

# H5N1 highly pathogenic avian influenza-biology of the recent outbreak

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# H5Nx clade 2.3.4.4c HPAI in the U.S. (2014-15)



American green-winged teal (*Anas carolinensis*)



American wigeon (*Anas americana*)



Mallard (*Anas platyrhynchos*)



Northern pintail (*Anas acuta*)



Gadwall (*Anas strepera*)



Wood duck (*Aix sponsa*)



Cinnamon Teal (*Anas cyanoptera*)



Lesser Snow Goose (*Chen caerulescens*)



Canada Goose (*Branta canadensis*)



Northern Shoveler (*Anas clypeata*)



Ring-necked Duck (*Aythya collaris*)



Snow geese (*Chen caerulescens*)



Cooper's hawk (*Accipiter cooperii*)



Gyrfalcon (*Falco rusticolus*)



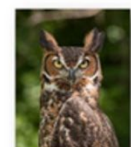
Peregrine falcon (*Falco peregrinus*)



Bald eagle (*Haliaeetus leucocephalus*)



Red-tailed Hawk (*Buteo jamaicensis*)

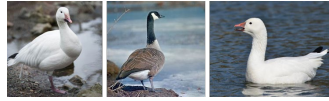


Great horned owl (*Bubo virginianus*)

- 50.4M poultry. **21** backyard and 211 commercial flocks, 21 states affected.
- A total of **98** HPAIV positive wild birds detected between December 2014 and June 2015.
- The majority were hunter-harvested waterfowl collected in the Pacific Flyway, but 16 were associated with three wild bird mortality events involving snow geese and ringed-necked ducks.
- Virus was eradicated from poultry (6/2015) and disappeared from wild birds in North America by end of 2016.

# H5N1 clade 2.3.4.4b HPAI in the US

- 190 Commercial flocks, **218** BY flocks, 40.18M birds affected, 39 states (August 24,2022)
- **2,276** wild bird detections; >35 species
- **Anseriformes** represent more than 70% of the positive wild bird cases (Dabbling ducks, diving ducks, geese, and swans)
- **Corvidae**
  - American crows, magpies, raven
- **Raptors** (Birds of Prey)
  - Accipitriformes: Black and Turkey vultures, Bald Eagle, Red-shoulder Hawk, Red-tailed Hawk, Coopers Hawk
  - Strigiformes: Snowy Owl, Great-horned Owl
- **Charadriiformes**
  - Scolopacidae: Sanderling
  - Laridae: Herring Gull, Ringed-billed Gull
- **Pelecaniformes**
  - Pelecanidae: Brown Pelican, White Pelican
  - Ardeidae -Great Blue Heron



Common Name	# of detections	Mortality	Hunter harvest or live
Snow goose	225	+++	++
Mallard	209	+	+++
Canada goose	200	+++	++
American wigeon	66	+	+++
Ross's goose	63	+++	++
American green-winged teal	47		+++
Gadwall	33		+++
American black duck	32		+++
Wood duck	48	++	+
Lesser scaup	24	++	+
Hooded merganser	14	+++	
Northern shoveler	14		++
Blue-winged teal	11		++
Redhead duck	9	+++	+
Cinnamon Teal	9		+++
Lesser snow goose	8	++	
Muscovy duck	7	++	
Ring-necked duck	5	+	
Tundra swan	5	+	
Trumpeter swan	4	+	
Northern pintail	4		+
Mute swan	3	+	
Ruddy duck	1	+	
Greater white-fronted goose	1		

