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**Function: Head of the clinical vaccine research (CIC Cochin Pasteur) and I Reivac (French clinical research network dedicated to vaccinology)**

**Main expertise: vaccination in particular populations, respiratory infections, emerging infections**



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DI SCIENZE DELLA SALUTE



# mRNA Covid-19 vaccines for elderly

Odile Launay

AIB technical meeting  
Warsaw, 8 May 2025

# Disclosures

- MSD, GSK bio, Sanofi Pasteur, Janssen, Pfizer, AstraZeneca, Moderna

# Covid-19 mortality

## 2 years after the start of the pandemic

Estimating excess mortality due to the COVID-19 pandemic:  
a systematic analysis of COVID-19-related mortality, 2020–21

COVID-19 Excess Mortality Collaborators\*



- Estimated excess mortality:  
**18.2 million (95% CI: 17.1-19.6)**
- Rate of 120.3/100,000 person-years
- Highest rates (> 400/100,000 PY): India, USA, Russia, Mexico, Brazil, Indonesia, Pakistan

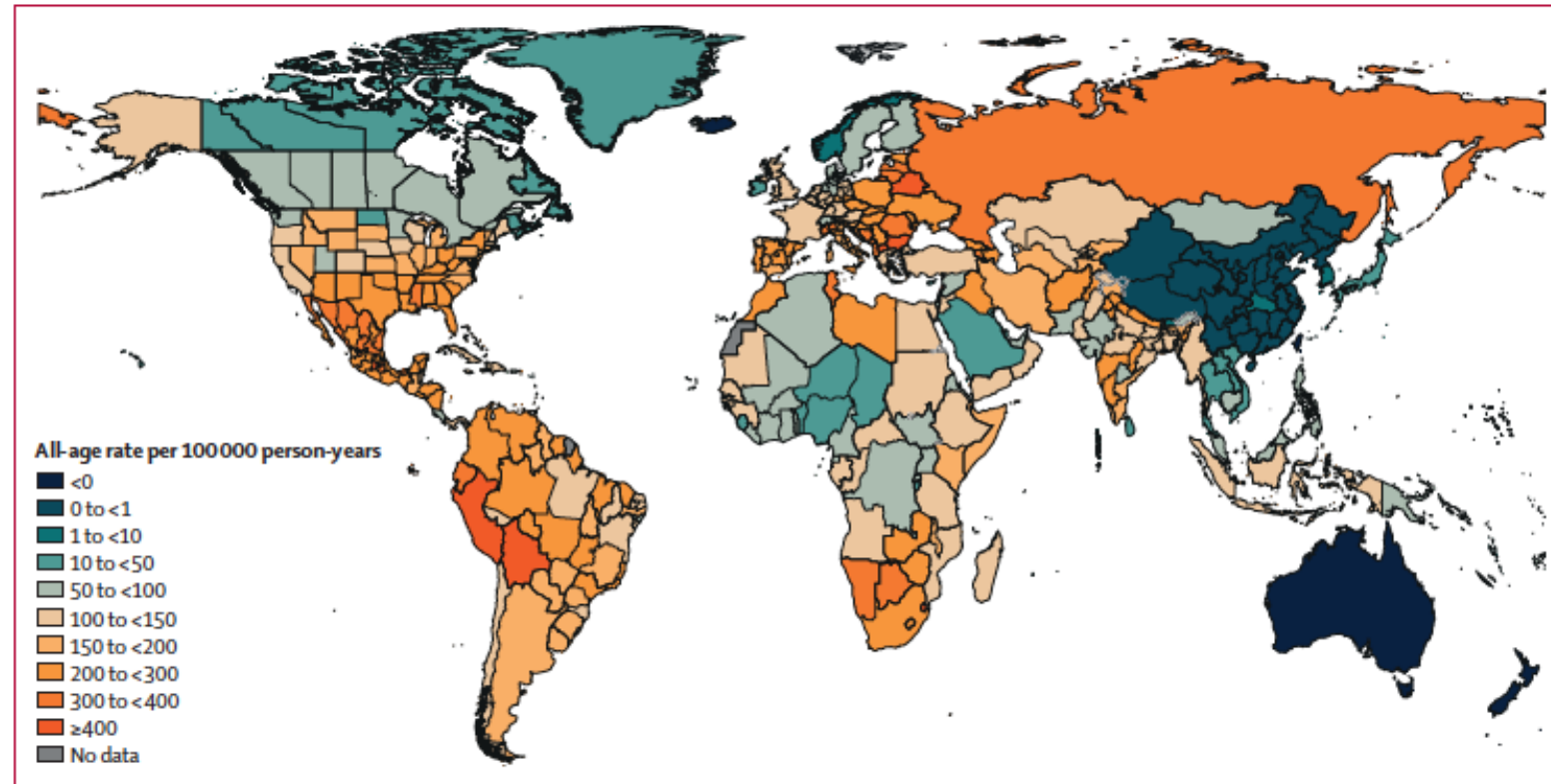
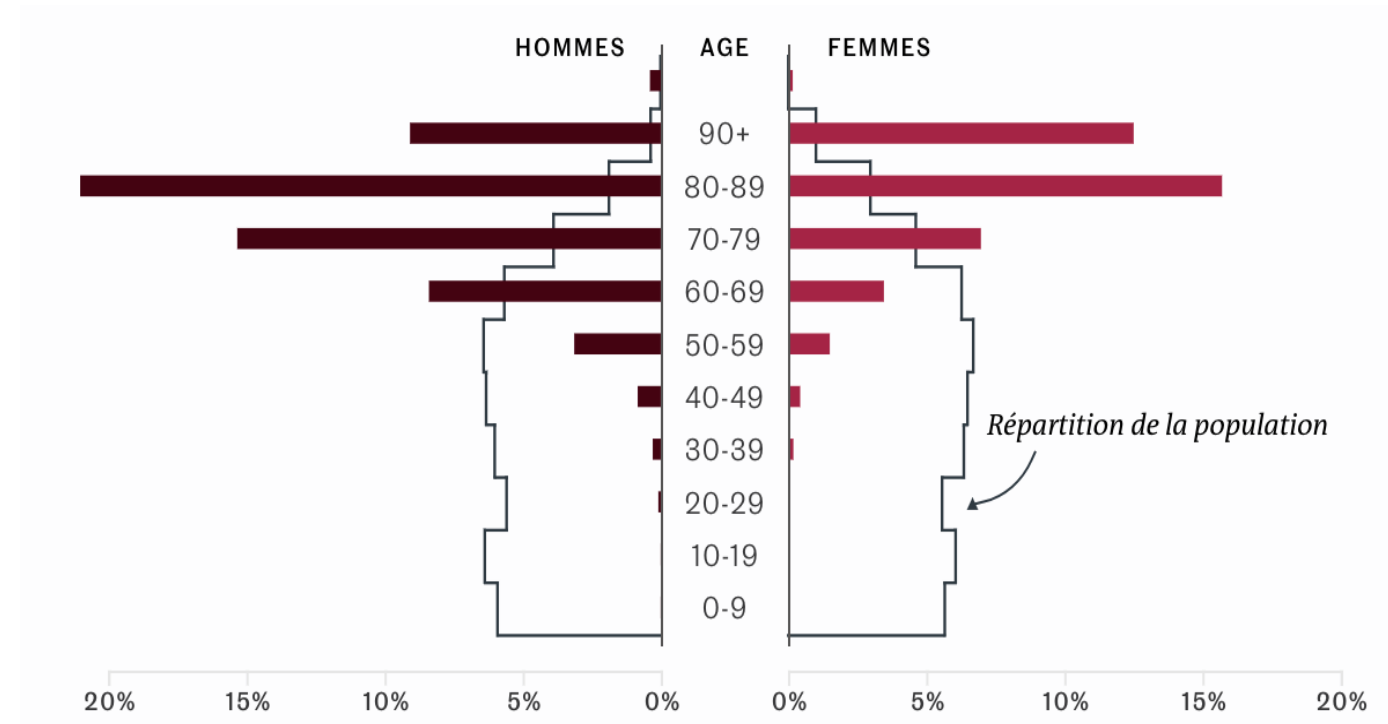


