

Global situation in humans

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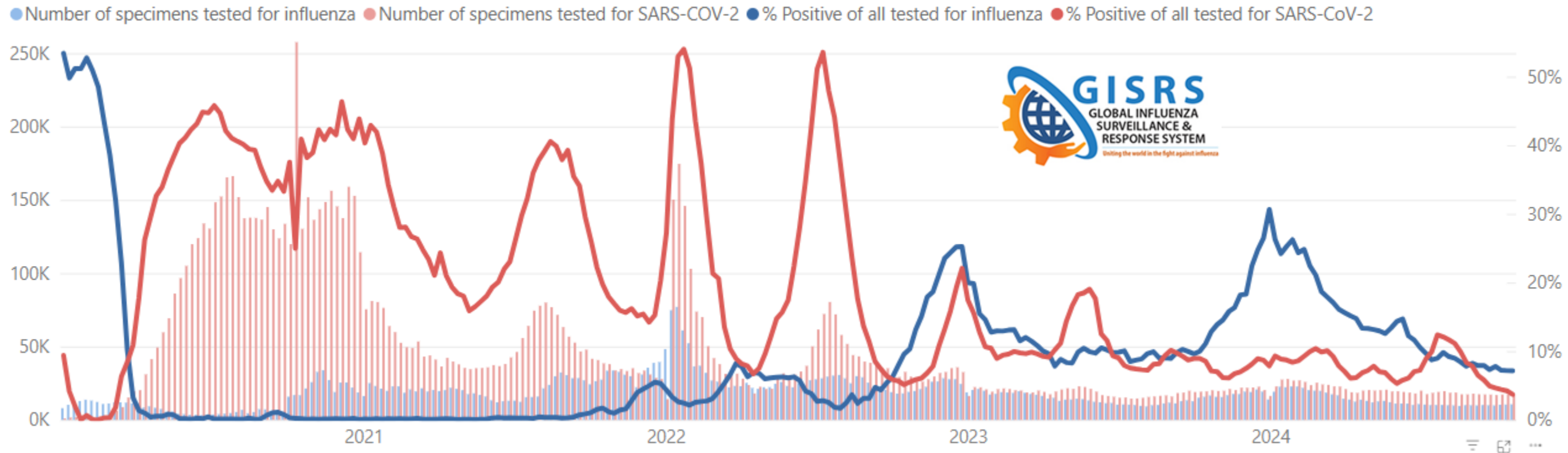
14 November 2024

Preparing for containment and mitigation of pandemic H5N1 influenza
Uses of Statistical and Mathematical Modeling
- a virtual meeting

