainty evidence that <u>2 doses</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from severe disease from VOC **Omicron** (range of mean estimates: <u>41 to 94%</u> -2 Obs [27][30]) in children age 5 to 11 years
- We have low certainty evidence that <u>2 doses</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from symptomatic infection from VOC **Omicron** (range of mean estimates: 48 to 71% -4 Obs [<u>22</u>][<u>25</u>][<u>28</u>][<u>30</u>]) in children age 5 to 11 years
- We have low certainty evidence that <u>2 doses</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from infection from VOC **Omicron** (range of mean estimates: <u>34 to 83</u>% 4 Obs [11][13][26][35]) in adolescents age 12 to 17 years
- We have low certainty evidence that <u>2 doses</u> of **BNT162b2 (Pfizer)** did not reache threshold for protection against severe disease from VOC **Omicron** (75.6% [95% CI, 58.1 to 85.8] 1 Obs [<u>23</u>]), in adolescents age 12 to 17 years
- We have moderate certainty evidence that <u>2 doses</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from symptomatic infection from VOC **Omicron** (range of mean estimates: 55 to 83% 4 Obs [<u>5][22][23][26]</u>) in adolescents age 12 to 17 years
- We have low certainty evidence that <u>2 doses</u> of **BNT162b2 (Pfizer)** reached threshold for protection against MIS-C from VOC **Omicron** (92% [95% CI, 71 to 98] 1 Obs [7]), in adolescents age 12 to 18 years
- We have low certainty evidence that <u>2 doses</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from infection from VOC **Omicron BA.2** (54.9% [95% CI, 38.9 to 66.8] 1 Obs [<u>29</u>]) in adolescents age 12 to 18 years
- We have low certainty evidence that <u>3 doses</u> of BNT162b2 (Pfizer) did not reach threshold for protection from infection from VOC Omicron (range of mean estimates: <u>64 to 72</u>% 2 Obs <u>[26][35]</u>) in adolescents age 12 to 17 years
- We have moderate certainty evidence that <u>3 doses</u> of **BNT162b2 (Pfizer)** did not reach threshold for protection from symptomatic infection from VOC **Omicron** (range of mean estimates: 62 to 87% 4 Obs [8][16][22][32]) in adolescents age 12 to 17 years
- We have low certainty evidence that <u>3 doses</u> of **BNT162b2 (Pfizer)** reached threshold for protection against infection from VOC **Omicron BA.2** (86.8% [95% CI, 80.5 to 91.1] 1 Obs [<u>29]</u>) in adolescents age 12 to 18 years
- We have low certainty evidence that <u>2 doses</u> of BNT162b2 (Pfizer) followed by mRNA vaccine did not reache threshold for protection against symptomatic infection from VOC Omicron (62.9% [95% CI, 60.5 to 65.1] 1 Obs [34]) in adolescents age 12 to 17 years

Moderna [mRNA-1723]

• VOC Omicron

We have low certainty evidence that <u>2 doses</u> of mRNA-1723 did not reach threshold for protection from infection from VOC Omicron (range of mean estimates: <u>55 to 78%</u> - 1 Obs – [<u>35]</u>) in adolescents age 12 to 17 years

Sinovac [CoronaVac]

VOC Omicron

- We have low certainty evidence that <u>1 dose</u> of CoronaVac did not reach threshold for protection from symptomatic infection from VOC Omicron (21.2% [95% CI, 18.6 to 23.8] 1 Obs [21]) in children age 6 to 11 years
- We have low certainty evidence that <u>1 dose</u> of CoronaVac did not reach threshold for protection from infection from VOC Omicron BA.2 (-14.7% [95% CI, -54.7 to 14.6] 1 Obs [29]) in children age 3 to 11 years

- We have low certainty evidence that <u>1 dose</u> of CoronaVac did not reach threshold for protection from infection from VOC Omicron BA.2 (21.5% [95% CI, -7.7 to 42.7] 1 Obs [<u>29</u>]) in adolescents age 12 to 18 years
- We have low certainty evidence that 1 dose of CoronaVac did not reach threshold for protection from infection from VOC Omicron BA.2 (22.7% [95% CI, -38.3 to 56.8] 1 Obs [33]) in persons age 5 to 17 years
- We have low certainty evidence that <u>1 dose</u> of **CoronaVac** did not reach threshold for protection from ICU admission from VOC **Omicron BA.2** (41.9% [95% CI, -10.4 to 72.2] 1 Obs [21]) in children age 6 to 11 years
- We have low certainty evidence that <u>2 doses</u> of CoronaVac did not reach threshold for protection from symptomatic infection from VOC Omicron (<u>39.8% [95% CI, 33.7 to 45.4</u>] 1 Obs [<u>21]</u>) in children age 6 to 11 years
- We have low certainty evidence that <u>2 doses</u> of CoronaVac did not reach threshold for protection from ICU admission from VOC Omicron (20.9% [95% CI, -177.2 to 85] 1 Obs [21]) in children age 6 to 11 years
- We have low certainty evidence that <u>2 doses</u> of **CoronaVac** did not reach threshold for protection from symptomatic infection from VOC **Omicron BA.1** (38.2% [95% CI, 36.5 to 39.9] 1 Obs [<u>12</u>]) in children age 3 to 5 years
- We have low certainty evidence that <u>2 doses</u> of CoronaVac did not reach threshold for protection from ICU admission from VOC Omicron BA.1 (69% [95% CI, 18.6 to 88.2] 1 Obs [<u>12</u>]) in children age 3 to 5 years
- We have low certainty evidence that <u>2 doses</u> of **CoronaVac** did not reach threshold for protection from infection from VOC **Omicron BA.2** (40.8% [95% CI, 12.8 to 59.5] 1 Obs [<u>29</u>]) in children age 3 to 11 years
- We have low certainty evidence that <u>2 doses</u> of **CoronaVac** did not reach threshold for protection from infection from VOC **Omicron BA.2** (55% [95% CI, 38.2 to 67.2] 1 Obs [<u>29</u>]) in adolescents age 12 to 18 years
- We have low certainty evidence that 3 doses of CoronaVac reached threshold for protection against infection from VOC Omicron BA.2 (92% [95% CI, 86.7 to 95.2] 1 Obs [29]) in adolescents age 12 to 18 years

Table 1: Visual summary of evidence for COVID-19 vaccines for variants of concern (up to 28 days after 2 doses)

Percentages indicate <u>level of effectiveness</u> from 0% (no effect) to 100% (full protection): ranges of estimated means are provided when ≥ 1 study is available; estimated mean value is provided for single studies

Colour indicates level of certainty based on the evidence*

^{*}Please note: prior to LES 8.9 moderate certainty evidence was coloured orange and low certainty evidence was coloured yellow

_	High certainty evidence	Moderate certainty evidence	Low certainty evidence
	pooling of low to moderate risk	single RCT with low to moderate	single RCT or observational
	of bias RCTs or pooling of	risk of bias or >one	study with serious risk of bias or
	observational studies with low	observational study with low to	multiple low to serious risk of
	risk of bias and consistent	moderate risk of bias and at least	bias observational studies with
	findings	partially consistent findings	inconsistent findings

Outcome (and vaccine)	Vaccine Effectiveness (2 doses unless otherwise stated) up to 28 days after last dose each combination of vaccine, variant, and outcome					
	0	verall		elta	Omi	cron
Age	5 to 11 y	12 to 18 y	5 to 11 y	12 to 18 y	5 to 11 y	12 to 18 y
Any Infection						
Pfizer		91%		81 - 98%	29 – 70%	34 - 83%
Moderna				90 to 96%		55 – 78%
CoronaVac						
Johnson & Johnson						
Symptomatic Infect	ion					
Pfizer				81 - 97%	48 – 71%	55 - 83%
Moderna				98%		
CoronaVac					40%	
Johnson & Johnson				58%*		
ICU Admission						
Pfizer				98%	21%	
Moderna						
CoronaVac						69%
Johnson & Johnson						
Severe disease (may	include deat	h for some stud	ies)			
Pfizer					41 – 94%	76%
Moderna						
CoronaVac						
Johnson & Johnson						
Death						
Pfizer						
Moderna						
CoronaVac						
Johnson & Johnson						

^{*}Single dose

Table 2: Visual summary of evidence for COVID-19 vaccines for variant of concern – Delta and Omicron [2 doses > 28 days since last dose; 3 doses: > 1 days since last dose]

Percentages indicate <u>level of effectiveness</u> from 0% (no effect) to 100% (full protection): ranges of estimated means are provided when ≥ 1 study is available; estimated mean value is provided for single studies

Colour indicates <u>level of certainty</u> based on the evidence*

*Please note: prior to LES 8.9 moderate certainty evidence was coloured orange and low certainty evidence was coloured yellow

High certainty evidence	Moderate certainty evidence	Low certainty evidence
pooling of low to moderate	single RCT with low to moderate	single RCT or observational
risk of bias RCTs or pooling of	risk of bias or >one observational	study with serious risk of bias or
observational studies with low	study with low to moderate risk of	multiple low to serious risk of
risk of bias and consistent	bias and at least partially consistent	bias observational studies with
findings	findings	inconsistent findings

Outcome (and vaccine)	Variant	Number of doses	Time since Last Dose (days)	Age (years)	Vaccine Effectiveness
Any Infection					
Pfizer	Delta	1	21 to 48	12 to 17	63 to 68
			28 to 56		86.4% (95% CI, 83.5 to 88.7)
			49 to 76		47 to 56
			56 to 84		61.5% (95% CI, 43.5 to 73.7)
			77		29 to 49
		2	28 to 55	12 to 18	90 to 97
			35 to 69		83% (95% CI, 79 to 87)
			56 to 83		95 to 96
			84 to 111		94 to 95
			70		82% (95% CI, 74 to 88)
			91 to 119		83% (95% CI, 34 to 95)
			112 to 139		91 to 92
			35 to 62	16 to 17	92.8% (95% CI, 89.8 to 94.9)
			63		83.7% (95% CI, 75.9 to 89)
			14 to 149	12 to 15	87% (95% CI, 49 to 97)
	Omicron	Omicron 1	21 to 48	12 to 17	16 to 34
			28 to 56		57.9% (95% CI, 50.9 to 63.9)
			49 to 76		-1 to 17
			77		-13 to -5
			56 to 84		63.7% (95% CI, 59 to 67.9)
			60	5 to 11	4% (95% CI, -12 to 18)

29 to 84	5% CI, 9 to 48)
	21 to 23
30 to 59 28.5% (95	% CI, 26.3 to 30.7
60 25.6% (95	% CI, 19.3 to 31.5)
70	5% CI, 20 to 26)
35 to 62 16 to 17 45.7% (95	% CI, 34.8 to 54.7)
63 23.3% (95	5% CI, 2.7 to 39.5)
	5% CI, 22 to 79)
28 to 55 12 to 17	59 to 63
35 to 69 23% (9.	5% CI, 19 to 27)
56 to 83	48 to 58
84 to 111	41 to 51
112 to 139	38 to 46
70 8% (9.	5% CI, 5 to 11)
3 14	4 to 72%
7 - 13	5% CI, 78 to 82)
35 to 69 30% (9.	5% CI, 27 to 33)
Moderna Omicron 2 35 to 69 12 to 17 29% (9.	5% CI, 23 to 35)
70 20% (9.	5% CI, 15 to 24)
CoronaVac	
Symptomatic Infection	
	% CI, 45.5 to 49.8)
	1 to 63%
	6 to 58%
	4 to 54%
	6 to 48%
	5 to 46%
	29 to 53%
	% CI, 58.8 to 60)
	% CI, 25.4 to 36.0)
2 35 to 69 16 to 17 91.5% (95	% CI, 89.9 to 93.0)
	% CI, 72.0 to 90.5)
	<u> </u>
14 to 149 12 to 17	5 to 92%
14 to 149 12 to 17 8 91.8% (95	% CI, 91.2 to 92.3)
14 to 149 12 to 17 8 91.8% (95	
14 to 149 12 to 17 8 91.8% (95 56 to 112	% CI, 91.2 to 92.3)
14 to 149 12 to 17 14 to 98 56 to 112 60 to 119 12 to 17 91.8% (95) 96% (9.88)	% CI, 91.2 to 92.3) 6 to 68%
14 to 149 12 to 17 14 to 98 91.8% (95) 56 to 112 60 to 119 96% (9) 31 to 60 12 to 19	% CI, 91.2 to 92.3) 6 to 68% 5% CI, 94 to 97)

		2 doses + mRNA vaccine	14 to 98	12 to 17	96% (95% CI, 92.2 to 97.9)
	Omicron	1	28 to 34	12 to 17	33 to 42%
			35 to 41]	36 to 49%
			42 to 55]	29 to 40%
			56 to 69]	23 to 27%
			70 to 83]	16 to 27%
			84		17 to 26%
			14 to 98		18.8% (95% CI, 17.2 to 20.3)
			105	16 to 17	12.5% (95% CI, 96.9 to 17.8)
		2	7 to 59	12 to 17	51% (95% CI, 38 to 61)
			14 to 149		34 to 45%
			56 to 112		35 to 38%
			60 to 119	1	31% (95% CI, 20 to 41)
			14 to 98		64.5% (95% CI, 63.3 to 65.4)
			35 to 69	16 to 17	49.5% (95% CI, 45.7 to 53.0)
			70		22.6% (95% CI, 14.5 to 29.9)
			30 - 90	5 to 11	28.9% (95% CI, 24.5 to 33.1)
			30 - 59		60.2% (95% CI, 54.1 to 65.5)
			60		42.7% (95% CI, 12 to 62.7)
			90		35% (95% CI, 21 to 46)
			30 - 90	12 to 15	16.6% (95% CI, 8.1 to 24.3)
			60 - 120	1	9.6% (95% CI, -0.1 to 18.3)
		3	7	12 to 17	62 to 87
			0 to 60		56% (95% CI, 34 to 70)
		2 doses +	14 to 98		62.9% (95% CI, 60.5 to 65.1)
		mRNA vaccine		12 to 17	
Moderna	Delta	2	31 to 60	16 to 19	91% (95% CI, 87 to 94)
			61 to 90	1	85% (95% CI, 82 to 88)
			91 to 120]	85% (95% CI, 87 to 87)
CoronaVac					
Johnson &	Delta	1	31 to 60	16 to 19	52% (95% CI, 27 to 69)
Johnson			61 to 90]	63% (95% CI, 43 to 75)
			91 to 120		58% (95% CI, 45 to 68)
Transmission					
Pfizer					
Moderna					
CoronaVac					
ICU Admission					

Pfizer					
Moderna					
CoronaVac					
MIS-C					
Pfizer	Delta	1	28	12 to 18	94% (95% CI, 83 to 98)
		2			91% (78 to 97)
	Omicron	2			92% (95% CI, 71 to 98)
Moderna					
CoronaVac					
Severe Disease	(may incl	ude death	for some studies)		
Pfizer	Omicron	2	7 to 60	12 to 17	76 to 84%
			60 to 120		82 to 86%
			60		74% (95% CI, 44 to 88)
			98		82.7 % (95% CI, 68.8 to 90.4)
Moderna					
CoronaVac					
Death					
Pfizer					
Moderna					
CoronaVac					

Table 3a: Key findings about vaccine effectiveness for VOC Omicron (Revised 20 Jun 2022)

	Omicron – 1 dose				
Vaccine	Time frame	Findings			
Pfizer/	Omicron	BNT162b2 provided protection against VOC Omicron for the			
BioNTech		following outcomes at least 14 days after 1st dose in adolescents age			
	At least 14 days	12 to 17:			
Comirnaty	after 1st dose	• 53.7% (95% CI, 43.3 to 62.2) from infection (1 Obs - [10])			
		• 25 to 53% (RME) from symptomatic infection (2 Obs - [5][23])			
[BNT162b2]		• 56.3% (95% CI, 45.9 to 64.6) from severe disease (1 Obs - [23])			
		BNT162b2 provided protection against VOC Omicron for the			
		following outcomes at least 14 days after 1st dose in children age 5			
		to 11:			
		• 17 to 27% (RME) from infection (2 Obs - [25][27])			
		• 13 to 23% (RME) from symptomatic infection (2 Obs – [23] [25])			
		• 38.1% (95% CI, 20.9 to 51.5) from severe disease (1 Obs - [23])			
		BA. 2			
		BNT162b2 provided protection against VOC Omicron for the			
		following outcomes at least 14 days after 1st dose in children age 3			
		to 11:			
		• 33.3% (95% CI, 3 to 53.3) from infection (1 Obs - [29])			

	Omicron >30 days after 1 st dose	BNT162b2 provided protection against VOC Omicron for the following outcomes at least 14 days after 1st dose in adolescents age 12 to 18: • 26.1% (95% CI, -0.3 to 45.6) from infection (1 Obs - [29]) BNT162b2 provided protection against VOC Omicron for the following outcomes at least 14 days after 1st dose in persons age 5 to 17: • 32.4% (95% CI, -29 to 64.6) from infection (1 Obs - [33]) (7 Obs) [5][10][23][23][23] last update 2022-09-13 BNT162b2 provided protection against infection by VOC Omicron the following number of days after 1st dose in adolescents age 12 to 17: • 57.9% (95% CI, 50.9 to 63.9) – at 28 to 56 days (1 Obs - [10]) • 63.7% (95% CI, 59 to 67.9) – at 56 to 84 days (1 Obs - [10]) • 10 17 (RME) – at 49 to 76 days (1 Obs - [13]) • 16 to 34 (RME) – at least 77 days (1 Obs - [13]) • 16 to 34 (RME) – at 21 to 48 days (1 Obs - [13]) BNT162b2 provided protection against symptomatic infection by VOC Omicron the following number of days after 1st dose in children age 5 to 11: • 4% (95% CI, -12 to 18) – at least 60 days (1 Obs - [30]) BNT162b2 provided protection against symptomatic infection by VOC Omicron the following number of days after 1st dose in adolescents age 12 to 17: • 33 to 42% (RME) – at 28 to 34 days (1 Obs - [5]) • 33 to 42% (RME) – at 28 to 34 days (1 Obs - [5]) • 23 to 27% (RME) – at 35 to 41 days (1 Obs - [5]) • 23 to 27% (RME) – at 56 to 69 days (1 Obs - [5]) • 16 to 27% (RME) – at 70 to 83 days (1 Obs - [5]) • 17 to 26% (RME) – at 1east 84 days (1 Obs - [5]) • 18.8% (95% CI, 17.2 to 20.3) – at 14 to 98 days (1 Obs - [34]) BNT162b2 provided protection against symptomatic infection by VOC Omicron the following number of days after 1st dose in adolescents age 16 to 17:
Sinovac [CoronaVac]	Omicron At least 14 days after 1 st dose	CoronaVac provided protection against VOC Omicron for the following outcomes at least 14 days after 1st dose in children age 6 to 11: • 21.2% (95% CI, 18.6 to 23.8) from symptomatic infection-(1 Obs-[21]) BA. 2
		BNT162b2 provided protection against VOC Omicron for the following outcomes at least 14 days after 1st dose in children age 3 to 11: • -14.7% (95% CI, - 54.7 to 14.6) from infection (1 Obs - [29]) BNT162b2 provided protection against VOC Omicron for the following outcomes at least 14 days after 1st dose in children age 6 to 11:

		 47.1% (95% CI, 26.6 to 62.7) from hospitalization (1 Obs - [21]) 41.9% (95% CI, -10.4 to 72.2) from ICU admission (1 Obs - [21]) BNT162b2 provided protection against VOC Omicron for the following outcomes at least 14 days after 1st dose in adolescents age 12 to 18: 21.5% (95% CI, -7.7 to 42.7) from infection (1 Obs - [29])
		BNT162b2 provided protection against VOC Omicron for the following outcomes at least 14 days after 1st dose in adolescents age 5 to 17:
		• 22.7% (95% CI, -38.3 to 56.8) from infection (1 Obs - [33])
		(2 Obs) [21][29][33]; last update 2022-09-13
DG===/	Omicron	Omicron – 2 doses DNIT1(2)2 provided protection assinct VOC Omicron for the
Pfizer/ BioNTech	At least 7 days after	BNT162b2 provided protection against VOC Omicron for the following outcomes at least 7 days after 2 nd dose in children age 5 to 11:
Comirnaty	2 nd dose	 29 to 70% (RME) from infection (4 Obs – [25][27][28][35]) 68 to 88% (RME) from hospitalization (2 Obs - [15][28])
[BNT162b2]		 48 to 71% (RME) from symptomatic infection (4 Obs - [22][25][28][30]) 41 to 94% (RME) from severe disease (2 Obs - [27][30]) BNT162b2 provided protection against VOC Omicron for the following outcomes at least 7 days after 2nd dose in adolescents age
		12 to 17: • 34 to 83% (RME) from infection (4 Obs - [11][13][26][35]) • 55 to 83% (RME) from symptomatic infection (4 Obs - [5][22][23][26]) • 75.6% (95% CI, 58.1 to 85.8) from severe disease (1 Obs - [23])
		BA. 2 BNT162b2 provided protection against VOC Omicron for the following outcomes at least 14 days after 2 nd dose in adolescents age 12 to 18: • 54.9% (95% CI, 38.9 to 66.8) from infection (1 Obs - [29]) BA.1, BA. 2, BA.4, BA.5
		BNT162b2 provided protection against VOC Omicron for the following outcomes at least 14 days after 2 nd dose in children age 5 to 11:
		• 25.7% (95% CI, 10 to 38.6) from infection (1 Obs - [31]) BNT162b2 provided protection against VOC Omicron for the following outcomes at least 14 days after 2 nd dose in adolescents age 12 to 18:
		• 30.6% (95% CI, 26.9 to 34.1) from infection (1 Obs - [31]) (14 Obs) [10][11][13][15][22][23][25][26][27][28][29][30][31][35]]; last update 2022-09-13
	Omicron	BNT162b2 provided protection against infection by VOC Omicron for the following number of days after 2 nd dose in
	>30 days after 2 nd dose	children age 5 to 11: • 31% (95% CI, 9 to 48) - at 14 to 82 days (1 Obs - [11]) • 21 to 23% (RME) - at 29 to 84 days (2 Obs - [27][35]) • 28.5% (95% CI, 26.3 to 30.7) - at 30 to 59 days (1 Obs - [28])
		• 25.6% (95% CI, 19.3 to 31.5) - at least 60 days (1 Obs - [28]) • 23% (95% CI, 20 to 26) - at least 70 days (1 Obs - [35])

BNT162b2 provided protection against infection by VOC Omicron for the following number of days after 2nd dose in adolescents age 12 to 15:

- 59% (95% CI, 22 to 79) at 14 to 149 days (1 Obs [11]) BNT162b2 provided protection against infection by VOC Omicron for the following number of days after 2nd dose in adolescents age 16 to 17:
- 45.7% (95% CI, 34.8 to 54.7) at 35 to 62 days (1 Obs [13])
- 23.3% (95% CI, 2.7 to 39.5) at least 63 days (1 Obs [13]) BNT162b2 provided protection against infection by VOC Omicron for the following number of days after 2nd dose in adolescents age 12 to 17:
- 59 to 63% (RME) at 28 to 55 days (1 Obs [26])
- 23% (95% CI, 19 to 27) at 35 to 69 days (1 Obs [35])
- \bullet 48 to 58% (RME) at 56 to 83 days (1 Obs [26])
- 41 to 51% (RME) at 84 to 111 days (1 Obs [26])
- 38 to 46% (RME) at 112 to 139 days (1 Obs [26])
- 8% (95% CI, 5 to 11) at least 70 days (1 Obs [35])

BNT162b2 provided protection against MIS-C by VOC Omicron for the following number of days after 2nd dose in adolescents age 12 to 18:

- 92% (95% CI, 71 to 98) at least 28 days (1 Obs [7]) BNT162b2 provided protection against symptomatic infection from VOC Omicron for the following number of days after 2nd dose in adolescents age 16 to 17:
- 49.5% (95% CI, 45.7 to 53) at 35 69 days (1 Obs [5])
- 22.6% (95% CI, 14.5 to 29.9) at least 70 days (1 Obs [5])

BNT162b2 provided protection against symptomatic infection by VOC Omicron for the following number of days after 2nd dose in children age 5 to 11:

- 51% (95% CI, 30 to 65) at 14 to 67 days (1 Obs [8])
- 28.9% (95% CI, 24.5 to 33.1) at 30 to 90 days (1 Obs [22])
- 60.2% (95% CI, 54.1 to 65.5)- at 30 to 59 days (1 Obs [28])
- 42.7% (95% CI, 12 to 62.7)- at least 60 days (1 Obs [28])
- 35% (95% CI, 21 to 46)- at least 90 days (1 Obs [30])

BNT162b2 provided protection against symptomatic infection by VOC Omicron for the following number of days after 2nd dose in adolescents age 12 to 15:

- 16.6% (95% CI, 8.1 to 24.3)- at 30 to 90 days (1 Obs [22])
- 9.6% (95% CI, -0.1 to 18.3) at 60 to 120 days (1 Obs [22])

BNT162b2 provided protection against symptomatic infection by VOC Omicron for the following number of days after 2nd dose in adolescents age 12 to 17:

- 51% (95% CI, 38 to 61) at 7 to 59 days (1 Obs [16])
- 34 to 45% (RME) at 14 to 149 days (1 Obs [8])
- 35 to 38% (RME) at 56 to 112 days (1 Obs [32])
- 31% (95% CI, 20 to 41) at 60 to 119 days (1 Obs [16])
- 64.5% (95% CI, 63.3 to 65.4) at 14 to 98 days (1 Obs [34])

		BNT162b2 provided protection against hospitalization by VOC Omicron for the following number of days after 2 nd dose in children age 5 to 11: • 80.4% (95% CI, 67 to 88.4) - at 30 to 59 days (1 Obs - [28]) BNT162b2 provided protection against hospitalization by VOC Omicron for the following number of days after 2 nd dose in adolescents age 12 to 18: • 43% (95% CI, -1 to 68) - at 14 to 67 days (1 Obs - [15]) BNT162b2 provided protection against severe disease by VOC Omicron for the following number of days after 2 nd dose in adolescents age 12 to 17: • 76 to 84% (RME) - at 7 to 60 days (2 Obs - [16][23])
		 82 to 86% (RME) - at 60 to 120 days (2 Obs - [16][23]) 74% (95% CI, 44 to 88) - at least 60 days (1 Obs - [30]) 82.7 % (95% CI, 68.8 to 90.4) - at least 98 days (1 Obs - [23])
		BA. 2 BNT162b2 provided protection against VOC Omicron for the following outcomes at 14 - 84 days after 2 nd dose in children age 5 to 17: ■ 3.2% (95% CI, -220.7 to 70.8) from infection-(1 Obs - [33]) (16 Obs) [5][7][8][11][13][15][16][22][23][26][28][30][32][33][34][35]; last update 2022-
		09-36
Moderna	Omicron	mRNA-1723 provided protection against VOC Omicron for the
Spilrovay	At least 7 days after	following outcomes at least 7 days after 2 nd dose in adolescents age 12 to 17:
Spikevax	At least 7 days after 2 nd dose	• 55 to 78% (RME) - from infection-(1 Obs - [35])
[mRNA-1723]	2 4030	(1 Obs) [35]; last update 2022-09-13
[]	Omicron	mRNA-1723 provided protection against infection by VOC
		Omicron for the following number of days after 2 nd dose in
	>30 days after 2 nd	adolescents age 12 to 17:
	dose	• 29% (95% CI, 23 to 35) - at 35 to 69 days (1 Obs - [35])
		• 20% (95% CI, 15 to 24) - at least 70 days (1 Obs - [35])
0:		(1 Obs) [35]; last update 2022-09-13
Sinovac	Omicron	CoronaVac provided protection against VOC Omicron for the
[CoronaVac]	At least 7 days after	following outcomes at least 14 days after 2 nd dose in children age 6 to 11:
	2 nd dose	• 39.8% (95% CI, 33.7 to 45.4) from symptomatic infection-(1 Obs -
		[21])
		• 59.2% (95% CI, 11.3 to 84,5) from hospitalization-(1 Obs - [21])
		• 20.9% (95% CI, -177.2 to 85) from ICU admission-(1 Obs - [21])
		BA. 1
		CoronaVac provided protection against VOC Omicron for the
		following outcomes at least 14 days after 2 nd dose in children age 3
		to 5:
		• 38.2% (95% CI, 36.5 to 39.9) from symptomatic infection-(1 Obs -
		[12]) • 64.6% (95% CI, 49.6 to 75.2) from hospitalization-(1 Obs - [12])
		• 69% (95% CI, 18.6 to 88.2) from ICU admission-(1 Obs - [12])
		07/0 (73/0 C1, 10.0 to 66.2) Holli ICO adillission-(1 obs - 12])

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	Omicron >30 days after 2 nd dose	BA. 2 CoronaVac provided protection against VOC Omicron for the following outcomes at least 14 days after 2 nd dose in children age 3 to 11: • 40.8% (95% CI, 12.8 to 59.5) from infection-(1 Obs - [29]) CoronaVac provided protection against VOC Omicron for the following outcomes at least 14 days after 2 nd dose in adolescents age 12 to 18: • 55% (95% CI, 38.2 to 67.2) from infection-(1 Obs - [29]) (3 Obs) [12][21][29]; last update 2022-98-13 BA. 2 CoronaVac provided protection against infection by VOC Omicron for the following number of days after 2 nd dose in persons age 5 to 17: • 55.6% (95% CI, -50.3 to 86.9) – at 14 to 84 days (1 Obs - [33])
		(1 Obs) [<u>33</u>]; last update <u>2022-98-13</u>
D.C.		Omicron – 3 doses
Pfizer/	Omicron	BNT162b2 provided protection against infection by VOC
BioNTech	A parties of the man	Omicron the following number of days after <u>3rd dose</u> in adolescents age 12 to 17:
Comirnaty	Any time frame after 3 rd dose	e 12 to 17. ■ 64 to 72% (RME) at least 14 days (2 Obs - [26][35])
Commutaty	after 5 dose	• 80% (95% CI, 78 to 82) — at 7 to 13 days (1 Obs - [35])
[BNT162b2]		• 30% (95% CI, 27 to 33) – at 35 to 69 days (1 Obs - [35])
		BNT162b2 provided protection against Symptomatic infection by
		VOC Omicron the following number of days after 3 rd dose in
		adolescents age 12 to 17:
		• 56% (95% CI, 34 to 70) – at 0 to 6 days (1 Obs - [16])
		• 62 to 87% (RME) at least 7 days (3 Obs - [8][16][32])
		BNT162b2 provided protection against Symptomatic infection by
		VOC Omicron the following number of days after 3 rd dose in
		adolescents age 12 to 15:
		• 71.1% (95% CI, 65.5 to 75.7) – at 14 to 45 days (1 Obs - [22])
		RA 2
		BA. 2 BNT162b2 provided protection against VOC Omicron for the
		following outcomes at least 14 days after 3 rd dose in adolescents age
		12 to 18:
		• 86.8% (95% CI, 80.5 to 91.1) from infection-(1 Obs - [29])
		(6 Obs) [8][16][22][26][29][35]; last update 2022-09-13
	Omicron	BNT162b2 (2 doses) followed by mRNA vaccine provided
		protection against VOC Omicron for the following outcomes after
	(2 doses followed	3 rd dose in adolescents age 12 to 17:
	by mRNA vaccine)	• 62.9% (95% CI, 60.5 to 65.1) - from symptomatic infection at 14
	(1)	to 98 days (1 Obs - [34])
C:	(Any time frame)	(1 Obs) [<u>34</u>]; last update <mark>2022-09-13</mark>
Sinovac	Omicron	BA. 2 Corona Vac provided protection against VOC Omicron for the
[CoronaVac]	Any time frame	CoronaVac provided protection against VOC Omicron for the following outcomes at least 14 days after 3 rd dose in adolescents age
	after 3 rd dose	12 to 18:
	arter 3 dose	14 tO 10.

		• 92% (95% CI, 86.7 to 95.2) from infection-(1 Obs - [29])			
		(1 Obs) [29]; last update 2022-08-16			
Omicron – Relative VE					
Any vaccine	Relative VE for	Study populations that received booster doses are commonly very different from populations who did not receive or were			
	primary series	not yet eligible for booster doses which increases the risk of			
	vaccine doses	bias			
	compared to				
	primary series plus	No data yet			
	booster vaccine				
	doses (instead of an				
	unvaccinated				
	group)				
		micron - Hybrid Immunity			
Pfizer/	Omicron	BNT162b2 (1 dose + prior infection) provided protection against			
BioNTech		VOC Omicron for the following outcomes after 1st dose in			
	Protection provided	adolescents age 12 to 17:			
Comirnaty	by previous	• 85.3% (95% CI, 83.7 to 86.8) from symptomatic infection at 14			
	infection plus	to 98 days (wild type prior infection) -(1 Obs - [34])			
[BNT162b2]	vaccination	• 81.5 % (95% CI, 80.0 to 82.9) from symptomatic infection at 14			
		to 98 days (Alpha prior infection) -(1 Obs - [34])			
		• 78.8 % (95% CI, 77.9 to 79.5) from symptomatic infection at 14			
		to 98 days (Delta prior infection) -(1 Obs - [34])			
		• 79.6 % (95% CI, 44.9 to 92.4) from symptomatic infection at 14			
		to 98 days (Omicron) prior infection -(1 Obs - [34])			
		BNT162b2 (2 doses + prior infection) provided protection against			
		VOC Omicron for the following outcomes after 2 nd dose in			
		adolescents age 12 to 17:			
		• 84.7% (95% CI, 82.6 to 86.5) from symptomatic infection at 14			
		to 98 days (wild type prior infection) -(1 Obs - [34])			
		• 85.5 % (95% CI, 84 to 86.9) from symptomatic infection at 14 to			
		98 days (Alpha prior infection) -(1 Obs - [34])			
		• 83.5 % (95% CI, 82.5 to 84.5) from symptomatic infection at 14			
		to 98 days (Delta prior infection) -(1 Obs - [34])			
		(1 Obs) [34]; last update 2022-09-13			
	Omicron	BNT162b2 (2 doses + prior infection) followed by mRNA vaccine			
	(2.1 (.11 1	provided protection against VOC Omicron for the following			
	(2 doses followed	outcomes after 3 rd dose in adolescents age 12 to 17:			
	by mRNA vaccine	• 79.8% (95% CI, 70.4 to 86.3) from symptomatic infection at 14			
	plus prior infection)	to 98 days (wild type prior infection) -(1 Obs - [34])			
	(Apy time frame)	• 79.6 % (95% CI, 71.4 to 85.5) from symptomatic infection at 14			
	(Any time frame)	to 98 days (Alpha prior infection) -(1 Obs - [34])			
		• 80.7 % (95% CI, 71.1 to 87.1) from symptomatic infection at 14			
		to 98 days (Delta prior infection) -(1 Obs - [34])			
		(1 Obs) [<u>34</u>]; last update <u>2022-09-13</u>			

Pan American Health Organization/World Health Organization. Pharmacovigilance for COVID-19 Vaccines. https://covid-19pharmacovigilance.paho.org

Table 3b: Key findings about vaccine effectiveness for VOC Delta (Revised 20 Jun 2022)