# HPAI in chickens and turkeys

## Field cases

### 2015 H5N2 HPAIV outbreak

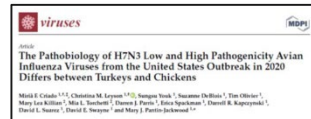
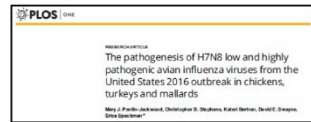
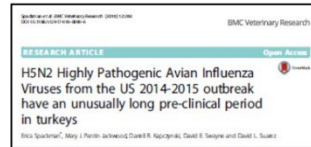
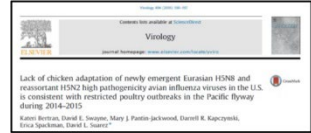
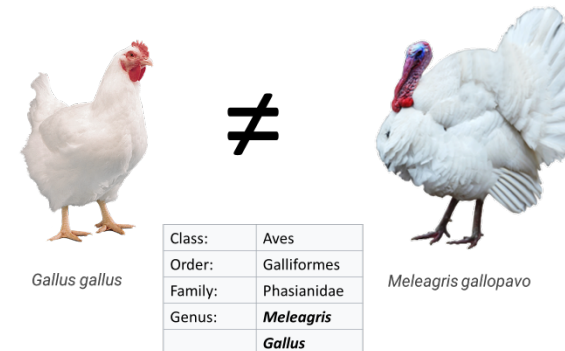
- 211 commercial poultry farms were affected. Most were commercial turkey operations (n = 160)

### 2022 H5N1 HPAIV detections (as of 7/25/22)

Commercial Chickens	# of incidents	Commercial Turkeys	# of incidents
Table Egg Layer	21	Meat	120
Table Egg Pullets	4	Breeder Hens	10
Table Egg Breeder	2	Breeder Toms	1
Broiler Production	12	Poult Supplier	1
Broiler Breeder	2		
<b>Total incidents</b>	<b>41</b>		<b>132</b>

## Experimental studies

- HPAIVs were highly infectious and transmitted easily in turkeys. In contrast, chickens needed higher amounts of the virus to become infected and transmission was poor.
- Infected turkeys and chickens had high mortality and birds excreted high amounts of virus from the oropharyngeal and cloacal route.



Infectivity and  
transmissibility of Gs/GD  
H5Nx clade 2.3.4.4 HPAIV in  
different avian species

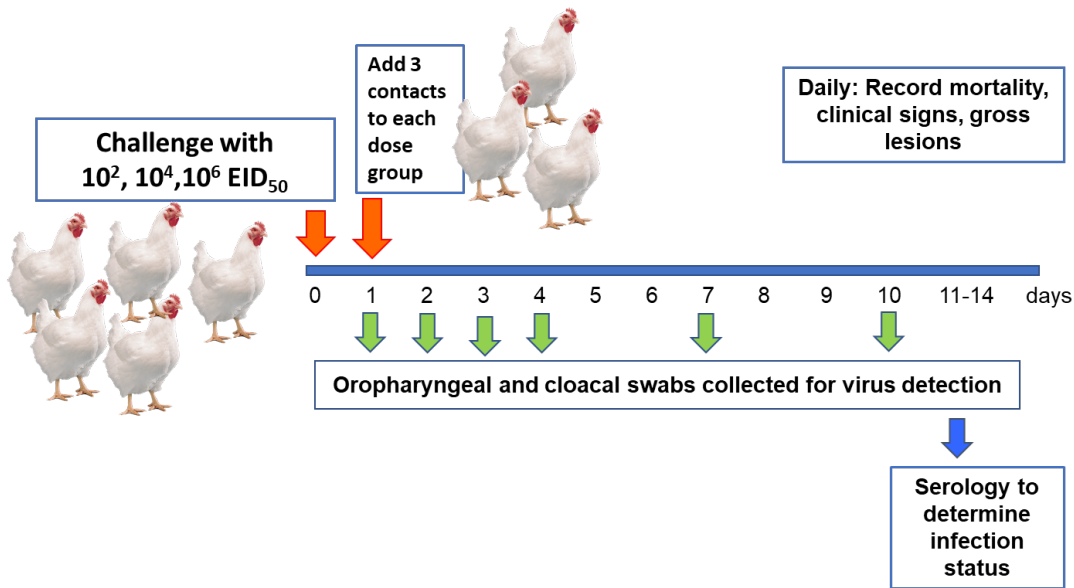
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# Studying the infectivity and transmissibility of H5N1 HPAIV

