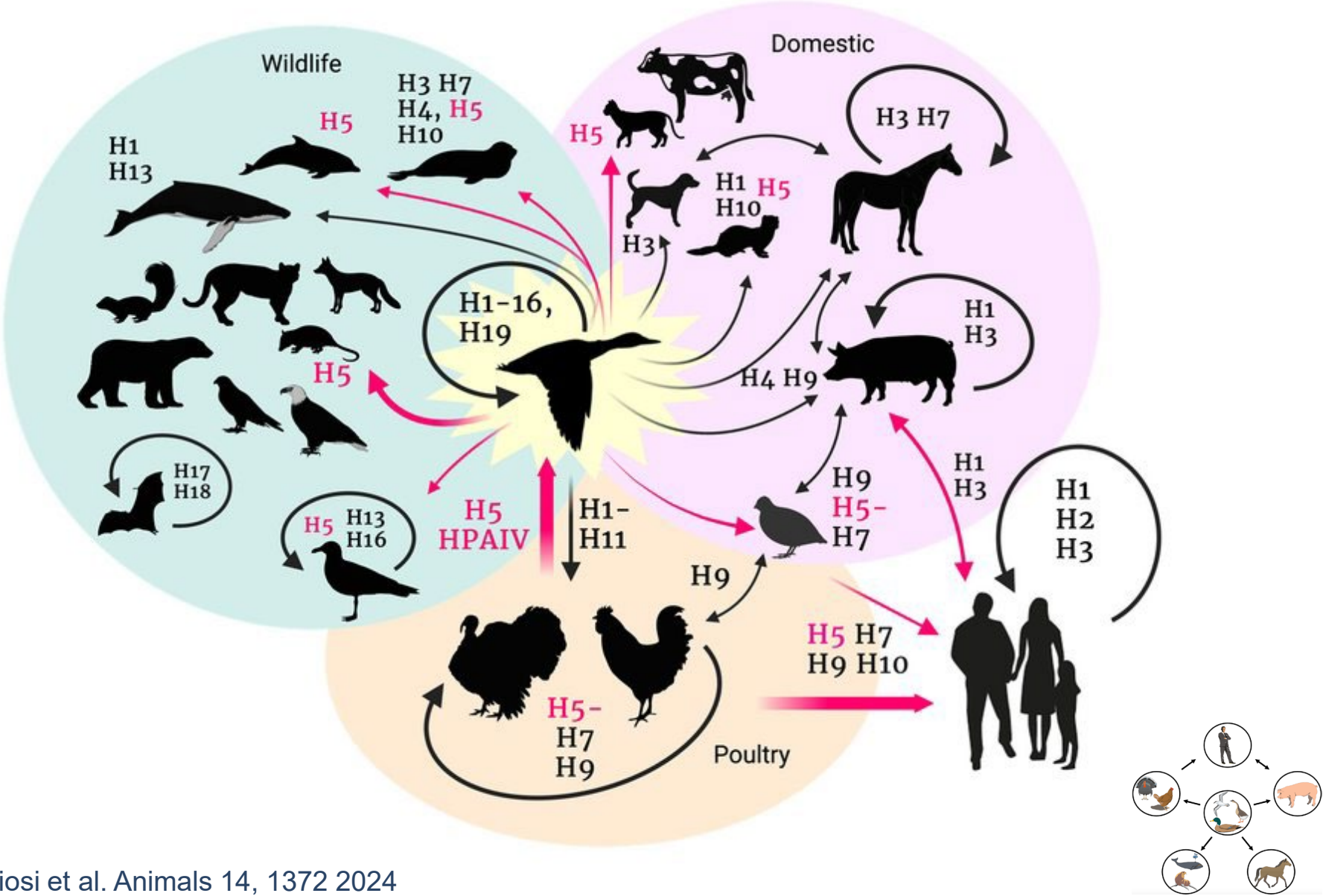




HPAI surveillance in vaccinated poultry & public health gains

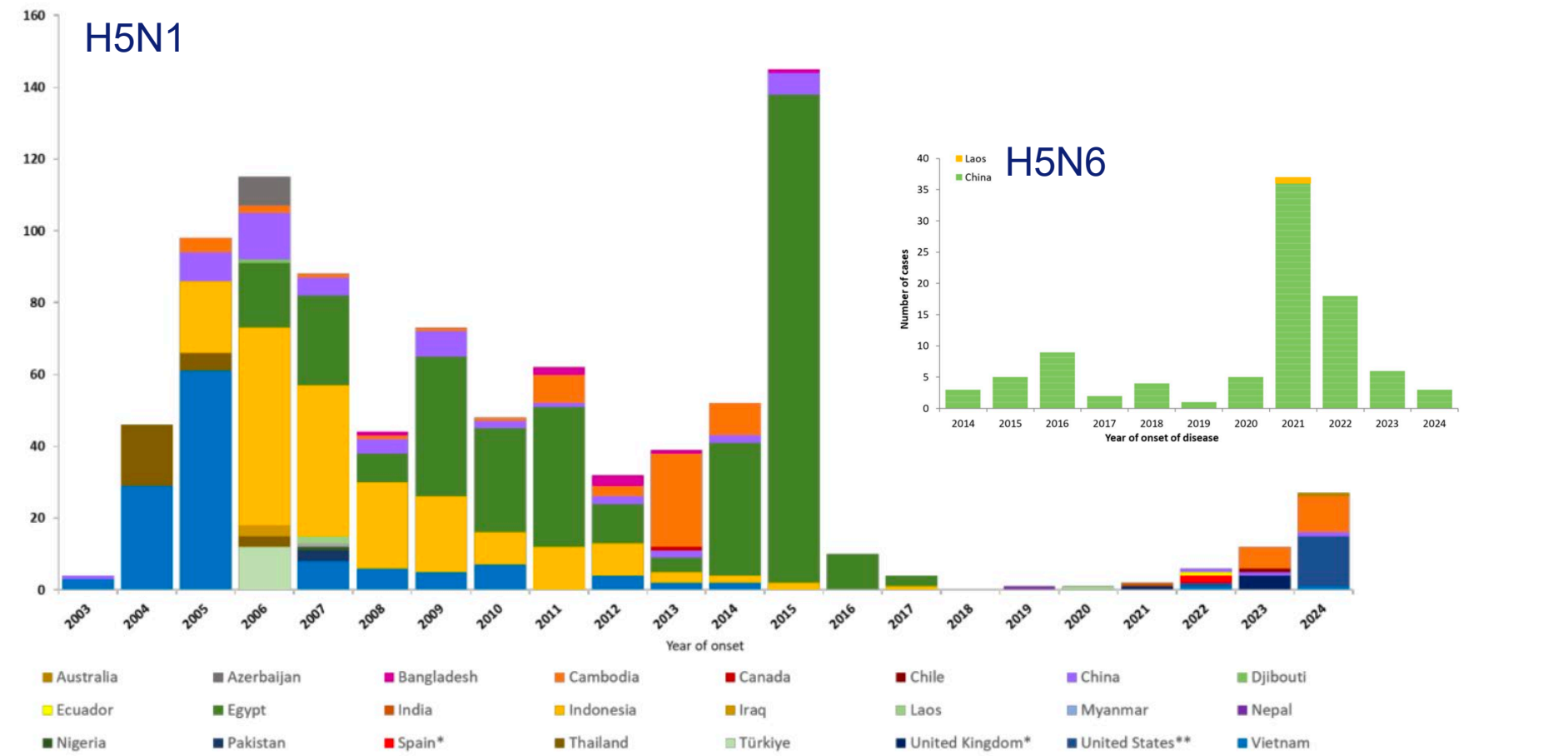
Ron A.M. Fouchier, PhD

Host switching of (H5) influenza virus



Human cases of A(H5) virus infection

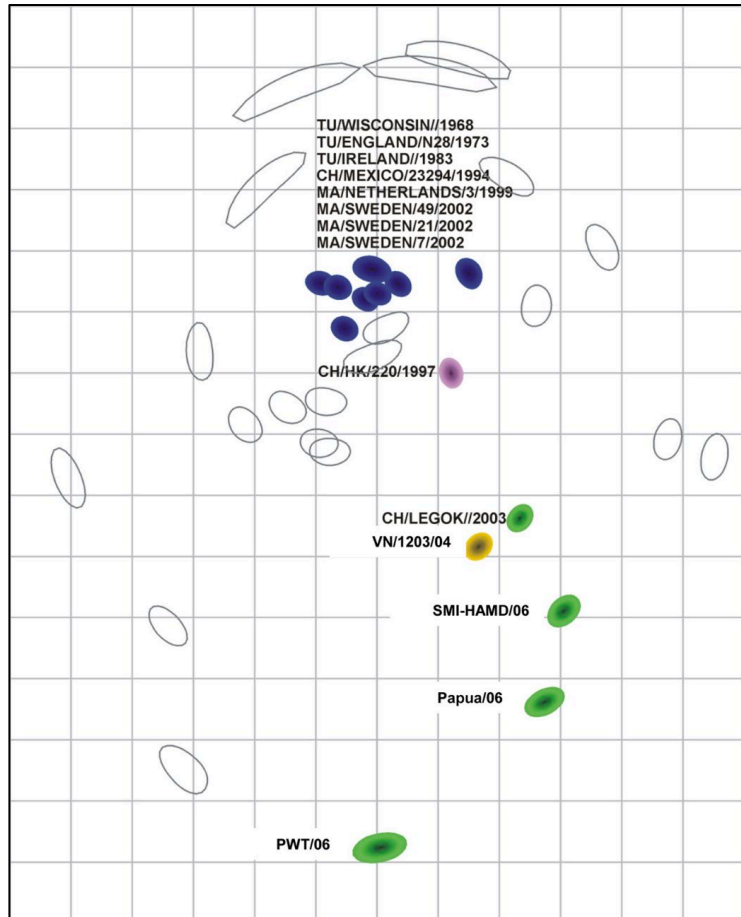
Number of cases



Pros and cons of poultry vaccine use

- Vaccination of poultry may improve animal health and welfare and food security
- Prevention of human exposure to HPAI A(H5) may be an added indirect benefit
- But: poultry vaccines may not provide sterilizing immunity
 - chance of dormant infection, virus excretion, risk of zoonoses
 - chance of onward spread to other (unvaccinated) animals
- And: vaccination can cause natural selection of antigenically drifted variants:
 - possible reduced effectiveness of H5 vaccines in poultry
 - possible increased need to update prepandemic H5 Candidate Vaccine Viruses

Antigenic drift & vaccines; escape



Antibody Titer Has Positive Predictive Value for Vaccine Protection against Challenge with Natural Antigenic-Drift Variants of H5N1 High-Pathogenicity Avian Influenza Viruses from Indonesia

David E. Swayne,^a David L. Suarez,^a Erica Spackman,^a Samadhan Jadhao,^{ax} Gwenaelle Dauphin,^b Mia Kim-Torchetti,^{bx} James McGrane,^c John Weaver,^{cx} Peter Daniels,^d Frank Wong,^d Paul Selleck,^d Agus Wiyono,^e Risa Indriani,^e Yuni Yupiana,^{fx} Elly Sawitri Siregar,^{gx} Teguh Prajitno,^h Derek Smith,ⁱ Ron Fouchier^l