Delta – 1 dose					
Vaccine	Time frame	Findings			
Pfizer/ BioNTech	Delta	BNT162b2 provided protection against VOC Delta for the following outcomes at least 14 days after 1st dose in adolescents age			
Comirnaty	At least 14 days after 1st dose	12 to 18: • 55 to 80% from infection (RME) (4 Obs - [2][10][17][18]) • 52 to 76% from symptometric infection (RME) (4 Obs - [2][10][17][18])			
[BNT162b2]		• 52 to 76% from symptomatic infection(RME) (4 Obs - [5][9][18][23]) BNT162b2 provided protection against VOC Delta for the following outcomes at 0 to 27 days after 1st dose in adolescents age			
		12 to 15: • 14.2% (95% CI, - 25.6 to 41.4) against hospitalization (1 Obs - [5])			
		BNT162b2 provided protection against VOC Delta for the following outcomes at 0 to 27 days after 1st dose in adolescents age			
		16 to 17: • 64.6% (95% CI, 40.7 to 78.9) from hospitalization (1 Obs - [5])			
		(7 Obs). [2][5][9][10][17][18]][23]; last update 2022-08-16			
	Delta	BNT162b2 provided protection against infection by VOC Delta the following number of days after 1 st dose in adolescents age 12 to			
	>30 days after 1 st	17:			
	dose	• 47.7% (95% CI, 45.5 to 49.8) – at least 28 days (1 Obs - [23])			
		• 86.4% (95% CI, 83.5 to 88.7) – at 28 to 56 days (1 Obs - [10])			
		• 61.5% (95% CI, 43.5 to 73.7) – at 56 to 84 days (1 Obs - [10])			
		• 63 to 68% (RME) – at 21 to 48 days (1 Obs - [13])			
		 47 to 56% (RME) – at 49 to 76 days (1 Obs - [13]) 29 to 49% (RME) – at least 77 days (1 Obs - [13]) 			
		· · · · · · · · · · · · · · · · · · ·			
		BNT162b2 provided protection against symptomatic infection by			
		VOC Delta the following number of days after 1 st dose in adolescents age 12 to 17:			
		• 61 to 63% (RME) – at 28 to 34 days (1 Obs - [5])			
		• 56 to 58% (RME) – at 35 to 41 days (1 Obs - [5])			
		• 44 to 54% (RME) – at 42 to 55 days (1 Obs - [5])			
		• 36 to 48% (RME) – at 56 to 69 days (1 Obs - [5])			
		• 35 to 46% (RME) – at 70 to 83 days (1 Obs - [5])			
		• 29 to 53% (RME) – at 84 to 104 days (1 1 Obs - [5])			
		• 59.4% (95% CI, 58.8 to 60) – at 14 to 98 days (1 Obs - [34])			
		BNT162b2 provided protection against symptomatic infection by			
		VOC Delta the following number of days after 1st dose in			
		adolescents age 16 to 17:			
		• 30.9% (95% CI, 25.4 to 36.0) – at least 105 days (1 Obs - [5])			
		BNT162b2 provided protection against hospitalization by VOC			
		Delta the following number of days after 1st dose in adolescents age 12 to 17:			
		• 76 to 83% (RME) - at least 28 days (1 Obs - [5])			
		(5 Obs) [5][10][13][23][34] ; last update 2022-09-13			
Johnson &	Delta	AD26.COV2.S provided protection against VOC Delta for the			
Johnson		following outcomes at least 14 days after dose in adolescents age 16			
[AD26.COV2.S]	Up to 30 days	to 19:			
	after dose	• 58% (95% CI, 19 to 79) from symptomatic infection-(1 Obs - [19])			

		(1 Obs) [19]; last update 2022-05-09		
	Delta	AD26.COV2.S provided protection against symptomatic infection		
	Delta	by VOC Delta for the following number of days after dose in		
	>30 days after	adolescents age 16 to 19:		
	dose	S		
	uose	• 52% (95% CI, 27 to 69) - at 31 to 60 days (1 Obs - [19])		
		• 63% (95% CI, 43 to 75) - at 61 to 90 days (1 Obs - [19])		
		• 58% (95% CI, 45 to 68)- at 91 to 120 days (1 Obs - [19])		
		(1 Obs) [19]; last update 2022-05-09		
DC. /	Du	Delta – 2 doses		
Pfizer/	Delta	BNT162b2 provided protection against VOC Delta for the		
BioNTech	A . 1 7 . 1	following outcomes at least 7 days after 2 nd dose in adolescents age		
	At least 7 days	12 to 18:		
Comirnaty	after 2 nd dose	• 81 to 98% against infection (RME) (9 Obs – [1][2][6][9][11][13][17][26][35])		
FD3 17714 (Q1 Q1		• 81 to 97% against symptomatic infection (RME) (6 Obs – [5][9][16][19]		
[BNT162b2]		[23][26]) BNT162b2 provided protection against VOC Delta for the		
		following outcomes at least 14 days after 2 nd dose in adolescents age		
		12 to 18:		
		• 94% (95% CI, 90 to 96) from hospitalization (1 Obs – [4])		
		• 98% (95% CI, 93 to 99) from ICU admission (1 Obs - [4])		
		(14 Obs) [1][2][4][5][6][9][11][13][16][17][19][23][26][35]; last update 2022-98-13		
	Delta	BNT162b2 provided protection against infection by VOC Delta for		
	Dena	the following number of days after 2^{nd} dose in adolescents age 12 to		
	>30 days after 2 nd	18:		
	dose	• 83% (95% CI, 34 to 95) - at 34 to 95 days (1 Obs - [2])		
	uose	• 83% (95% CI, 79 to 87)- at 35 to 69 days (1 Obs - [2])		
		• 90 - 97% (RME) - at 28 to 55 days (2 Obs - [2][26])		
		• 95 to 96% (RME) - at 56 to 83 days (2 Obs - [2][26])		
		• 94 to 95% (RME) - at 84 to 111 days (1 Obs - [26])		
		` '		
		• 91 to 92% (RME) - at 112 to 139 days (1 Obs - [26])		
		• 82% (95% CI, 74 to 88) - at least 70 days (1 Obs - [35])		
		BNT162b2 provided protection against infection by VOC Delta for		
		the following number of days after 2 nd dose in adolescents age 12 to		
		15:		
		• 87% (95% CI, 49 to 97) - at 14 to 149 days (1 Obs - [11])		
		BNT162b2 provided protection against infection by VOC Delta for		
		the following number of days after 2 nd dose in adolescents age 16 to 17:		
		• 92.8% (95% CI, 89.8 to 94.9) - at 35 to 62 days (1 Obs - [13])		
		• 83.7% (95% CI, 75.9 to 89) - at least 63 days (1 Obs - [13])		
		BNT162b2 provided protection against MIS-C by VOC Delta the		
		following number of days after 2nd dose in adolescents age 12 to 18:		
		• 94% (95% CI, 83 to 98) - at least 28 days (1 Obs - [7])		
		BNT162b2 provided protection against symptomatic infection by		
		VOC Delta for the following number of days after 2 nd dose in		
		adolescents age 16 to 17:		
		• 91.5% (95% CI, 89.9 to 93.0) - at 35 to 69 days (1 Obs - [5])		
		• 83.7% (95% CI, 72.0 to 90.5) - at least 70 days (1 Obs - [5])		

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		BNT162b2 provided protection against symptomatic infection by			
		VOC Delta for the following number of days after 2 nd dose in			
		adolescents age 12 to 17:			
		• 85 to 92% (RME) - at 14 to 149 days (1 Obs - [8])			
		• 66 to 68% (RME) - at 56 to 112 days (1 Obs - [32])			
		• 96% (95% CI, 94 to 97) - at 60 to 119 days (1 Obs - [16])			
		• 91.8% (95% CI, 91.2 to 92.3) - at 14 to 98 days (1 Obs – [34])			
		BNT162b2 provided protection against symptomatic infection by			
		VOC Delta for the following number of days after 2 nd dose in			
		adolescents age 12 to 19:			
		• 87 to 93% (RME) - at 31 to 60 days (1 Obs - [19])			
		• 86 to 92% (RME) - at 61 to 90 days (1 Obs - [19])			
		• 82 to 92% (RME) - at 91 to 120 days (1 Obs - [19])			
		BNT162b2 provided protection against hospitalization by VOC			
		Delta for the following number of days after 2 nd dose in adolescents			
		age 12 to 18:			
		• 93% (95% CI, 89 to 95)- at 14 to 154 days (1 Obs - [13])			
		(12 Obs) [5][7][8][9][11][13][16][19][26][32][34][35]); last update 2022-09-13			
Moderna	Delta	mRNA-1723 provided protection against VOC Delta for the			
		following outcomes at least 14 days after 2 nd dose in adolescents age			
Spikevax	At least 7 days	16 to 19:			
	after 2 nd dose	• 98% (95% CI, 92 to 99) from symptomatic infection-(1 Obs - [19])			
[mRNA-1723]		mRNA-1723 provided protection against VOC Delta for the			
[]		following outcomes at least 7 days after 2 nd dose in adolescents age			
		12 to 17:			
		• 90 to 96% (RME) - from infection-(1 Obs - [35])			
		(2 Obs) [19][35]; last update 2022-09-13			
	Delta	mRNA-1723 provided protection against symptomatic infection by			
		VOC Delta for the following number of days after 2 nd dose in			
	>30 days after 2 nd	adolescents age 16 to 19:			
	dose	• 91% (95% CI, 87 to 94) - at 31 to 60 days (1 Obs - [19])			
		• 85% (95% CI, 82 to 88) - at 61 to 90 days (1 Obs - [19])			
		• 85% (95% CI, 82 to 87)- at 91 to 120 days (1 Obs - [19])			
		(1 Obs) [19]; last update 2022-05-09			
		Delta – 3 doses			
Pfizer/	Delta	BNT162b2 (2 doses) followed by mRNA vaccine provided			
BioNTech		protection against VOC Delta for the following outcomes after 3 rd			
	(2 doses followed	dose in adolescents age 12 to 17:			
Comirnaty	by mRNA	• 96% (95% CI, 92.2 to 97.9) - from symptomatic infection at 14			
	vaccine)	to 98 days (1 Obs - [34])			
[BNT162b2]		(1 Obs) [34]; last update 2022-09-13			
[,]	(Any time frame)	(), war ry war			
		Delta – Relative VE			
Any vaccine	Delta	The results in this section should be reviewed with caution.			
		Study populations that received booster doses are commonly			
	Relative VE for	very different from populations who did not receive or were			
	primary series	not yet eligible for booster doses which increases the risk of			
	vaccine doses	bias			
	compared to				
	primary series plus	No data yet			
1	I Paris Journey Plus	i i o anan jou			

	booster vaccine				
	doses (instead of				
	an unvaccinated				
	group)				
	Delta - Hybrid Immunity				
Pfizer/	Delta	BNT162b2 (1 dose + prior infection) provided protection against			
BioNTech		VOC Delta for the following outcomes after 1^{st} dose in adolescents			
	Protection	age 12 to 17:			
Comirnaty	provided by	• 98.1% (95% CI, 97.6 to 98.6) from symptomatic infection at 14			
	previous infection	to 98 days (wild type prior infection) -(1 Obs - [34])			
[BNT162b2]	plus vaccination	• 95.5 % (95% CI, 94.8 to 96.1) from symptomatic infection at 14			
		to 98 days (Alpha prior infection) -(1 Obs - [34])			
		• 97.5% (95% CI, 97 to 97.9) from symptomatic infection at 14 to			
		98 days (Delta prior infection) -(1 Obs - [34])			
		BNT162b2 (2 doses + prior infection) provided protection against			
		VOC Delta for the following outcomes after 2 nd dose in			
		adolescents age 12 to 17:			
		• 98.8% (95% CI, 96.7 to 98.8) from symptomatic infection at 14			
		to 98 days (wild type prior infection) -(1 Obs - [34])			
		• 99.2 % (95% CI, 97.8 to 99.7) from symptomatic infection at 14			
		to 98 days (Alpha prior infection) -(1 Obs - [34])			
		• 98.7 % (95% CI, 96.8 to 99.4) from symptomatic infection at 14			
		to 98 days (Delta prior infection) -(1 Obs - [34])			
		(1 Obs) [<u>34</u>]; last update <u>2022-09-13</u>			

Pan American Health Organization/World Health Organization. Pharmacovigilance for COVID-19 Vaccines. https://covid-19pharmacovigilance.paho.org

Table 3c: Key findings about vaccine effectiveness in studies covering more than one VOC (Revised 20 Jun 2022)

More than one VOC – 1 dose				
Vaccine	Time frame	Findings		
Pfizer/	Overall	BNT162b2 provided protection for the following outcomes at least		
BioNTech		14 days after 1st dose in adolescents age 12 to 15:		
		• 67% (95% CI, 50 to 78) from infection (1 Obs – [3])		
Comirnaty		• 100% (95% CI, 100 to 100) from hospitalization (1 Obs - [3])		
		(1 Obs) [3]; last update 2021-12-13		
[BNT162b2]	Delta to	BNT162b2 provided protection against VOC Delta to Omicron for		
	Omicron	the following outcomes at least 14 days after 1st dose in adolescents		
		age 12 to 17:		
	At least 14 days	• 38% (95% CI, -51 to 79) from hospitalization (1 Obs – [14])		
	after 1st dose	BNT162b2 provided protection against VOC Delta to Omicron for		
		the following outcomes at least 14 days after 1st dose in children age		
		4 to 11:		
		• 32% (95% CI, -49 to 72) from hospitalization (1 Obs – [14])		
		BNT162b2 provided protection against VOC Delta to Omicron for		
		the following outcomes at least 14 days after 1st dose in children		
		and adolescents age 4 to 17:		

		270/ (050/ CI 12 + (7) C 1 '- 1' + (4 O) - (4 O)		
		• 37% (95% CI, -13 to 67) from hospitalization (1 Obs – [14])		
	D 1	(1 Obs) [14]; last update 2022-04-11		
	Delta to	BNT162b2 provided protection against infection by VOC Delta		
	Omicron	Omicron the following number of days after 1st dose in adolescents		
		age 12 to 17:		
	>30 days after 1 st	• 62 to 65 (RME) — at 21 to 48 days (1 Obs - [13])		
	dose	• 48 to 57 (RME) — at 49 to 76 days (1 Obs - [13])		
		• 48 to 70 (RME) — at least 77 days (1 Obs - [13])		
		(1 Obs) - [13]; last update 2022-04-11		
	M	ore than one VOC – 2 doses		
Pfizer/	Overall	BNT162b2 provided protection for the following outcomes at least		
BioNTech		7 days after 2 nd dose in adolescents age 12 to 15:		
		• 91% (95% CI, 88 to 93) from infection (1 Obs - [3])		
Comirnaty		• 81% (95% CI, -55 to 98) from hospitalization (1 Obs - [3])		
		(1 Obs) [3]; last update 2021-12-13		
[BNT162b2]	Delta to	BNT162b2 provided protection against VOC Delta to Omicron for		
[D11110202]	Omicron	the following outcomes at least 7 days after 2^{nd} dose in adolescents		
	Officion	,		
	A + 1 + 7 J	age 12 to 17:		
	At least 7 days	• 83 to 91% (RME) from infection (2 Obs - [13][26])		
	after 2 nd dose	BNT162b2 provided protection against VOC Delta to Omicron for		
		the following outcomes at least 14 days after 2 nd dose in adolescents		
		age 12 to 18:		
		• 82 to 83% (RME) from hospitalization (1 Obs - [15])		
		• 87.9% (95% CI, 86.1 to 89.5) from symptomatic infection (1 Obs -		
		[26]) RNT162h2 provided protection against VOC Delta to Omigran for		
		BNT162b2 provided protection against VOC Delta to Omicron for the following outcomes at least 14 days after 2 nd dose in adolescents		
		age 12 to 17:		
		• 59% (95% CI, 23 to 82) from hospitalization (1 Obs - [14])		
		BNT162b2 provided protection against VOC Delta to Omicron for		
		the following outcomes at least 14 days after 2 nd dose in adolescents		
		age 4 to 17:		
		• 59% (95% CI, 23 to 79) from hospitalization (1 Obs - [14])		
		(4 Obs) [13][14][15][26]; last update 2022-07-19		
	Delta to	BNT162b2 provided protection against infection by VOC Delta to		
	Omicron	Omicron for the following number of days after 2 nd dose in		
		adolescents age 12 to 17:		
	>30 days after 2 nd	• 88 to 95% (RME) - at 28 to 62 days (2 Obs - [13][26])		
	dose	• 84 to 88% (RME) - at 56 to 83 days (2 Obs - [13][26])		
		• 88 to 92% (RME) - at 84 to 111 days (1 Obs - [26])		
		• 83 to 87% (RME) - at 112 to 139 days (1 Obs - [26])		
		BNT162b2 provided protection against MIS-C by VOC Delta to		
		Omicron for the following number of days after $\frac{2^{nd}}{dose}$ in		
		children age 5 to 11:		
		• 78% (95% CI, 48 to 90) - at least 28 days (1 Obs - [7])		
		BNT162b2 provided protection against MIS-C by VOC Delta to		
		Omicron for the following number of days after 2 nd dose in		
		adolescents age 12 to 18:		
		S		
		• 90% (95% CI, 81 to 95) - at least 28 days (1 Obs - [7])		

	1			
		BNT162b2 provided protection against hospitalization by VOC Delta to Omicron for the following number of days after 2 nd dose		
		in children age 5 to 11:		
		• 74% (95% CI, -35 to 95) - at 14 to 67 days (1 Obs - [8])		
		BNT162b2 provided protection against hospitalization by VOC		
		Delta to Omicron for the following number of days after 2 nd dose		
		in adolescents age 12 to 17:		
		• 92 to 94% (RME) - at 14 to 149 days (1 Obs - [8])		
		BNT162b2 provided protection against symptomatic infection by		
		VOC Delta to Omicron for the following number of days after 2 nd		
		dose in children age 5 to 11:		
		• 46% (95% CI, 24 to 61) - at 14 to 67 days (1 Obs - [8])		
		BNT162b2 provided protection against symptomatic infection by		
		VOC Delta to Omicron for the following number of days after 2^{nd}		
		dose in adolescents age 12 to 17:		
		• 76 to 83% (RME) - at 14 to 149 days (1 Obs - [8])		
		(4 Obs) [7][8][13][26]; last update 2022-08-16		
		ore than one VOC – 3 doses		
Pfizer/	Delta to	BNT162b2 provided protection against VOC Delta to Omicron for		
BioNTech	Omicron	the following outcomes at least 7 days after 3 rd dose in adolescents		
		age 16 to 17:		
Comirnaty	Any time frame	• 86% (95% CI, 73 to 93) from symptomatic infection (1 Obs - [8])		
FD3 1/T/4 < 01 01	after 3 rd dose	(1 Obs) [8]; last update 2022-03-14		
[BNT162b2]	3.6	1 VOC D1 C VE		
A		e than one VOC – Relative VE		
Any vaccine	More than one VOC	The results in this section should be reviewed with caution.		
	VOC	Study populations that received booster doses are commonly very different from populations who did not receive or were		
	Relative VE for	not yet eligible for booster doses which increases the risk of		
	primary series	bias		
	vaccine doses	DIAG		
	compared to	No data yet		
	primary series plus			
	booster vaccine			
	doses (instead of			
	an unvaccinated			
	group)			
	10 17	nan one VOC - Hybrid Immunity		
Any vaccine	More than one	No data yet		
	VOC			
	Protection			
	provided by			
	previous infection			
	plus vaccination			

Pan American Health Organization/World Health Organization. Pharmacovigilance for COVID-19 Vaccines. https://covid-19pharmacovigilance.paho.org

Flórez ID^{1,2}, Velásquez-Salazar P¹, Martínez JC¹, Linkins L³, Abdelkader W³, Iorio A³, Lavis J³, Patiño-Lugo DF¹. COVID-19 living evidence synthesis #8 (version 16): What is the effectiveness of available COVID-19 vaccines in children and adolescents in general and specifically for variants of concern? Evidence and Deliberation Unit for Decision Making (UNED), University of Antioquia & Health Information Research Unit (HIRU), McMaster University, 13 Sep 2022.