**Virus:** A/American widgeon/SC/22-000345-001/2022 (H5N1) - H5 clade 2.3.4.4b – A/AW/SC/2022 (H5N1)

- SPF White Leghorn chickens
- Commercial broad-breasted white turkeys

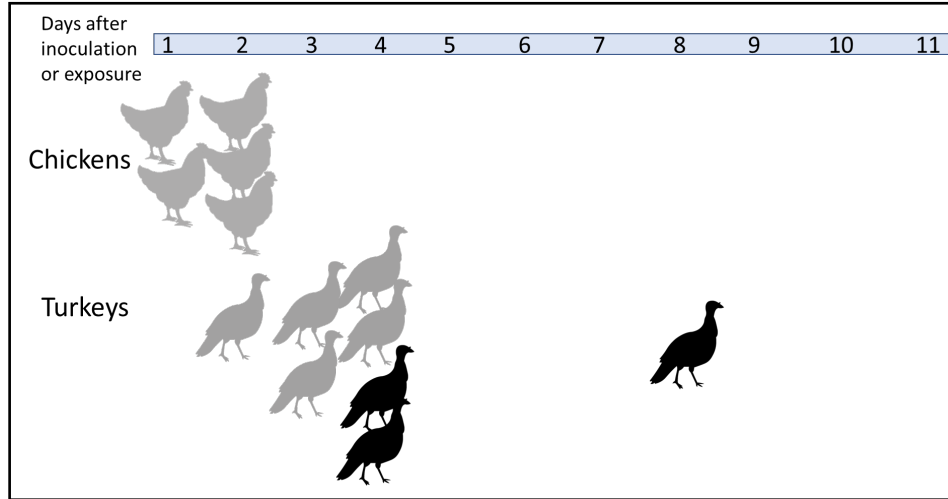


	Virus dose	Inoculated dead/total (MDT)	BID <sub>50</sub> (log <sub>10</sub> EID <sub>50</sub> )	Contact dead/total
Chickens	low	3/5 (2.7)	≤3.5	0/3
	medium	5/5 (2)		0/3
	high	5/5 (1)		0/3
Turkeys	low	5/5 (4.6)	<3.5	3/3
	medium	5/5 (3.4)		3/3
	high	5/5 (2.6)		3/3

MDT = mean death time, BID<sub>50</sub> = 50% bird infective dose.  
 EID<sub>50</sub> = 50% egg infective dose.