Figure 2: Global distribution of estimated excess mortality rate due to the COVID-19 pandemic, for the cumulative period 2020–21

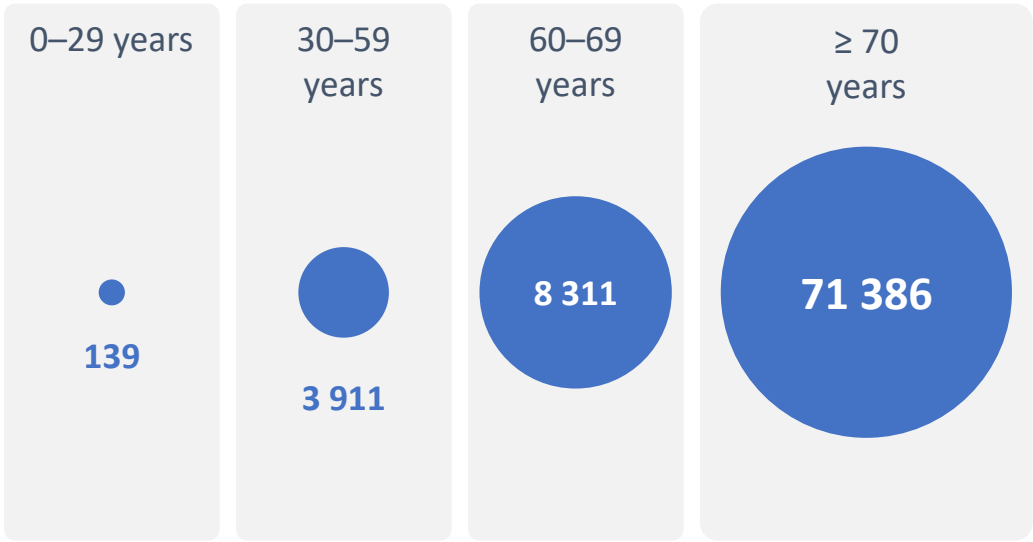
# Covid-19 mortality in older people (France)

Percentages of death according to age and gender in France

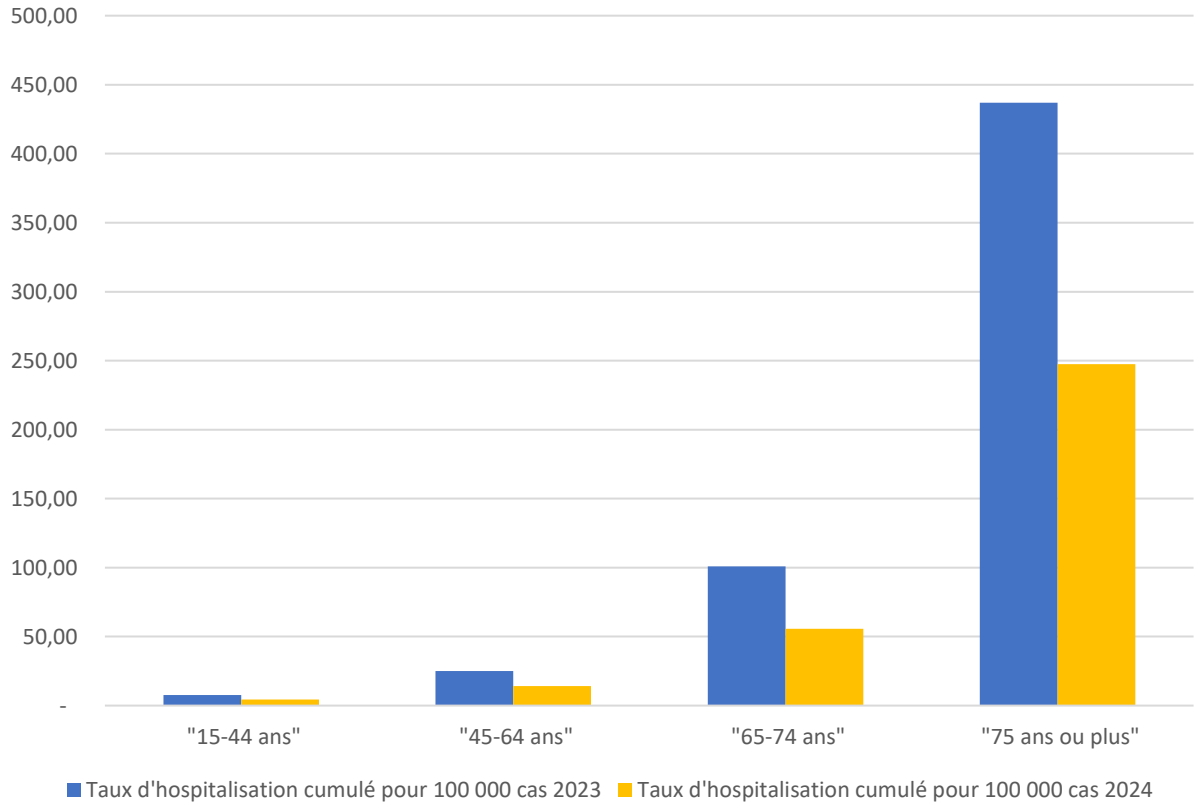


Role of ageing related immune alterations / immune ageing?

# Cumulative number of deaths (2019-2023) and cumulative rates of hospitalizations (2023-2024) associated with Covid-19 by age group (France)



Cumulative number of deaths associated with Covid-19 by age group (2019–2023)



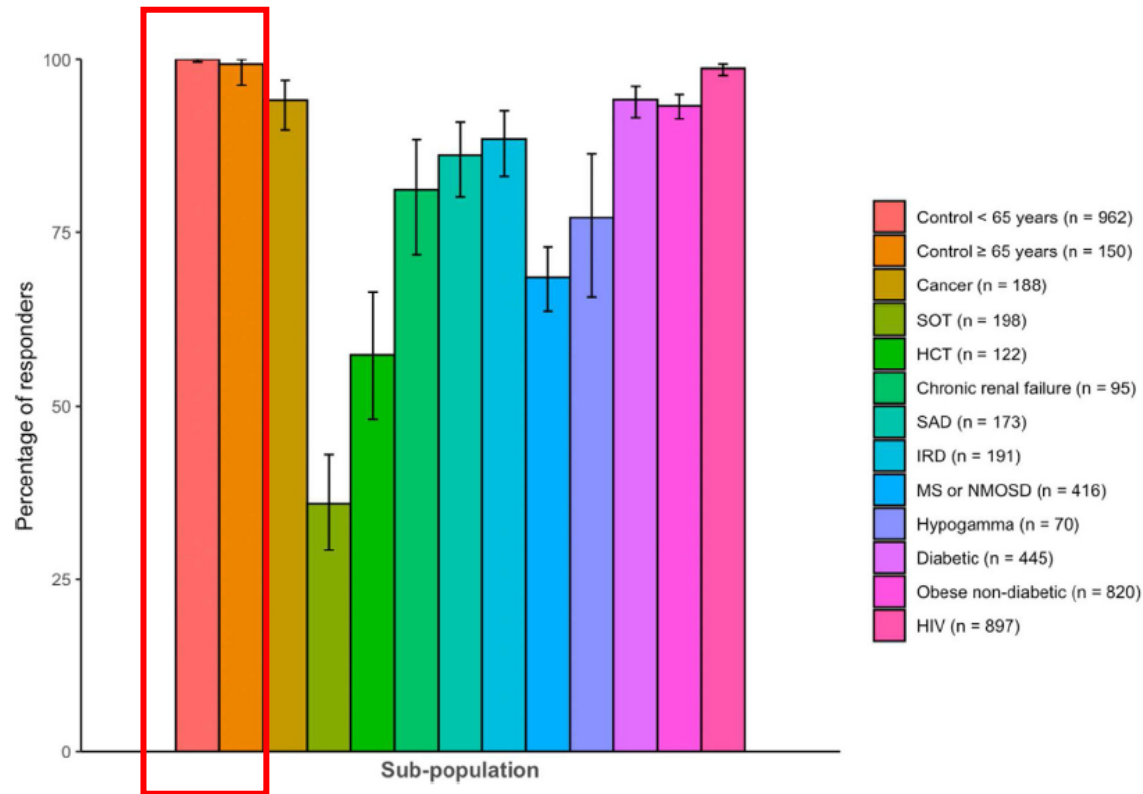
Cumulative rates of hospitalizations associated with Covid-19 by age group (2023 blue bars, 2024, yellow bars)

