

Wisconsin State Laboratory of Hygiene

UNIVERSITY OF WISCONSIN-MADISON









Important Update!



The WSLH Communicable Disease Division will be moving to the Sta Agriculture Drive, Madison, effective Monday, October, 10, 2016.

On and after October 10, 2016, please send clinical specimens as follows:

	WSLH Unit	Address
•	Communicable Diseases (surveillance, emergency response, outbreak, etc.)	2601 Agriculture Drive, P.O. Box 7904 Madison, WI 53718
•	Rabies	(For directions, see map below. Please
•	Cytology	deliver to loading dock off Vondron
		Road.)



WSLH Contact Information

- Email addresses will remain the same.
- ➤ CDD Customer Service # 800-862-1013
- Phone & FAX numbers are changing



WSLH will communicate new key contact phone #'s when they become available



Influenza and other Respiratory Viruses Update--2016

Pete Shult, PhD

CDD Director & Emergency Laboratory Response

and

Erik Reisdorf, MPH, M(ASCP)^{CM}

Surveillance and Virology Lab-Team Lead



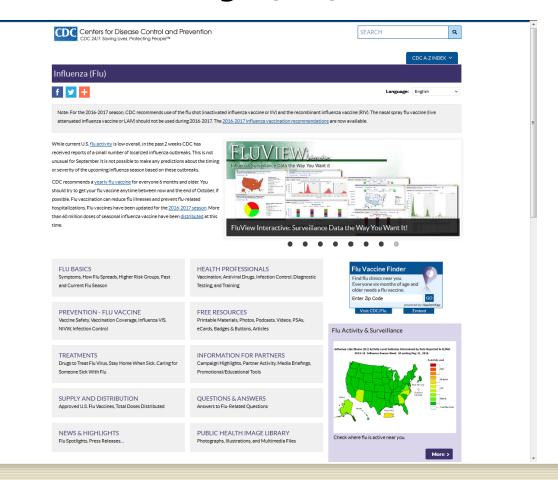
Learning Objectives

- Moving Day!
- Review of the 2015-201 influenza season.
- Vaccine strain selection and efficacy.
- Influenza "variant" virus update.
- Non-influenza respiratory pathogens.
- Technology update.
- Discuss surveillance strategy for 2016-2017



Influenza The latest information

www.cdc.gov/flu/index.htm



What We've Been Dealing With: 2016



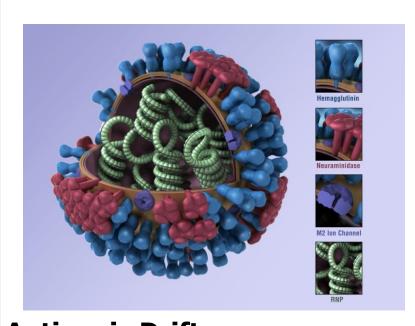
- Zika virus
- Dengue fever
- Chikungunya
- Elizabethkingia spp.
- Measles, mumps, pertussis
- Food-borne illnesses

... So what's the big deal with influenza?

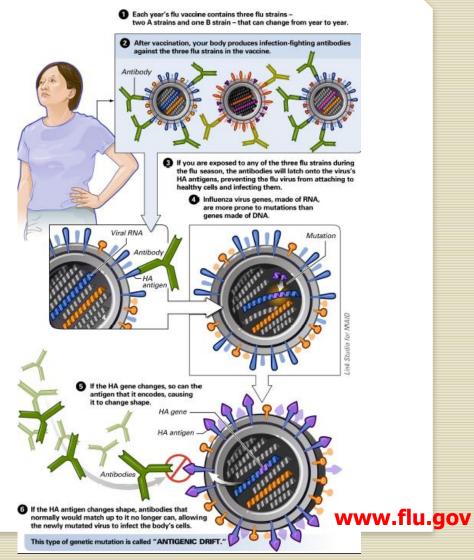


The Changeability of Influenza Antigenic Drift → Seasonal Influenza



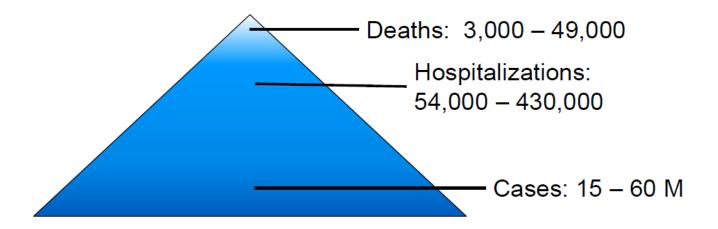


Antigenic Drift
Manifests in HA and NA as a result of continuous and gradual accumulation of point mutations in the HA and NA genes