# A/AW/SC/2022 (H5N1) $5.5 \log_{10} \text{EID}_{50}$

## Mortality



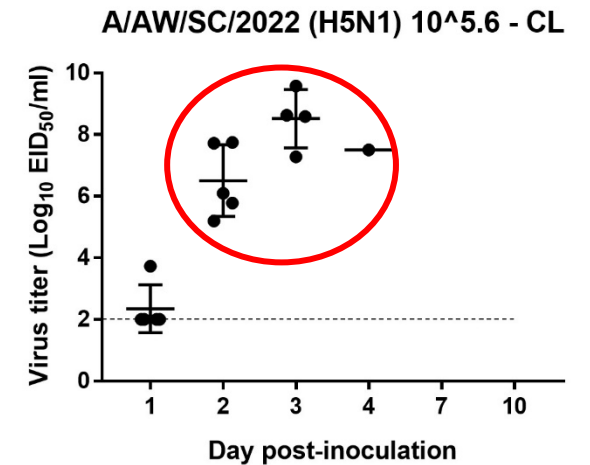
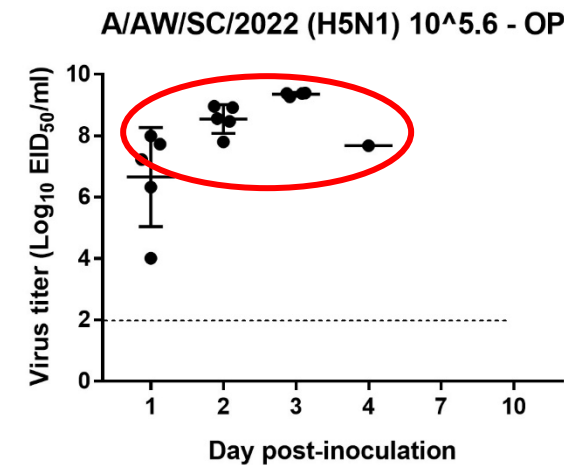
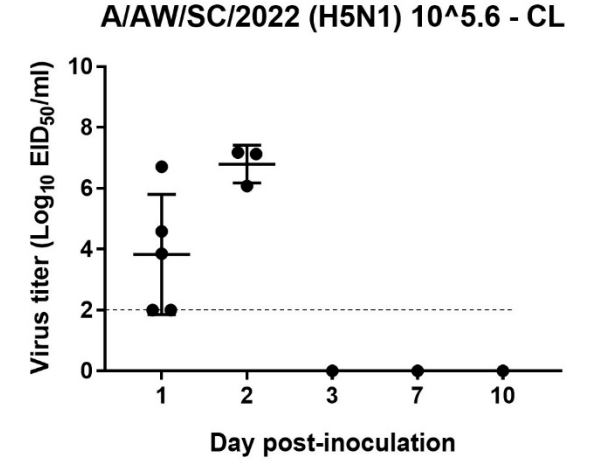
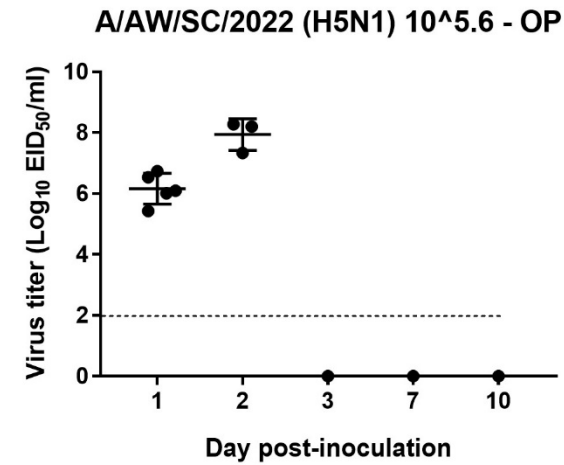
= inoculated      = contact



## Virus shedding

### Oropharyngeal

### Cloacal







# Comparing A/AW/SC/2022 (H5N1) with A/Northern pintail/WA/2014 (H5N2)



	Virus	Dead/total (MDT)	BID <sub>50</sub> (log <sub>10</sub> EID <sub>50</sub> )	Transmission
Chickens	H5N1	5/5 (1.4)	≤3.5	No
	H5N2 <sup>a</sup>	3/5 (3.0)	5.7	No
Turkeys	H5N1	5/5 (3.2)	<3.5	Yes, with all doses
	H5N2 <sup>b</sup>	5/5 (5.3)	5.0	Only in the high dose group

MDT = mean death time, in 5.6-6.6 log<sub>10</sub> EID<sub>50</sub> dose group  
 EID<sub>50</sub> = 50% egg infective dose. BID<sub>50</sub> = 50% bird infective dose

<sup>a</sup>Bertran et al. Vet Res (2019)

<sup>b</sup>Spackman et al. BMC Veterinary Research (2016)