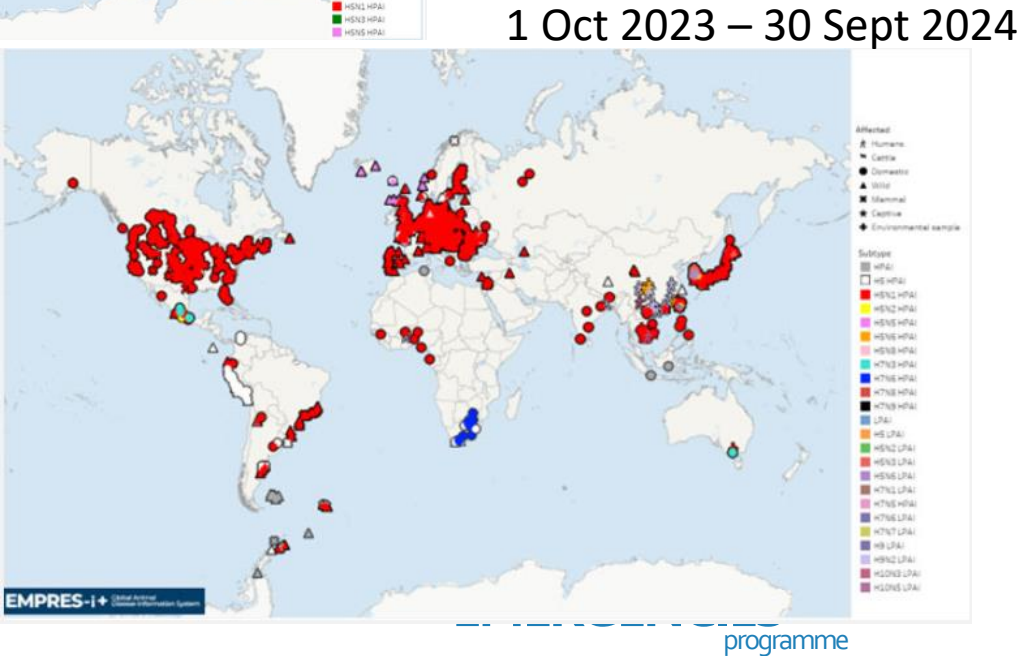
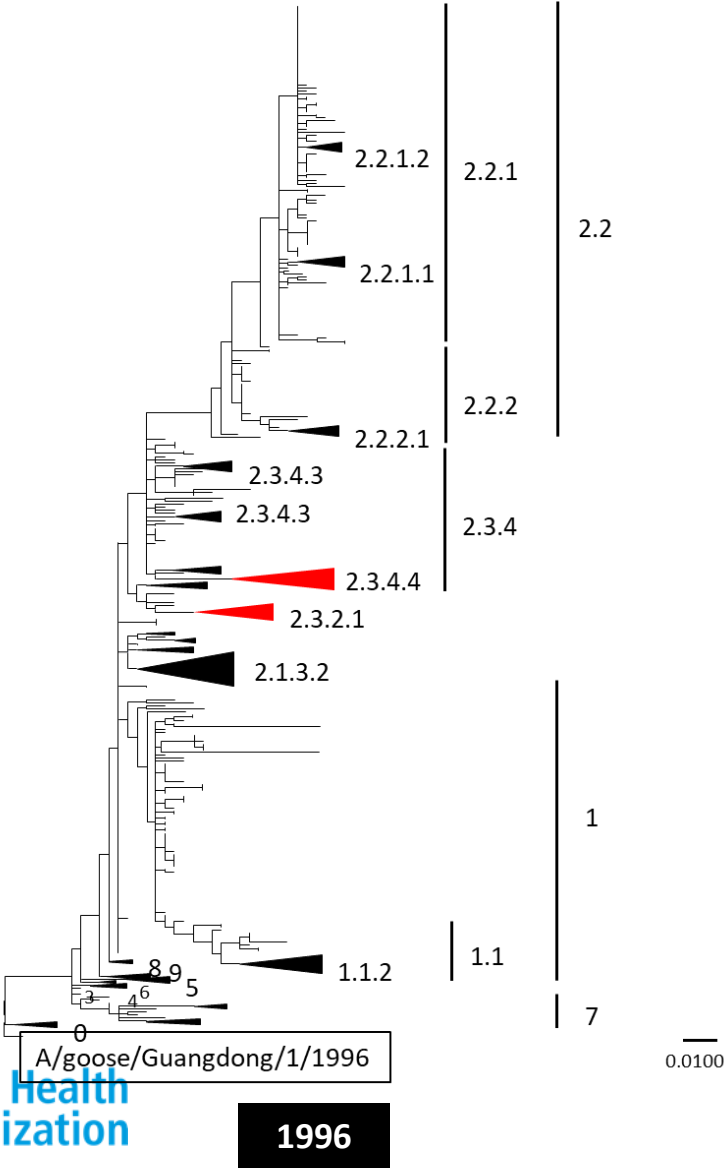


Global co-circulation of influenza and SARS-CoV-2

2003 – 7 Nov 2024



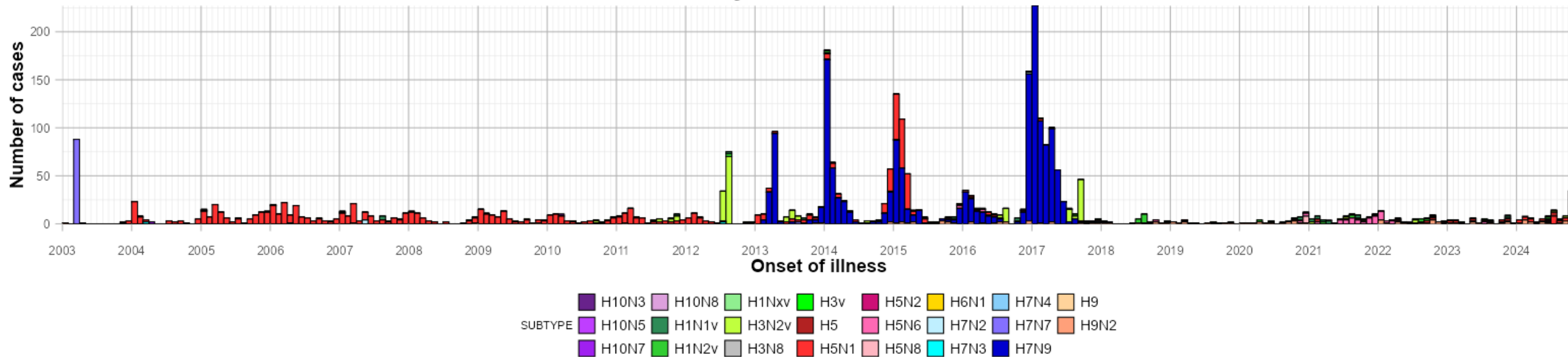
Evolution and spread of avian influenza viruses