Estimated Annual Burden of Seasonal Influenza in the United States



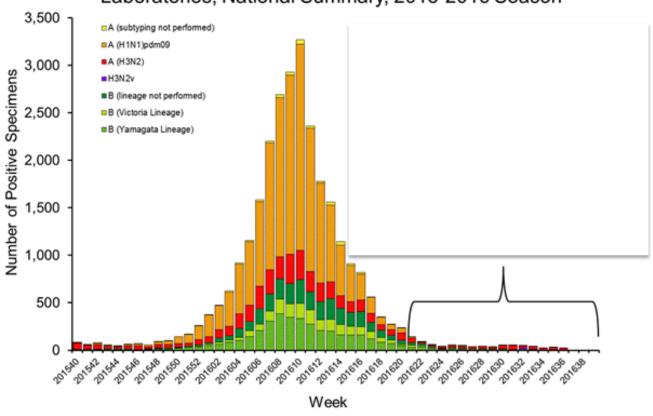
Direct medical costs: \$10.4 billion





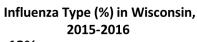
Influenza in the U.S. 2015-16

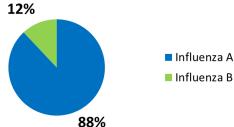
Influenza Positive Tests Reported to CDC by U.S. Public Health Laboratories, National Summary, 2015-2016 Season



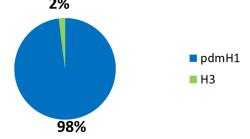


Influenza in WI, 2015-2016

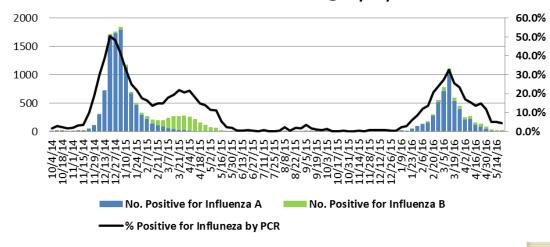




Influenza Subtype (%) in Wisconsin, 2015-2016



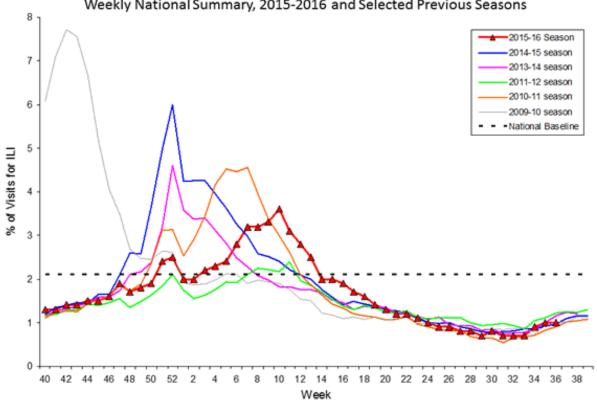
% Positive for Influenza by PCR (Wisconsin), Week Ending 5/14/16





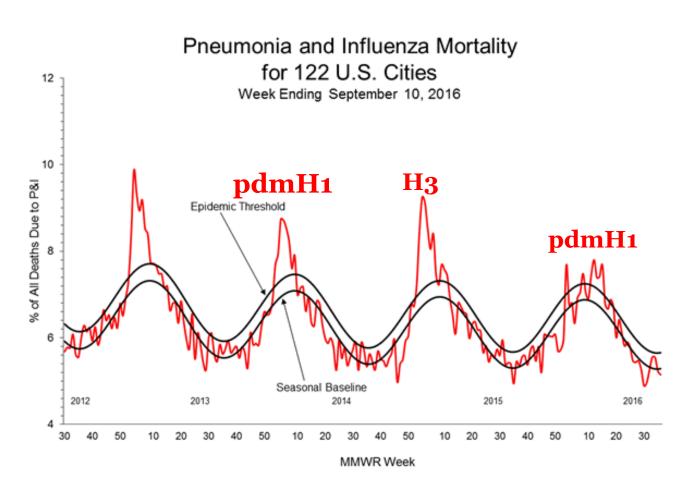
Influenza in the U.S. 2015-16

Percentage of Visits for Influenza-like Illness (ILI) Reported by the U.S. Outpatient Influenza-like Illness Surveillance Network (ILINet), Weekly National Summary, 2015-2016 and Selected Previous Seasons