# 2024-2025: Covid-19 mortality still high

Characteristics of patients hospitalized in intensive care unit in France  
(unconsolidated data)

	Influenza (N= 1849)	Covid-19 (N=326)	RSV (N= 321)
Men (N,%)	1014 (55%)	198 (61%)	165 (52%)
Age ≥ 65 years (N,%)	846 (46%)	221 (68%)	206 (64%)
Comorbidities (N,%)	1564 (86%)	287 (88%)	307 (96%)
Deaths (N,%)	266 (14%)	70 (21%)	43 (13%)
<b>Age ≥ 65 years (N,%)</b>	<b>162 (61%)</b>	<b>63 (90%)</b>	<b>34 (79%)</b>

# Humoral response to primary immunization with SARS-CoV-2 Spike mRNA LNP vaccine



Percentage of responders with anti-Spike antibodies 1 month after the second dose (



## Original article

One-month humoral response following two or three doses of messenger RNA coronavirus disease 2019 vaccines as primary vaccination in specific populations in France: first results from the Agence Nationale Recherche contre le Sida (ANRS)0001S COV-POPART cohort

*P. Loubet et al. / Clinical Microbiology and Infection 29 (2023) 388.e1–388.e8*



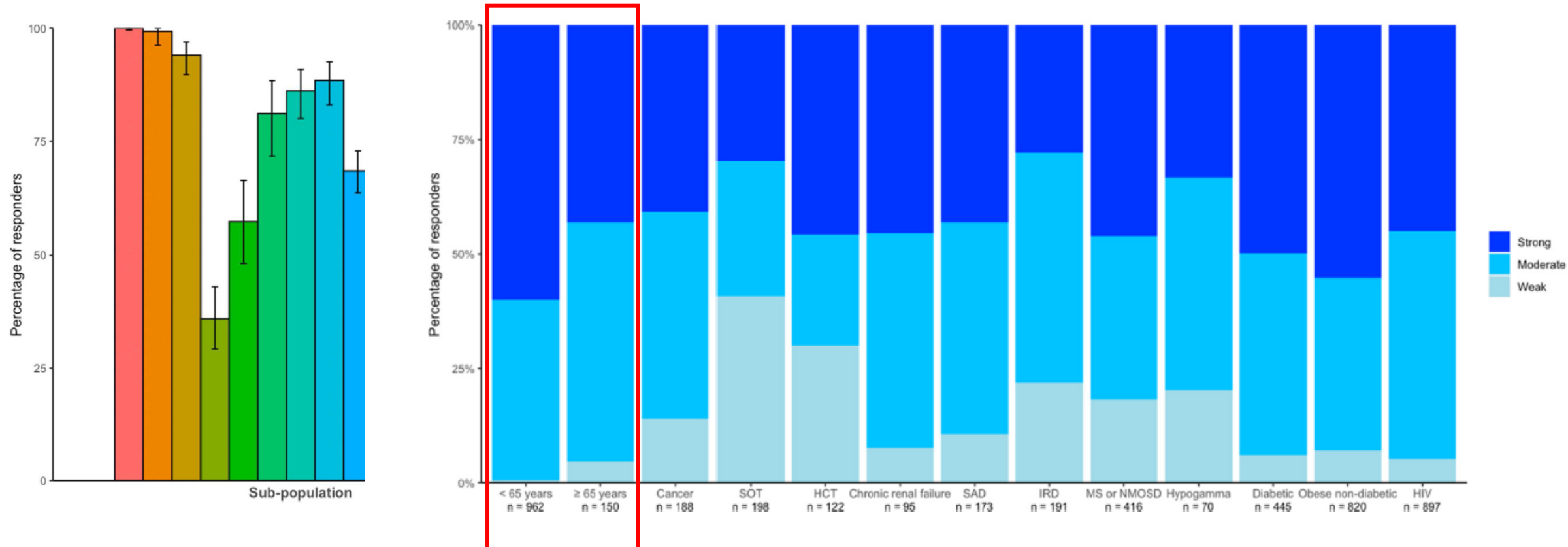
# Humoral response to primary immunization with SARS-CoV-2 spike mRNA LNP vaccine



Original article

One-month humoral response following two or three doses of messenger RNA coronavirus disease 2019 vaccines as primary

*P. Loubet et al. / Clinical Microbiology and Infection 29 (2023) 388.e1–388.e8*



Distribution of responders according to the strength of anti-Spike antibodies response

# Waning in immune humoral response to Covid-19 vaccine

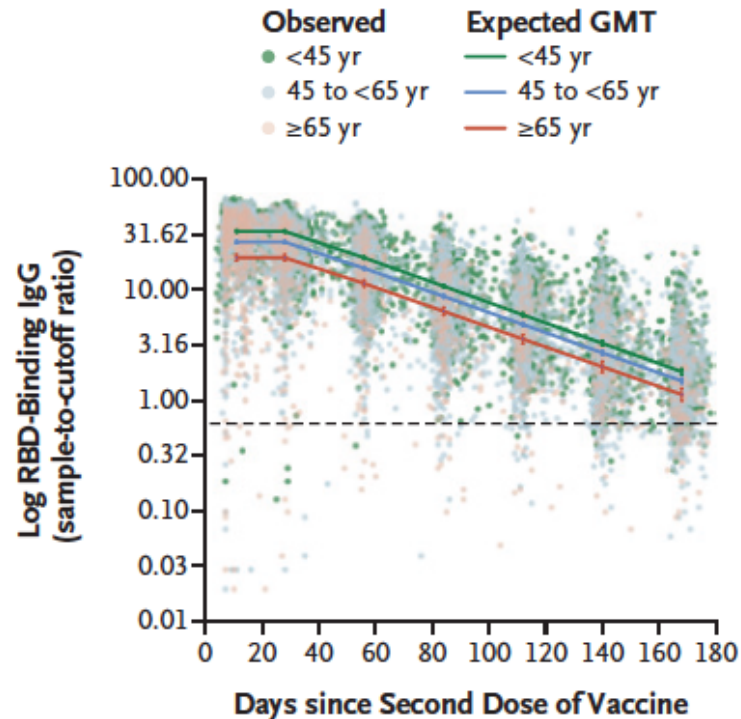
ORIGINAL ARTICLE

## Waning Immune Humoral Response to BNT162b2 Covid-19 Vaccine over 6 Months

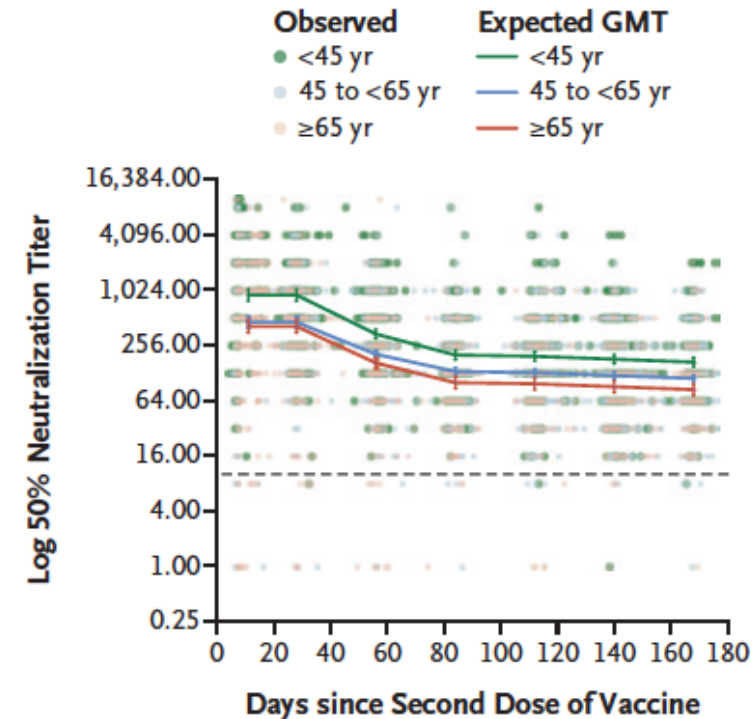
Einav G. Levin, M.D., Yaniv Lustig, Ph.D., Carmit Cohen, Ph.D.,