Human infections* with zoonotic influenza A viruses

2003 to 7 Nov 2024

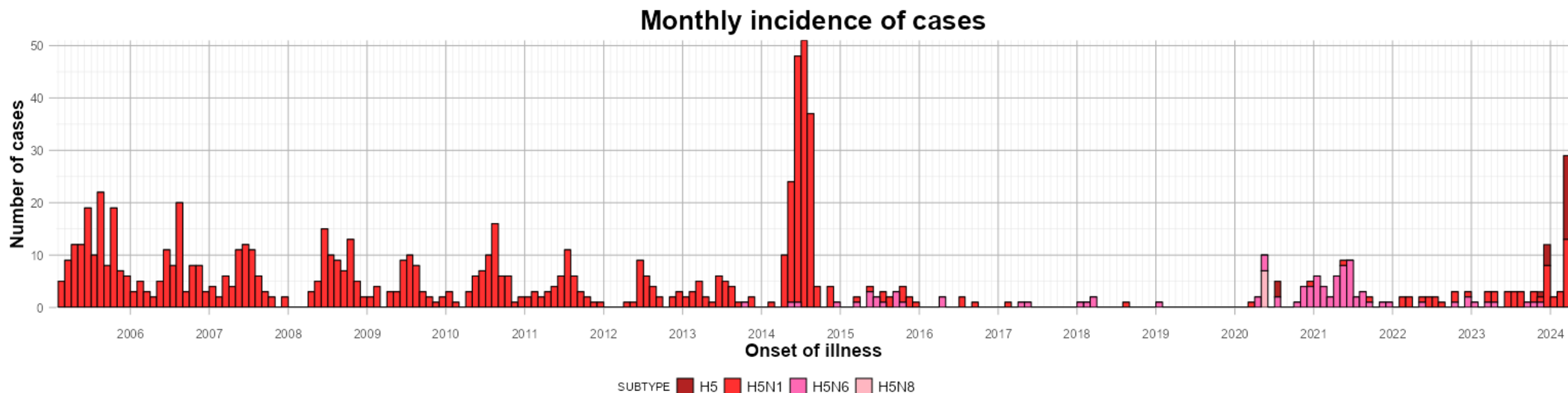
Monthly incidence of cases



*Not including H3N2v cases, USA, 2012 (309).

