

Pandemic Update: Monkeypox and COVID-19

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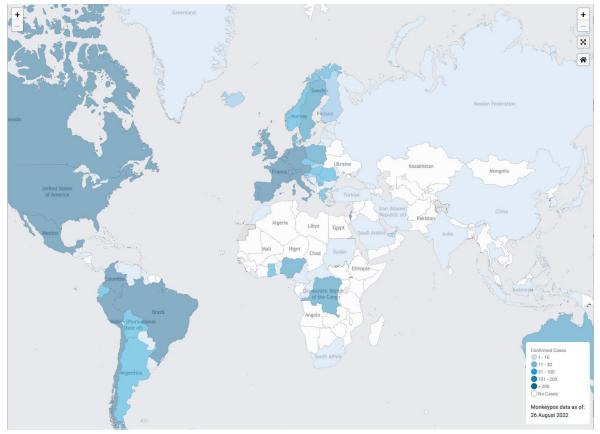
> IRB Brown Bag Lunch 31 August 2022

Disclosures

- Research Support[°]
 - $_{\circ}\,$ GlaxoSmithKline, Pulmocide
- Paid Consultation
 - Adagio, ADMA Biologics, AlloVir, Cidara, Genentech/Roche, Janssen, Shionogi, Takeda, Viracor Eurofins
- Royalties
 - UpToDate
- Data & Safety Monitoring Board Participation
 - o Adamis, Allovir, CSL Behring, Janssen, Merck, Sequiris, Takeda, Talaris



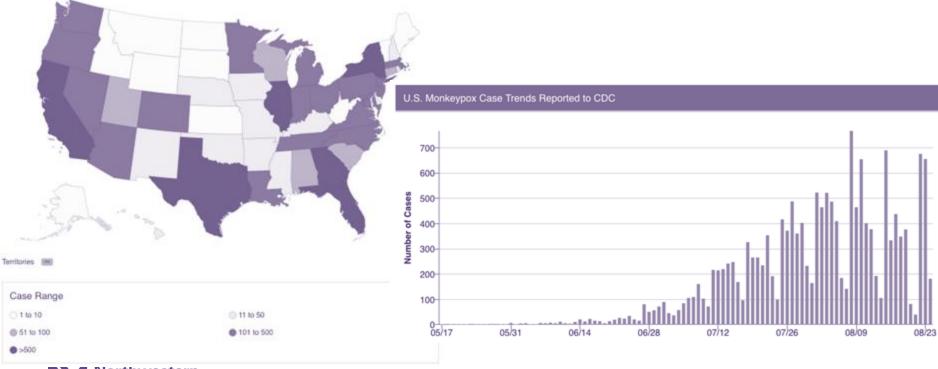




Northwestern Medicine https://extranet.who.int/publicemergency/#

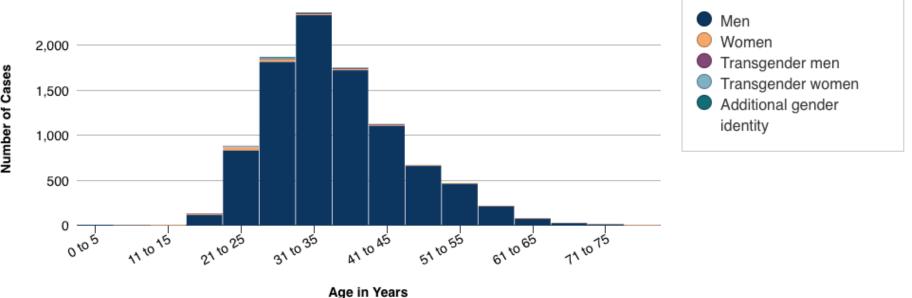
16,926 Total confirmed monkeypox/orthopoxvirus cases

"One Florida case is listed here but included in the United Kingdom case counts because the individual was tested while in the UK.



Northwestern Medicine* https://www.cdc.gov/poxvirus/monkeypox/response/2022/mpx-trends.html

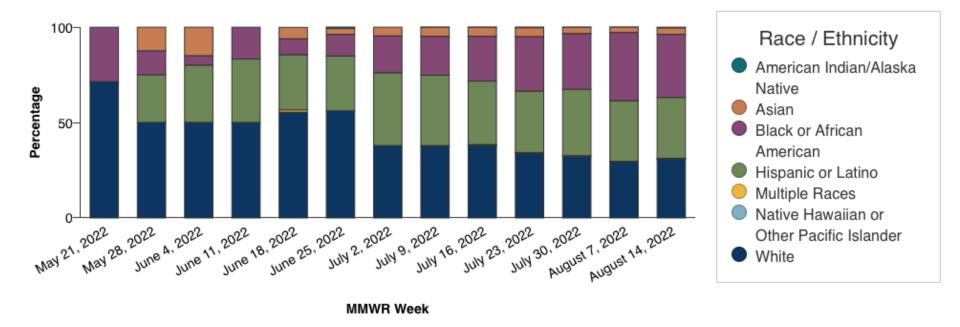
Monkeypox cases reported to CDC: Age and Gender



Northwestern Medicine[®] https://www.cdc.gov/poxvirus/monkeypox/response/2022/mpx-trends.html

Number of Cases

Monkeypox cases reported to CDC: Race/Ethnicity



Northwestern Medicine* https://www.cdc.gov/poxvirus/monkeypox/response/2022/mpx-trends.html

Monkeypox: Clinical Presentation

Incubation Period

- $_{\odot}$ Monkeypox symptoms usually start within 3 weeks of exposure to the virus
- If someone has flu-like symptoms, they will usually develop a rash 1-4 days later
- Monkeypox can be spread from the time symptoms start until the rash has healed, all scabs have fallen off, and a fresh layer of skin has formed
- $_{\odot}$ The illness typically lasts 2-4 weeks
- Clinical Presentation
 - $_{\odot}$ Often a flu-like prodrome before rash
 - $_{\odot}$ Some presents with rash at onset
 - Other symptoms: Fever, chills, swollen lymph nodes, fatigue, muscle aches, headache, respiratory symptoms







