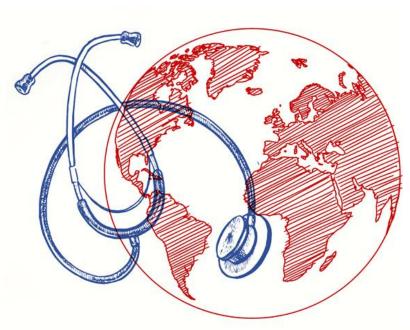
Global Health Cast 60 February 19, 2024





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What we talk about today

- Mild flu tied to 2x risk of heart attack, stroke in older patients
- > 1st fatal case of Alaskapox: man undergoing cancer treatment
- Ebola vaccine cut deaths in half during DRC outbreak
- Vaccine-specific Adverse Event or side effects of COVID vaccination 1 Bell's Palsy (BP)
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Mild flu tied to 2x risk of heart attack, stroke in older patients

The Journal of Infectious Diseases

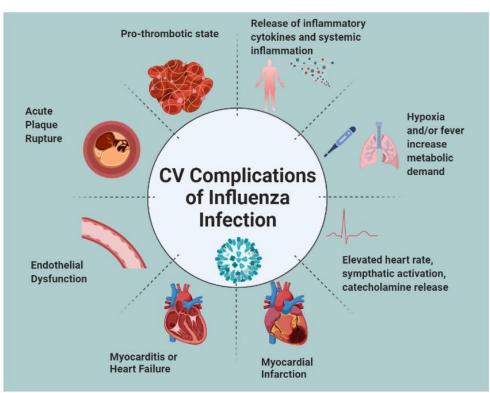
MAJOR ARTICLE

Risk of cardiovascular events after influenza: a population-based Self Controlled Case Series study, Spain 2011-2018

Cintia Muñoz-Quiles^{a, b, *}, Mónica López-Lacort^{a, b, *}, Arantxa Urchueguía^{a, b}, Javier Díez-Domingo^{a, b, c}, Alejandro Orrico-Sánchez^{a, b, c}

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"This work reinforces the official recommendations for influenza prevention in at-risk groups and should also increase the awareness of even milder influenza infection and its possible complications in the general population."





State of Alaska Epidemiology



Bulletin

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> Bulletin No. 2 February 9, 2024

Fatal Alaskapox Infection in a Southcentral Alaska Resident

Background

Orthopoxviruses are double-stranded DNA viruses, and many are zoonotic, occurring in a range of mammalian taxa.¹ Alaskapox virus (AKPV) is a recently discovered orthopoxvirus that was first identified in an adult living near Fairbanks in 2015.²

Seven AKPV infections to date have been reported to the Alaska Section of Epidemiology (SOE). Until December 2023, all reported infections occurred in residents of the Fairbanks area and involved self-limiting illness consisting of a localized rash and lymphadenopathy.³ Small mammal testing in the Fairbanks area identified evidence of current or prior AKPV infection in four different species (though mostly in red-backed voles).⁴ Evidence suggestive of prior AKPV infection has also been documented in at least one domestic pet linked to a patient. The extent of AKPV's geographic distribution and animal reservoirs remain unknown. This *Bulletin* describes a recently reported fatal case of Alaskapox in a resident of the Kenai Peninsula.

Case Report

In mid-September 2023, an elderly man from the Kenai Peninsula with a history of drug-induced immunosuppression secondary to cancer treatment noted a tender red papule in his right axilla. Over the next 6 weeks, he presented to his primary provider and the local emergency department (ED) several times for clinical evaluation of the lesion and was prescribed multiple antibiotic regimens. A punch biopsy revealed no evidence of malignancy or bacterial infection. Despite antibiotic therapy, the patient experienced fatigue and increasing induration and pain in the right axilla and shoulder. On November 17, he was hospitalized due to extensive progression of presumed infectious cellulitis that impacted the range of motion of his right arm. The patient was subsequently transferred to a hospital in Anchorage.

that regularly hunted small mammals and frequently scratched the patient, including one notable scratch near his right axilla in the month prior to rash onset. The patient did not report other recent contact with small mammals but did report gardening in his backyard through September 2023. Serum and mucosal swabs collected from the stray cat were submitted to CDC for antibody and orthopoxyirus testing; all tests were negative.