Human infections with zoonotic influenza A(H5) viruses

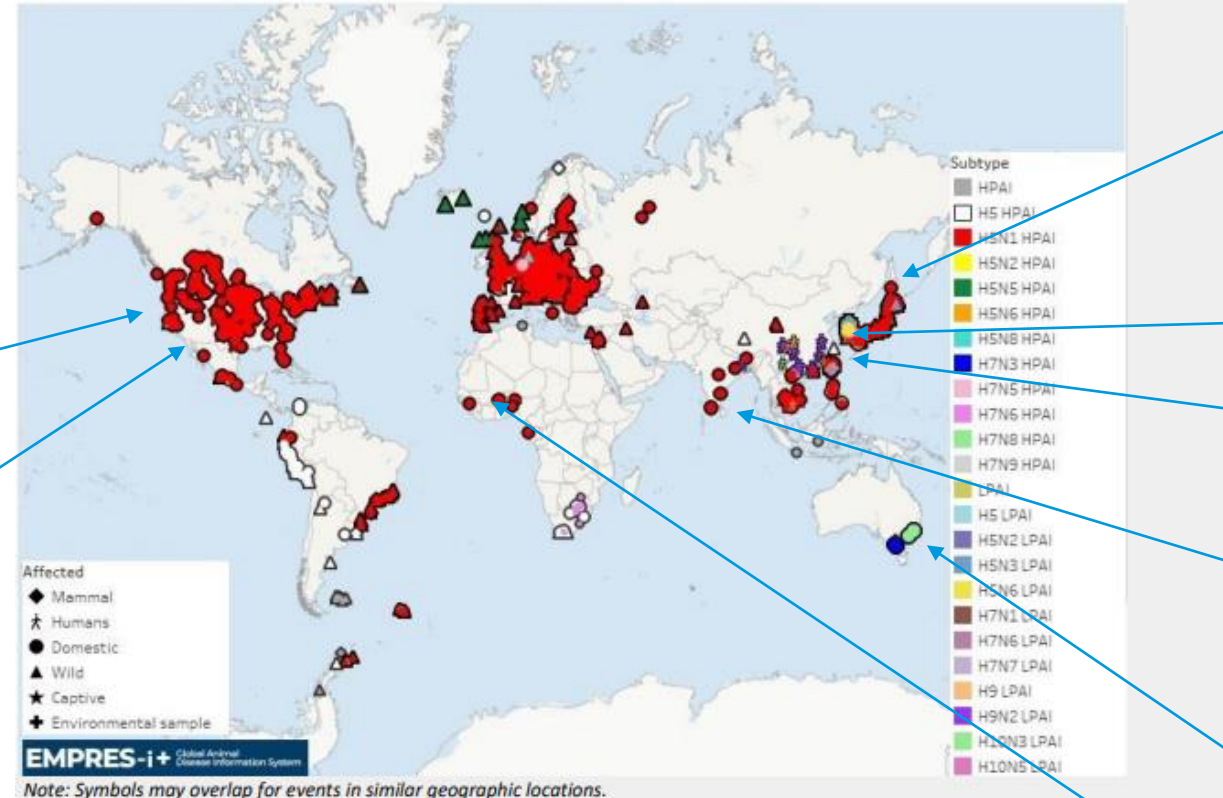
2003 to 2024



Avian influenza viruses with zoonotic potential and human cases

Oct 2023-2024

Map 1. Global distribution of AIV with zoonotic potential observed since 01 October 2023 (i.e. current wave)
Confirmed Avian Influenza events globally from 1 October 2023 to date



USA: H5 (46)

Mexico:
H5N2 (1)

China:
H5N1 (1),
H5N6 (5),
H9N2 (19)
H10N3 (1),
H10N5 (1),

Cambodia:
H5N1 (14)

Viet Nam:
H5N1 (1)
H9N2 (1)

India:
H9N2 (1)

Australia:
H5N1 (1)

Ghana: H9N2 (1)

<https://www.fao.org/animal-health/situation-updates/global-aiv-with-zoonotic-potential>

Genetic diversity of avian influenza viruses infecting humans

(2.3.4.4b)



Health officials in US conducted sero-surveys

- Recent HPAI A(H5) infections among dairy workers in two states.
- 7% of participating dairy workers had evidence of recent infection with HPAI A(H5) virus.
- All were either mild or no symptoms.
- None of the positive ones used respiratory protection.

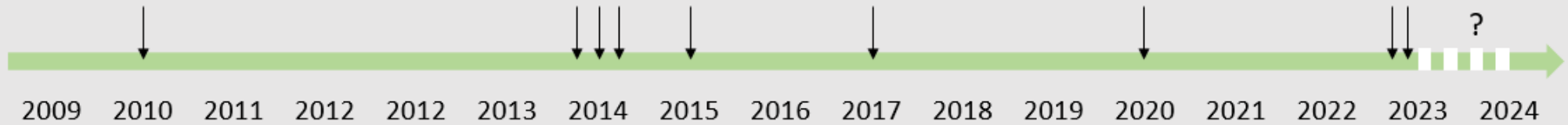


Nguyen et al bioRxiv
<https://doi.org/10.1101/2024.05.01.591751>; t
Phylogenetic tree including WA cases – Courtesy of T. Davis