Monkeypox: Rash

 \mathbf{N}

Stage	Stage Duration	Characteristics
Enanthem		• Sometimes, lesions first form on the tongue and in the mouth.
Macules	1–2 days	Macular lesions appear.
Papules	1–2 days	• Lesions typically progress from macular (flat) to papular (raised).
Vesicles	1–2 days	• Lesions then typically become vesicular (raised and filled with clear fluid).
Pustules	5–7 days	 Lesions then typically become pustular (filled with opaque fluid) – sharply raised, usually round, and firm to the touch (deep seated). Finally, lesions typically develop a depression in the center (umbilication). The pustules will remain for approximately 5 to 7 days before beginning to crust.
Scabs	7–14 days	By the end of the second week, pustules have crusted and scabbed over.Scabs will remain for about a week before beginning to fall off.

Northwestern Medicine* https://www.cdc.gov/poxvirus/monkeypox/clinicians/clinical-recognition.html

Monkeypox cases reported to CDC: Symptoms

Deeb					98.6%
Rash				72.7%	
Malaise	e (tiredness or not feeling well)			12.170	
Fever			•	72.1%	
01.111			68	9%	
Chills			65.2%		
Headac	che		05.2 /0		
Enlarge	ed lymph nodes (swollen glands)		64.3%		
Muslaia	(musele sekee)		61.8%		
Myaigia	a (muscle aches)		60.2%		
Pruritis	(itching)		00.270		
Pruritis Rectal p	pain	• 43.9%			
	blood on stools				
1 43 61 1	25%				
Rectal b	bleeding				
Tenesm	 23.1% nus (pain when wanting to pass stool) 				
Abdomi	 19% inal pain (stomach ache) 				
	—— 18%				
	g or nausea 16.3%				
	s (swelling, soreness in the rectal area)				
	 6% ctivitis (redness or pain in the eye) 				
6	20	40	60	80	100



https://www.cdc.gov/poxvirus/monkeypox/response/2022/mpx-trends.html

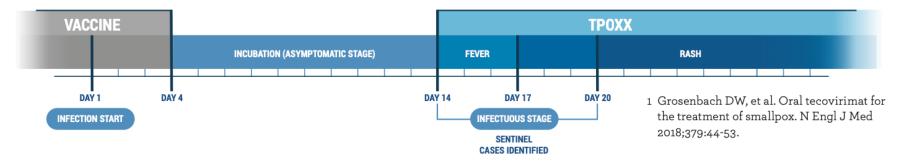
Monkeypox: Transmission

- Close or Intimate Contact
 - Monkeypox can spread to anyone through close, personal, often skin-to-skin contact, including:
 - Direct contact with monkeypox rash, scabs, or body fluids from a person with monkeypox.
 - Touching objects, fabrics (clothing, bedding, or towels), and surfaces that have been used by someone with monkeypox.
 - $\circ~$ Contact with respiratory secretions.
- This direct contact can happen during intimate contact, including:
 - Oral, anal, and vaginal sex or touching the genitals (penis, testicles, labia, and vagina) or anus (butthole) of a person with monkeypox.
 - $_{\odot}~$ Hugging, massage, and kissing.
 - $_{\circ}~$ Prolonged face-to-face contact.
 - Touching fabrics and objects during sex that were used by a person with monkeypox and that have not been disinfected, such as bedding, towels, fetish gear, and sex toys.
- Not transmitted through routine day-to-day exposures (riding CTA, going to the bathroom)



Monkeypox: Approach

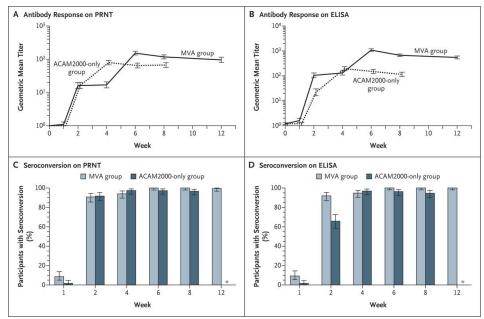
Antiviral Treatment (FDA Approved)





Monkeypox: Vaccines

- Two vaccines are available
 - ACAM2000: Live Smallpox Vaccine, skin prick (85% effective against MPV)
 - ∘ Jynneos: Non-replicating virus, SubQ injection 4 weeks apart





Pittman et al. N Engl J Med. 2019;381:1897-1908.

Monkeypox: Vaccines

- Jynneos Use
 - $_{\odot}\,$ The sooner it is given the better
 - Use within 4 days of exposure is associated with prevention of onset of disease
 - Use between 4 and 14 days may be associated with reduced severity of infection
 - $_{\odot}\,$ Protection 2 weeks after 2nd dose
 - If prior smallpox vaccine >3 years ago, would get revaccinated



AS OF 8.18.22 You are eligible for a MPV vaccine if you:

- Live anywhere in Chicago or Illinois and have not previously been infected with MPV **AND meet the criteria below:**
- 2. Have had close contact (e.g., household members with close physical contact or intimate partners) with someone diagnosed with MPV regardless of sex, gender, or sexual orientation **OR**

Are a gay, bisexual, or other man who has sex with men and/or transgender person **who is sexually active.**

If you meet eligibility criteria, especially consider getting vaccinated if you met recent partners through online applications or social media platforms (such as Grindr, Tinder or Scruff), or at clubs, raves, sex parties, saunas, or exchange good or services for sex. At this time, MPV vaccine is NOT recommended for the general public. As vaccine supply increases, guidance may evolve.