To help Canadian decision-makers as they respond to unprecedented challenges related to the COVID-19 pandemic, COVID-END in Canada is preparing rapid evidence responses like this one. The development and continued updating of this living evidence synthesis has been funded by the Canadian Institutes of Health Research (CIHR) and the Public Health Agency of Canada. The opinions, results, and conclusions are those of the team that prepared the living evidence synthesis, and independent of the Government of Canada, CIHR and the Public Health Agency of Canada. No endorsement by the Government of Canada, CIHR or Public Health Agency of Canada is intended or should be inferred.

¹ Faculty of Medicine, University of Antioquia, Colombia

² School of Rehabilitation Science, McMaster University, Canada

³ Faculty of Health Sciences, McMaster University, Canada

Appendix 1: Summary of Study Findings and Appraisals

	Section 1: included studies						
Ref	Author	Bottom line	ROBINS- I*	Design, Notes			
		*Note: ROBINS-I score risk of bias: Low risk	x of bias indicat	tes high quality			
1	Glatman- Freedman	BNT162b2 showed VE 91.5% (95% CI, 88.2 to 93.9) against infection at least 8 days after 2 nd dose in adolescents age 12 to 15 years. There were no deaths in either group.	Serious	Population cohort in Israel of adolescents age 12 to 15 years; 2,034,591 vaccinated persondays and 13,623,714 unvaccinated person-days; time and setting for VOC Delta <i>Included in LES 8.1</i>			
2	Reis	BNT162b2 showed VE 59% (95% CI, 52 to 65) against infection 14 to 20 days after 1st dose in adolescents age 12 to 18. BNT162b2 showed VE 90% (95% CI, 88 to 92) against infection 7 to 21 days after 2nd dose in adolescents age 12 to 18.	Moderate	Case-control study in Israel; 94,354 vaccinated matched to 94,354 unvaccinated adolescents age 12 to 18; time and setting for VOC Delta Included in LES 8.1			
3	Tartof	BNT162b2 showed VE 67% (95% CI, 50 to 78) against infection and VE 100% (95% CI, 100 to 100) against hospitalization at least +14 days after 1st dose in adolescents age 12 to 15 years. BNT162b2 showed VE 91% (95% CI, 88 to 93) against infection and VE 81% (95% CI, -55 to 98) against hospitalization at least +7 days after 2nd dose in adolescents age 12 to 15 years.	Moderate	Retrospective Cohort in USA of 3,436,957 Kaiser Permanente Southern California (KPSC) healthcare system members ≥12 years of age between Dec 14, 2020 – Aug 8, 2021. The cohort included 122,779 adolescents age 12 to 15 years. The primary exposure was being fully vaccinated, defined as receiving 2 doses of BNT162b2 with ≥ 7 days after the second dose. Over the study period, 28.4% of 9,147 specimens sent for whole genome sequencing (WGS) and viral lineage designation were Delta. <i>Included in LES 8.1</i>			
4	Olson	BNT162b2 showed VE 94% (95% CI, 90 to 96) against hospitalization at least +14 days after 2 nd dose in adolescents age 12 to 18 years. BNT162b2 showed VE 95% (95% CI, 88 to 97) in adolescents age 12 to 15 years and VE 94% (95% CI, 88 to 97) in adolescents	Moderate	Test-negative study in U.S of adolescents age 12 to 18 years between Jun 1–Oct 25, 2021; 299 fully vaccinated (receipt of 2 doses of BNT162b2 vaccine, with the second dose administered ≥14 days before illness onset), 55 partially vaccinated (had received only			

age 16 to 18 years against hospitalization at least +14 days after 2 nd dose.	one dose of vaccine or who had received a second dose less than 14 days before illness onset) and
BNT162b2 showed VE 98% (95% CI, 93 to 99) against ICU admission at least +14 days after 2 nd dose in adolescents age 12 to 18 years. 5 Powell BNT162b2 showed after 1 st dose VE Moderate	868 unvaccinated (no receipt of any COVID-19 vaccine before illness onset), time and setting for VOC Delta. Included in LES 8.2 last update in LES 8.3 Test-negative case-control
74.5% (95% CI, 73.2 to 75.6) at 14-20 days, VE 63.4% (95% CI, 61.7 to 65.1) at 28-34 days, VE 47.5% (95% CI, 44.9 to 49.9) at 56-69 days, and VE 53.1% (95% CI, 41.6 to 62.4) at least 84 days, in adolescents age 12 to 15 years against infection. (VOC Delta) BNT162b2 showed after 1st dose VE 49.6% (95% CI, 43.9 to 54.8) at 14-20 days, VE 42.1% (95% CI, 36.7 to 46.9) at 28-34 days, VE 22.5% (95% CI, 19.1 to 25.8) at 56-69 days, and VE 17.2% (95% CI, 12.0 to 22.1) at least 84 days, in adolescents age 12 to 15 years against infection. (VOC Omieron) BNT162b2 showed after 1st dose VE 75.9% (95% CI, 74.3 to 77.3) at 14-20 days, VE 60.6% (95% CI, 58.1 to 62.9) at 28-34 days, VE 36.3% (95% CI, 33.1 to 39.3) at 56-69 days, VE 29.3% (95% CI, 25.9 to 32.6) at 84-104 days, and VE 30.9% (95% CI, 25.4 to 36.0) at least 105 days, in adolescents age 16 to 17 years against infection. (VOC Delta) BNT162b2 showed after 1st dose VE 51.4% (95% CI, 42.7 to 58.8) at 14-20 days, VE 33.9% (95% CI, 17.4 to 34.8) at 56-69 days, VE 20.5% (95% CI, 17.4 to 34.8) at 56-69 days, VE 20.5% (95% CI, 17.4 to 34.8) at 56-69 days, VE 20.5% (95% CI, 17.4 to 34.8) at 56-69 days, VE 20.5% (95% CI, 17.8 at least 105 days, in adolescents age 16 to 17 years against infection. (VOC Omieron) BNT162b2 showed after 2st dose VE 12.5% (95% CI, 6.9 to 17.8) at least 105 days, in adolescents age 16 to 17 years against infection. (VOC Omieron) BNT162b2 showed after 2st dose VE 12.5% (95% CI, 6.9 to 17.8) at least 105 days, in adolescents age 16 to 17 years against infection. (VOC Omieron)	design in England of adolescents age 12-17 years from week 37, 2021 onwards; there were 617,259 eligible tests for 12-15-year-olds and 225,670 for 16-17-year-olds. Symptomatic 12-15-year-olds and 16-17-year-olds with PCR-confirmed SARS-COV-2 infection was compared with vaccination status in symptomatic adolescents in the same age-groups who had a negative SARS-COV-2 PCR test. All cases prior to week 48 were defined as Delta, unless S gene target failure (SGTF), genotyping or sequencing information confirmed otherwise. Tests were defined as Omicron from week 48 onwards using SGTF, genotyping or sequencing information. Included in LES 8.2 Updated in LES 8.6 Link Updated in LES 8.8

				_
		least 14 days in adolescents age 12 to 15		
		years against infection. (VOC Delta)		
		BNT162b2 showed after 2 nd dose VE		
		83.1% (95% CI, 78.2 to 86.9) at 7-13 days		
		and VE 73% (95% CI, 66.4 to 78.3) at		
		least 14 days in adolescents age 12 to 15		
		years against infection. (VOC Omicron)		
		BNT162b2 showed after 2 nd dose VE		
		93.1% (95% CI, 91.6 to 94.4) at 7-13 days,		
		VE 96.1% (95% CI, 95.2 to 96.8) at 14-34		
		days, VE 91.5% (95% CI, 89.9 to 93.0) at		
		35-69 days, and VE 83.7% (95% CI, 72.0		
		to 90.5) at least 70 days in adolescents age		
		16 to 17 years against infection. (VOC		
		Delta)		
		BNT162b2 showed after 2 nd dose VE		
		76.1% (95% CI, 73.4 to 78.6) at 7-13 days,		
		VE 71.3% (95% CI, 69.3 to 73.1) at 14-34		
		days, VE 49.5% (95% CI, 45.7 to 53.0) at		
		35-69 days, and VE 22.6% (95% CI, 14.5		
		to 29.9) at least 70 days in adolescents age		
		16 to 17 years against infection. (VOC		
		Omicron)		
		BNT162b2 showed after 1st dose VE		
		14.2% (95% CI, -25.6 to 41.4) at 0-27 days,		
		and VE 83.4% (95% CI, 54.0 to 94.0) at		
		least 28 days in adolescents age 12 to 15		
		years against hospitalization. (VOC Delta)		
		BNT162b2 showed after 1st dose VE		
		64.6% (95% CI, 40.7 to 78.9) at 0-27 days,		
		and VE 76.3% (95% CI, 61.1 to 85.6) at		
		least 28 days in adolescents age 16 to 18		
		years against hospitalization. (VOC Delta)		
6	<u>Lutrick</u>	BNT162b2 showed VE 92% (95% CI, 79	Moderate	Prospective cohort in Arizona,
		to 97) against infection at least +14 days		of 243 adolescents aged 12–17
		after 2 nd dose in adolescents age 12 to 17		years between Jul 25 - Dec 4,
		years.		2021; 21,693 vaccinated person-
				days and 4,288 unvaccinated
				person-days; time and setting
				for VOC Delta.
7	71	DNTT4 (21.2 -1 - 1 VIE 040/ /050/ CL 74	· ·	Included in LES 8.3
7	<u>Zambrano</u>	BNT162b2 showed VE 84% (95% CI, 74	Serious	Test-negative case-control
		to 90) against MIS-C at least +28 days		design in 29 hospitals in 22
		after 2 nd dose in persons age 5 to 18 years.		states of U.S among hospitalized
		(VOC Delta to Omicron)		patients aged 5–18 years

	BNT162b2 showed VE 78% (95% CI, 48 to 90) against MIS-C at least +28 days after 2 nd dose in children age 5 to 11 years. (VOC Delta to Omicron) BNT162b2 showed VE 90% (95% CI, 81 to 95) against MIS-C at least +28 days after 2 nd dose in adolescents age 12 to 18 years. (VOC Delta to Omicron) BNT162b2 showed VE 94% (95% CI, 83 to 98) against MIS-C at least +28 days after 2 nd dose in adolescents age 12 to 18 years. (VOC Delta) BNT162b2 showed VE 92% (95% CI, 71 to 98) against MIS-C at least +28 days after 2 nd dose in adolescents age 12 to 18 years. (VOC Delta)		between Jul 1, 2021–Apr 7, 2022; 806 participants; VE was assessed by comparing the odds of being fully vaccinated with two doses of BNT162b2 vaccine versus being unvaccinated in MIS-C (case patients) compared to controls; time and setting for VOC Delta to VOC Omicron. Included in LES 8.3 Updated in LES 8.15
8 Klein	BNT162b2 showed after 2nd dose VE 74% (95% CI, -35 to 95) at 14-67 days, in children age 5 to 11 years against hospitalization. (VOC Delta to Omicron) BNT162b2 showed after 2nd dose VE 92% (95% CI, 79 to 97) at 14-149 days, in adolescents age 12 to 15 years against hospitalization. (VOC Delta to Omicron) BNT162b2 showed after 2nd dose VE 94% (95% CI, 87 to 97) at 14-149 days, in adolescents age 16 to 17 years against hospitalization. (VOC Delta to Omicron) BNT162b2 showed after 2nd dose VE 46% (95% CI, 24 to 61) at 14-67 days, in children age 5 to 11 years against symptomatic infection. (VOC Delta to Omicron) BNT162b2 showed after 2nd dose VE 83% (95% CI, 80 to 85) at 14-149 days, in adolescents age 12 to 15 years against symptomatic infection. (VOC Delta to Omicron) BNT162b2 showed after 2nd dose VE 76% (95% CI, 71 to 80) at 14-149 days, in adolescents age 16 to 17 years against	Serious	Test-negative case-control design in 10 states of the U.S among 39,217 emergency department (ED) and urgent care (UC) encounters and 1,699 hospitalizations among persons aged 5–17 years with COVID-19–like illness during April 9, 2021– January 29, 2022. VE was estimated comparing the odds of a positive SARS-CoV-2 test result between vaccinated (received at least 2 doses ≥14 days earlier or 3 doses ≥7 days earlier) and unvaccinated (received no doses) patients; time and setting for VOC Delta and VOC Omicron. Included in LES 8.7

				<u></u>
		symptomatic infection. (VOC Delta to		
		Omicron)		
		BNT162b2 showed after 3 rd dose VE 86% (95% CI, 73 to 93) at least 7 days, in adolescents age 16 to 17 years against symptomatic infection. (VOC Delta to Omicron)		
		BNT162b2 showed after 2 nd dose VE 92% (95% CI, 89 to 94) at 14-149 days, in adolescents age 12 to 15 years against symptomatic infection. (VOC Delta)		
		BNT162b2 showed after 2 nd dose VE 85% (95% CI, 81 to 89) at 14-149 days, in adolescents age 16 to 17 years against symptomatic infection. (VOC Delta)		
		BNT162b2 showed after 2 nd dose VE 51% (95% CI, 30 to 65) at 14-67 days, in children age 5 to 11 years against symptomatic infection. (VOC Omicron)		
		BNT162b2 showed after 2 nd dose VE 45% (95% CI, 30 to 57) at 14-149 days, in adolescents age 12 to 15 years against symptomatic infection. (VOC Omicron)		
		BNT162b2 showed after 2 nd dose VE 34% (95% CI, 8 to 53) at 14-149 days, in adolescents age 16 to 17 years against symptomatic infection. (VOC Omicron)		
		BNT162b2 showed after 3 rd dose VE 81% (95% CI, 59 to 91) at least 7 days, in adolescents age 16 to 17 years against symptomatic infection. (VOC Omicron)		
9	<u>Oliveira</u>	BNT162b2 showed after 1st dose VE 74% (95% CI, 18 to 92) at least 14 days, in adolescents age 12 to 18 years against infection. (VOC Delta)	Moderate	Matched case-control study in Connecticut (US) of 542 adolescents aged 12-18 years, including 186 case participants and 356 matched control
		BNT162b2 showed after 2 nd dose VE 90% (95% CI, 79 to 95) at least 14 days, VE 91% (95% CI, 33 to 99) at 7-28 days, VE 90% (95% CI, 67 to 97) at 35-56 days, VE 95% (95% CI, 79 to 99) at 63-84 days, and VE 83% (95% CI, 34 to 95) at 91-119		participants, between Jun 1 - Aug 15, 2021; time and setting for VOC Delta. Included in LES 8.8

		days, in adolescents age 12 to 18 years		
		against infection. (VOC Delta)		
		<u>.</u>		
		BNT162b2 showed after 2 nd dose VE 93%		
		(95% CI, 81 to 97) at least 14 days, in		
		adolescents age 12 to 18 years against		
		symptomatic infection. (VOC Delta)		
10	Molteni	BNT162b2 showed after 1st dose VE	Serious	Prospective cohort in the
		80.4% (95% CI, 78.5 to 82.2) at 14-30		United Kingdom using data
		days, VE 86.4% (95% CI, 83.5 to 88.7) at		from the Covid Symptom Study
		1-2 months (28 to 56 days), and VE 61.5%		(CSS), of 101,076 adolescents
		(95% CI, 43.5 to 73.7) at 2-3 months (56		aged 12-17 years, between Aug
		to 84 days), in adolescents age 12 to 17		05, 2021–Feb 14, 2022; time and
		years against infection. (VOC Delta)		setting for VOC Delta to VOC
		years against infection. (VOC Delta)		Omicron.
		BNT162b2 showed after 1st dose VE		
		53.7% (95% CI, 43.3 to 62.2) at 14-30		In the article, the effectiveness is
		days, VE 57.9% (95% CI, 50.9 to 63.9) at		presented as an adjusted relative
		1-2 months (28 to 56 days), and VE 63.7%		risk reduction obtained by
		3 //		RRR = (RR - 1) * 100, in the
		(95% CI, 59 to 67.9) at 2-3 months (56 to		present report it is transformed
		84 days), in adolescents age 12 to 17 years		for the reader's understanding.
		against infection. (VOC Omicron)		Included in LES 8.8
11	<u>Fowlkes</u>	BNT162b2 showed after 2 nd dose VE 81%	Moderate	Prospective cohort in four states
		(95% CI, 51 to 93) at least 14 days, and VE		of US (Arizona, Florida, Texas,
		87% (95% CI, 49 to 97) at 14-149 days, in		and Utah), of 1,364 participants
		adolescents age 12 to 15 years against		between Jul 2021–Feb 2022; the
		infection. (VOC Delta)		PROTECT cohort included
		,		1,052 children aged 5–11 years
		BNT162b2 showed after 2 nd dose VE 31%		and 312 adolescents aged 12–15
		(95% CI, 9 to 48) at 14-82 days, in children		years that were tested weekly for
		age 5 to 11 years against infection. (VOC		SARS-CoV-2; viral whole
		Omicron)		genome sequencing was
		3.2.2.3.3.4		assessed, time and setting for
		BNT162b2 showed after 2 nd dose VE 59%		VOC Delta to VOC Omicron.
		(95% CI, 24 to 78) at least 14 days, and VE		
		59% (95% CI, 22 to 79) at 14-149 days, in		Included in LES 8.8
		adolescents age 12 to 15 years against		
		infection. (VOC Omicron)		
12	Lawa	CoronaVac showed after 2 nd dose VE	Moderate	Dopulation based solvert in
14	<u>Jara</u>		Moderate	Population based cohort in
		38.2% (95% CI, 36.5 to 39.9) at least 14		Chile, of 490,694 children aged
		days, in children age 3 to 5 years against		3–5 years, between Dec 06,
		symptomatic infection. (VOC Omicron,		2021 - Feb 26, 2022; to estimate
		BA.1 sub-lineage)		the effectiveness of the
				complete primary immunization
1		0 II 1 1 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2		
		CoronaVac showed after 2 nd dose VE		schedule (two doses, 28 days
		64.6% (95% CI, 49.6 to 75.2) at least 14		schedule (two doses, 28 days apart) of an inactivated SARS-
		64.6% (95% CI, 49.6 to 75.2) at least 14 days, in children age 3 to 5 years against		schedule (two doses, 28 days apart) of an inactivated SARS-CoV-2 vaccine, CoronaVac;
		64.6% (95% CI, 49.6 to 75.2) at least 14		schedule (two doses, 28 days apart) of an inactivated SARS-