NEJM 2021

C IgG According to Age Group



D Neutralizing Antibody According to Age Group



# First-generation' COVID-19 vaccines: VE in real life

## ORIGINAL ARTICLE

## BNT162b2 mRNA Covid-19 Vaccine in a Nationwide Mass Vaccination Setting

Noa Dagan, M.D., Noam Barda, M.D., Eldad Kepten, Ph.D., Oren Miron, M.A., Shay Perchik, M.A., Mark A. Katz, M.D., Miguel A. Hernán, M.D., Marc Lipsitch, D.Phil., Ben Reis, Ph.D., and Ran D. Balicer, M.D.

- Study conducted in Israel using a database of 4.7 million people (53% of the population)
- Case (vaccinated)/control (unvaccinated) study
- 596,618 people in each group

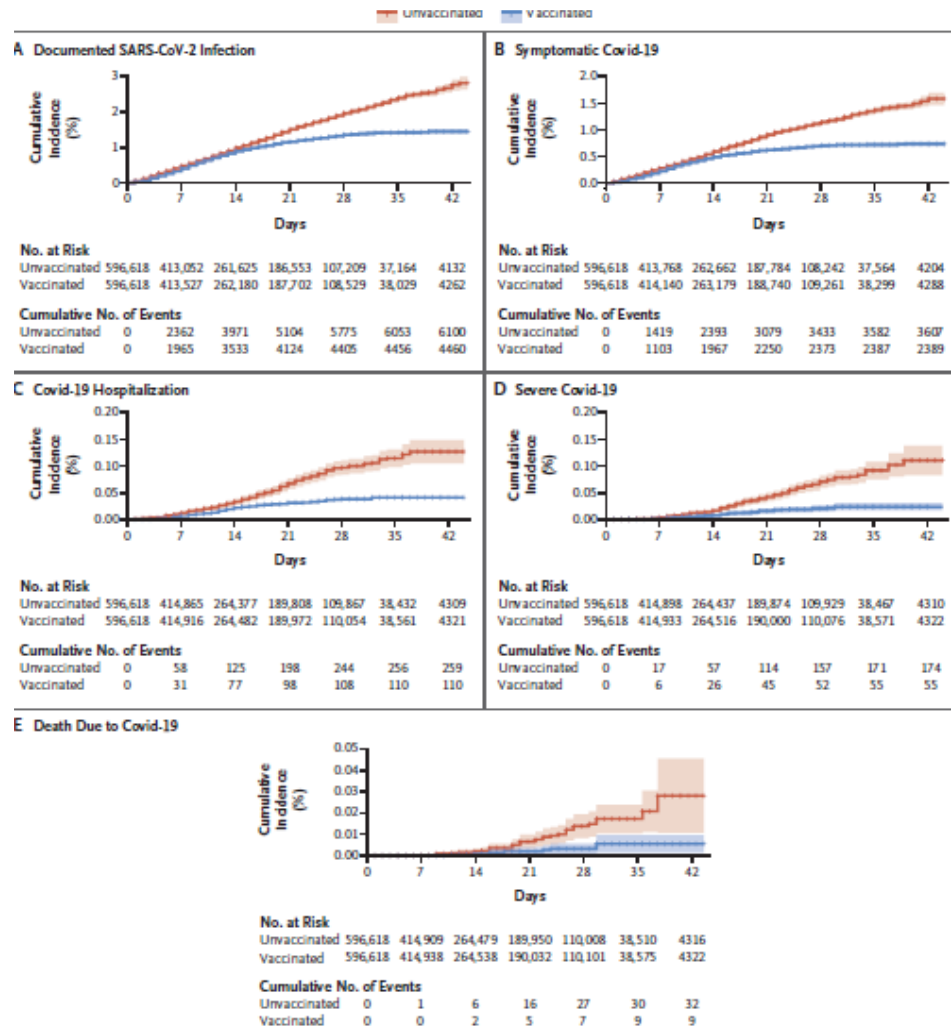


Figure 2. Cumulative Incidence of the Five Outcomes.

Cumulative incidence curves (1 minus the Kaplan–Meier risk) for the various outcomes are shown, starting from the day of administration of the first dose of vaccine. Shaded areas represent 95% confidence intervals. The number at risk at each time point and the cumulative number of events are also shown for each outcome. Graphs in which all data are shown with a y axis scale from 0 to 100 (along with the data shown, as here, on an expanded y axis) are provided in Figure S8 in the Supplementary Appendix.

Table 1. Demographic and Clinical Characteristics of Vaccinated Persons and Unvaccinated Controls at Baseline.\*

Characteristics	Unvaccinated Controls (N=596,618)	Vaccinated Persons (N=596,618)
Median age (IQR) — yr	45 (35–62)	45 (35–62)
Age group — no. (%)		
16 to 39 yr	213,090 (35.7)	213,090 (35.7)
40 to 49 yr	130,752 (21.9)	130,752 (21.9)
50 to 59 yr	85,609 (14.3)	85,609 (14.3)
60 to 69 yr	88,153 (14.8)	88,153 (14.8)
70 to 79 yr	56,946 (9.5)	56,946 (9.5)
≥80 yr	22,068 (3.7)	22,068 (3.7)

# 'First-generation' COVID-19 vaccines: VE in real life

## ORIGINAL ARTICLE

## BNT162b2 mRNA Covid-19 Vaccine in a Nationwide Mass Vaccination Setting

Noa Dagan, M.D., Noam Barda, M.D., Eldad Kepten, Ph.D., Oren Miron, M.A., Shay Perchik, M.A., Mark A. Katz, M.D., Miguel A. Hernán, M.D., Marc Lipsitch, D.Phil., Ben Reis, Ph.D., and Ran D. Balicer, M.D.

**Table 3. Estimated Vaccine Effectiveness against Covid-19 Outcomes in Subpopulations According to Characteristics at Baseline.\***