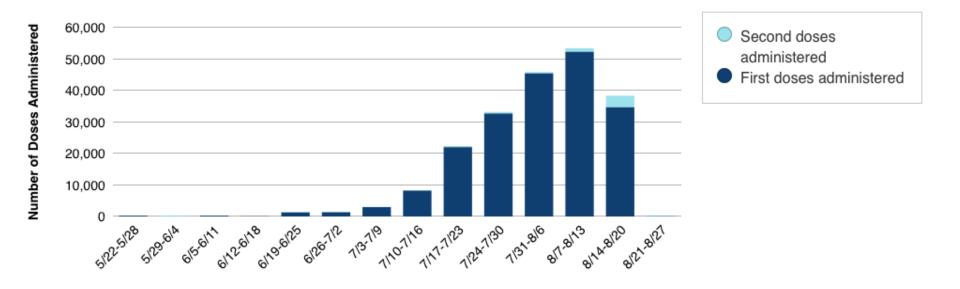
Monkeypox: Vaccines

Event				MVA Group	0			ACAM2000- Only Group
	Period 1 MVA (N=220)	P Value†	Period 2 MVA (N=208)	P Value†	Periods 1 and 2 MVA (N=220)	Period 3 ACAM2000 (N=196)	P Value†	Period 1 (N=213)
	no. (%)		no. (%)		no. (%)			no. (%)
Documented adverse event	169 (76.8)	<0.001	135 (64.9)	< 0.001	184 (83.6)	181 (92.3)	0.008	209 (98.1)
Nonserious adverse event within 29 days after vaccination	168 (76.4)	<0.001	135 (64.9)	<0.001	183 (83.2)	181 (92.3)	0.008	209 (98.1)
Serious adverse event‡	2 (0.9)	1.0	0	1.0	2 (0.9)	0	1.0	1 (0.5)
Adverse event of special interest	2 (0.9)	0.44	2 (1.0)	0.68	4 (1.8)	2 (1.0)	0.69	4 (1.9)
Related adverse event within 29 days after vaccination§	112 (50.9)	<0.001	76 (36.5)	<0.001	130 (59.1)	61 (31.1)	<0.001	158 (74.2)
Adverse event grade ≥3 within 29 days after vaccination	13 (5.9)	<0.001	4 (1.9)	<0.001	17 (7.7)	10 (5.1)	<0.001	64 (30.0)
Related adverse event grade ≥3§	3 (1.4)	< 0.001	2 (1.0)	<0.001	5 (2.3)	3 (1.5)	<0.001	22 (10.3)
Related adverse event grade ≥3 within 29 days∬	3 (1.4)	<0.001	2 (1.0)	<0.001	5 (2.3)	3 (1.5)	<0.001	22 (10.3)
Adverse event leading to withdrawal from trial	2 (0.9)	0.5	0	NA	2 (0.9)	0	NA	0
Adverse event leading to withdrawal from vaccination	2 (0.9)	0.5	0	NA	2 (0.9)	0	NA	0

Morthwestern Medicine[®] Pittman *et al.* **N Engl J Med.** 2019;381:1897-1908.

Monkeypox: Vaccine Update

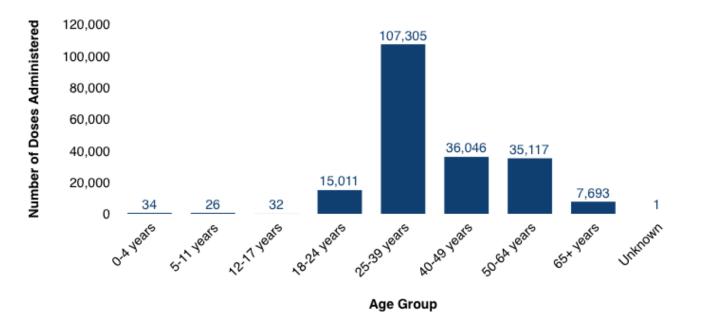
Total JYNNEOS Vaccine Doses Administered and Reported to CDC

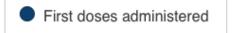


Northwestern Medicine* https://www.cdc.gov/poxvirus/monkeypox/response/2022/mpx-trends.html

Monkeypox: Vaccine Update

JYNNEOS Vaccine Doses Administered, by Age





Northwestern Medicine[®] https://www.cdc.gov/poxvirus/monkeypox/response/2022/mpx-trends.html

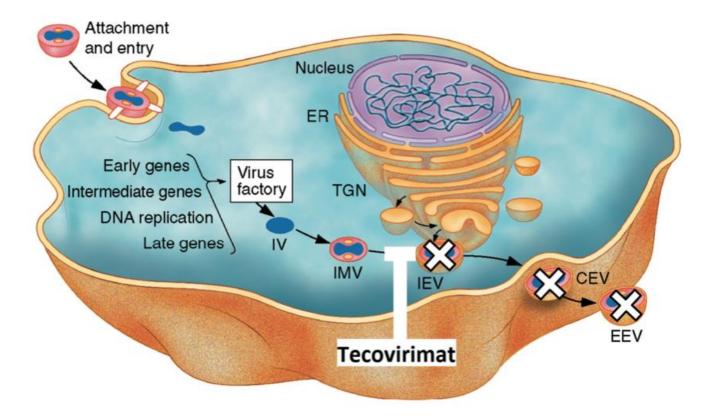




Table 6:Survival Rates in Tecovirimat Treatment Studies in Cynomolgus Macaques
and NZW Rabbits Exhibiting Clinical Signs of Orthopoxvirus Disease