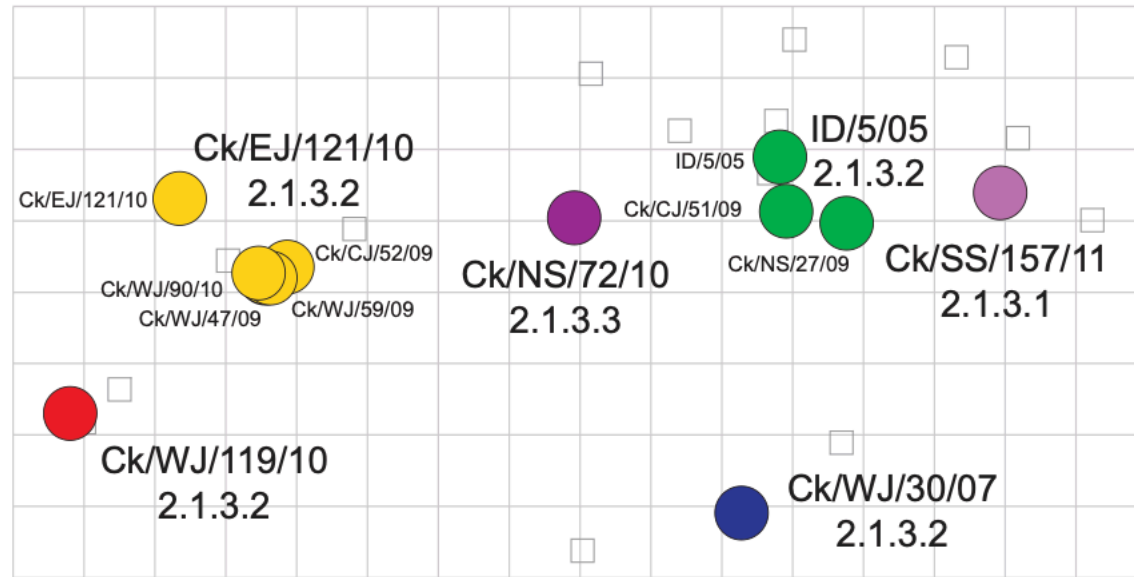
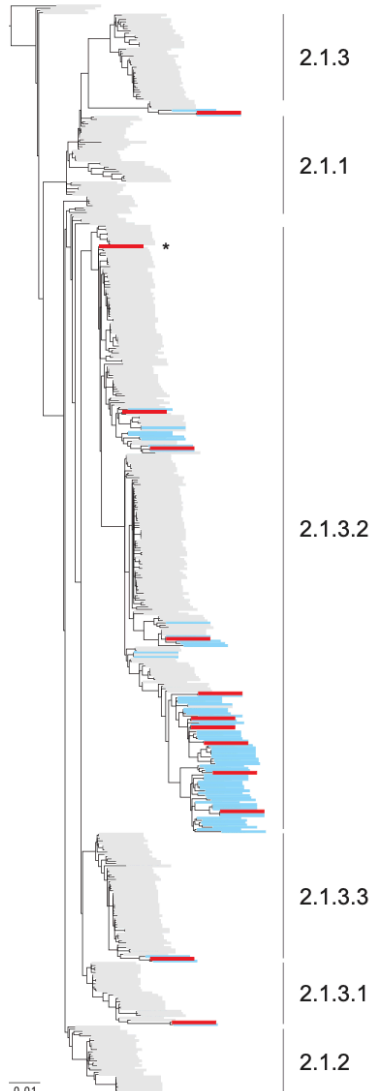
TABLE 5 Detection of H5N1 HPAI virus shedding from oropharyngeal cavity at 2 DPC in chickens vaccinated with six different vaccine seed strains and challenged with one of three H5N1 HPAI viruses from Indonesia (Table view)

Vaccine seed strain	Vaccine code	SMI-HAMD/06		Papua/06		PWT/06	
		MDT ^a	Oral shedding at 2 DPC ^b	MDT	Oral Shedding at 2 DPC	MDT	Oral shedding at 2 DPC
Legok/03	K	2 ^c	0/10 ^A ($\leq 3.9^A$)		0/10 ^A ($\leq 4.7^A$)	6.6 ^{AB}	1/10 ^A (5.68 ^A)
VN/04	O		1/10 ^A (4.0 ^A)	6.0 ^d	0/10 ^A ($\leq 4.7^A$)	5 ^{ABC}	2/10 ^A (4.5 ^B)
rgGD/96	N		0/10 ^A ($\leq 3.9^A$)	4.0 ^d	0/10 ^A ($\leq 4.7^A$)	3.1 ^C	ND ^e
Eng/73	B	3.4 ^A	1/10 ^A (4.0 ^A)	5.8 ^A	3/10 ^A (5.1 ^A)	4 ^{BC}	8/10 ^{BC} (6.13 ^{AC})
Mex/94	E		1/10 ^A (4.0 ^A)		2/10 ^A (4.8 ^A)	6.7 ^{AC}	0/10 ^A (<4.50 ^B)
WI/68	F		0/10 ^A ($\leq 3.9^A$)	5.0 ^{AB}	1/10 ^A (4.9 ^A)	3.7 ^C	4/10 ^{AC} (5.40 ^B)
Sham	U	2.0 ^{cA}	10/10 ^B (7.1 ^B)	1.6 ^{cB}	10/10 ^B (7.1 ^B)	1.7 ^{cD}	10/10 ^B (6.3 ^C)