Characteristic and Period	Documented Infection		Symptomatic Illness	
	1-RR	Risk Difference	1-RR	Risk Difference
	% (95% CI)	no./1000 persons (95% CI)	% (95% CI)	no./1000 persons (95% CI)
<b>Male sex</b>				
14 to 20 days after first dose	41 (32 to 50)	1.71 (1.22 to 2.21)	52 (41 to 61)	1.26 (0.90 to 1.62)
21 to 27 days after first dose	57 (48 to 65)	2.25 (1.76 to 2.75)	62 (49 to 72)	1.30 (0.92 to 1.67)
7 days after second dose to end of follow-up	91 (80 to 96)	7.33 (4.48 to 10.84)	88 (71 to 98)	2.90 (1.87 to 4.02)
<b>Female sex</b>				
14 to 20 days after first dose	50 (41 to 57)	2.39 (1.84 to 2.86)	60 (52 to 68)	1.81 (1.43 to 2.19)
21 to 27 days after first dose	63 (55 to 71)	2.38 (1.91 to 2.91)	69 (58 to 78)	1.38 (1.02 to 1.71)
7 days after second dose to end of follow-up	93 (88 to 97)	9.75 (6.84 to 13.48)	96 (90 to 100)	6.22 (3.60 to 9.56)
<b>Age, 16 to 39 yr</b>				
14 to 20 days after first dose	49 (41 to 57)	2.29 (1.74 to 2.88)	57 (46 to 68)	1.38 (0.99 to 1.80)
21 to 27 days after first dose	64 (54 to 72)	2.80 (2.20 to 3.48)	67 (52 to 78)	1.27 (0.89 to 1.73)
7 days after second dose to end of follow-up	94 (87 to 97)	8.72 (5.72 to 12.69)	99 (96 to 100)	4.06 (2.76 to 5.66)
<b>Age, 40 to 69 yr</b>				
14 to 20 days after first dose	47 (40 to 55)	2.13 (1.69 to 2.66)	59 (50 to 67)	1.68 (1.32 to 2.05)
21 to 27 days after first dose	58 (49 to 67)	2.19 (1.67 to 2.70)	65 (53 to 74)	1.38 (1.03 to 1.80)
7 days after second dose to end of follow-up	90 (82 to 95)	8.96 (6.16 to 13.05)	90 (75 to 98)	5.01 (2.53 to 8.67)
<b>Age, ≥70 yr</b>				
14 to 20 days after first dose	22 (-9 to 44)	0.81 (-0.28 to 1.89)	44 (19 to 64)	1.36 (0.48 to 2.36)
21 to 27 days after first dose	50 (19 to 72)	1.40 (0.42 to 2.35)	64 (37 to 83)	1.35 (0.62 to 2.22)
7 days after second dose to end of follow-up	95 (87 to 100)	6.10 (3.43 to 9.61)	98 (90 to 100)	4.77 (2.14 to 7.70)

# COVID-19 vaccination and deaths avoided

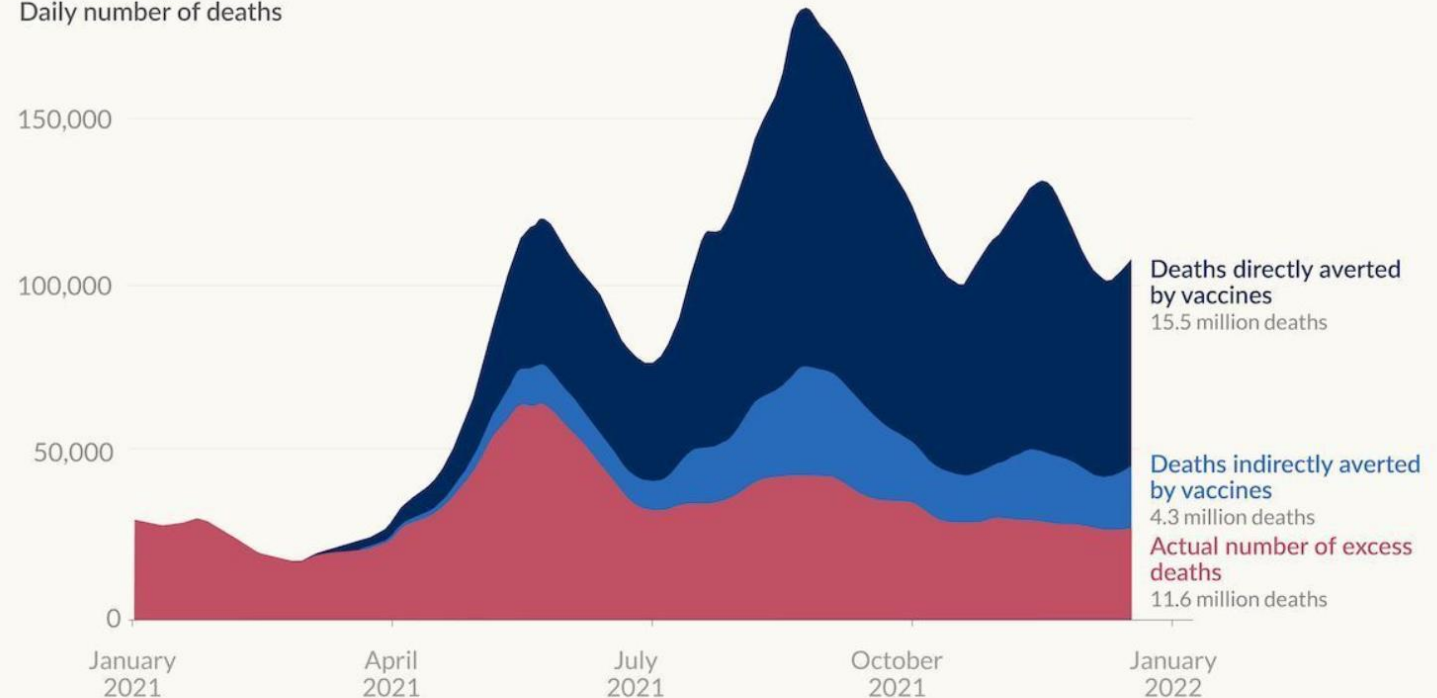
- **Globally: 15.5 million deaths directly averted during the first year of vaccination**

## COVID-19 vaccination saved millions of lives worldwide

Our World  
in Data

Estimated global number of deaths averted by COVID-19 vaccination, between December 2021 and December 2022. This compares the number of excess deaths to the number of averted deaths in people directly protected by vaccination and those indirectly protected by lower infection rates.

Daily number of deaths



**Note:** These estimates don't account for possible policies that might have been implemented to control COVID-19 if vaccines hadn't been available. They are based on models that consider COVID-19 spread, vaccination rates, demographics, healthcare capacity, excess mortality, and vaccine efficacy.

**Source:** Oliver J. Watson et al. (2022) Global impact of the first year of COVID-19 vaccination: a mathematical modelling study. *Lancet Infectious Diseases*.

[OurWorldinData.org](https://www.ourworldindata.org) — Research and data to make progress against the world's largest problems.

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# COVID-19 vaccination and deaths avoided

- **Globally: 15.5 million deaths directly averted during the first year of vaccination**
- **In Europe: estimated number of lives saved between December 2020 and March 2023**

57% reduction in deaths, or approximately 1.4 million lives saved among adults aged  $\geq 25$  (range, 0.7 to 2.6 million) (2.5 million deaths observed)