	Treatment Initiation ^a	Survival Percentage (# survived/n)		p-value ^b	Survival Rate Difference ^c	
		Placebo	Tecovirimat		(95% CI) ^d	
Cynomolgus Macaques						
Study 1	Day 4	0% (0/7)	80% (4/5)	0.0038	80% (20.8%, 99.5%)	
Study 2	Day 4	0% (0/6)	100% (6/6)	0.0002	100% (47.1%, 100%)	
Study 3	Day 4	0% (0/3)	83% (5/6)	0.0151	83% (7.5%, 99.6%)	
	Day 5		83% (5/6)	0.0151	83% (7.5%, 99.6%)	
	Day 6		50% (3/6)	0.1231	50% (-28.3%, 90.2%)	
NZW Rabbits						
Study 4	Day 4	0% (0/10)	90% (9/10)	< 0.0001	90% (50.3%, 99.8%)	
Study 5	Day 4	NA ^e	88% (7/8)	NA	NA	



Table 2:Adverse Reactions Reported in ≥ 2% of Healthy Adult Subjects Receiving at
Least One Dose of TPOXX 600 mg

Adverse Reaction	TPOXX 600 mg N = 359 (%)	Placebo N = 90 (%)
Headache	12	8
Nausea	5	4
Abdominal pain ^a	2	1
Vomiting	2	0

^a Includes abdominal pain, abdominal pain upper, abdominal distension, abdominal discomfort, abdominal pain lower, epigastric pain

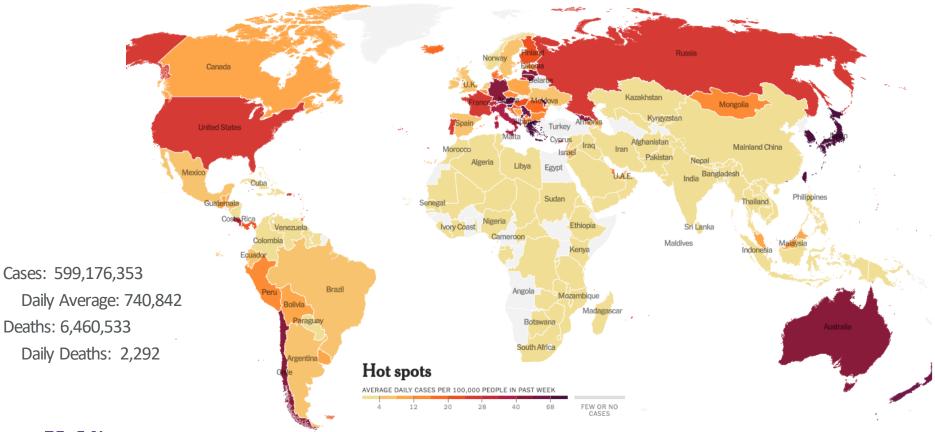


Tecovirimat may be considered for treatment in people infected with *Monkeypox virus*:

- With severe disease (e.g., hemorrhagic disease, confluent lesions, sepsis, encephalitis, or other conditions requiring hospitalization)
- Who are at high risk of severe disease:
 - People with immunocompromising conditions
 - Pediatric populations, particularly patients younger than 8 years of age
 - Pregnant or breastfeeding women
 - People with a history or presence of atopic dermatitis, people with other active exfoliative skin conditions (e.g., eczema, burns, impetigo, varicella zoster virus infection, herpes simplex virus infection, severe acne, severe diaper dermatitis with extensive areas of denuded skin, psoriasis, or Darier disease)
 - People with one or more complication (e.g., secondary bacterial skin infection; gastroenteritis with severe nausea/vomiting, diarrhea, or dehydration; bronchopneumonia; concurrent disease or other comorbidities)
- With aberrant infections involving accidental implantation in eyes, mouth, or other anatomic areas where *Monkeypox virus* infection might constitute a special hazard (e.g., the genitals or anus)

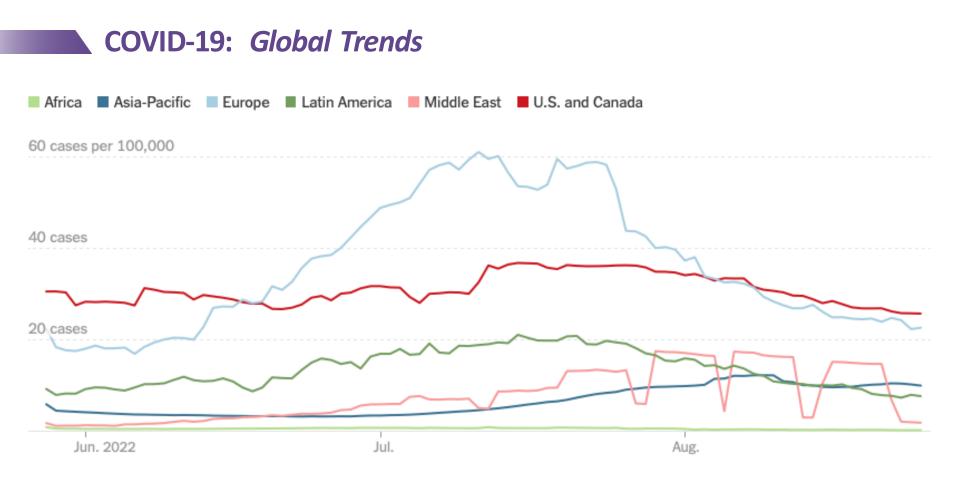


COVID-19: *Global Trends*



Northwestern https://www.nytimes.com/inter

https://www.nytimes.com/interactive/2021/world/covid-cases.html Accessed 26 August 2022

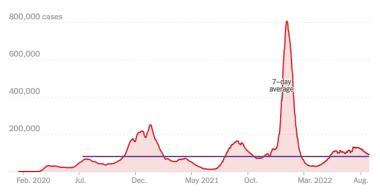


https://www.nytimes.com/interactive/2021/world/covid-cases.html Accessed 8/26/22

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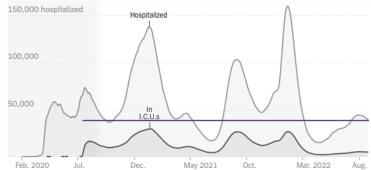
COVID-19: US Trends

New reported cases by day



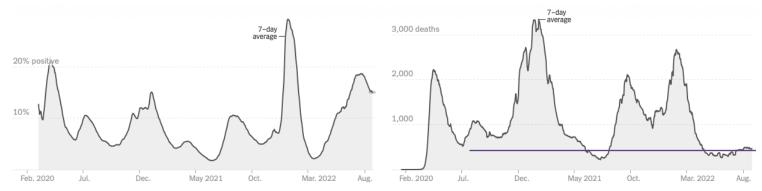
Covid patients in hospitals and I.C.U.s

Early data may be incomplete.