		CoronaVac showed after 2 nd dose VE 69% (95% CI, 18.6 to 88.2) at least 14 days, in children age 3 to 5 years against ICU admission. (VOC Omicron, BA.1 sub-		Included as <u>Araos</u> in LES 8.8 Updated in LES 8.13
13	Veneti	BNT162b2 showed after 1st dose VE 67.9 % (95% CI, 64.0 to 71.4) at 21-48 days, VE 55.8% (95% CI, 52.7 to 58.8) at 49-76 days, and VE 48.8% (95% CI, 46 to 51.5) at least 77 days, in adolescents age 12 to 15 years against infection. (VOC Delta) BNT162b2 showed after 1st dose VE 62.6 % (95% CI, 56.2 to 68) at 21-48 days, VE 47.3% (95% CI, 40 to 53.8) at 49-76 days, and VE 29.3% (95% CI, 20.4 to 37.1) at least 77 days, in adolescents age 16 to 17 years against infection. (VOC Delta) BNT162b2 showed after 2nd dose VE 90.8% (95% CI, 89.1 to 92.3) at 7-34 days, VE 92.8% (95% CI, 89.8 to 94.9) at 35-62 days, and VE 83.7% (95% CI, 75.9 to 89) at least 63 days, in adolescents age 16 to 17 years against infection. (VOC Delta) BNT162b2 showed after 1st dose VE 16.2 % (95% CI, -2.4 to 31.3) at 21-48 days, VE -1.3% (95% CI, -22.4 to 16.2) at 49-76 days, and VE -12.8% (95% CI, -21.7 to -4.6) at least 77 days, in adolescents age 12 to 15 years against infection. (VOC Omicron) BNT162b2 showed after 1st dose VE 33.7% (95% CI, -88.3 to 5.1) at 21-48 days, VE 16.8% (95% CI, -87.3 to 27.1) at 49-76 days, and VE -5.3% (95% CI, -32.9 to 16.6) at least 77 days, in adolescents age 12 to 15 years against infection. (VOC Omicron) BNT162b2 showed after 2nd dose VE 33.7% (95% CI, -87.3 to 27.1) at 49-76 days, and VE -5.3% (95% CI, -32.9 to 16.6) at least 77 days, in adolescents age 16 to 17 years against infection. (VOC Omicron) BNT162b2 showed after 2nd dose VE 53.1% (95% CI, 42.6 to 61.7) at 7-34 days, VE 45.7% (95% CI, 34.8 to 54.7) at 35-62 days, and VE 23.3% (95% CI, 2.7 to 39.5) at least 63 days, in adolescents age 16 to 17 years against infection. (VOC Omicron)	Moderate	Population-based cohort in Norway, of 372,179 adolescents aged 12-17 years, between Aug 25, 2021 – Jan 16, 2022; to estimate BNT162b2 one dose effectiveness for individuals 12-15 years old and one or two doses effectiveness for individuals 16-17 years old against SARS-CoV-2 infections; time and setting for VOC Delta to Omicron. Included in LES 8.9

15	<u>Price</u>	BNT162b2 showed after 2 nd dose VE 59% (95% CI, 23 to 79) at least 14 days in children and adolescents age 4 to 17 years against hospitalization. (VOC Delta to Omicron) BNT162b2 showed after 2 nd dose VE 93% (95% CI, 89 to 95) at 2–22 weeks in	Serious	Test-negative case-control design in 23 states of the U.S
		BNT162b2 showed after 2 nd dose VE 59% (95% CI, 23 to 82) at least 14 days in adolescents age 12 to 17 years against hospitalization. (VOC Delta to Omicron)		
		BNT162b2 showed after 1 st dose VE 37% (95% CI, -13 to 67) at least 14 days in children and adolescents age 4 to 17 years against hospitalization. (VOC Delta to Omicron)		and setting for VOC Delta to VOC Omicron. Included in LES 8.9
		BNT162b2 showed after 1 st dose VE 38% (95% CI, -51 to 79) at least 14 days in adolescents age 12 to 17 years against hospitalization. (VOC Delta to Omicron)		years, between May 28, 2021- Jan 10, 2022; to estimate the effectiveness of one and two mRNA vaccine doses at preventing hospitalization; time
14	Simmons	BNT162b2 showed after 1 st dose VE 32% (95% CI, -49 to 72) at least 14 days in children age 4 to 11 years against hospitalization. (VOC Delta to Omicron)	Serious	Age and time-matched nested case-control design in Ontario, Canada of 1,441 pediatric and adolescent patients aged 4-17
		BNT162b2 showed after 2 nd dose VE 90.7% (95% CI, 87.4 to 93.1) at 7-34 days, VE 92.3% (95% CI, 82.9 to 96.6) at 35-62 days, and VE 87.8% (95% CI, 78.8 to 92.9) at least 63 days, in adolescents age 16 to 17 years against infection. (VOC Delta to Omicron)		
		BNT162b2 showed after 1st dose VE 61.5% (95% CI, 57.1 to 65.5) at 21-48 days, VE 48% (95% CI, 43.3 to 52.4) at 49-76 days, and VE 47.5% (95% CI, 39 to 54.9) at least 77 days, in adolescents age 16 to 17 years against infection. (VOC Delta to Omicron)		
		BNT162b2 showed after 1st dose VE 65 % (95% CI, 62.3 to 67.6) at 21-48 days, VE 57.3% (95% CI, 54.4 to 60) at 49-76 days, and VE 70.2% (95% CI, 45.9 to 83.6) at least 77 days, in adolescents age 12 to 15 years against infection. (VOC Delta to Omicron)		

adolescents age 12 to 18 years against hospitalization. (VOC Delta) BNT162b2 showed after 2 nd dose VE 96% (95% CI, 90 to 98) at least 14 days in adolescents age 12 to 18 years against critical COVID-19. (VOC Delta) BNT162b2 showed after 2 nd dose VE 43% (95% CI, -1 to 68) at 2–22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 48% (95% CI, -2 to 68) at 2–22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against hospitalization. (VOC Omicron)	
BNT162b2 showed after 2nd dose VE 96% (95% CI, 90 to 98) at least 14 days in adolescents age 12 to 18 years against critical COVID-19. (VOC Delta) BNT162b2 showed after 2nd dose VE 43% (95% CI, -1 to 68) at 2-22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2nd dose VE 43% (95% CI, -1 to 68) at 2-22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	ged
BNT162b2 showed after 2nd dose VE 96% (95% CI, 90 to 98) at least 14 days in adolescents age 12 to 18 years against critical COVID-19. (VOC Delta) BNT162b2 showed after 2nd dose VE 43% (95% CI, -1 to 68) at 2–22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2nd dose VE 43% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	
(95% CI, 90 to 98) at least 14 days in adolescents age 12 to 18 years against critical COVID-19. (VOC Delta) BNT162b2 showed after 2nd dose VE 43% (95% CI, -1 to 68) at 2–22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2nd dose VE 43% case patients as compared controls; time and setting to the control of th	ainst
adolescents age 12 to 18 years against critical COVID-19. (VOC Delta) BNT162b2 showed after 2 nd dose VE 43% (95% CI, -1 to 68) at 2–22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	
critical COVID-19. (VOC Delta) BNT162b2 showed after 2 nd dose VE 43% (95% CI, -1 to 68) at 2–22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	
antecedent vaccination (further product of the second of t	ated
BNT162b2 showed after 2 nd dose VE 43% (95% CI, -1 to 68) at 2–22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	
(95% CI, -1 to 68) at 2–22 weeks in adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	ly
adolescents age 12 to 18 years against hospitalization. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	l) in
hospitalization. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	vith
BNT162b2 showed after 2 nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	or
BNT162b2 showed after 2 nd dose VE 68% (95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	cron.
(95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	
(95% CI, 42 to 82) at least 14 days, in children age 5 to 11 years against	
children age 5 to 11 years against	
BNT162b2 showed after 2 nd dose VE 79%	
(95% CI, 51 to 91) at least 14 days in	
adolescents age 12 to 18 years against	
critical COVID-19. (VOC Omicron)	
BNT162b2 showed after 2 nd dose VE 83%	
(95% CI, 77 to 88) at least 14 days in	
adolescents age 12 to 15 years against	
hospitalization. (VOC Delta to Omicron)	
BNT162b2 showed after 2 nd dose VE 82%	
(95% CI, 74 to 88) at least 14 days in	
adolescents age 16 to 18 years against	
hospitalization. (VOC Delta to Omicron)	
16 <u>Buchan</u> BNT162b2 showed after 2 nd dose VE 97% Moderate Test-negative case-control	
(95% CI, 94 to 99) at 7-59 days, and VE design in Ontario, Canada	
96% (95% CI, 94 to 97) at 60-119 days in among adolescents aged 12	_17
adolescents age 12 to 17 years against years during Nov 22, 2021	
symptomatic infection. (VOC Delta) 6, 2022, including 9,902	1,141
Omicron-positive cases wi	h
BNT162b2 showed after 2 nd dose VE 51% 19,953 test-negative control	
(95% CI, 38 to 61) at 7-59 days, and VE and 502 Delta-positive	-0,
31% (95% CI, 20 to 41) at 60-119 days in Cases with 17,930 test-neg	itive
adolescents age 12 to 17 years against controls. VE against	, .
symptomatic infection. (VOC Omicron) symptomatic infection and	
symptomatic infection. (VOC Officion) severe outcomes (i.e.,	
BNT162b2 showed after 3 rd dose VE 56% hospitalization or death) w	ıs
(95% CI, 34 to 70) at 0-6 days, and VE estimated over time since	~~
62% (95% CI, 49 to 72) at least 7 days in second or third dose receipt	
adolescents age 12 to 17 years against of BNT162b2; time and se	t
symptomatic infection. (VOC Omicron) for VOC Delta and VOC	
symptomize integral (100 cinition)	

		BNT162b2 showed after 2 nd dose VE 100% at 7-59 days, and VE 100% at 60-119 days in adolescents age 12 to 17 years against severe outcomes. (VOC Delta) (there were no cases of patients that presented severe outcomes) BNT162b2 showed after 2 nd dose VE 76% (95% CI, -10 to 95) at 7-59 days, and VE 83% (95% CI, 55 to 93) at 60-119 days in adolescents age 12 to 17 years against severe outcomes. (VOC Omicron)		Omicron, Delta outcomes were assessed prior to Jan 2, 2022. Included in LES 8.10
17	<u>Kildegaard</u>	BNT162b2 showed after 1st dose VE 62% (95% CI, 59 to 65) at 0-20 days in adolescents age 12 to 17 years against infection. (VOC Delta) BNT162b2 showed after 2nd dose VE 93% (95% CI, 93 to 94) at 0-59 days in adolescents age 12 to 17 years against infection. (VOC Delta)	Serious	Population-based cohort in Denmark, of adolescents aged 12-17 years, who were vaccinated before or on 1 October 2021; vaccine effectiveness was assessed in 229,799 adolescents after a first dose and 175,176 after a second dose of BNT162b2; time and setting for VOC Delta. <i>Included in LES 8.10</i>
18	<u>Chadeau-</u> <u>Hyam</u>	BNT162b2 showed after 1st dose VE 54.94% (95% CI, 40.98 to 65.6) at least 14 days in adolescents age 12 to 17 years against infection. (VOC Delta) BNT162b2 showed after 1st dose VE 58.56% (95% CI, 41.52 to 70.64) at least 14 days in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)	Serious	Surveillance study in England; 100,112 participants, including 14,974 (14.96%) adolescents aged 12 to 17 years; vaccine effectiveness was assessed after a first BNT162b2 dose comparing swab positivity among vaccinated and unvaccinated individuals; time and setting for VOC Delta. <i>Included in LES 8.11 Updated LES 8.12</i>
19	Britton	BNT162b2 showed after 2 nd dose VE 97% (95% CI, 95 to 98) at 14 days, VE 94% (95% CI, 94 to 95) at 14 - 60 days, VE 96% (95% CI, 95 to 97) at 14 - 30 days, VE 93% (95% CI, 92 to 94) at 31 - 60 days, VE 92% (95% CI, 91 to 93) at 61 - 90 days and VE 90% (95% CI, 88 to 91) at 91-120 days in adolescents age 12 to 15 years against symptomatic infection. (VOC Delta) BNT162b2 showed after 2 nd dose VE 94% (95% CI, 92 to 95) at 14 days, VE 90% (95% CI, 89 to 91) at 14 - 60 days, VE 94% (95% CI, 92 to 95) at 14 - 30 days,	Serious	Test-negative case-control design in U.S with data from 6884 US COVID-19 testing sites in the pharmacy-based Increasing Community Access to Testing platform, including 180,112 laboratory-based SARS-CoV-2 nucleic acid amplification tests from adolescents aged 12–19 years during Mar 13, – Oct 17, 2021; time and setting for VOC Delta. <i>Included in LES 8.11</i>

		VE 87% (95% CI, 85 to 89) at 31 - 60 days, VE 86% (95% CI, 84 to 87) at 61 - 90 days and VE 82% (95% CI, 80 to 83) at 91-120 days in adolescents age 16 to 19 years against symptomatic infection. (VOC Delta)		
		mRNA-1273 showed after 2 nd dose VE 99% (95% CI, 96 to 99) at 14 days, VE 94% (95% CI, 92 to 96) at 14 - 60 days, VE 98% (95% CI, 92 to 99) at 14 - 30 days, VE 91% (95% CI, 87 to 94) at 31 - 60 days, VE 85% (95% CI, 82 to 88) at 61 - 90 days and VE 85% (95% CI, 82 to 87) at 91-120 days in adolescents age 16 to 19 years against symptomatic infection. (VOC Delta)		
		AD26.COV 2.S showed after dose VE 52% (95% CI, 6 to 75) at 14 days, VE 54% (95% CI, 38 to 70) at 14 - 60 days, VE 58% (95% CI, 19 to 79) at 14 - 30 days, VE 52% (95% CI, 27 to 69) at 31 - 60 days, VE 63% (95% CI, 46 to 75) at 61 - 90 days and VE 58% (95% CI, 45 to 68) at 91-120 days in adolescents age 16 to 19 years against symptomatic infection. (VOC		
20	Dorabawila	BNT162b2 showed after 2 nd dose VE 68% (95% CI, 63 to 72) at Dec. 13-19, VE 57% (95% CI, 48 to 52) at Dec. 20-26, VE 50% (95% CI, 48 to 52) at Dec. 27-Jan 2, VE 48% (95% CI, 47 to 50) at Jan. 3-9, VE 34% (95% CI, 31 to 36) at Jan. 10-16, VE 20% (95% CI, 16 to 23) at Jan. 17-23 and VE 12% (95% CI, 6 to 16) at Jan. 24-30 in children age 5 to 11 years against infection. (VOC Delta to Omicron) BNT162b2 showed after 2 nd dose VE 85% (95% CI, 84 to 86) at Nov. 29- Dec 05, VE 82% (95% CI, 81 to 83) at Dec. 6-12, VE 66% (95% CI, 64 to 67) at Dec. 13-19, VE 57% (95% CI, 56 to 58) at Dec. 20-26, VE 55% (95% CI, 54 to 56) at Dec. 27-Jan 2, VE 53% (95% CI, 54 to 56) at Dec. 27-Jan 2, VE 50% (95% CI, 48 to 51) at Jan. 10-16, VE 50% (95% CI, 48 to 52) at Jan. 17-23 and VE 51% (95% CI, 48 to 54) at Jan. 24-30 in adolescents age 12 to 17 years against	Serious	Data-linkage study in New York state, U.S; that included 1,539,762 person days of children aged 5-11 years and 151,005 person days of children aged 12-17 years, to estimate BNT162b2 vaccine effectiveness against COVID cases and hospitalizations during Dec, 2021- Jan, 2022; time and setting for VOC Omicron. Included in LES 8.11
		infection. (VOC Delta to Omicron)		