The 2022 H5N1 HPAIV is more infectious than the index 2014 H5N2 HPAIV

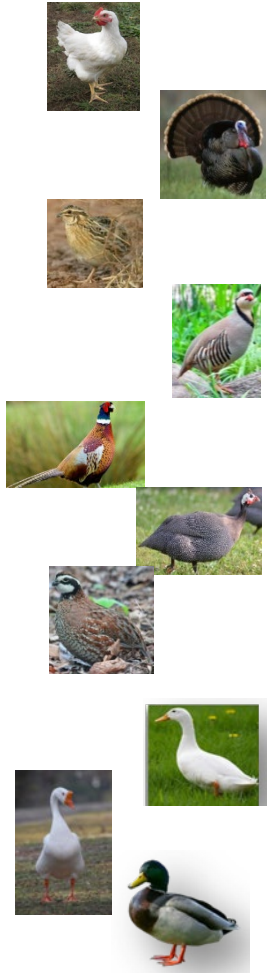
# Infectious dose in chickens: H5N2 HPAIVs (March-May 2015)

Virus	% Mortality	BID <sub>50</sub> (log <sub>10</sub> )
A/Northern Pintail/Washington/40964/2014	60	5.7
A/Tk/South Dakota/12511/2015	100	3.2
A/Tk/Minnesota/12582/2015	100	3.6
A/Ck/Iowa/13388/2015	100	3.5

- Midwest H5N2 viruses were better chicken-adapted (April-May 2015)

The longer a HPAIV circulates in poultry, the more infectious it becomes

# A/northern pintail/WA/2014 (H5N2)



	% Mortality	MDT at highest dose (days)	BID <sub>50</sub>	Transmission
Chickens	60	3	5.7	No
Turkeys	100	9	5.0	Only in the high dose group
Japanese quail		3.0	3.7	In the medium and high dose groups
Chukar partridges		5.2	3.6	
Ring-necked pheasants		4.8	3.6	
Guinea fowl		3.8	3.0	In low, medium, and high dose groups
Bobwhite quail		4.9	<2	
Pekin ducks	0	-	3	
White Chinese Geese	25	7	<2	
Mallards	0	-	<2	

# Pathogenicity of H5Nx HPAIV in ducks:

## Differences between species - *Wild ducks*

**Virus: A/Whooper Swan/Mongolia/244/ 2005 (H5N1)**

	Mortality	Oral*	Cloacal*
Wood Ducks	<b>3/6</b>	<b>4.96 / 5</b>	<b>3.30 / 3</b>
Mallards	0/6	2.57 / 2	0.97 / 1
Redheads	0/6	2.70 / 2	1.23 / 1
Northern Pintails	0/6	3.14 / 2	0.97 / 1
Blue-winged Teal	0/6	1.32 / 1	1.21 / 1

\*Avg. viral titers (log<sub>10</sub> EID<sub>50</sub>/ml) / Avg. duration (days)



*Aix sponsa*

*Anas platyrhynchos*

*Aythya americana*

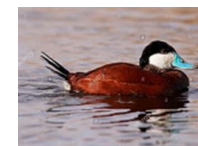
*Anas acuta*

*Anas discors*

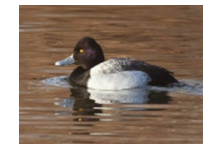
Brown, et al. J. Wildl Dis, EID, 2008

**A/GF/WA/14 (H5N8) HPAIV and  
A/NP/WA/14 (H5N2) HPAIV**

	Virus	Mortality	BID <sub>50</sub>
<b>Ruddy Duck -juvenile</b>	<b>H5N8</b>	3/5	<2
<b>Ruddy Duck -adult</b>	<b>H5N2</b>	1/5	
<b>Lesser Scaup-juvenile</b>	<b>H5N8</b>	0%	3
	<b>H5N2</b>	0%	<4



(*Aythya affinis*)



(*Oxyura jamaicensis*)

Spackman et al. J. Wildlife Diseases 2017

Infectivity and transmissibility of HPAIV can be affected by the species, age, and health status of the birds