**96% aged  $\geq 60$  years old**

**50% aged  $\geq 80$  years old**

– 1st booster dose: 51% of deaths averted

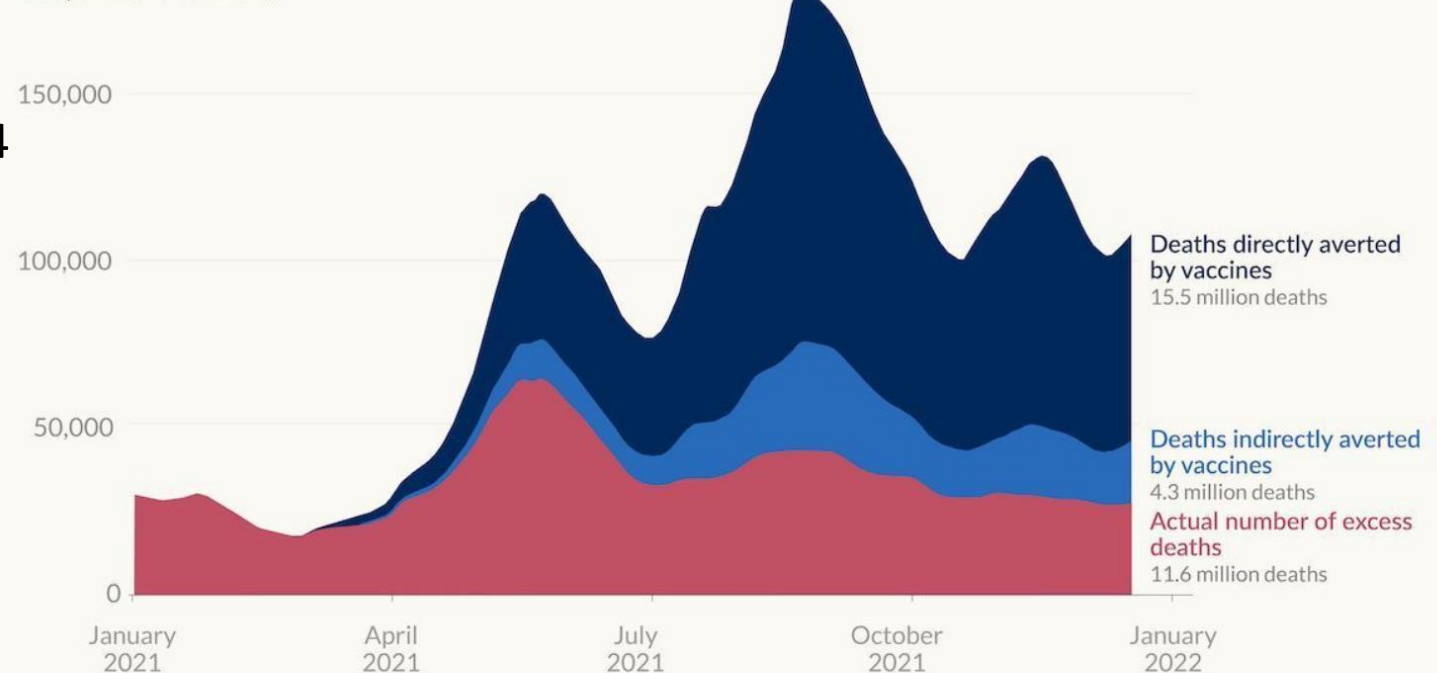
- 67% of deaths averted during the Omicron period

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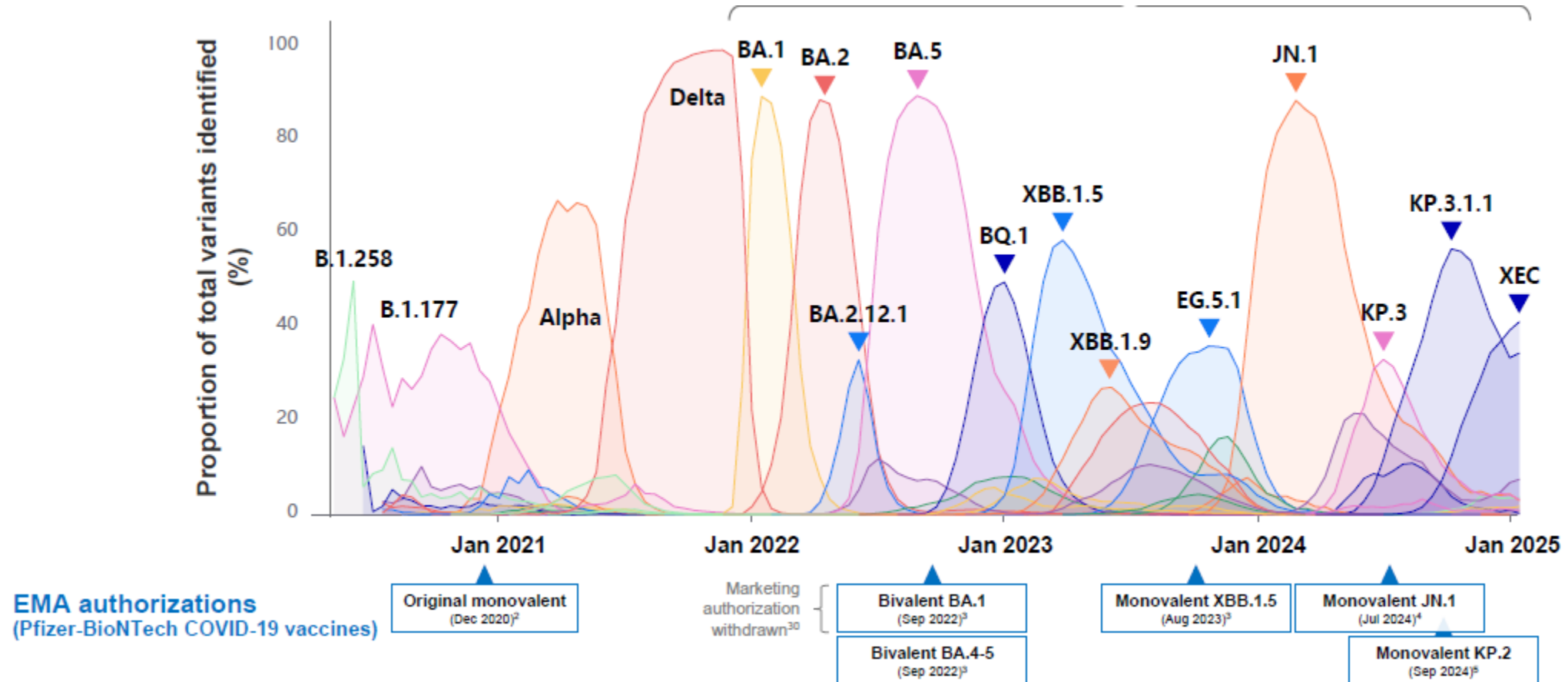
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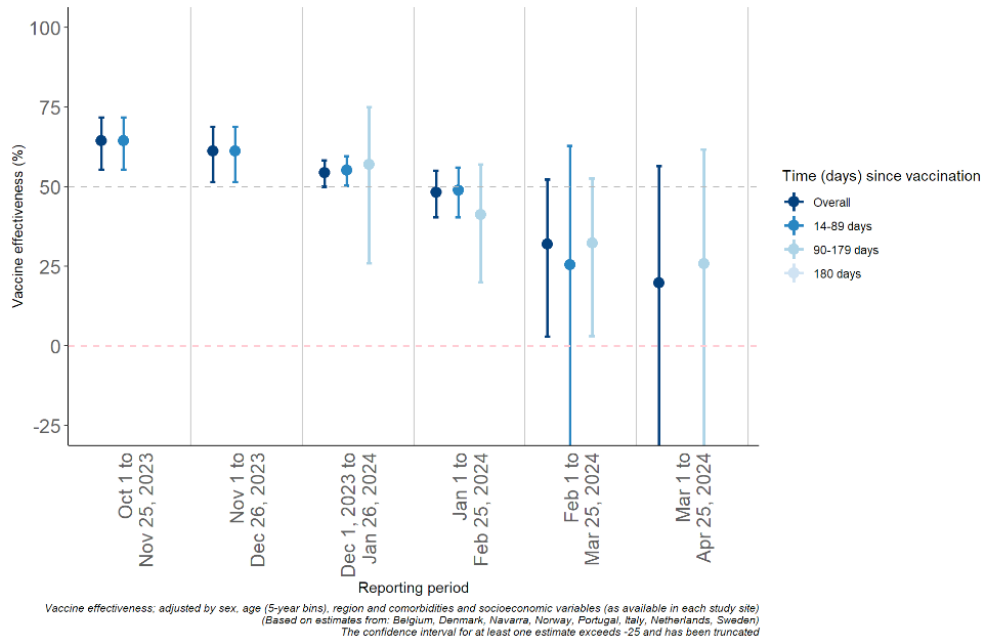
# Adapted mRNA Covid 19 vaccines