Test positivity rate

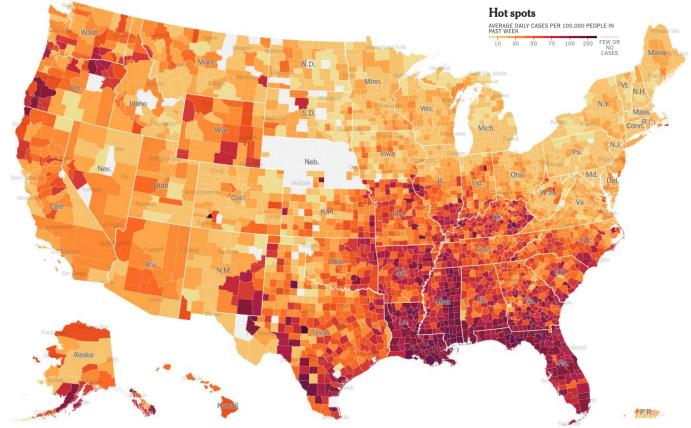
New reported deaths by day





https://www.nytimes.com/interactive/2021/us/covid-cases.html Accessed 8/26/22

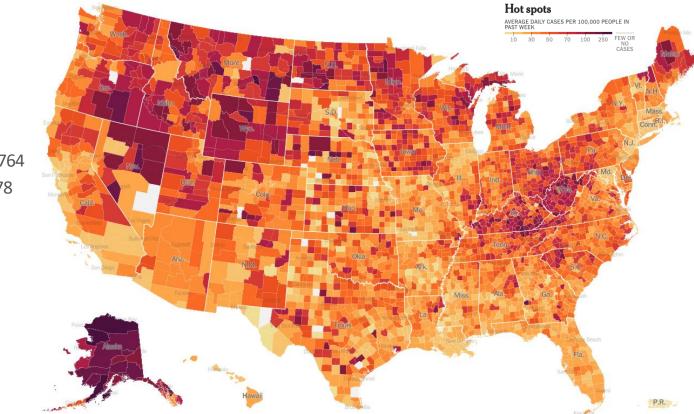
COVID-19: US Trends August 2021





https://www.nytimes.com/interactive/2021/us/covid-cases.html Accessed 8/16/21

COVID-19: US Trends October 2021



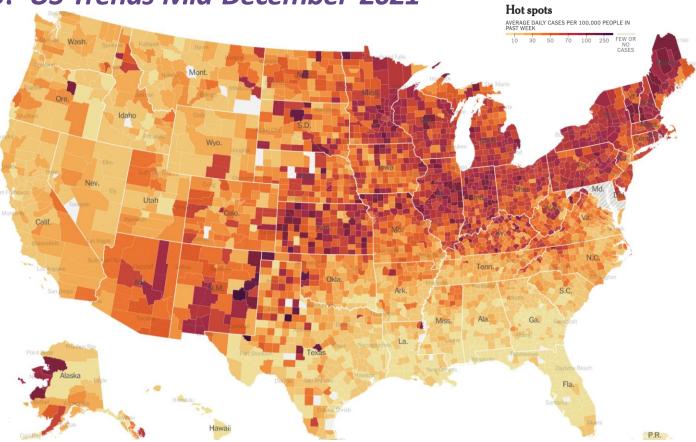
Cases: 46,683,764 Deaths: 701,178



https://www.nytimes.com/interactive/2021/us/covid-cases.html Accessed 10/4/21

COVID-19: US Trends Mid-December 2021

Cases: 46,683,764 Daily New Cases: 119,301 Hospitalizations: 65,962 Deaths: 800,922 Daily Deaths: 1,298

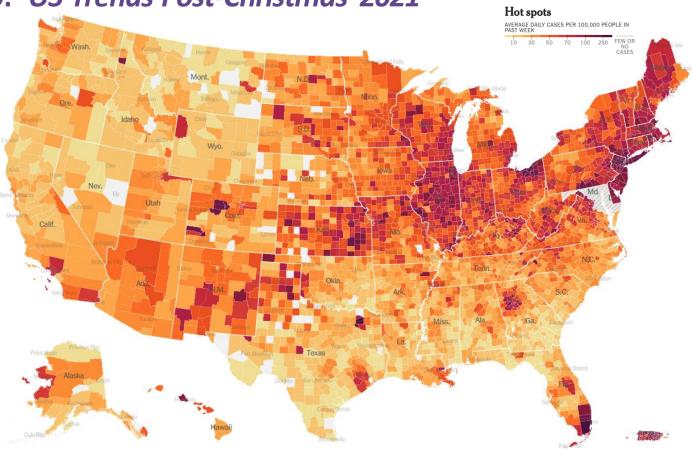




https://www.nytimes.com/interactive/2021/us/covid-cases.html Accessed 12/12/21

COVID-19: US Trends Post-Christmas 2021

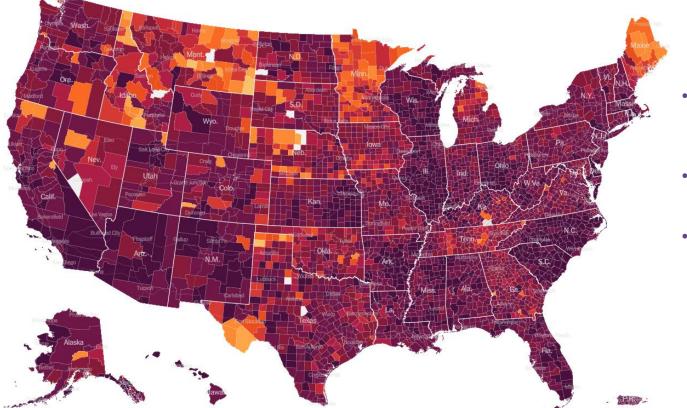
Cases: 52,244,696 Daily New Cases: 214,499 Hospitalizations: 71,034 Deaths: 814,970 Daily Deaths: 1,328