Swayne et al., JV 2015

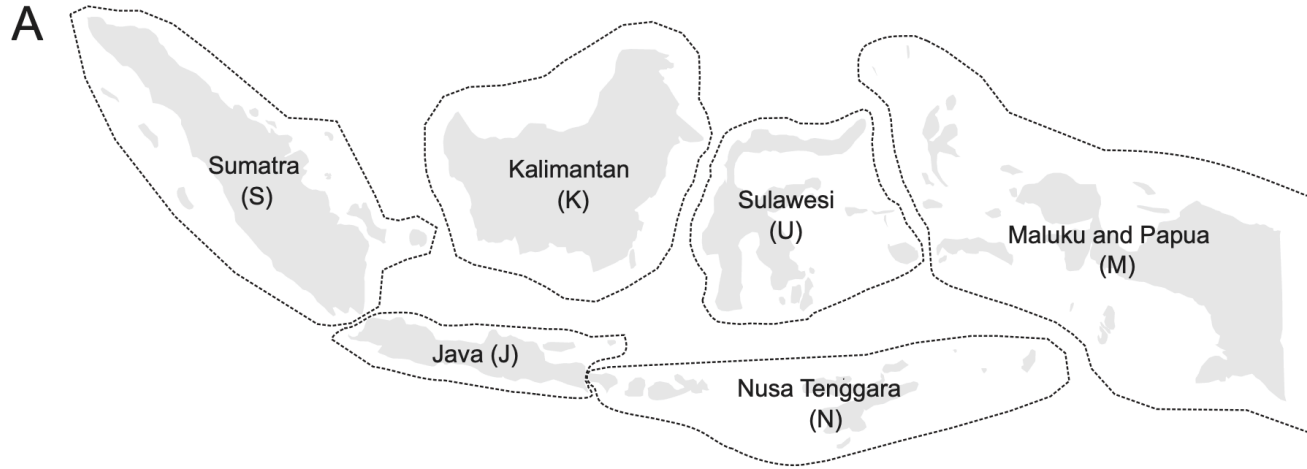


Vaccine-induced drift?



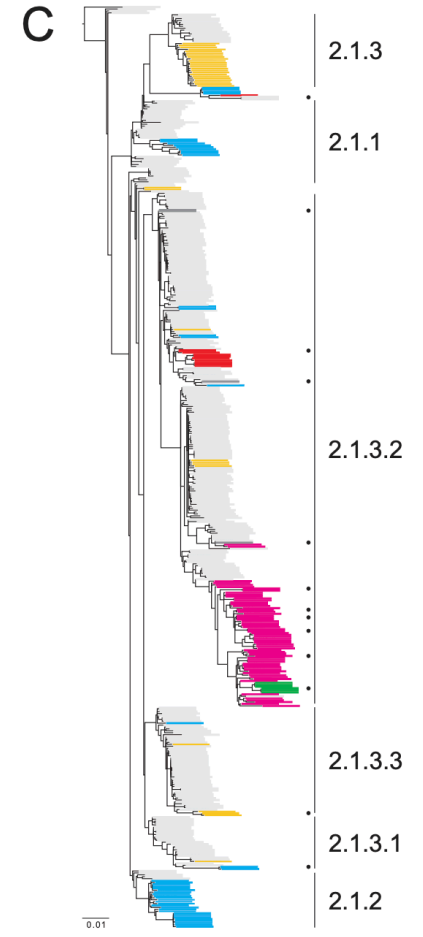
Koel et al., mBio 2014

Vaccine-induced or geo/temporal?