# Pathogenicity of H5N1 HPAIV in Geese and Swans

Species	Virus	Sick/Dead /Total	MDT	Ages and References
<b>Emden goose</b> ( <i>Anser anser domesticus</i> )	A/Ck/HK/220/97	5/0/11	-	2 wks-old; <i>Perkins, 2002</i>
<b>Chinese white goose</b> ( <i>Anser cygnoides</i> )	A/Gs/HK/437-4/99 A/Gs/HK/485-3/00	3/3/3 3/0/3	7-8 -	3-4 wks-old; <i>Webster, 2002</i>
<b>Chinese goose</b>	A/Dk/SH/04	5/5/5	7	5 wks-old <i>Tian, 2005</i>
<b>Cackling goose</b> ( <i>Branta hutchinsii</i> )	A/W.sw/Mong/05	4/3/4	6.0	12 wks-old; <i>Brown, 2008</i>
<b>Bar-headed goose</b> ( <i>Anser indicus</i> )	A/W.sw/Mong/05	5/2/5	6.5	
<b>Canada Goose</b> ( <i>Branta canadenziz</i> )	A/Ck/VN/14/05	5/5/5	5.0-20	13 wks-old, adults; <i>Pasick, 2007</i>
<b>Geylag goose</b> ( <i>Anser anser</i> )	A/Ck/S.Korea/06	2/0/2	-	7 wks-old <i>Kwon, 2010</i>
<b>Chinese goose</b>	A/Ck/Indonesia/07	9/9/12	5.1	4 wks-old <i>Eggert,2010</i>



Note: previous exposure to AIV can modulate the outcome of HPAIV infection

# Comparing clade 2.3.4.4 Gs/GD H5 HPAIV's in mallards



**BID<sub>50</sub> <2 log<sub>10</sub>**

*DeJesus et al. 2016*  
*Leyson et al. 2019*

Virus	% infected	% mortality	# of days virus positive	% of contacts infected
A/Gf/WA/2014 (H5N8)	100	0	11	100
A/Np/WA/2014 (H5N2)	100	0	>14	100
A/Tk/MN/2015 (H5N2)	100	10	>14	100
A/Ck/IA/2015 (H5N2)	100	0	11	100
A/Td/Denmark/2016 (H5N8)	100	88	5	100
<b>A/AW/SC/2022 (H5N1)**</b>	<b>100</b>	<b>23</b>	<b>&gt;14</b>	<b>100</b>

\*\*Clinical signs: 7/34 (20%) neurological signs (euthanized), 6/34 (18%) corneal opacity, 1/34 (3%) mortality. Shed high titers of virus orally and cloacally.

***Spackman et al. preliminary results***



A/northern pintail/WA/2014  
(H5N2 clade 2.3.4.4c)

0% mortality

A/American wigeon/SC/2022

(H5N1 clade 2.3.4.4b)

23% mortality

A/Tufted-duck/Denmark/2016

(H5N8 clade 2.3.4.4b)