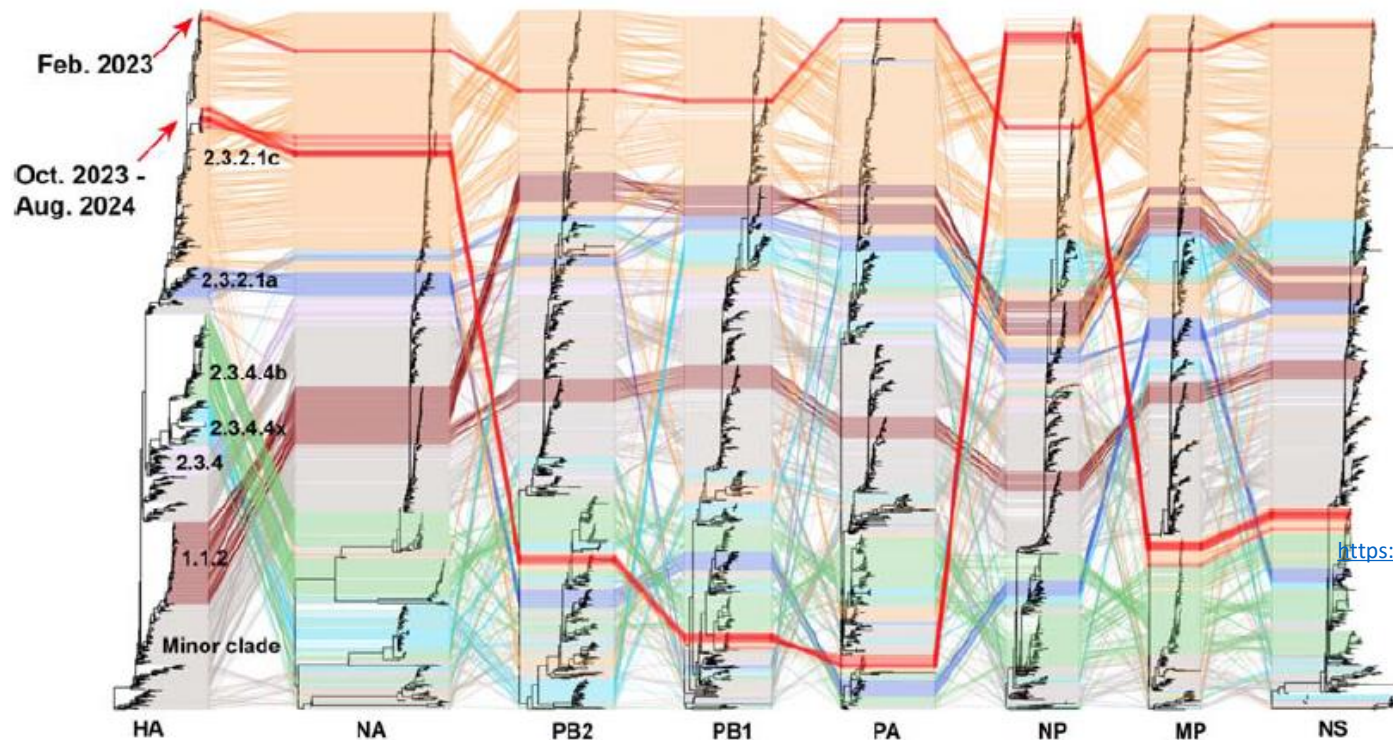
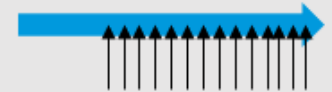
Genetic diversity of avian influenza viruses infecting humans

(2.3.2.1c)

1) Before introducing clade 2.3.4.4b A(H5)-derived gene segments



2) After introducing clade 2.3.4.4b A(H5)-reassortment (after Oct 2023)



<https://www.medrxiv.org/content/10.1101/2024.11.04.24313747v1.full.pdf>

Antigenicity of avian influenza viruses and candidate vaccine viruses (avian and swine zoonotic influenza viruses)

From WHO vaccine composition consultation Sept 2024:

2.3.4.4b

Haemagglutination inhibition assay of human derived A(H5) clade 2.3.4.4b viruses

Reference antigens	Subtype	Clade	Post-infection ferret antisera					TX/37
			VN/ 1203	CNIC- 21099	IDCDC- RG71A	IDCDC- RG78A	IDCDC- RG80A	
A/Viet Nam/1203/2004	H5N1	1	2560	<10	<10	<10	<10	<10
CNIC-21099 (A/Fujian-Sanyuan/21099/2017)	H5N6	2.3.4.4b	20	80	160	80	1280	40
IDCDC-RG71A (A/Astrakhan/3212/2020-like)	H5N8	2.3.4.4b	10	80	320	160	1280	80
IDCDC-RG78A (A/American Wigeon/South Carolina/22-000345-001/2021)	H5N1	2.3.4.4b	10	80	640	320	1280	80
IDCDC-RG80A (A/chicken/Ghana/AVL-763_21VIR7050-39/2021)*	H5N1	2.3.4.4b	<10	10	80	40	1280	80
A/Texas/37/2024	H5N1	2.3.4.4b	10	40	320	160	1280	80
Test antigens								
A/Colorado/109/2024	H5N1	2.3.4.4	10	10	320	320	1280	80
A/Colorado/134/2024	H5N1	2.3.4.4	10	10	160	160	1280	80
A/Colorado/137/2024	H5N1	2.3.4.4	10	10	160	320	1280	80
A/Colorado/138/2024	H5N1	2.3.4.4	10	40	160	160	1280	80
A/Michigan/90/2024	H5N1	2.3.4.4	10	40	160	320	1280	80