B

Antigenic Variant	Clade	2004	2005	2006	2007	2008	2009	2010	2011
Ck/SS/157/11	2.1.3.1		* S J N	* S J	* J N	*	* K	*	* U
Ck/WJ/119/10	2.1.3.2							J	
Ck/EJ/121/10	2.1.3.1					J	J	S J	S J U
Ck/NS/72/10	2.1.3.3	J		J K	S J N	U	S	U	
Ck/WJ/30/07	2.1.3.2			J	J	J	J		
ID/5/05	2.1.3.2		J				S J		



Koel et al., mBio 2014

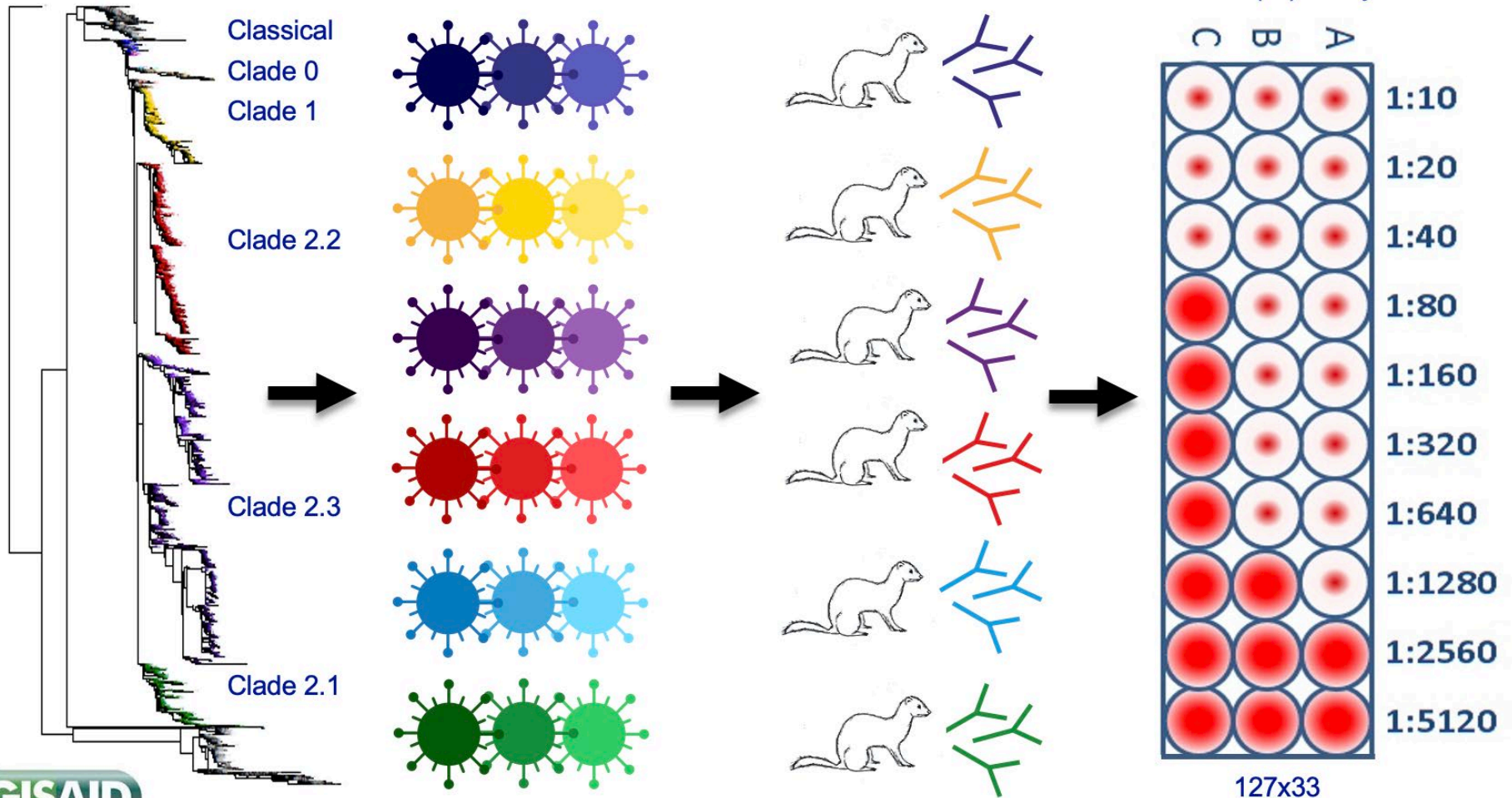
Antigenic diversity of influenza A(H5)