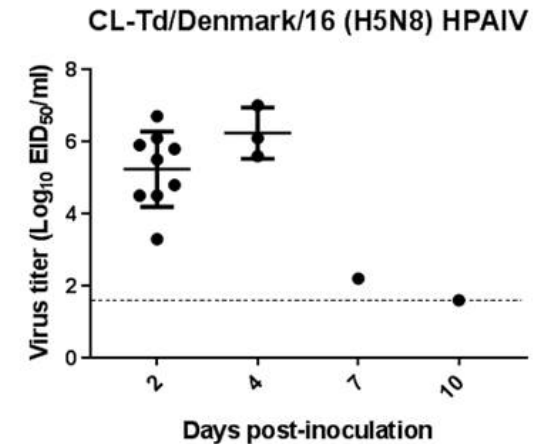
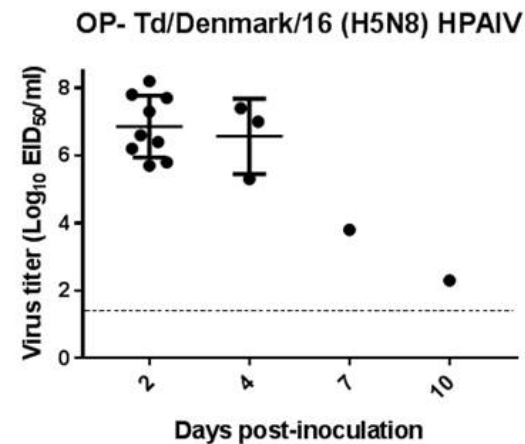
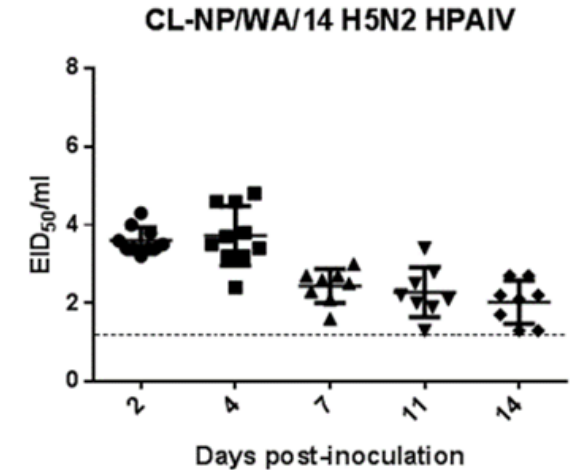
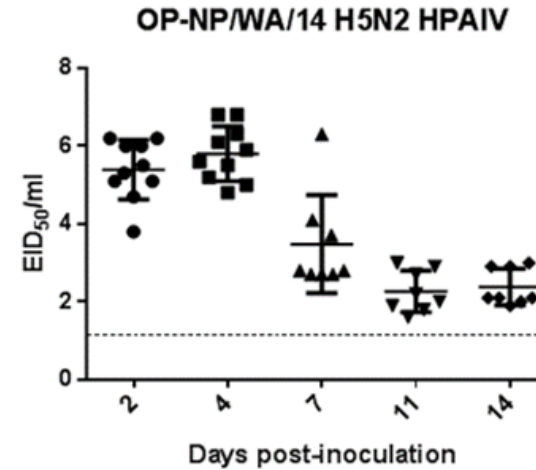
88 % mortality

Decreased disease severity compared to the 2016 virus could help the virus spread more easily in the wild waterbird population.

# Virus shedding

Oropharyngeal

Cloacal



# A/AmWigeon/SC/345/2022 H5N1 in Mallards

- $BID_{50} < 2\log_{10}$
- Transmitted to all contact exposed ducks
- Shed high titers orally and cloacally
  - Ducks still positive for shed 14DPC (mostly cloacal)
- Clinical signs:
  - Most ducks no signs
  - ~20% had neurological and corneal opacity, mild lethargy
- IHC and rRT-PCR on tissue: virus systemic replication- similar to other 2.3.4.4 HPAIVs