https://www.nytimes.com/interactive/2021/us/covid-cases.html Accessed 12/27/21

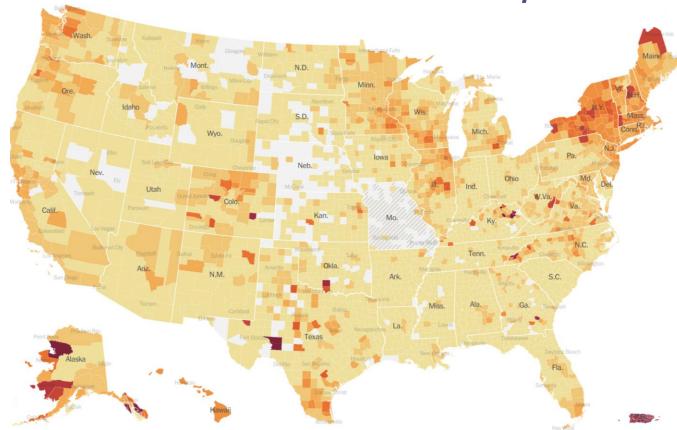
Current Situation: United States – Mid-January



- Cases:
- o Total: 67,705,330
- Daily: 756,752
- Hospitalizations • Daily: 156,894
- Deaths
 - Total: 853740
 - Daily: 1,889



https://www.nytimes.com/interactive/2021/us/covid-cases.html. Accessed 1/19/22



Current Situation: United States – April 2022

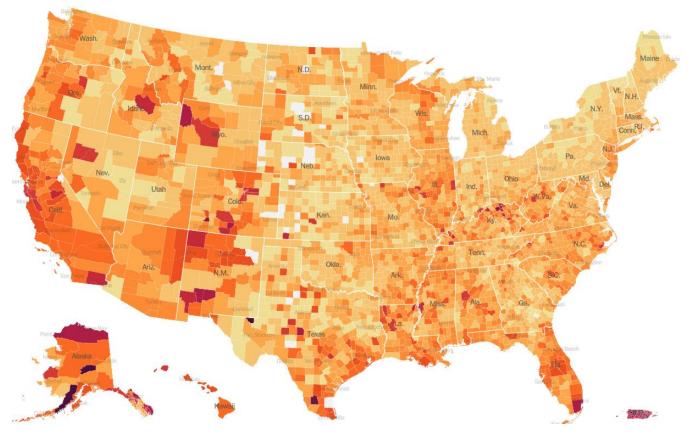
- Cases:
 - o Total: 81,237,905
 - Daily: 56,869
- Hospitalizations

 Daily: 16,897
 - ICU: 1,973
- Deaths
 - Total: 991,921
 - Daily: 316



https://www.nytimes.com/interactive/2021/us/covid-cases.html. Accesse 4/30/22

Current Situation: United States – Current Status

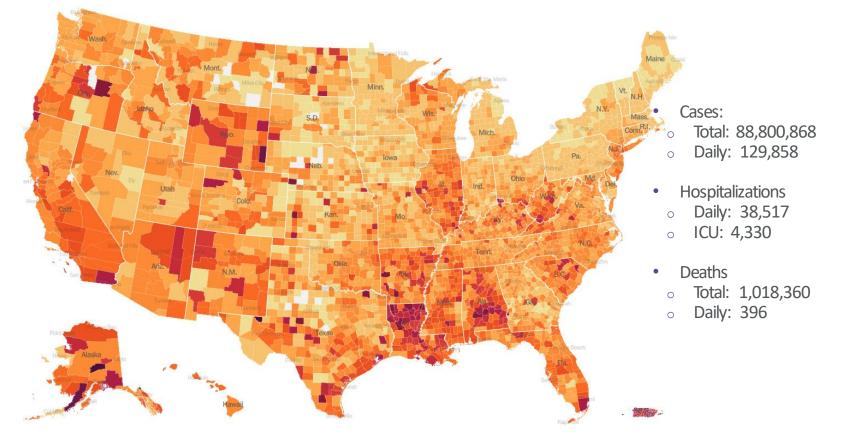


- Cases:
 - Total: 86,899,773
 - Daily: 102,818
- Hospitalizations
 - Daily: 31,650
 - o ICU: 3,498
- Deaths
 - Total: 1,012,486
 - Daily: 348



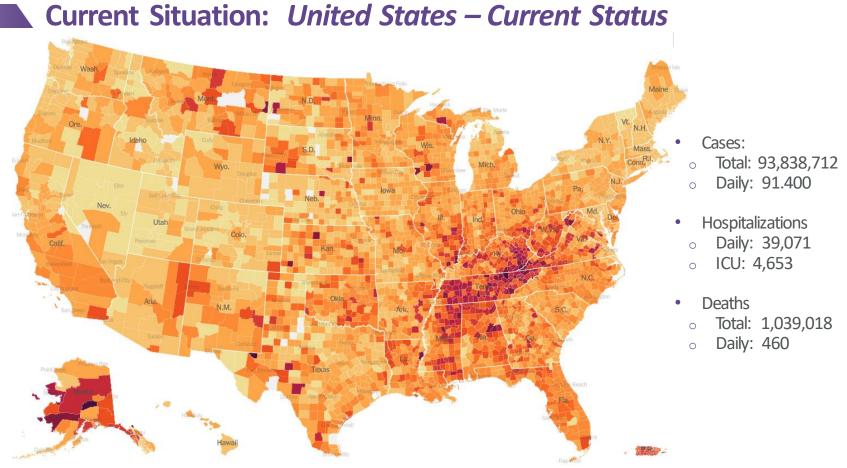
https://www.nytimes.com/interactive/2021/us/covid-cases.html. Accesse 6/27/22