BNT162b2 showed after 2nd dose VE 100% (95% CI, -189 to 100) at Dec. 13-19, VE 73% (95% CI, -7 to 97) at Dec. 20-26, VE 82% (95% CI, 36 to 96) at Jan. 3-9, VE 68% (95% CI, 28 to 91) at Jan. 10-16, VE 46% (95% CI, -15 to 77) at Jan. 17-23 and VE 48% (95% CI, -15 to 77) at Jan. 17-23 and VE 48% (95% CI, -15 to 77) at Jan. 17-23 and VE 48% (95% CI, -15 to 77) at Jan. 17-23 and VE 48% (95% CI, -15 to 77) at Jan. 17-23 and VE 48% (95% CI, -15 to 75) at Jan. 24-30 in children age 5 to 11 years against hospitalization. (VOC Delta to Omicron) BNT162b2 showed after 2nd dose VE 94% (95% CI, 76 to 99) at Nov. 29- Dec 05, VE 95% (95% CI, 64 to 100) at Dec. 6-12, VE 85% (95% CI, 63 to 95) at Dec. 13-19, VE 78% (95% CI, 63 to 84) at Dec. 20-26, VE 74% (95% CI, 63 to 84) at Dec. 27-Jan 2, VE 74% (95% CI, 63 to 82) at Jan. 3-9, VE 75% (95% CI, 64 to 86) at Jan. 10-16, VE 75% (95% CI, 61 to 83) at Jan. 17-23 and VE 73% (95% CI, 53 to 87) at Jan. 24-30 in adolescents age 12 to 17 years against hospitalization. (VOC Delta to Omicron) 21 Florentino CoronaVac showed after 1nd dose VE 99% (95% CI, -13.1 to -4.9) at 0 - 13 days, and VE 21.2% (95% CI, 18.6 to 23.8) at least 14 days in children age 6 to 11 years against symptomatic infection. (VOC Omicron) CoronaVac showed after 1nd dose VE 30.8% (95% CI, 24.2 to 36.8) at 0 - 13 days, and VE 39.8% (95% CI, 33.7 to 45.4) at least 14 days in children age 6 to 11 years against symptomatic infection. (VOC Omicron) CoronaVac showed after 1nd dose VE 27% (95% CI, -5.2 to 51.1) at 0 - 13 days, and VE 47.1% (95% CI, 26.6 to 62.7) at least 14 days in children age 6 to 11 years against hospitalization. (VOC Omicron) CoronaVac showed after 2nd dose VE 82.4% (95% CI, 44.2 to 97.1) at 0 - 13 days, and VE 47.1% (95% CI, 26.6 to 62.7) at least 14 days in children age 6 to 11 years against hospitalization. (VOC Omicron)	Moderate	Test-negative case-control design in Brazil, including 197,958 tests among children aged 6–11 years during Jan 21, 2022 – April 15, 2022, to assess CoronaVac effectiveness against symptomatic infection, hospitalization, and ICU admission; time and setting for VOC Omicron. Included in LES 8.11 Updated in LES 8.16
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		CoronaVac showed after 1st dose VE 20.2% (95% CI, -61.3 to 65.9) at 0 – 13 days, and VE 41.9% (95% CI, -10.4 to 72.2) at least 14 days in children age 6 to 11 years against ICU admission. (VOC Omicron) CoronaVac showed after 2nd dose VE 37.8% (95% CI, -147.7 to 93.2) at 0 – 13 days, and VE 20.9% (95% CI, -177.2 to 85) at least 14 days in children age 6 to 11 years against ICU admission. (VOC Omicron)		
22	Fleming- Dutra	BNT162b2 showed after 2 nd dose VE 60.1% (95% CI, 54.7 to 64.8) at 14 – 30 days, and VE 28.9% (95% CI, 24.5 to 33.1) at 30 - 90 days in children age 5 to 11 years against symptomatic infection. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 59.5% (95% CI, 44.3 to 70.6) at 14 – 30 days, VE 16.6% (95% CI, 8.1 to 24.3) at 30 - 90 days, and VE 9.6% (95% CI, -0.1 to 18.3) at 60 - 120 days in adolescents age 12 to 15 years against symptomatic infection. (VOC Omicron) BNT162b2 (3 doses) showed VE 71.1% (95% CI, 65.5 to 75.7) at 14 – 45 days in adolescents age 12 to 15 years against symptomatic infection. (VOC Omicron)	Serious	Test-negative case-control design in 49 states of the U.S among persons aged 5–15 years with COVID-19–like illness during Dec 26, 2021– Feb 21, 2022, including 74,208 tests from children 5 to 11 years of age and 47,744 tests from adolescents 12 to 15 years of age; VE was estimated comparing the odds of a positive SARS-CoV-2 test result between vaccinated (Two BNT162b2 doses 2 weeks or more before SARS-CoV-2 testing for children; 2 or 3 doses 2 weeks or more before testing for adolescents) and unvaccinated (received no doses) patients; time and setting for VOC Omicron. <i>Included in LES 8.12</i>
23	Florentino 1	BNT162b2 showed after 1st dose VE 52.4% (95% CI, 50.5 to 54.3) at least 14 days in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta, Brazil) BNT162b2 showed after 2nd dose VE 80.7% (95% CI, 77.8 to 83.3) at 14 – 27 days, VE 68% (95% CI, 63.2 to 72.3) at 28 – 41 days, VE 37.6% (95% CI, 27 to 46.7) at 42 – 55 days and VE 26.6% (95% CI, 4.1to 43.9) at 56 - 69 days in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta, Brazil)	Moderate	Test-negative case-control design in Brazil and Scotland among adolescents aged 12–17 years, including 503,776 adolescents from Brazil, and 127,168 adolescents from Scotland; VE was estimated comparing the odds of a positive SARS-CoV-2 test result between vaccinated and unvaccinated patients; time and setting for VOC Delta to VOC Omicron. Included in LES 8.12 Updated in LES 8.15

BNT162b2 showed after 1st dose VE 55.4% (95% CI, 53.4 to 57.3) at least 14 days in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta, Scotland)

BNT162b2 showed after 2nd dose VE 92.8% (95% CI, 85.7 to 96.4) at 14 – 27 days, VE 91.2% (95% CI, 81.8 to 95.8) at 28 – 41 days, VE 82.6% (95% CI, 63.9 to 91.6) at 42 – 55 days and VE 86.5% (95% CI, 72.2 to 93.4) at 56 - 69 days in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta, Scotland)

BNT162b2 showed after 1st dose VE 28% (95% CI, 26.3 to 29.7) at least 14 days in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron, Brazil)

BNT162b2 showed after 2nd dose VE 64.7% (95% CI, 63 to 66.3) at 14 – 27 days, VE 53% (95% CI, 51.3 to 54.7) at 28 – 41 days, VE 40.6% (95% CI, 38.8 to 42.4) at 42 – 55 days, VE 32% (95% CI, 30 to 33.9) at 56 - 69 days, VE 25.3% (95% CI, 22.9 to 27.6) at 70 - 83 days, VE 17% (95% CI, 13.8 to 20) at 84 - 97 days, and VE 5.9% (95% CI, 2.2 to 9.4) at least 98 days in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron, Brazil)

BNT162b2 showed after 1st dose VE 25.1% (95% CI, 21.3 to 28.7) at least 14 days in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron, Scotland)

BNT162b2 showed after 2nd dose VE 82.6% (95% CI, 80.6 to 84.5) at 14 – 27 days, VE 77.4% (95% CI, 74.7 to 79.8) at 28 – 41 days, VE 69.6% (95% CI, 66.3 to 72.6) at 42 – 55 days, VE 65.4% (95% CI, 61.9 to 68.7) at 56 - 69 days , VE 58% (95% CI, 52.9 to 62.6) at 70 - 83 days , VE 45.3% (95% CI, 37.2 to 52.4) at 84 - 97 days, and VE 50.6% (95% CI, 42.7 to 57.4)

Note: Due to the substantial heterogeneity found in the effectiveness data reported in this study, most of the results are only reported in this summary, not in the key findings tables.

24	Amir 1	at least 98 days in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron, Scotland) BNT162b2 showed after 1st dose VE 56.3% (95% CI, 45.9 to 64.6) at least 14 days in adolescents age 12 to 17 years against severe cases. (VOC Omicron, Brazil) BNT162b2 showed after 2nd dose VE 75.6% (95% CI, 58.1 to 85.8) at 14 – 27 days, VE 82.8% (95% CI, 72.1 to 89.4) at 28 – 41 days, VE 84.2% (95% CI, 76.3 to 89.5) at 42 – 55 days, VE 83.7% (95% CI, 76.3 to 89.5) at 42 – 55 days, VE 83.7% (95% CI, 76.6 to 88.9) at 56 – 69 days, VE 82% (95% CI, 72.6 to 88.2) at 70 – 83 days, VE 86.4% (95% CI, 75.2 to 92.6) at 84 – 97 days, and VE 82.7% (95% CI, 68.8 to 90.4) at least 98 days in adolescents age 12 to 17 years against severe cases. (VOC Omicron, Brazil) In children aged 5 to 10 years, being unvaccinated showed RR 2.4 (95% CI, 2.2, 2.6) of infection compared to BNT162b2 14 to 35 days after 2nd dose. (VOC Omicron, BA.1 sub-lineage) In children aged 5 to 10 years, BNT162b2 3 to 7 days after 1st dose showed RR 2.3	Moderate	Prospective cohort in the Israel using data from the Israeli Ministry of Health, of 1,444,406 Children aged 5-11 years and adolescents aged 12-17 years, between Dec 26, 2021-Jan 8, 2022; time and setting for VOC Omicron (BA.1 sub-lineage).
24	Amir 1	98 days in adolescents age 12 to 17 years against severe cases. (VOC Omicron, Brazil) In children aged 5 to 10 years, being unvaccinated showed RR 2.4 (95% CI, 2.2, 2.6) of infection compared to BNT162b2 14 to 35 days after 2 nd dose. (VOC Omicron, BA.1 sub-lineage)	Moderate	using data from the Israeli Ministry of Health, of 1,444,406 Children aged 5-11 years and adolescents aged 12-17 years, between Dec 26, 2021-Jan 8,
		In adolescents aged 12 to 15 years, BNT162b2 60 to 120 days after 2 nd dose showed RR 3.8 (95% CI, 3.3, 4.5) of		

	T	<u>, </u>		
		infection compared to BNT162b2 14 to 60		
		days after <u>3rd dose</u> . (VOC Omicron, BA.1		
		sub-lineage)		
		140 . 45		
		In adolescents aged 12 to 15 years,		
		BNT162b2 3 to 7 days after 3 rd dose		
		showed RR 3.3 (95% CI, 2.8, 4) of		
		infection compared to BNT162b2 14 to 60		
		days after <u>3rd dose</u> . (VOC Omicron, BA.1		
25	C 1 C:	sub-lineage)	· ·	D .: 1 .: 1 T 1
25	Cohen-Stavi	BNT162b2 showed after 1 st dose VE 17%	Serious	Prospective cohort in the Israel
		(95% CI, 7 to 25) at 14 – 27 days in		using data from the Clalit
		children age 5 to 11 years against infection.		Health Services and the Israeli
		(VOC Omicron)		Ministry of Health, of 136,127
		DNT140h2 showed - 6 2nd 1 - 3777 [4407		Children aged 5-11 years,
		BNT162b2 showed after 2 nd dose VE 51%		between Nov 23, 2021-Jan 7,
		(95% CI, 39 to 61) at 7 – 21 days, in		2022; time and setting for VOC
		children age 5 to 11 years against infection.		Omicron. Included in LES 8.14
		(VOC Omicron)		1nuucu in LES 0.14
		BNT162b2 showed after 1st dose VE 18%		
		(95% CI, -2 to 34) at 14 – 27 days in		
		children age 5 to 11 years against		
		symptomatic infection. (VOC Omicron)		
		symptomatic infection. (VOC Officion)		
		BNT162b2 showed after 2 nd dose VE 48%		
		(95% CI, 29 to 63) at 7 – 21 days, in		
		children age 5 to 11 years against		
		symptomatic infection. (VOC Omicron)		
26	Ionescu	BNT162b2 showed after 2 nd dose VE	Serious	Test-negative design in two
20	10110500	95.5% (95% CI, 95 to 96) at least 14 days,	Selious	provinces of Canada (Quebec
		VE 97.7% (95% CI, 96.2 to 98.6) at 14 –		and British Columbia) among
		27 days, VE 97% (95% CI, 96.3 to 97.6) at		adolescents aged 12–17 years,
		28 – 55 days, VE 96.1% (95% CI, 95.3 to		including 60,903 positive test
		96.7) at 56 – 83 days, VE 93.8% (95% CI,		and 193,899 controls, between
		92.7 to 94.8) at 84 - 111 days, and VE		Sep 05, 2021-Apr 30, 2022; VE
		92.4% (95% CI, 90.4 to 94) at 112 - 139		was estimated comparing the
		days in adolescents age 12 to 17 years		odds of a positive SARS-CoV-2
		against infection. (VOC Delta, Quebec)		test result between vaccinated
				and unvaccinated patients; time
		BNT162b2 showed after <u>2nd dose</u> VE		and setting for VOC Delta to
		95.7% (95% CI, 95.1 to 96.2) at least 14		VOC Omicron.
		days, VE 96.8% (95% CI, 94.4 to 98.2) at		Included in LES 8.14
		14 – 27 days, VE 96.7% (95% CI, 95.7 to		
		97.5) at 28 – 55 days, VE 96.2% (95% CI,		
		94.1 to 96.2) at 56 – 83 days, VE 95.2%		
		(95% CI, 94.1 to 96.2) at 84 - 111 days,		
		and VE 90.9% (95% CI, 87.7 to 93.2) at		
		112 - 139 days in adolescents age 12 to 17		

years against infection. (VOC Delta, British Columbia)

BNT162b2 showed after 2nd dose VE 97.3% (95% CI, 96.8 to 97.7) at least 14 days, in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta, Quebec)

BNT162b2 showed after 2nd dose VE 82.8% (95% CI, 81 to 84) at least 14 days, VE 83.1% (95% CI, 68.9 to 90.8) at 14 – 27 days, VE 88.2% (95% CI, 82.3 to 92.1) at 28 – 55 days, VE 84.3% (95% CI, 79.6 to 87.9) at 56 – 83 days, VE 87.6% (95% CI, 85.1 to 89.7) at 84 - 111 days, VE 82.7% (95% CI, 80.7 to 84.6) at 112 - 139 days, and VE 75.4% (95% CI, 72.1 to 78.4) at 140 - 167 days in adolescents age 12 to 17 years against infection. (VOC Delta to Omicron, Quebec)

BNT162b2 showed after 2nd dose VE 88% (95% CI, 85.1 to 90.3) at least 14 days, VE 94.8% (95% CI, 83.7 to 98.4) at 28 – 55 days, VE 87.8% (95% CI, 76.6 to 93.6) at 56 – 83 days, VE 91.6% (95% CI, 85.4 to 95.2) at 84 - 111 days, VE 86.5% (95% CI, 82.5 to 89.5) at 112 - 139 days, and VE 84.2% (95% CI, 77.8 to 88.8) at 140 - 167 days in adolescents age 12 to 17 years against infection. (VOC Delta to Omicron, British Columbia)

BNT162b2 showed after 2nd dose VE 87.9% (95% CI, 86.1 to 89.5) at least 14 days, in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta to Omicron, Quebec)

BNT162b2 showed after 2nd dose VE 41.9% (95% CI, 37.7 to 45.8) at least 14 days, VE 75.6% (95% CI, 65.8 to 82.6) at 14 – 27 days, VE 59.3% (95% CI, 50.9 to 66.3) at 28 – 55 days, VE 48.1% (95% CI, 39.9 to 55.1) at 56 – 83 days, VE 50.9% (95% CI, 44.9 to 56.3) at 84 - 111 days, VE 46% (95% CI, 40.9 to 50.7) at 112 - 139 days, VE 44.6% (95% CI, 40 to 49) at 140 - 167 days, and VE 33.9% (95% CI, 27.4