Haemagglutination inhibition assay of A(H5) clade 2.3.2.1c viruses

Reference antigens	Subtype	Clade	Post-infection ferret antisera		
			SJ001	duck/VN/ 1584	NIBRG- 301
SJ001 (A/duck/Bangladesh/19097/2013-like)	H5N1	2.3.2.1a	80	80	160
A/duck/Viet Nam/NCVD-1584/2012	H5N1	2.3.2.1c	20	80	160
NIBRG-301 (A/duck/Viet Nam/NCVD-1584/2012)	H5N1	2.3.2.1c	80	160	640
Test antigens					
A/Cambodia/24020155/2024 (January 2024)	H5N1	2.3.2.1c	<10	40	80
A/Cambodia/24020179/2024 (February 2024)	H5N1	2.3.2.1c	<10	20	80
A/Cambodia/SVH240441/2024 (July 2024)	H5N1	2.3.2.1c	<10	20	40
A/Viet Nam/Khanh Hoa/RV1-005/2024 x PR8	H5N1	2.3.2.1c	<10	20	80

2.3.2.1c

Antigenicity of avian influenza viruses and candidate vaccine viruses

(avian and swine zoonotic influenza viruses)

Table 12. Status of influenza A(H3N2)v candidate vaccine virus development

Candidate vaccine viruses (like viruses)	Lineage	Type	Institution*	Available
NYMC X-203 (A/Minnesota/11/2010)	3.1888	Conventional	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	3.1888	Conventional	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	3.1888	Conventional	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	3.1888	Conventional	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	3.1888	Conventional	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	3.1888	Conventional	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	3.1888	Conventional	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	3.1888	Conventional	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	3.1888	Conventional	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	3.1888	Conventional	CDC	Yes

H3N8

Table 5. Status of influenza A(H5) candidate vaccine virus development*

Candidate vaccine viruses (like virus)	Clade	Institution*	Available
CDC-RG (A/Viet Nam/1203/2004)	1	CDC	Yes
SJRG-161052 (A/Viet Nam/1203/2004)	1	CDC	Yes
NIBRG-14 (A/Viet Nam/1194/2004)	1	CDC	Yes
NIBRG-88 (A/Cambodia/R0405050/2007)	1.1	CDC	Yes
IDCDC-RG34B (A/Cambodia/X0810301/2013)	1.1.2	CDC	Yes
SJRG-166614 (A/duck/Hunan/795/2002)	2.1.1	CDC	Yes
CDC-RG2 (A/Indonesia/5/2005)	2.1.3.2	CDC	Yes
NIIDRG-9 (A/Indonesia/NIHRD11771/2011)	2.1.3.2a	CDC	Yes
SJRG-163222 (A/bar-headed goose/Qinghai/1A/2005)	2.2	CDC	Yes
IBCDC-RG7 (A/chicken/India/NIV33487/2006)	2.2	CDC	Yes
SJRG-163243 (A/whooper swan/Mongolia/244/2005)	2.2	CDC	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	CDC	Yes
NIBRG-23 (A/Egypt/109/2007)	2.2.1	MHRA	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	MHRA	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	MHRA	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	MHRA	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	MHRA	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	MHRA	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	MHRA	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	MHRA	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	MHRA	Yes
IDCDC-RG11 (A/Egypt/109/2007)	2.2.1	MHRA	Yes

H9N2

H10Nx

Table 13. Status of influenza A(H1)v candidate vaccine virus development

Candidate vaccine viruses (like viruses)	Clade	Type	Institution*	Available
NYMC X-213 (A/Indiana/10/2011)	1C.2.3	Conventional	CCDC	Yes
NYMC X-213 (A/Indiana/10/2011)	1A.3.3.3	Reverse genetics	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	1B.2.1	Reverse genetics	CDC	Yes
NYMC X-213 (A/Indiana/10/2011)	1C.2.1	Reverse genetics	MHRA	Pending
NYMC X-213 (A/Indiana/10/2011)	1C.2.1	Reverse genetics	MHRA	Pending
NYMC X-213 (A/Indiana/10/2011)	1C.2.1	Reverse genetics	MHRA	Pending
NYMC X-213 (A/Indiana/10/2011)	1C.2.1	Reverse genetics	MHRA	Pending
NYMC X-213 (A/Indiana/10/2011)	1C.2.1	Reverse genetics	MHRA	Pending
NYMC X-213 (A/Indiana/10/2011)	1C.2.1	Reverse genetics	MHRA	Pending
NYMC X-213 (A/Indiana/10/2011)	1C.2.1	Reverse genetics	MHRA	Pending