Current Situation: United States – Current Status





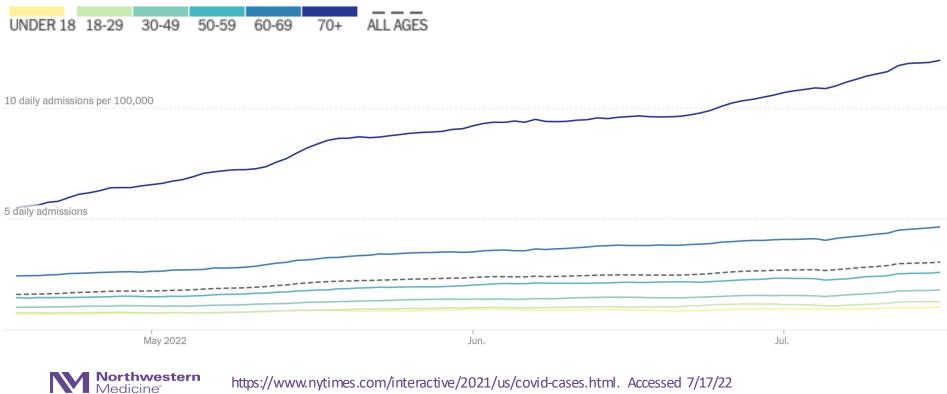
https://www.nytimes.com/interactive/2021/us/covid-cases.html. Accesse 7/13/22



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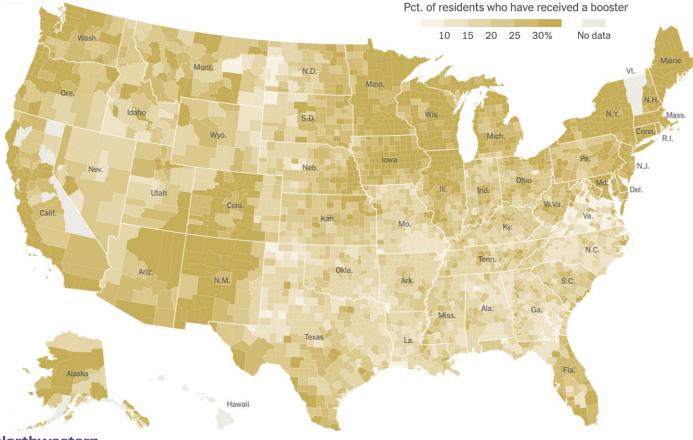
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https://www.nytimes.com/interactive/2021/us/covid-cases.html. Accessed 7/17/22

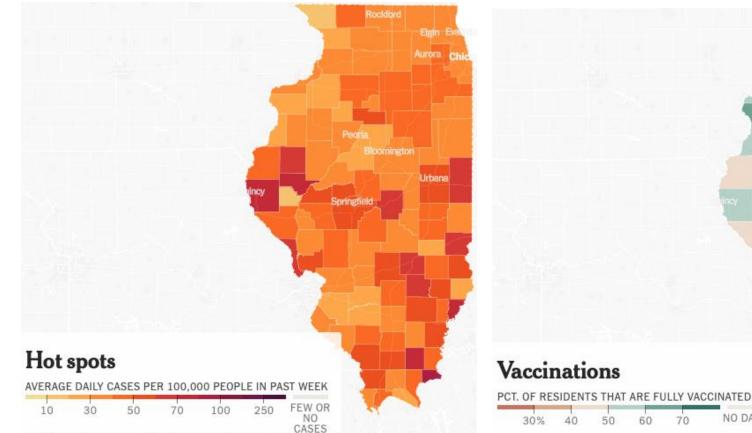
COVID-19: Booster Uptake is Slow



Northwestern Medicine*

https://www.nytimes.com/interactive/2021/us/covid-cases.html. Accessed 7/17/22

COVID-19: Illinois Trends and Vaccination





https://www.nytimes.com/interactive/2021/us/covid-cases.html Accessed 7/13/22

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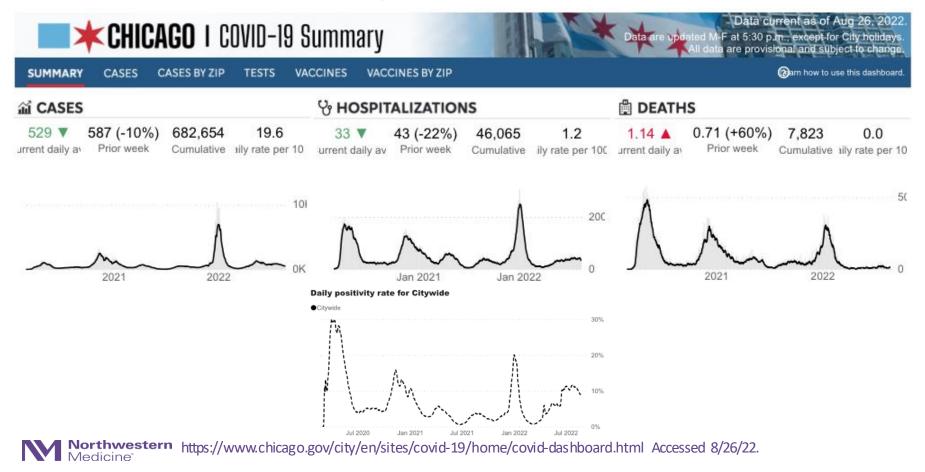
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Urban