Phylogenetic tree HA

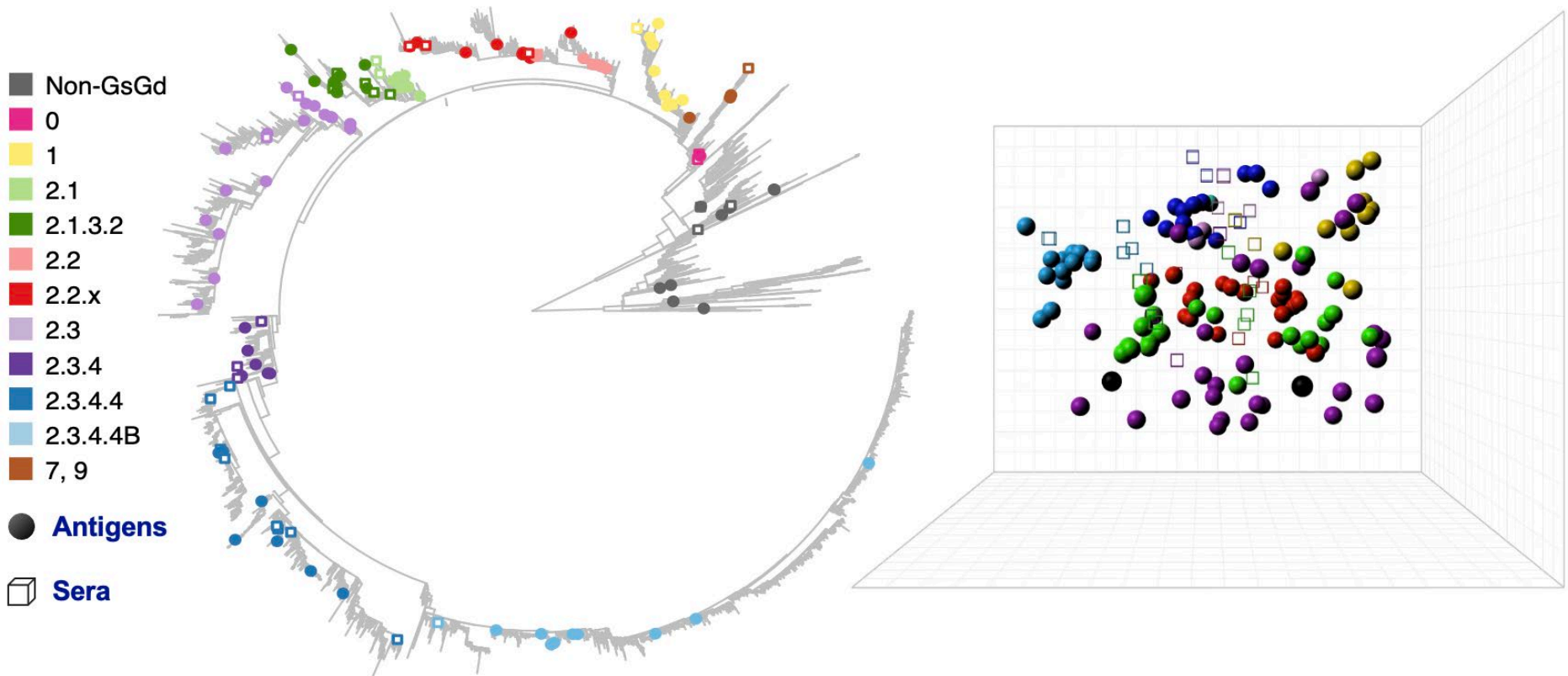
127 antigens

33 sera

Hemagglutination inhibition (HI) assay



Antigenic diversity of influenza A(H5)