Discussion

This is the first case of severe Alaskapox infection resulting in hospitalization and death. The patient's immunocompromised status likely contributed to illness severity. Moreover, being the first case of Alaskapox identified outside of the Interior region, it indicates that AKPV appears to be more geographically widespread in Alaska's small mammals than previously known and warrants increased statewide awareness among clinicians. The route of exposure in this case remains unclear, although scratches from the stray cat represent a possible source of inoculation through fomite transmission. SOE is working with the University of Alaska Museum and CDC to test small mammals for AKPV outside of the Interior region.

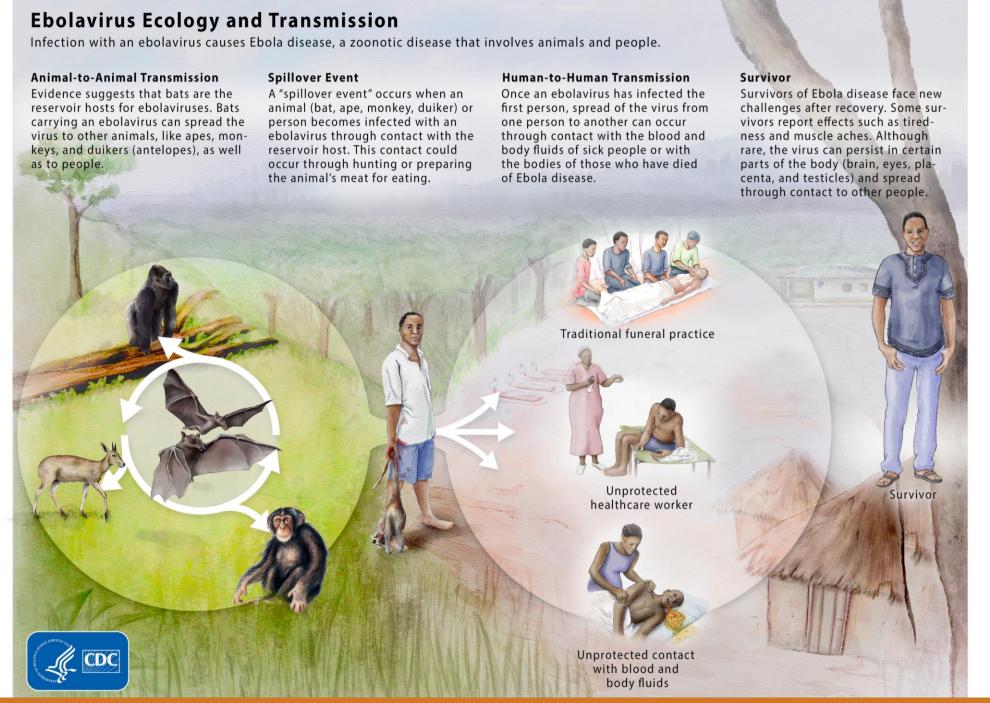
Recommendations

- Clinicians should become familiar with the clinical features of Alaskapox and consider testing for orthopoxvirus infection in patients with a clinically compatible illness.²⁻⁴
- Promptly report suspected Alaskapox cases to SOE at 907-269-8000; SOE staff can help facilitate testing.
- Advise outpatients with suspected Alaskapox to avoid touching lesions, keep lesions dry and covered, practice good hand hygiene, avoid sharing cloth that might have been in contact with lesions, and launder clothing and linens separately from other household items.⁵