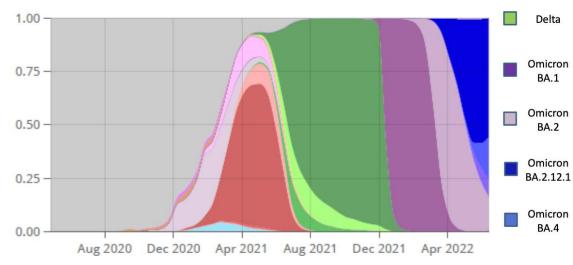
Current Situation: Chicago



SARS-CoV-2 Key Variant

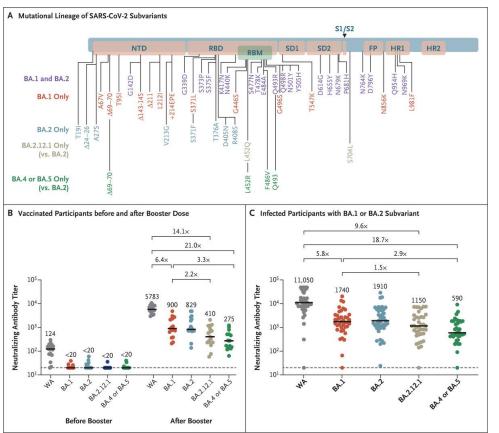
- Omicron Variant:
 - 30 mutations and deletions in the spike protein
 - Associated with increase rates of infection, reduced protection from 2 doses of vaccine Deletions

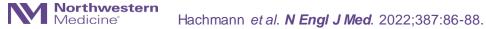
Mutations



Key mutations in the Omicron spike (top view) Northwestern Medicine

Omicron BA.5: What Do We Know?





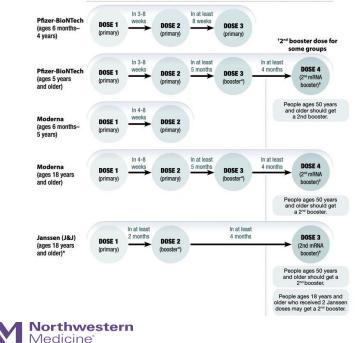
COVID-19 Vaccine: Schedules

AT-A-GLANCE

COVID-19 Vaccination Schedules

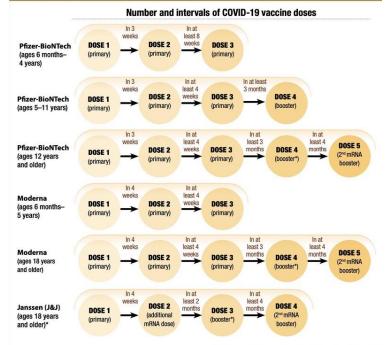
Use the schedules below to determine how many total COVID-19 vaccine doses are recommended based on primary series product, age, and immune status. This schedule does not include clinical details necessary for administering COVID-19 vaccines. For clinical details, see the resources at the end of this document.

COVID-19 Vaccination Schedule for Most People



Number and intervals of COVID-19 vaccine doses

COVID-19 Vaccination Schedule for People Who Are Moderately or Severely Immunocompromised



*Age-appropriate mRNA COVID-19 vaccines are preferred over Janssen COVID-19 Vaccine for primary and booster vaccination. Janssen COVID-19 Vaccine should only be used in limited situations. See: https://www.cdc.gov/vaccines/covid-19/clinical-considerations/interimconsideration-us.htmi/econsiderations-anseen.



For more specific clinical guidance, see:

- Interim COVID-19 Immunization Schedule
- Interim Clinical Considerations for Use of COVID-19 Vaccines Currently Approved or Authorized in the United States

cdc.gov/coronavirus

SARS-CoV-2 Boosters: Bivalent Vaccine

- Same Moderna and Pfizer vaccine technology
- Contains mRNA for 2 different spike proteins
 - $_{\odot}$ Original Wuhan-like virus
 - $_{\rm O}$ Omicron virus
 - Studies of the BA.1 virus showed improvement in protection but less protection vs. BA.5
 - $_{\odot}$ New vaccine to include BA.4/BA.5 spike protein
 - Using studies to suggest what the antibody levels against BA.5 will be with this approach
 - Clinical studies will follow authorization



Numbers are Going Back Up: Have a Plan to Stay Safe

- •Generally anything outdoors is safe
- •Masking in any public indoor space is the safest
- •Many with symptoms are mild (even allergy like) test if unsure
- •Get Evusheld if you are eligible
- •Get your booster and stay Up To Date on vaccine
- •Call your doctor if sick, especially if you have COVID-19
- More than COVID-19 is circulating!

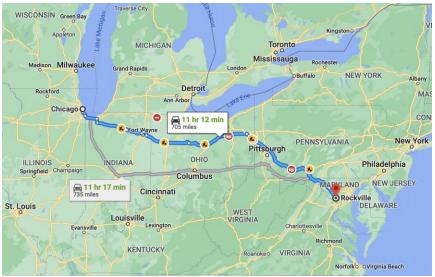


What to Expect for the Fall

- There will be more variants that emerge
- There will be a bivalent COVID-19 booster available SOON
- There will be a heavy and early flu season this year (along with lots and lots of other viruses in addition to COVID-19)
- I suspect that most people will get COVID-19 before the end of the year
- What can you do?
 - $_{\circ}~$ Keep your guard up
 - Get your boosters, wear your mask and keep others that are sick away
 - $_{\odot}\,$ Get your flu shot (and there is likely to be an RSV shot in the near future)
 - Call your doctor if you are sick and have plans to access medications if you will be traveling Northwestern Medicine*

What Else to Expect for the Fall: Dr. Ison to Move to NIH!







Are you a registered organ donor?

Jorthwestern University

Translational Sciences Institute

I am!

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Questions?

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