H3N2 variant

Table 7. Status of influenza A(H10N3) candidate vaccine virus development

Candidate vaccine viruses (like virus)	Lineage	Institution*	Available
A/Jiangsu/428/2021	Eurasian	CDC	Pending

H1Nx variant

Table 6. Status of influenza A(H9N2) candidate vaccine virus development

Candidate vaccine viruses (like virus)	Lineage	Type	Institution*	Available
A/Hong Kong/1073/99	G-like (G1)	Reverse genetics	MHRA	Yes
NIBRG-91 (A/chicken/Hong Kong/G9/97)	B-like (Y280/G9)	Reverse genetics	MHRA	Yes
IBCDC-2 (A/chicken/Hong Kong/G9/97)	B-like (Y280/G9)	Conventional	CDC	Yes
IDCDC-RG26 (A/Hong Kong/33982/2009)	G4 (G1)	Reverse genetics	CDC	Yes
IDCDC-RG26 (A/Hong Kong/33982/2009)	G5.7 (G1)	Reverse genetics	CDC	Yes
IDCDC-RG26 (A/Hong Kong/33982/2009)	B4.7 (Y280/G9)	Reverse genetics	SJCRH	Yes
IDCDC-RG26 (A/Hong Kong/33982/2009)	B4.7.4 (Y280/G9)	Reverse genetics	CDC/CCDC	Yes
IDCDC-RG26 (A/Hong Kong/33982/2009)	G5.5 (G1)	Reverse genetics	CDC	Yes

H7Nx

Antigenic prototype	Clade	Type of virus or reassortant	Developing Institute	Available from
A/Henan/4-10CNIC/2022	Eurasian Lineage	Reverse genetics	CDC/CCDC	Pending

Candidate vaccine viruses (like virus)	Clade	Institution*	Available
A/Cambodia/SVH240441/2024	7.1	CDC	Yes
IDCDC-RG75A (A/chicken/Ghana/FJ21099/2018)	7.1	CDC	Yes
CNIC-FJ21099 (A/Fujian-Sanyuan/1/2018)	7.1	CDC	Yes
A/chicken/Ghana/AVL-76321V/2018	7.1	CDC	Yes
CNIC-HB29578 (A/Hubei/29578/2018)	7.1	CDC	Yes
SJ1010 (A/chicken/Vietnam/NCV-03/2008)	7.1	CDC	Yes
A/Guangdong/18SF020/2018-like (H5N6)	2.3.4.4h	CCDC	Pending

<https://www.who.int/teams/global-influenza-programme/vaccines/who-recommendations/zoonotic-influenza-viruses-and-candidate-vaccine-viruses>

GISRS-WHOHQ@WHO.int



Subtype	Antigenic prototype	Clade	Candidate vaccine virus	Type of virus or reassortant	Developing Institute	Available from
H7N1	A/mallard/Netherlands/12/2000*	Eurasian	NIBRG-63*	Reverse genetics	MHRA, UK	MHRA, UK
H7N2	A/turkey/Italy/3889/99	Eurasian	Wild type virus			MHRA, UK
	A/turkey/Virginia/452/2002	American	IBCDC-5*	Classical reassortant	CDC, USA	CDC, USA
	A/New York/107/2003	American	NIBRG-109*	Reverse genetics	MHRA, UK	MHRA, UK
				Wild type virus		WHO CCs

Assessment of risk associated with the viruses (as of Nov 2024)

TIPRA results on 2.3.2.1c (2024)