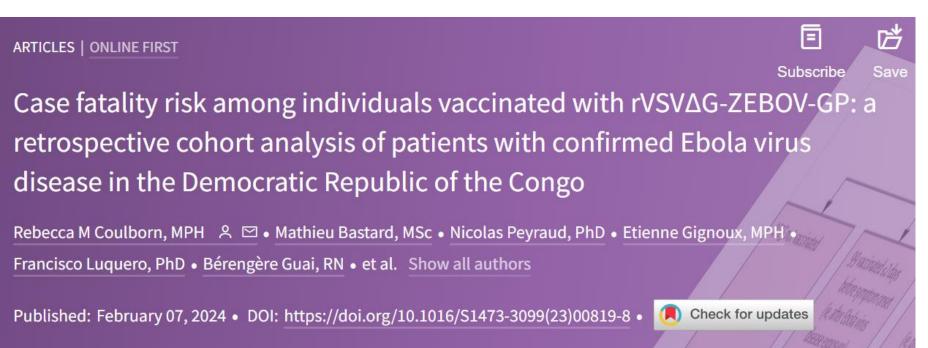


Rare virus Alaskapox 1st reported fatal case





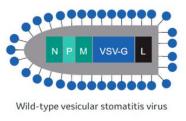


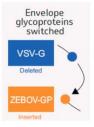


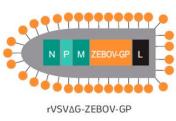
ERVEBO® (Ebola Zaire Vaccine, Live also known as V920, rVSVΔG-ZEBOV-GP or rVSV-ZEBOV) is approved by the U.S. Food and Drug Administration (FDA) for the prevention of disease caused by Ebola virus (EBOV; species Zaire ebolavirus) in individuals 12 months of age and older as a single dose administration.

Interpretation

To our knowledge, this is the **first observational study describing the protective effect of rVSVΔG-ZEBOV-GP vaccination against death among patients with confirmed Ebola virus disease** admitted to an Ebola health facility. Vaccination was protective against death for all patients, even when adjusted for Ebola virus disease-specific treatment, age group, and time from symptom onset to admission.









SARS-CoV-2 Vaccine Platforms: Vaccine-specific AE

Coincidental or Causal Association? Frequency?

Whole Virus (inactivated)

Virus-like Particles

Viral Vector

Subunit

Viral Vector

DNA

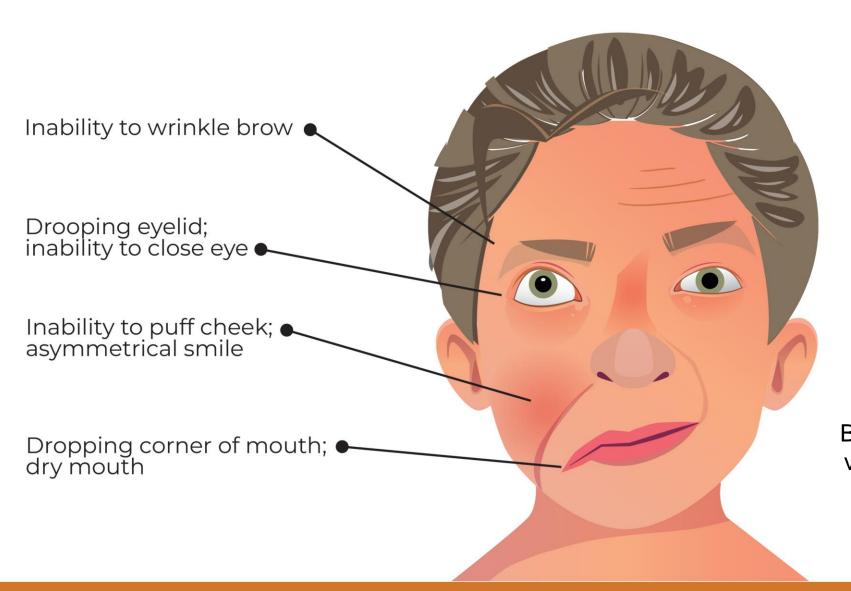
RNA



- 1. Facial Paralysis (Bell's Palsy, BP)
- 2. Diseases from activation of the clotting system
- 3. Myocarditis