Omicron

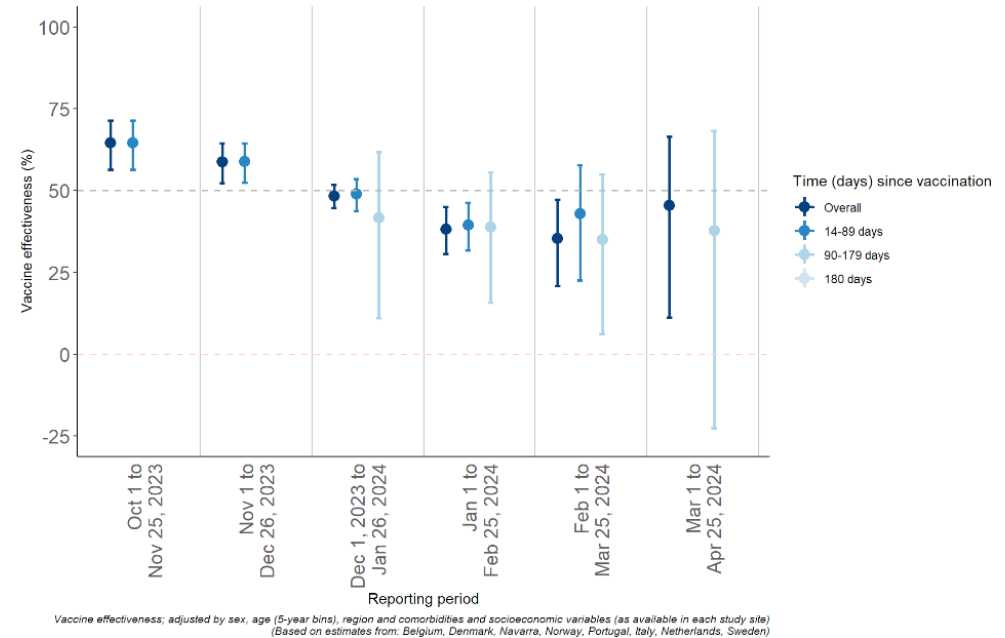


# XBB1.5 Omicron Covid 19 vaccine effectiveness (2023-24) (hospitalisations)

- 65-79 years



- $\geq 80$  years





# KP2 Omicron Covid 19 vaccine effectiveness (2024-25)

- Negative-design studies in the US
- Two networks
  - VISION network (adults, > 200 hospitals)
  - IVY network (> 65 years old, 26 hospitals)
- Data collection from registries
- Preliminary data

## Interim Estimates of 2024–2025 COVID-19 Vaccine Effectiveness Among Adults Aged ≥18 Years — VISION and IVY Networks, September 2024–January 2025

Ruth Link-Gelles, PhD<sup>1</sup>; Sean Chickery, DHSc<sup>2</sup>; Alexander Webber, MPH<sup>1</sup>; Toan C. Ong, PhD<sup>3</sup>;

Characteristic	Vaccine effectiveness network and setting, no. (column %)								
	VISION ED/UC encounters, all adults aged ≥18 years			VISION hospitalizations, all adults aged ≥65 years			IVY hospitalizations, immunocompetent adults aged ≥65 years		
	Total	COVID-19 case- patients	COVID-19 control- patients	Total	COVID-19 case- patients	COVID-19 control- patients	Total	COVID-19 case- patients	COVID-19 control- patients
Total	137,543	10,459	127,084	34,411	2,846	31,565	1,929	683	1,246
Median age	53 [34, 72]	58 [37, 74]	53 [34, 71]	78 [72, 84]	79 [73, 86]	78 [71, 84]	77 [71, 84]	78 [72, 85]	76 [70, 83]
Age group									
18-64 years	88,858 (65)	6,113 (58)	82,745 (65)	--	--	--	--	--	
≥65 years	48,685 (35)	4,346 (42)	44,339 (35)	34,411 (100)	2,846 (100)	31,565 (100)	1,929 (100)	683 (100)	1,246 (100)
Immunocompromised*	--	--	--	8,192 (24)	598 (21)	7,594 (24)	--	--	

# KP2 Omicron Covid 19 vaccine effectiveness (2024-25)

U.S. Centers for Disease Control and Prevention

# MMWR

Weekly / Vol. 74 / No. 6

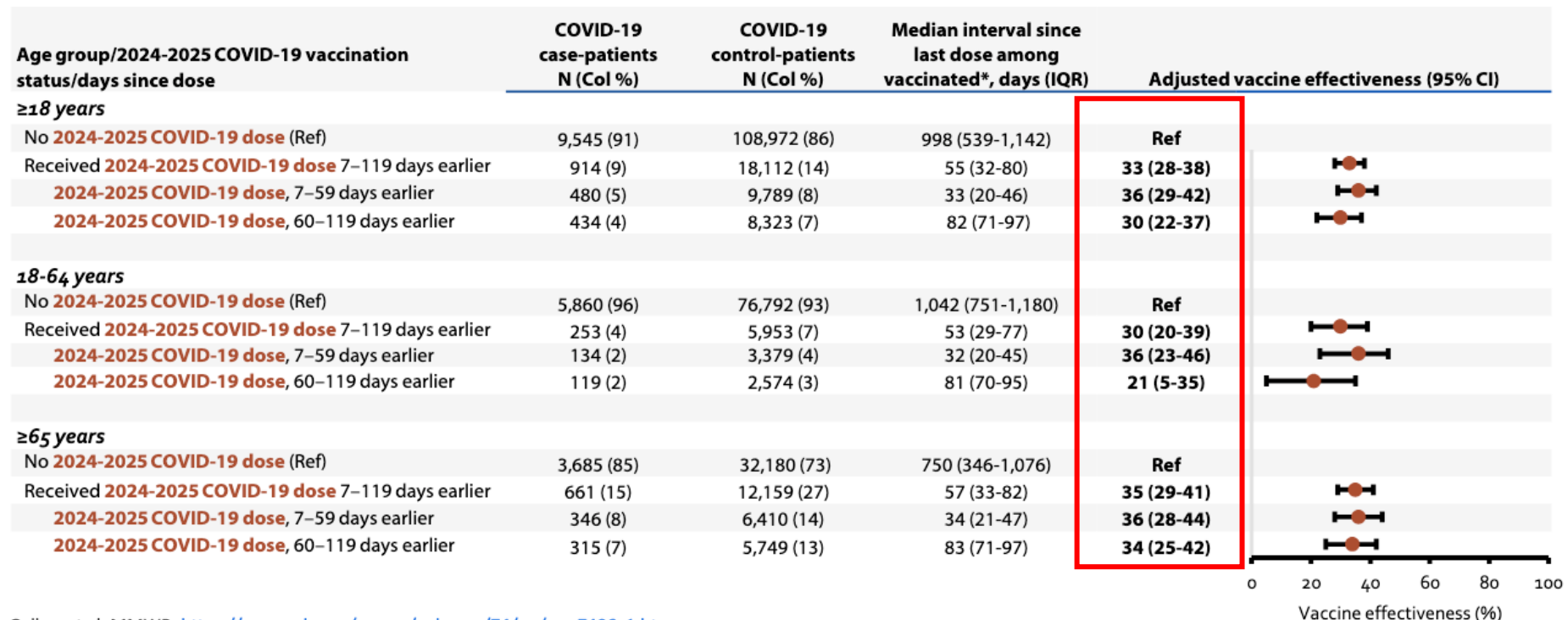
Morbidity and Mortality Weekly Report

February 27, 2025

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Ruth Link-Gelles, PhD<sup>1</sup>; Sean Chickery, DHSc<sup>2</sup>; Alexander Webber, MPH<sup>1</sup>; Toan C. Ong, PhD<sup>3</sup>;