		,		
		to 39.9) at 168 - 195 days in adolescents		
		age 12 to 17 years against infection. (VOC		
		Omicron, Quebec)		
		DNT1(2h2 showed - 6 2nd 1 VIII		
		BNT162b2 showed after 2 nd dose VE		
		33.9% (95% CI, 25.7 to 41.1) at least 14		
		days, VE 63.4% (95% CI, 21.4 to 83) at 28		
		- 55 days, VE 57.7% (95% CI, 37.2 to		
		71.6) at 56 – 83 days, VE 40.8% (95% CI,		
		23.2 to 54.4) at 84 - 111 days, VE 37.7%		
		(95% CI, 22.7 to 49.7) at 112 - 139 days, VE 33.9% (95% CI, 24.1 to 42.2) at 140 -		
		167 days, and VE 22.2% (95% CI, 8.4 to		
		33.9) at 168 - 195 days in adolescents age		
		12 to 17 years against infection. (VOC		
		Omicron, British Columbia)		
		Officion, British Columbia)		
		BNT162b2 showed after 2 nd dose VE		
		55.2% (95% CI, 49.5 to 60.3) at least 14		
		days, in adolescents age 12 to 17 years		
		against symptomatic infection. (VOC		
		Omicron, Quebec)		
		3		
		BNT162b2 (<u>3 doses</u>) showed VE 63.7%		
		(95% CI, 41.1 to 77.7) at least 14 days in		
		adolescents age 12 to 17 years against		
		infection. (VOC Omicron, British		
		Columbia)		
27	Sacco	BNT162b2 showed after 1 st dose VE	Moderate	Data-linkage study in Italy; that
		27.4% (95% CI, 26.4 to 28.8) at least 14		included 2,965,918 children
		days in children age 5 to 11 years against		aged 5-11 years, to estimate
		infection. (VOC Omicron)		BNT162b2 vaccine
		_		effectiveness against SARS-
		BNT162b2 showed after <u>2nd dose</u> VE		CoV-2 infection and severe
		29.4% (95% CI, 28.5 to 30.2) at least 14		disease (Hospitalization or
		days, VE 38.7% (95% CI, 37.7 to 39.7) at 0		death) during Jan 17- Apr 13,
		- 14 days, VE 29.3% (95% CI, 28.1 to		2022; time and setting for VOC
		30.4) at 15 – 28 days, VE 23.1% (95% CI,		Omicron.
		21.7 to 24.5) at 29 – 42 days, and VE		Included in LES 8.14
		21.2% (95% CI, 19.7 to 22.7) at 43 – 84		
		days, in children age 5 to 11 years against infection. (VOC Omicron)		
		intection. (voc Officion)		
		BNT162b2 showed after 1 st dose VE		
		38.1% (95% CI, 20.9 to 51.5) at least 14		
		days in children age 5 to 11 years against		
		severe disease. (VOC Omicron)		
		(
1				
		BNT162b2 showed after 2 nd dose VE		

		days, in children age 5 to 11 years against		
28	<u>Tan</u>	severe disease. (VOC Omicron) BNT162b2 showed after 2 nd dose VE 36.8% (95% CI, 35.3 to 38.2) at least 7 days, VE 35.7% (95% CI, 33 to 38.2) at 1 - 6 days, VE 48.8% (95% CI, 46.9 to 50.8) at 7 – 14 days, VE 37.6% (95% CI, 35.7 to 39.3) at 15 – 29 days, VE 28.5% (95% CI, 26.3 to 30.7) at 30 – 59 days, and VE 25.6% (95% CI, 19.3 to 31.5) at least 60 days, in children age 5 to 11 years against infection. (VOC Omicron)	Serious	National cohort in Singapore, of 255,936 Children aged 5-11 years, to estimate BNT162b2 vaccine effectiveness against SARS-CoV-2 infection and hospitalization between Jan 21-Apr 8, 2022; time and setting for VOC Omicron. Included in LES 8.15
		BNT162b2 showed after 2 nd dose VE 65.3% (95% CI, 62 to 68.3) at least 7 days, VE 58.1% (95% CI, 51.9 to 63.5) at 1 - 6 days, VE 70.6% (95% CI, 65.9 to 74.7) at 7 – 14 days, VE 66.3% (95% CI, 61.7 to 70.2) at 15 – 29 days, VE 60.2% (95% CI, 54.1 to 65.5) at 30 – 59 days, and VE 42.7% (95% CI, 12 to 62.7) at least 60 days, in children age 5 to 11 years against symptomatic infection. (VOC Omicron)		
		BNT162b2 showed after 2 nd dose VE 82.7% (95% CI, 74.8 to 88.2) at least 7 days, VE 64.7% (95% CI, 37.3 to 80.2) at 1 - 6 days, VE 87.8% (95% CI, 72.2 to 94.7) at 7 – 14 days, VE 84.5% (95% CI, 72.7 to 91.2) at 15 – 29 days, and VE 80.4% (95% CI, 67 to 88.4) at 30 – 59 days in children age 5 to 11 years against hospitalization. (VOC Omicron)		
29	Lau	BNT162b2 showed after 1st dose VE 33.3% (95% CI, 3 to 53.3) at least 14 days in children age 3 to 11 years against infection. (VOC Omicron, BA.2 sublineage) BNT162b2 showed after 1st dose VE 26.1% (95% CI, -0.3 to 45.6) at least 14 days in adolescents age 12 to 18 years against infection. (VOC Omicron, BA.2 sub-lineage) BNT162b2 showed after 2nd dose VE	Serious	Ecological study in Hong Kong; of 953,400 participants between Jan 01–Apr 19, 2022; including 506,100 children aged 3–11 years and 447,300 adolescents aged 12–18 years; time and setting for VOC Omicron BA.2 <i>Included in LES 8.15</i>
		54.9% (95% CI, 38.9 to 66.8) at least 14 days in adolescents age 12 to 18 years against infection. (VOC Omicron, BA.2 sub-lineage)		

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mes (death or ime and setting on.
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21	Chamaitalle	BNT162b2 showed after 2nd dose VE 81% (95% CI, 64 to 90) at least 7 days, VE 94% (95% CI, 56 to 99) at 7 - 29 days, and VE 74% (95% CI, 44 to 88) at least 60 days, in children age 5 to 11 years against severe outcomes. (VOC Omicron)	Moderate	Droppostive cohert in Octor - 5
31	Chemaitelly	BNT162b2 showed after 2 nd dose VE 25.7% (95% CI, 10 to 38.6) at least 14 days, in children age 5 to 11 years against infection. (VOC Omicron, BA.1, BA.2, BA.4, BA.5 sub-lineages) BNT162b2 showed after 2 nd dose VE 30.6% (95% CI, 26.9 to 34.1) at least 14 days, in adolescents age 12 to 17 years against infection. (VOC Omicron, BA.1, BA.2, BA.4, BA.5 sub-lineages) BNT162b2 showed after 2 nd dose VE 87.6% (95% CI, 84 to 90.4) at least 14 days, in adolescents age 12 to 17 years against infection. (VOC Alpha, Beta and especially Delta)	Moderate	Prospective cohort in Qatar, of 119,896 persons, including 37,456 Children aged 5-11 years and 82,440 adolescents, to estimate BNT162b2 vaccine effectiveness against SARS-CoV-2 infection; time and setting for VOC Alpha to VOC Omicron. Included in LES 8.15
32	Tartof 1	BNT162b2 showed after 2nd dose VE 89% (95% CI, 69 to 96) at 56 days, VE 68% (95% CI, 46 to 81) at 56-112 days, VE 71% (95% CI, 57 to 81) at 112-168 days, and VE 49% (95% CI, 27 to 65) at least 168 days in adolescents age 12 to 17 years against Emergency Department or Urgent Care Encounters (Without Subsequent Hospitalization). (VOC Delta) BNT162b2 showed after 2nd dose VE 88% (95% CI, 68 to 96) at 56 days, VE 66% (95% CI, 44 to 80) at 56-112 days, VE 70% (95% CI, 56 to 80) at 112-168 days, and VE 47% (95% CI, 23 to 63) at least 168 days in adolescents age 12 to 17 years against Emergency Department or Urgent Care Encounters (Without Subsequent Hospitalization) Without Prior Documented SARS-CoV-2 Infection. (VOC Delta) BNT162b2 showed after 2nd dose VE 73% (95% CI, 54 to 84) at 56 days, VE 38% (95% CI, 54 to 84) at 56 days, VE 38% (95% CI, 14 to 56) at 56-112 days, VE 45% (95% CI, 28 to 57) at 112-168 days, and VE 16% (95% CI, -7 to 34) at least	Moderate	Test-negative design in USA among 3,168 adolescents aged 12 –17 years, members of Kaiser Permanente Southern California (KPSC) healthcare system between Nov 01, 2021 – Mar 18, 2022; VE was estimated against Emergency Department or Urgent Care Encounters; time and setting for VOC Delta to VOC Omicron. Included in LES 8.15

		168 days in adolescents age 12 to 17 years		
		against Emergency Department or Urgent		
		Care Encounters (Without Subsequent Hospitalization). (VOC Omicron)		
		Trospitalizationy. (VOC Chilefon)		
		BNT162b2 showed after 2 nd dose VE 72%		
		(95% CI, 52 to 84) at 56 days, VE 35% (95% CI, 9 to 54) at 56-112 days, VE 46%		
		(95% CI, 29 to 59) at 112-168 days, and		
		VE 18% (95% CI, -6 to 36) at least 168		
		days in adolescents age 12 to 17 years against Emergency Department or Urgent		
		Care Encounters (Without Subsequent		
		Hospitalization) Without Prior		
		Documented SARS-CoV-2 Infection. (VOC Omicron)		
		(VOC Officion)		
		BNT162b2 (<u>3 doses</u>) showed VE 87%		
		(95% CI, 72 to 94) at a median follow up of 19 days, in adolescents age 12 to 17		
		years against Emergency Department or		
		Urgent Care Encounters (Without		
		Subsequent Hospitalization). (VOC Omicron)		
		BNT162b2 (<u>3 doses</u>) showed VE 87%		
		(95% CI, 71 to 95) at a median follow up of 19 days, in adolescents age 12 to 17		
		years against Emergency Department or		
		Urgent Care Encounters (Without		
		Subsequent Hospitalization) Without Prior Documented SARS-CoV-2 Infection.		
		(VOC Omicron)		
33	<u>Tsang</u>	BNT162b2 showed after 1 st dose VE 32.4% (95% CI, -29 to 64.6) at least 14	Moderate	Prospective cohort in Hong Kong, China, of 8,636 persons,
		days in persons age 5 to 17 years against		including 886 Children and
		infection. (VOC Omicron, BA.2 sub-		adolescents aged 5-17 years,
		lineage)		between Mar 01 - Apr 15, 2022; to estimate BNT162b2 and
		BNT162b2 showed after <u>2nd dose</u> VE		CoronaVac vaccine
		3.2% (95% CI, -220.7 to 70.8) at 14 - 84		effectiveness against SARS-
		days in persons age 5 to 17 years against infection. (VOC Omicron, BA.2 sub-		CoV-2 infection; time and setting for VOC Omicron (BA.2
		lineage)		sub-lineage).
		,		Included in LES 8.16
		CoronaVac showed after 1 st dose VE 22.7% (95% CI, -38.3 to 56.8) at least 14		
		days in persons age 5 to 17 years against		
		infection. (VOC Omicron, BA.2 sub-		
		lineage)		

	1			
		CoronaVac showed after 2 nd dose VE 55.6% (95% CI, -50.3 to 86.9) at 14 - 84 days in persons age 5 to 17 years against infection. (VOC Omicron, BA.2 sublineage)		
34	Powell 1	BNT162b2 showed after 1st dose VE 8.5% (95% CI, 6.7 to 10.3) at 0-7 days (0 – 1 weeks), VE 59.4% (95% CI, 58.8 to 60) at 14-98 days (2-14 weeks), VE 23.5% (95% CI, 18.3 to 28.3) at 105-168 days (15-24 weeks), and VE 57.4% (95% CI, 39 to 70.2) at 175 – 273 days (25-39 weeks), in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)	Moderate	Test-negative design in England among 1,161,704 tests performed in adolescents aged 12 –17 years, between Aug 09, 2021 – Mar 31, 2022; VE was estimated against symptomatic infection; time and setting for VOC Delta to VOC Omicron. <i>Included in LES 8.16</i>
		BNT162b2 showed after 1st dose plus wild type prior infection VE 92.6% (95% CI, 90.4 to 94.3) at 0-7 days (0 – 1 weeks), VE 98.1% (95% CI, 97.6 to 98.6) at 14-98 days (2-14 weeks), and VE 98.6% (95% CI, 90.3 to 99.8) at 105-168 days (15-24 weeks), in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)		
		BNT162b2 showed after 1st dose plus Alpha prior infection VE 90.3% (95% CI, 88.4 to 92) at 0-7 days (0 – 1 weeks), VE 95.5% (95% CI, 94.8 to 96.1) at 14-98 days (2-14 weeks), and VE 94.2% (95% CI, 85.9 to 97.6) at 105-168 days (15-24 weeks), in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)		
		BNT162b2 showed after 1st dose plus Delta prior infection VE 91.3% (95% CI, 89.2 to 93) at 0-7 days (0 – 1 weeks), VE 97.5% (95% CI, 97 to 97.9) at 14-98 days (2-14 weeks), and VE 99% (95% CI, 92.8 to 99.9) at 105-168 days (15-24 weeks), in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)		
		BNT162b2 showed after 2 nd dose VE 71% (95% CI, 68.9 to 73) at 0-7 days (0 – 1 weeks), VE 91.8% (95% CI, 91.2 to 92.3) at 14-98 days (2-14 weeks), VE 80.9% (95% CI, 79.4 to 82.3) at 105-168 days (15-24 weeks), and VE 71.9% (95% CI, 67.9		

to 75.4) at 175 – 273 days (25-39 weeks), in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)

BNT162b2 showed after 2nd dose plus wild type prior infection VE 98.8% (95% CI, 96.7 to 98.8) at 14-98 days (2-14 weeks), VE 98.6% (95% CI, 94.3 to 99.7) at 105-168 days (15-24 weeks), and VE 94.8% (95% CI, 78.4 to 98.8) at 175 – 273 days (25-39 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)

BNT162b2 showed after 2nd dose plus Alpha prior infection VE 98.4% (95% CI, 95 to 99.5) at 0-7 days (0 – 1 weeks), VE 99.2% (95% CI, 97.8 to 99.7) at 14-98 days (2-14 weeks), VE 97% (95% CI, 92.7 to 98.8) at 105-168 days (15-24 weeks), and VE 97.3% (95% CI, 80.2 to 99.6) at 175 – 273 days (25-39 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)

BNT162b2 showed after 2nd dose plus Delta prior infection VE 99.6% (95% CI, 97.1 to 99.9) at 0-7 days (0 – 1 weeks), and VE 98.7% (95% CI, 96.8 to 99.4) at 14-98 days (2-14 weeks), in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)

BNT162b2 (2 doses) followed by any mRNA vaccine showed VE 84.8% (95% CI, 77.6 to 89.7) at 0-7 days (0 – 1 weeks), and VE 96% (95% CI, 92.2 to 97.9) at 14-98 days (2-14 weeks), in adolescents age 12 to 17 years against symptomatic infection. (VOC Delta)

BNT162b2 showed after 1st dose VE 15.2% (95% CI, 9.9 to 20.1) at 0-7 days (0 – 1 weeks), VE 18.8% (95% CI, 17.2 to 20.3) at 14-98 days (2-14 weeks), VE 17.9% (95% CI, 14.9 to 20.7) at 105-168 days (15-24 weeks), and VE 12.8% (95% CI, -1.6 to 25.1) at 175 – 273 days (25-39 weeks), in adolescents age 12 to 17 years

against symptomatic infection. (VOC Omicron)

BNT162b2 showed after 1st dose plus wild type prior infection VE 69.2 % (95% CI, 55.9 to 78.5) at 0-7 days (0 – 1 weeks), VE 85.3 % (95% CI, 83.7 to 86.8) at 14-98 days (2-14 weeks), VE 73.4 % (95% CI, 67.2 to 78.4) at 105-168 days (15-24 weeks), and VE 67.8 % (95% CI, 24.1 to 86.3) at 175 – 273 days (25-39 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)

BNT162b2 showed after 1st dose plus Alpha prior infection VE 77.6% (95% CI, 69.5 to 83.6) at 0-7 days (0 – 1 weeks), VE 81.5% (95% CI, 80.0 to 82.9) at 14-98 days (2-14 weeks), VE 69.5% (95% CI, 64.5 to 73.8) at 105-168 days (15-24 weeks), and VE 66.7% (95% CI, 35.2 to 82.9) at 175 – 273 days (25-39 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)

BNT162b2 showed after 1st dose plus Delta prior infection VE 79.3% (95% CI, 76.7 to 81.6) at 0-7 days (0 – 1 weeks), VE 78.8% (95% CI, 77.9 to 79.5) at 14-98 days (2-14 weeks), VE 67.2% (95% CI, 63.7 to 70.3) at 105-168 days (15-24 weeks), and VE 55.8% (95% CI, 17.2 to 76.4) at 175 – 273 days (25-39 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)

BNT162b2 showed after 1st dose plus Omicron prior infection VE 79.6 % (95% CI, 44.9 to 92.4) at 14-98 days (2-14 weeks), in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)

BNT162b2 showed after 2nd dose VE 52.2 % (95% CI, 50.4 to 53.9) at 0-7 days (0 – 1 weeks), VE 64.5% (95% CI, 63.6 to 65.4) at 14-98 days (2-14 weeks), VE 29.8% (95% CI, 24.9 to 34.2) at 105-168 days (15-24 weeks), and VE 19.4% (95% CI, 11.7 to 26.4) at 175 – 273 days (25-39 weeks), in

adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)

BNT162b2 showed after 2nd dose plus wild type prior infection VE 87.4% (95% CI, 83.5 to 90.4) at 0-7 days (0 – 1 weeks), VE 84.7% (95% CI, 82.6 to 86.5) at 14-98 days (2-14 weeks), VE 53.4% (95% CI, 32.7 to 67.7) at 105-168 days (15-24 weeks), and VE 28.9% (95% CI, -15.5 to 56.3) at 175 – 273 days (25-39 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)

BNT162b2 showed after 2nd dose plus Alpha prior infection VE 84.9% (95% CI, 81.3 to 87.8) at 0-7 days (0 – 1 weeks), VE 85.5% (95% CI, 84 to 86.9) at 14-98 days (2-14 weeks), VE 64.3% (95% CI, 52.4 to 73.3) at 105-168 days (15-24 weeks), and VE 63.6% (95% CI, 46 to 75.5) at 175 – 273 days (25-39 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)

BNT162b2 showed after 2nd dose plus Delta prior infection VE 82.1% (95% CI, 80.1 to 83.9) at 0-7 days (0 – 1 weeks), VE 83.5% (95% CI, 82.5 to 84.5) at 14-98 days (2-14 weeks), and VE 75.5% (95% CI, 65.6 to 82.5) at 105-168 days (15-24 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)

BNT162b2 (2 doses) followed by any mRNA vaccine showed VE 55.1% (95% CI, 50.7 to 59.1) at 0-7 days (0 – 1 weeks), VE 62.9% (95% CI, 60.5 to 65.1) at 14-98 days (2-14 weeks), and VE 33.6% (95% CI, 14.6 to 48.3) at 105-168 days (15-24 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)

BNT162b2 (2 doses) followed by any mRNA vaccine dose plus Wild type prior infection showed VE 77.7% (95% CI, 55.7 to 88.8) at 0-7 days (0 – 1 weeks), and VE 79.8% (95% CI, 70.4 to 86.3) at 14-98 days (2-14 weeks) in adolescents age 12 to 17