FACIAL NERVE PALSY



BP after SARS-CoV-2 vaccination is usually mild and self-limited



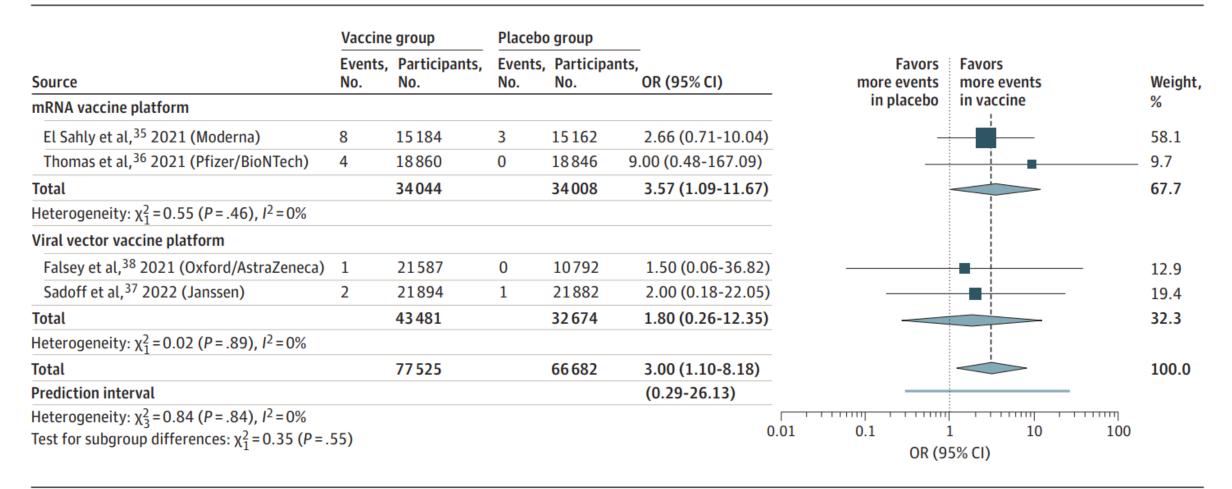
Facial paralysis (Bell's Palsy) following COVID vaccination Systematic Review and Meta-Analysis

	Cases	Total	Per 1,000,000
First dose	10,139	59,235,299	171.2
Second dose	206	4,888,784	42.1
Unspecified vaccines	3,445	12,386,275	278.1
Oxford/AstraZeneca ChAdOx1 nCoV-19	5,933	33,224,858	178.6
Sinovac	65	1,407,798	46.2
Pfizer/BioNTech BNT162b2	5,242	199,455,808	26.3
Janssen (Johnson & Johnson) Ad26.COV2-S	809	128,085,700	6.3
Moderna	1,923	307,350,232	6.3
Total	17,417	687,371,182	25.3

BP after SARS-CoV-2 vaccination is usually mild and self-limited



Figure 1. Bell Palsy Events in Groups of Vaccine Recipients vs Saline Placebo Recipients, With Data From Randomized Clinical Trials



Dashed line indicates the point estimate of the overall effect; dotted line, no effect; diamonds, overall effects. OR indicates odds ratio.



Figure 2. Bell Palsy Events in Groups of mRNA-Vaccinated Participants vs Unvaccinated Participants, With Data From Observational Studies

	Vaccinated group		Unvaccinated group				
Source	Events, No.	Participants/ doses, No.	Events, No.	Participants, No.	OR (95% CI)	Favors Favors more events	Weight,
Cohort subgroup						in unvaccinated in vaccinated	%
Klein et al, ⁴² 2021	543	11845128	2379	11845128	0.23 (0.21-0.25)		13.2
Lai et al, ⁴⁶ 2022 (first dose)	1	138141	3	136743	0.33 (0.03-3.17)		3.6
McMurry et al, 44 2021 (second dose)	10	41909	30	41909	0.33 (0.16-0.68)		10.5
McMurry et al, 44 2021 (first dose)	16	68266	45	68266	0.36 (0.20-0.63)		11.3
Shasha et al, ⁴³ 2022 (second dose)	8	131033	12	131033	0.67 (0.27-1.63)	i	9.4
Shasha et al, ⁴³ 2022 (first dose)	23	233159	24	233159	0.96 (0.54-1.70)	<u> </u>	11.3
Barda et al, ⁴¹ 2021	81	923692	59	923692	1.37 (0.98-1.92)	—	12.5
Bardenheier et al, ⁴⁵ 2021	1	16924	0	11072	1.96 (0.08-48.19)		2.1
Lai et al, ⁴⁶ 2022 (second dose)	4	119664	2	118300	1.98 (0.36-10.80)		5.3
Total		13517916		13509302	0.59 (0.34-1.01)		79.2
Heterogeneity: $\chi_8^2 = 133.4 \ (P < .001), I^2 = 94\%$							
Case-control subgroup							
Shemer et al, ³⁹ 2021	21	65	16	46	0.89 (0.40-1.99)		9.9
Wan et al,40 2022 (Pfizer/BioNTech subset)	14	45	256	1353	1.94 (1.01-3.69)		10.9
Total		110		1399	1.37 (0.64-2.90)		20.8
Heterogeneity: $\chi_1^2 = 2.17 (P = .14), I^2 = 54\%$							
Total		13518026		13510701	0.70 (0.42-1.16)		100.0
Prediction interval					(0.13-3.83)		
Heterogeneity: $\chi_{10}^2 = 175.75 (P < .001), I^2 = 949$							
Test for subgroup differences: $\chi_1^2 = 3.18$ ($P = .0$)	7)					0.05 0.1 1 10	100
						OR (95% CI)	