### ◆ Effectiveness on emergency department consultations (VISION)

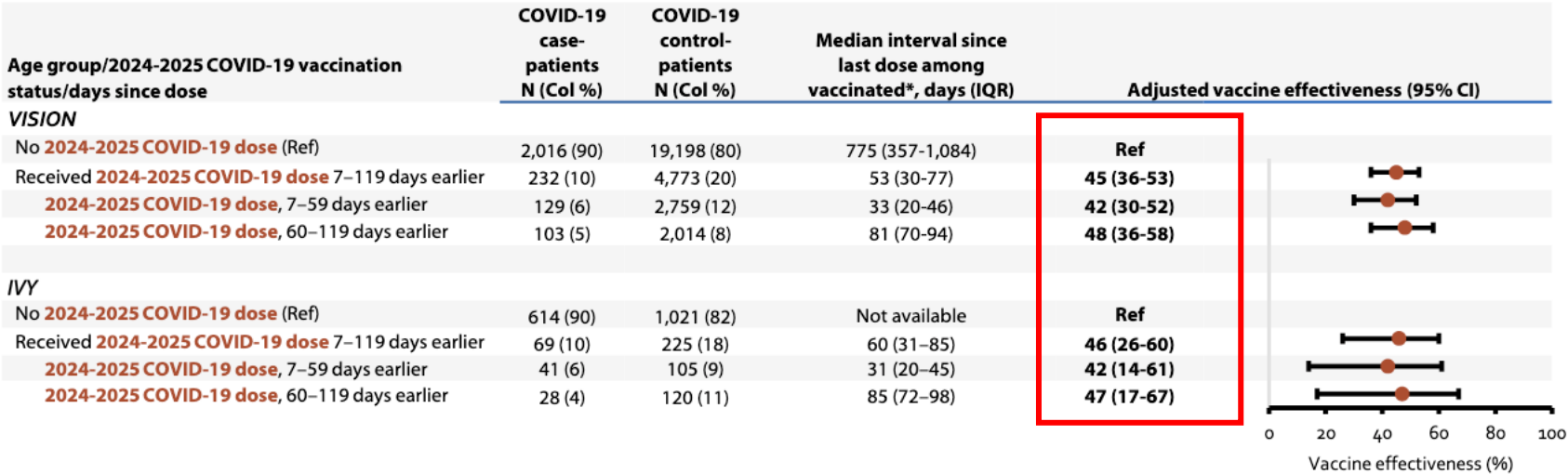


# KP2 Omicron Covid 19 vaccine effectiveness (2024-25)

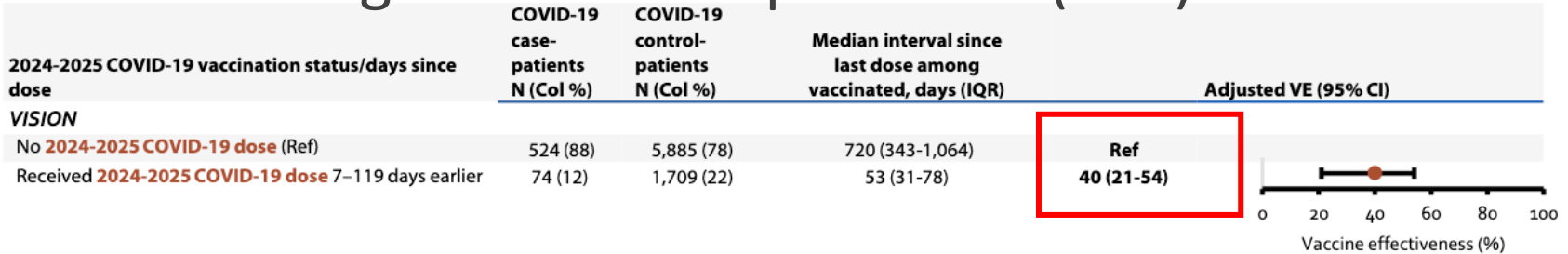
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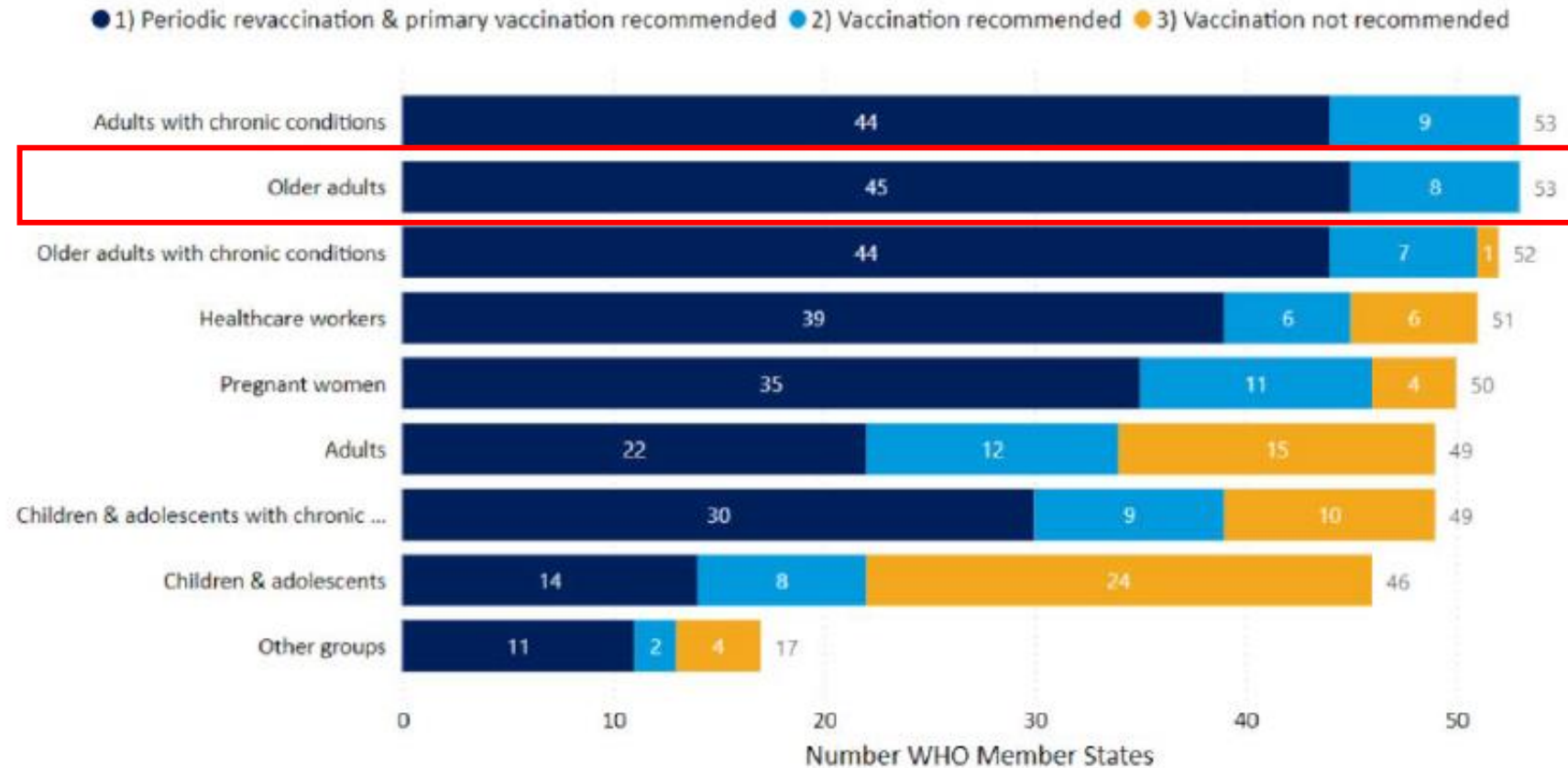
### ◆ Effectiveness on hospitalisation (65+)



### ◆ Effectiveness among immunocompromised (65+)



# WHO: National policies on Covid-19 vaccination



**85**

WHO MS reported on COVID-19 vaccination policies for at least one population group

**71**

WHO MS reported recommending periodic vaccination for at least one population group

**Older adults**

are most frequently reported as being recommended to be periodically revaccinated against COVID-19

Covid-19 vaccine coverage in France  
(French national agency of public health SPF, non published)

	2023-2024	Spring 2024	2024-2025
≥ 65 years old	30.2%	-	19.8%
≥ 80 years old	36.9%	8.6%	20.8%
Nursing homes	68.4%	-	-

# Covid-19 vaccination in elderly: conclusions, perspectives

- COVID-19 remains common and severe in the elderly
- Maintained mRNA vaccine efficacy despite the emergence of variants (adapted vaccines)
- Significance of infections outside the winter season justifies a spring campaign in specific populations, especially elderly ( $\geq 80$  years old)
- Challenge of vaccination coverage
- Place of the sub-unit adjuvanted vaccine?