	1			
		years against symptomatic infection. (VOC Omicron)		
		BNT162b2 (2 doses) followed by any mRNA vaccine dose plus Alpha prior infection showed VE 82.2% (95% CI, 68.1 to 90.1) at 0-7 days (0 – 1 weeks), and VE 79.6% (95% CI, 71.4 to 85.5) at 14-98 days (2-14 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)		
		BNT162b2 (2 doses) followed by any mRNA vaccine dose plus Delta prior infection showed VE 89.5% (95% CI, 81.7 to 94) at 0-7 days (0 – 1 weeks), and VE 80.7% (95% CI, 71.1 to 87.1) at 14-98 days (2-14 weeks) in adolescents age 12 to 17 years against symptomatic infection. (VOC Omicron)		
35	Cocchio	BNT162b2 showed after 2nd dose VE 83% (95% CI, 76 to 88) at 0-6 days, VE 84% (95% CI, 77 to 89) at 7-13 days, VE 88% (95% CI, 85 to 91) at 14-34 days, VE 83% (95% CI, 79 to 87) at 35 – 69 day, and VE 82% (95% CI, 74 to 88) at least 70 days in adolescents age 12 to 17 years against infection. (VOC Delta) BNT162b2 showed after 2nd dose VE 72% (95% CI, 69 to 74) at 0-6 days, VE 70% (95% CI, 67 to 72) at 7-13 days, VE 53% (95% CI, 51 to 55) at 14-34 days, VE 22% (95% CI, 19 to 24) at 35 – 69 day, and VE 23% (95% CI, 20 to 26) at least 70 days in children age 5 to 11 years against infection. (VOC Omicron) BNT162b2 showed after 2nd dose VE 81% (95% CI, 76 to 85) at 0-6 days, VE 83% (95% CI, 79 to 86) at 7-13 days, VE 59% (95% CI, 55 to 62) at 14-34 days, VE 23% (95% CI, 19 to 27) at 35 – 69 day, and VE 8% (95% CI, 5 to 11) at least 70 days in adolescents age 12 to 17 years against infection. (VOC Omicron)	Serious	Data-linkage study in Veneto region, Italy; of 430,584 participants, including 193,509 children aged 5-11 years and 237,075 adolescents aged 12 – 17 years, to estimate BNT162b2 and mRNA-1273 vaccine effectiveness against SARS-CoV-2 infection during two periods, Aug 01- Oct 21, 2021 and Feb 01- Apr 27, 2022; time and setting for VOC Delta and VOC Omicron. Included in LES 8.16
		BNT162b2 (3 doses) showed VE 79% (95% CI, 77 to 81) at 0-6 days, VE 80% (95% CI, 78 to 82) at 7-13 days, VE 72%		

(95% CI, 70 to 73) at 14-34 days, and VE 30% (95% CI, 27 to 33) at 35 – 69 days in adolescents age 12 to 17 years against infection. (VOC Omicron)		
mRNA-1273 showed after 2 nd dose VE 90% (95% CI, 69 to 97) at 0-6 days, VE 90% (95% CI, 68 to 97) at 7-13 days, and VE 96% (95% CI, 86 to 99) at 14-34 days in adolescents age 12 to 17 years against infection. (VOC Delta)		
mRNA-1273 showed after 2 nd dose VE 88% (95% CI, 81 to 92) at 0-6 days, VE 78% (95% CI, 69 to 84) at 7-13 days, VE 55% (95% CI, 49 to 61) at 14-34 days, VE 29% (95% CI, 23 to 35) at 35 – 69 day, and VE 20% (95% CI, 15 to 24) at least 70 days in adolescents age 12 to 17 years against infection. (VOC Omicron)		

Section 2: excluded studies					
Author	Reason for exclusion	Version of exclusion			
Tang	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.1			
Naleway	Did not report results according to vaccine type	Excluded in LES 8.1			
Chadeau-Hyam round 14	Vaccine effectiveness not reported	Excluded in LES 8.1			
de Gier	Did not report results according to vaccine type	Excluded in LES 8.2			
Delahoy	Did not report results according to vaccine type	Excluded in LES 8.2			
Lin	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.2*			
McLean	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.2			
<u>Amir</u>	Critical risk of bias	Excluded in LES 8.3			
Chung	Did not report the vaccine effectiveness in <18 years, Did not report results according to vaccine type	Excluded in LES 8.3*			
<u>Fisman</u>	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.3			
Lyngse	Did not report results according to vaccine type	Excluded in LES 8.3			
<u>Prunas</u>	Critical risk of bias	Excluded in LES 8.3			
Chiew	Critical risk of bias	Excluded in LES 8.3			
Elliot	Critical risk of bias	Excluded in LES 8.4			
New York State Department of Health	Did not report results according to vaccine type	Excluded in LES 8.4			
Andeweg	Did not report results according to vaccine type	Excluded in LES 8.5*			
<u>Jalali</u>	Did not report results according to vaccine type	Excluded in LES 8.5*			

<u>Choe</u>	Critical risk of bias	Excluded in LES 8.6
<u>Madhi</u>	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.6
<u>De Serres</u>	Did not report results according to vaccine type	Excluded in LES 8.7
Nyberg	Did not report results according to vaccine type	Excluded in LES 8.7
Hoeg	Clinical outcomes of interest for this LES not reported	Excluded in LES 8.7
<u>Levi</u>	Did not report results according to vaccine type	Excluded in LES 8.7
<u>Nygaard</u>	Critical risk of bias	Excluded in LES 8.8
Chemaitelly 1	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.8*
<u>AlHosani</u>	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.8
Ng	Vaccine effectiveness not reported	Excluded in LES 8.8
<u>Petrie</u>	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.10
<u>González</u>	Critical risk of bias	Excluded in LES 8.1 <mark>1*</mark>
Carazo	Did not report results according to vaccine type	Excluded in LES 8.11
Rennert	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.12
<u>Braeye</u>	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.12
<u>Fano</u>	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.13
<u>Topfner</u>	Vaccine effectiveness not reported	Excluded in LES 8.13
<u>Mattiuzzi</u>	Did not report results according to vaccine type	Excluded in LES 8.13
<u>Haile</u>	Vaccine effectiveness not reported	Excluded in LES 8.13
<u>Andrejko</u>	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.13
<u>Spicer</u>	Did not report results according to vaccine type	Excluded in LES 8.13
<u>Husin</u>	Critical risk of bias	Excluded in LES 8.13
<u>Lytras</u>	Did not report the vaccine effectiveness in <18 years	Excluded in LES 8.13
<u>Shi</u>	Vaccine effectiveness not reported	Excluded in LES 8.14
<u>Tonnara</u>	Did not report results according to vaccine type	Excluded in LES 8.14
<u>De Lemos</u>	Did not report results according to vaccine type	Excluded in LES 8.15
Ziv	Critical risk of bias	Excluded in LES 8.15
Westerhof	Vaccine effectiveness not reported	Excluded in LES 8.16
Sumner	Did not report results according to vaccine type	Excluded in LES 8.16
<u>Lau 1</u>	Vaccine effectiveness not reported	Excluded in LES 8.16
<u>Kim</u>	Critical risk of bias	Excluded in LES 8.16
Andeweg	Did not report results according to vaccine type	Excluded in LES 8.16

^{*} For this studies links have been updated after their exclusion

Appendix 2: Glossary (revised 13 Jan 2022)

AZ: AstraZeneca

Alpha: variant of concern B.1.1.7

Beta: variant of concern B.1.351

Delta: variant of concern B.1.617.2

Gamma: variant of concern P.1

Epsilon: variant of concern B.1.427/B.1.429

MIS-C: Multisystem inflammatory syndrome in children

MOD: Moderna

Obs: observational study

OR: odds ratio

PF: Pfizer

RME: range of mean estimates across 2 or more studies

VE (Vaccine effectiveness): measure of how well a vaccine protects people from getting the outcome of interest in real-world practice (For example: VE of 92% against infection means that 92% of people will be protected from becoming infected with COVID and 8% of people will still be at risk of becoming infected with COVID)

VET: vaccine effectiveness against transmission

VOC: variant of concern

VOI: variant of interest

Appendix 3: Data-extraction template (revised 13 Jan 2022)

Vaccine product	
Source	First author of study
Link	DOI or PubMed ID
Date published	in format YYYY/MM/DD or preprint
Country	
Funding	public or industry
Study details	
Study type	RCT/cohort/data-linkage/test-negative/case-control/other
Surveillance	routine screening Y or N
Intervention	Pfizer/Comirnaty [BNT162b2]/Moderna/Spikevax [mRNA-1273]/AstraZeneca/Vaxzevria [ChAdOx1]/Johnson & Johnson [AD26.COV2.S]/Sinovac [CoronaVac]/Sinopharm (Wuhan) [WIV04]/Novavax [NVX-CoV2373]/FBRI [EpiVacCorona]/Bharat Biotech [Covaxin] [BBV152]/Gamaleya [Sputnik V] [Gam-COVID-Vac]
Dose and timing	
Control group	not vaccinated, <7day vaccinated internal control, none, other
Total (N)	number of all study participants
Female	number or %
< 12 years	number or %
≥ 12 years	number or %
Outcomes	outcomes separated by VOC type
Outcomes	confirmed infection/asymptomatic/mild symptomatic/severe symptoms/hospitalized/ICU/death/MIS-C
1st Dose VE	VE with 95% CI
Days post 1st dose	days post 1st dose when VE provided
2nd Dose VE	VE with 95% CI
Days post 2nd dose	days post 2nd dose when VE provided
Rates per X person- days/years	vaccinated vs control
HR	vaccinated vs control
RR	vaccinated vs control
Adjusted	Regression, stratification, matching and associated variables
Transmission	infection rates in unvaccinated contacts of vaccinated individuals
Critical appraisal	See Appendix 5

Appendix 4: Process for assigning Variant of Concern to studies

A Variant of Concern is considered to be the dominant (≥50%) strain in a study if any of the following conditions apply:

- i) the authors make a statement about prevalence of VOC during the study time frame
- ii) time and setting of the study is consistent with a VOC being dominant according to the following open tracking sources:

Nextstrain. Real-time tracking of pathogen evolution. https://nextstrain.org/ Outbreak Info. https://outbreak.info/location-reports

Appendix 5: Research question and critical appraisal process (revised 13 Jan 2022)

Review question:

Participants	People aged under 18 years at risk of COVID-19 (usually without but sometimes with previous COVID-19 infection)
	sometimes with previous COVID-19 infection)
Intervention	COVID-19 Vaccine
Comparator	Unvaccinated children and adolescents (*)
Outcomes	PCR-diagnosis of COVID-19 infection; symptomatic disease; hospital/ICU
	admission; death; transmission; MIS-C

^(*) Eligible studies must have a comparison group (unvaccinated; non-immune period; time since vaccination; 2 doses vs 3 doses); before-after studies, where the infection rate in the first 2 weeks after the vaccination are used as control are commonly performed and may be appraised

Key exclusion criteria

Studies that address the question of interest but from which the information of children cannot be separated from that of adults.

Comparison of one vaccine vs another (e.g., relative effectiveness) is not eligible. Studies reporting only antibody responses are excluded.

Critical Appraisal Process

We appraise the quality of the individual studies using an adapted version of ROBINS-I. This tool classifies the Risk of Bias of a study as **Low, Moderate, Serious, Critical, or No Information**. <u>Low Risk of Bias indicates High Quality, and Critical Risk of Bias indicates Very Low (insufficient) Quality</u>. ROBINS-I appraises 7 bias domains and judges each study against an ideal reference randomized controlled trial. To improve the utility of ROBINS-I for assessing studies reporting vaccine effectiveness, we have focused on study characteristics that introduce bias as reported in the vaccine literature. (WHO. Evaluation of COVID-19 vaccine effectiveness. Interim Guidance. 17 March 2021). Studies rated as "critical" risk of bias will not be included in the Summary statements on Page 1-2 (exception: if limited data available for an outcome for a VOC). An overall judgement of "serious" or "critical" is given when the study is judged to be at serious or critical risk of bias in at least one domain or "serious" in 3 separate ROBINS-I domains.

VE Study	Description
Characteristics that	
may introduce bias	
Study design	In cohort studies, people who get vaccinated may differ in health-seeking
	behaviour from people who do not get vaccinated; using a test-negative study
ROBINS-I: Bias in	design minimizes this type of bias
selection of participants	
into study	Examples and typical judgement:
	• test-negative design with a clearly defined symptomatic study population (low)
	• test-negative design (mixed or unclear study population) or case-control
	or cohort design or data-linkage with no concerns (moderate)
	• cross-sectional design or case-control (concerns about whether controls
	had same access to vaccines/risk of exposure to COVID or unclear) or
	cohort design (concerns that exposed and non-exposed were not drawn
	from the same population) (serious)

Method for confirming	Questionnaires are prone to recollection bias; Population databases
vaccination	
vaccination	developed for purpose of tracking COVID vaccines minimize this type of
DODING L D.	bias
ROBINS-I: Bias in	
classification of	Examples and typical judgement:
interventions	database linkage study (low)
	• Questionnaire with confirmation by an additional method (e.g., registry)
	of at least a subset of study population (moderate)
	• Questionnaire without confirmation by an additional method (serious)
	Estimating vaccination status based on surveillance data alone (critical)
Databases used for	Databases developed for collecting data on COVID are less prone to bias
retrieval of COVID test	due to missing information and misclassification
results, participant	
prognostic factors, and	Examples and typical judgement:
clinical outcomes	database for non-COVID purpose but with individual level data
	(moderate)
ROBINS-I: Bias in	• database for non-COVID purpose without individual level data (serious)
classification of	• no or unclear description of database type (critical)
interventions	
Assignment of	Using date of symptom onset (if within 10 days of testing) as infection start
infection start date	date reduces risk of misclassification bias (e.g., vaccinated participant who is
	reported as COVID+ may have been infected prior to receiving the vaccine
ROBINS-I: Bias in	or during non-immune period) and sensitivity of assays decreases over time
classification of	
interventions	Examples and typical judgement:
	using a PCR positive test that was part of an ongoing standardized
	monitoring system (e.g., within a health network) (low)
	• using sample date without interview or documented confirmation of
	symptoms ≤ 10 days (relevant for symptomatic disease only) (serious)
Verification of	Prospective, standardized collection of symptoms from patients reduces risk
symptoms	of missing information bias; testing within 10 days after symptom onset
-y P	reduces risk of false-negative COVID test
ROBINS-I: Bias in	3000000
classification of	Examples and typical judgement:
interventions	 using sample date without patient report/ documented confirmation of
	symptoms ≤ 10 days (relevant for symptomatic disease only) (serious)
	 if symptomatic COVID is not an outcome (no information)
Accounting for non-	Reported absence of vaccine effect during non-immune period reduces risk
immune period (first 14	of residual confounding bias
days after first vaccine	or residual confounding only
dose)	Example/common case:
	 presence of an effect during non-immune period or result not reported
ROBINS-I: Bias due to	(moderate)
confounding	 unclear that non-immune period was considered (serious)
Inclusion of	Exclusion (or separate analysis) of participants with prior COVID infection
participants with prior	reduces concern about differences in infectivity as well as risk-taking and
COVID infection	health-seeking behaviour
COVID IIIICCIOII	incarui-seekiiig beliavioui
ROBINS-I: Bias due to	Examples and typical judgement:
confounding	• inclusion of prior infection status as a covariate in the models (moderate)

	previously infected not excluded or analyzed separately (serious)
Accounting for	Accounting for calendar time reduces bias due to differences in vaccine
calendar time	accessibility and risk of exposure over time
ROBINS-I: Bias due to	Examples and typical judgement:
confounding (time-	• use of time-varying statistics without explicit mention of adjustment for
varying confounding)	calendar time (moderate)
	• not taken into account but short-time frame (e.g., ≤2 months) (serious)
	• not taken into account and time frame >2 months (critical)
Adjustment for	Adjustment for prognostic factors for COVID infection, severity of disease,
prognostic factors	and vaccination, such as age, gender, race, ethnicity, socioeconomic factors,
	occupation (HCW, LTC), and chronic medical conditions
ROBINS-I: Bias due to	
confounding	Examples and typical judgement:
	• no or insufficient adjustment for occupation (or number of tests as a
	surrogate for exposure risk) -exception age>65 or LTCF resident
	(moderate)
	• no or insufficient adjustment for socioeconomic factors (or neighborhood
	or income as a surrogate), race, ethnicity (serious)
	• no or insufficient adjustment for age (any study population) or chronic
Taking Conservation	medical conditions (LTC)(critical)
Testing frequency	Similar frequency of testing between groups reduces risk of bias introduced
ROBINS-I: Bias in	by detecting asymptomatic infection in one group but not in another (e.g.,
measurement of	when only one group undergoes surveillance screening)
outcomes	Examples and typical judgement:
outcomes	 no systematic screening but consistent methods for detection in one
	group vs. the other, e.g., within health networks (moderate)
	 screening performed for a subset of both study groups (serious)
	 screening performed routinely in one study group but not in the other
	(critical)
	(Cirucar)

Appendix 6: Detailed description of the narrative summary statement (revised 20 Jun 2022)

We include studies with the following clinical outcomes: prevention of infection, MIS-C, severe disease (as defined by the study investigators), hospitalization, death, and prevention of transmission. These outcomes were selected because they are less susceptible to bias, or they are important for parents and patients. If data are not available for these specific outcomes, but are available for symptomatic infection, data for these additional outcomes are provided temporarily.

We aim at providing a lay language, standardized summary statement for each combination of vaccine and VOC for which we found evidence.

Where more than one study was found, we will provide a summary statement with a <u>range of the</u> estimates across the studies.

Where a <u>single study</u> provided data, we will provide the <u>estimate plus 95% confidence interval</u> for that study. As additional studies are added, the estimate plus confidence interval will be replaced by a range as described above.

In the summaries, "prevented" or "protects" will be applied to mean estimates that are greater than or equal to 70% with the lower 95% $CI \ge 50\%$, or range of mean estimates that are greater than or equal to 70% for infection and, mean estimates that are greater than or equal 90% with lower limit of 95% $CI \ge 70\%$, or range of mean estimates that are greater than or equal to 90% for severe disease (the lowest acceptable limit for vaccine effectiveness as determined by WHO); otherwise "did not reach threshold for protection" will be applied.