Figure 4. Bell Palsy in Groups of SARS-CoV-2 Infection vs SARS-CoV-2 Vaccine Recipients, With Data From Observational Studies

Source	Total No. of infected patients	Total No. of vaccine recipients/doses	RR (95% CI)	Favors more events in SARS-CoV-2 vaccine	Favors more events in SARS-CoV-2 infection	Weight, %
Patone et al, ⁵⁶ 2021	2005280	32552534	1.84 (1.45-2.33)			26.5
Tamaki et al, ⁵⁵ 2021	348088	63551	6.80 (3.50-13.21)			22.2
Li et al, ⁵⁷ 2022 (SIDIAP database)	288030	4372633	5.71 (4.50-7.25)			26.5
Barda et al, ⁴¹ 2021	180674	923692	1.64 (1.06-2.55)			24.8
Total	2822072	37912410	3.23 (1.57-6.62)			100.0
Prediction interval			(0.11-96.91)			
Heterogeneity: $\chi_2^2 = 57.14 \ (P < .001), I^2 = 95\%$					· · · · · · · · · · · · · · · · · · ·	
-				0.1	1 10	100
					RR (95% CI)	

Dotted line indicates no effect; diamond, overall effect. RR indicates risk ratio; SIDIAP, Spanish database of Information System for Research in Primary Care.



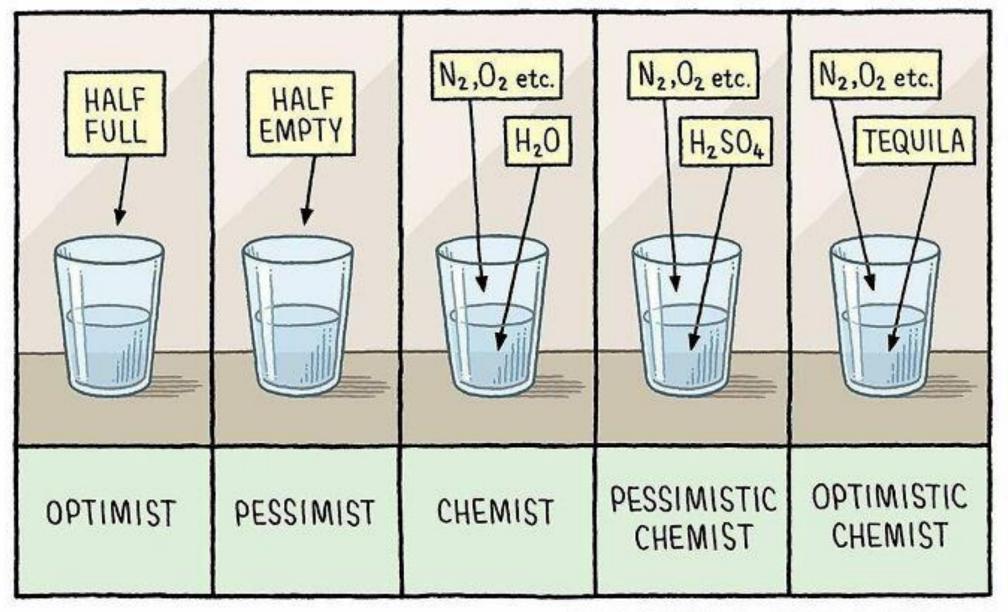
Conclusion

- Observational studies: mRNA SARS-CoV-2—vaccinated participants had
 - no significant increase in BP incidence vs the unvaccinated participants.
- Current study: Strong association between the SARSCoV-2 vaccine and BP in 4 RCTs,
 - Conclusion: BP is a result of SARS-CoV-2 vaccine exposure.
- SARS-CoV2 <u>vaccination</u> does cause BP; BUT:
- SARS-CoV-2 <u>infection</u> has 3.23- fold higher BP risk vs. <u>vaccination</u>;
- This favors a protective role of vaccination in reducing the incidence of BP associated with exposure to SARS-CoV-2.



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TOM GAULD for NEW SCIENTIST