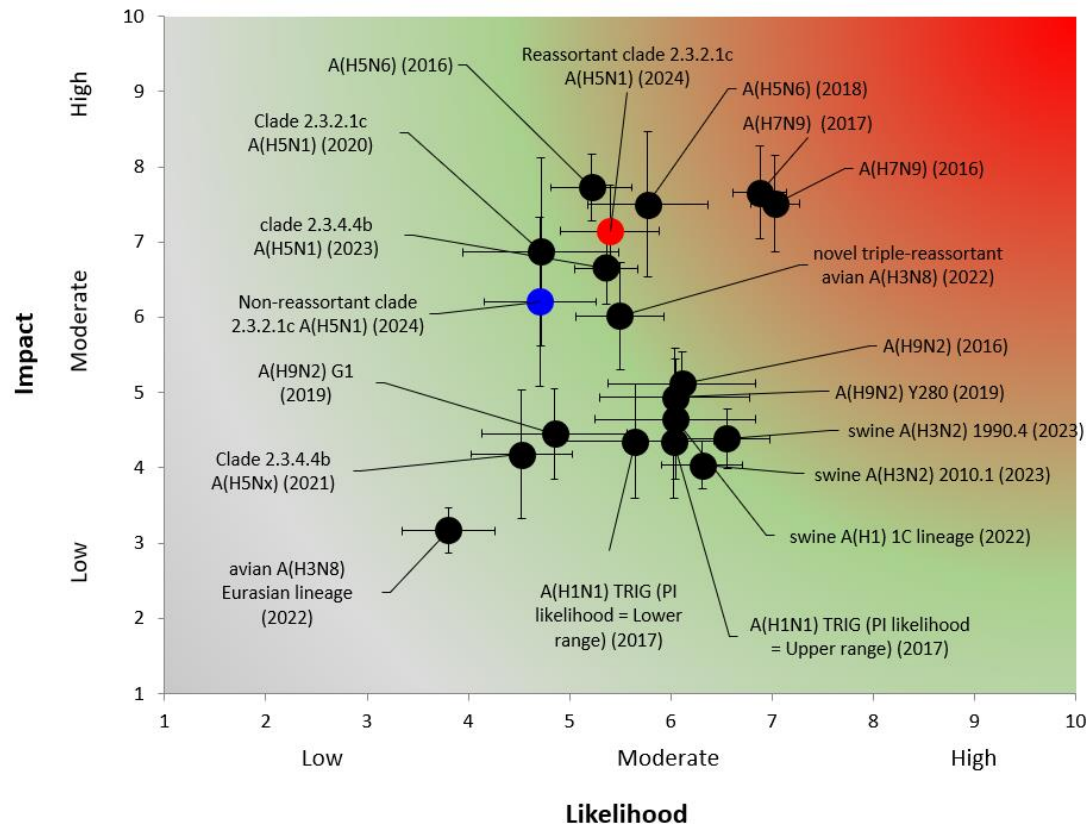
Food and Agriculture
Organization of the
United Nations



World Health
Organization



World Organisation
for Animal Health
Founded as OIE



Updated joint FAO/WHO/WOAH assessment of recent influenza A(H5N1) virus events in animals and people

- **Current risk**
 - Low – global public health risk
 - Low-moderate – risk of infection for personnel with occupational exposure
- **Human-to-human transmission**
 - Current circulating viruses do not have such ability

Summary of H5 human infections

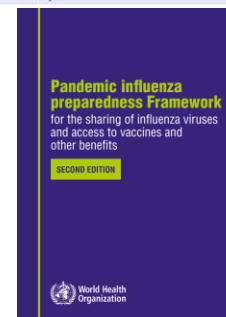
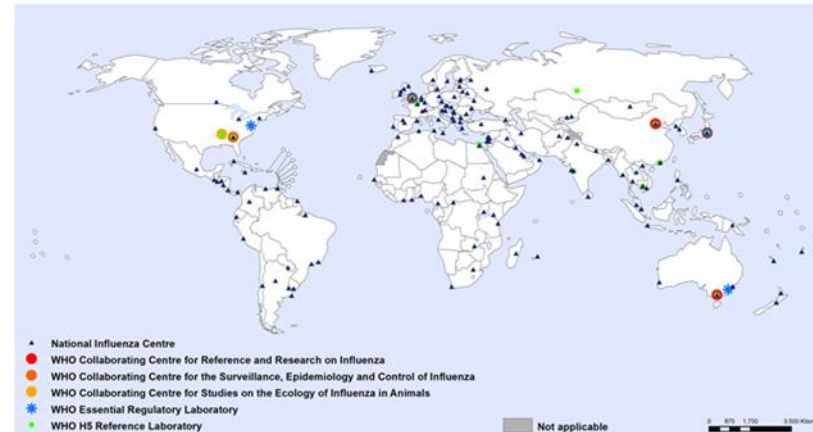
- **Sporadic** with direct or indirect **exposure** to infected animals or contaminated environments.
- Infections **range** from asymptomatic/mild, to severe and even fatal.
- Available genetic sequences of the virus from the human cases are similar to those from local **animals**.
- Based on evidence available to date, **no further spread** of the virus to close contacts and no cases detected in the community.
- **Situation can evolve quickly**
 - **Prompt & thorough investigations** are critical, **early identification** of any unusual events that could signal person-to-person transmission of the virus and **timely reporting** are key to optimal outcome of response.

WHO response & preparedness to avian & pandemic influenza

HEPR – health Emergency Preparedness, Response & Resilience Framework



Influenza specific resources



Thank you