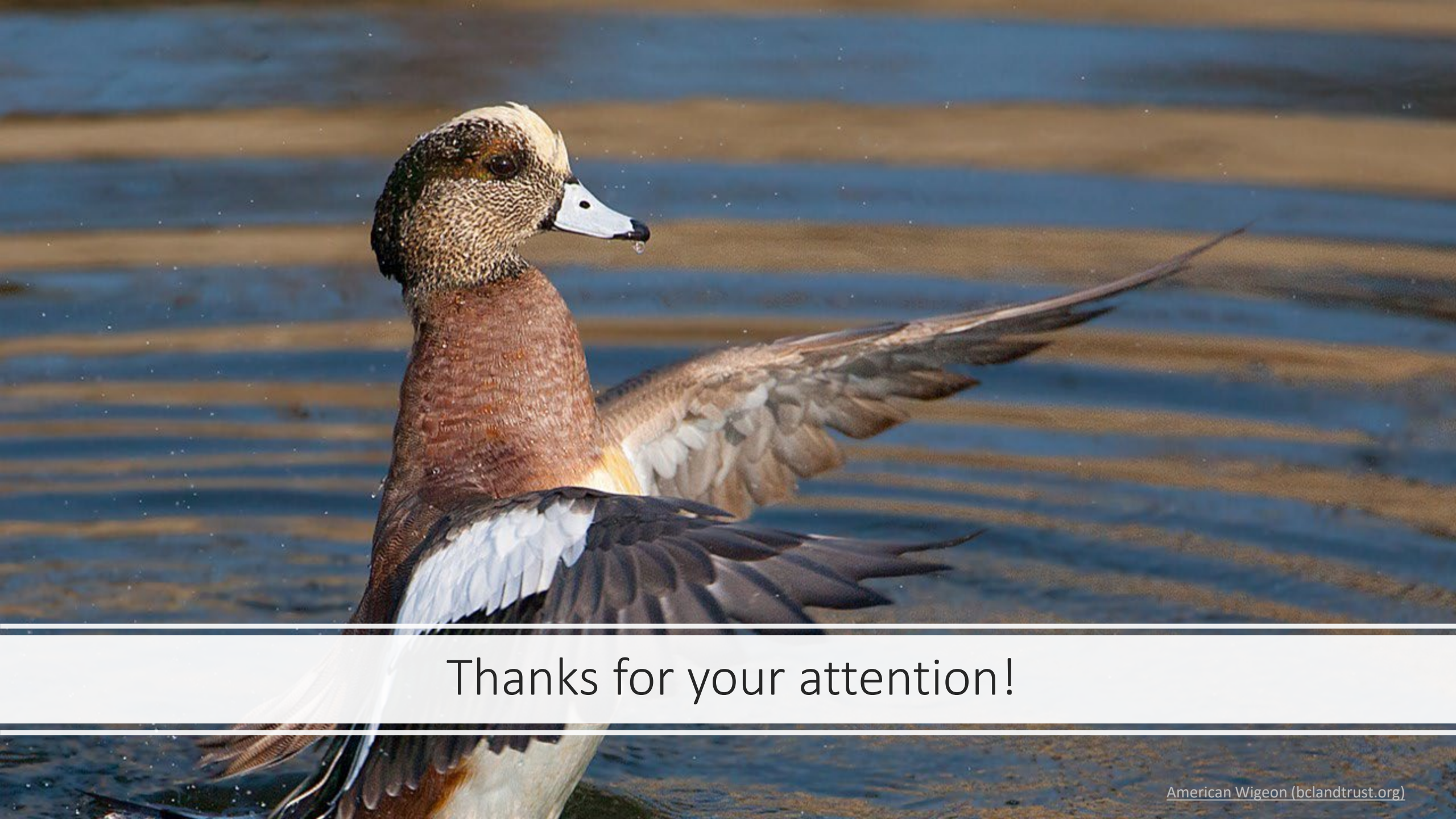
# Summary

- The infectivity, transmissibility, and pathogenicity of HPAIVs in avian species is affected by the strain of the virus and the species, age, health, and immune status of the birds.
- Once infected, Gallinaceous species will become severely sick. Turkeys are highly susceptible to HPAIV infections. This high susceptibility, coupled with a longer mean death time and high amounts and duration of virus excreted, can favor HPAIV spread.
- Wild birds show variable susceptibility to disease, with some wild migratory waterfowl being key contributors in the spread of H5N1 HPAIV.
- Some infected wild bird species can show clinical signs and mortality, which would limit spread of the virus, but these birds might act as sentinels of viral circulation.
- This information is important for surveillance efforts and help clarify epidemiological data from outbreaks of H5Nx HPAIV in poultry and wild bird populations.



Photograph by Lyuba Filatova (fineartamerica.com)





Thanks for your attention!