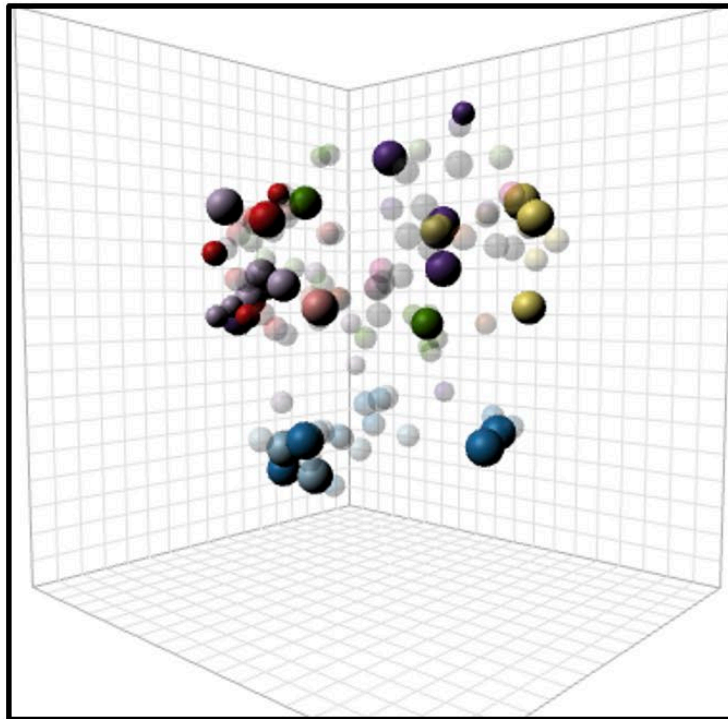
Mathilde Richard (PI)



Adinda Kok (PhD student)

Antigenic diversity of influenza A(H5)

WHO candidates (N = 44)



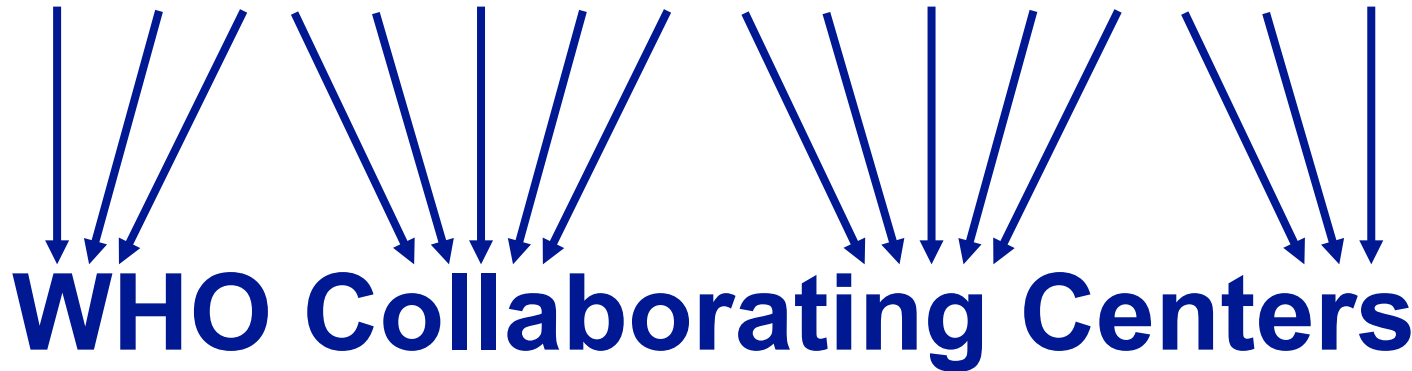
- Non-GsGd
- 0
- 1
- 2.1
- 2.1.3.2
- 2.2
- 2.2.x
- 2.3
- 2.3.4
- 2.3.4.4
- 2.3.4.4B
- 7, 9

- Antigens
- ▣ Sera

WHO GISRS

(Global influenza surveillance and response system)

National Influenza Centers



Conclusions, discussion

- Vaccination of poultry may improve animal health and welfare and food security
- Prevention of human exposure to HPAI A(H5) may be an added indirect benefit
- But: poultry vaccines may not provide sterilizing immunity
 - chance of dormant infection, virus excretion, risk of zoonoses
 - chance of onward spread to other (unvaccinated) animals
- And: vaccination can cause natural selection of antigenically drifted variants:
 - possible reduced effectiveness of H5 vaccines in poultry
 - possible increased need to update prepandemic H5 Candidate Vaccine Viruses
- **Post-vaccination monitoring:**
 - **Detection of dormant infections, monitor virus evolution and antigenic drift**
 - **Consider systematic random sampling on vaccinated farms**
 - **React to signs of dormant infection in vaccinated and other animals**
 - **Link post-vaccination surveillance to WHO GISRS system (update CVVs)**