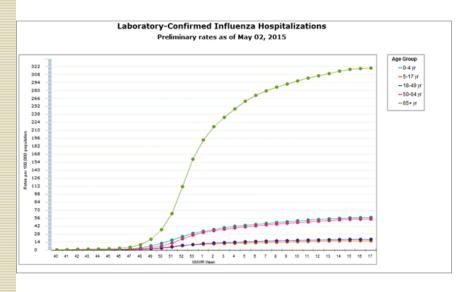
Influenza in the U.S. 2015-16



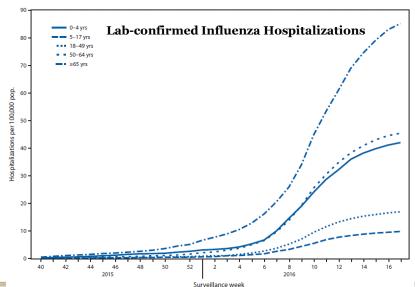


Influenza Hospitalizations

2014-15



In contrast, 2015-16





Influenza 2015-16

What was expected...

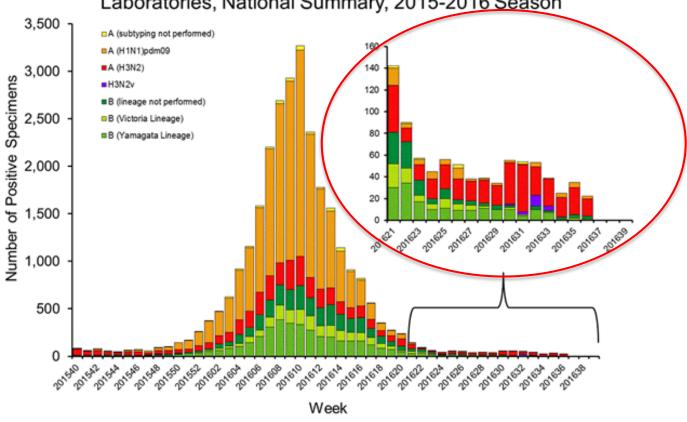
- A/Switzerland/9715293/2013(H3N2)
- A/California/7/2009
- B/Phuket/3073/2013 (B/Yamagata-lineage)
- B/Brisbane/60/2008 (B/Victoria-lineage)

... and that's what we got@



Influenza in the U.S. Early in 2016-17

Influenza Positive Tests Reported to CDC by U.S. Public Health Laboratories, National Summary, 2015-2016 Season





Early 2016-2017 Season....

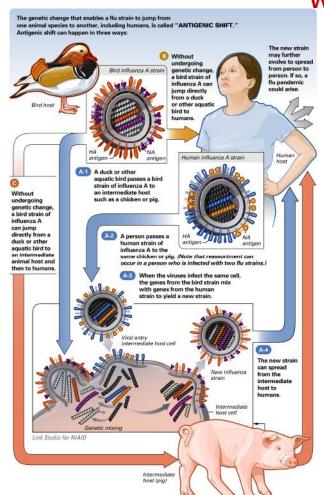


PH Region	Date Received	Influenza type
?	7/14/2016	Flu A (H3)
?	7/28/2016	Flu A (H3)
Travel (AK)	8/10/2016	Flu A (H3)
Travel (India)	8/30/2016	Flu A (H3), FluB
?	9/15/2016	Flu A (H3)
?	9/21/2016	Flu A (H3)
Monroe Co.	June 2016	Flu A H1N2v





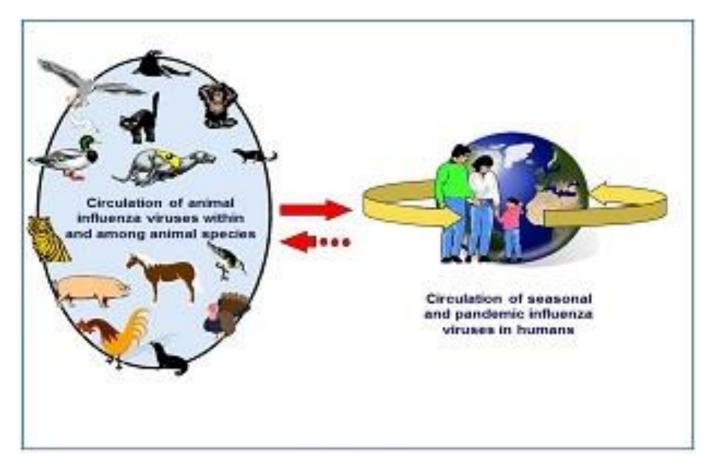
www.flu.gov



Antigenic Shift
When a new
subtype (a novel
HA and/or NA) of
influenza A
emerges in the
host (humans)



Infectious Diseases at the Human-Animal Interface Influenza as an Example



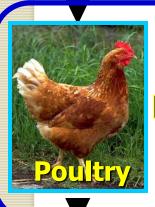
Influenza at the Human-Animal Interface



Influenza A

- H1 H17
- N1 N10









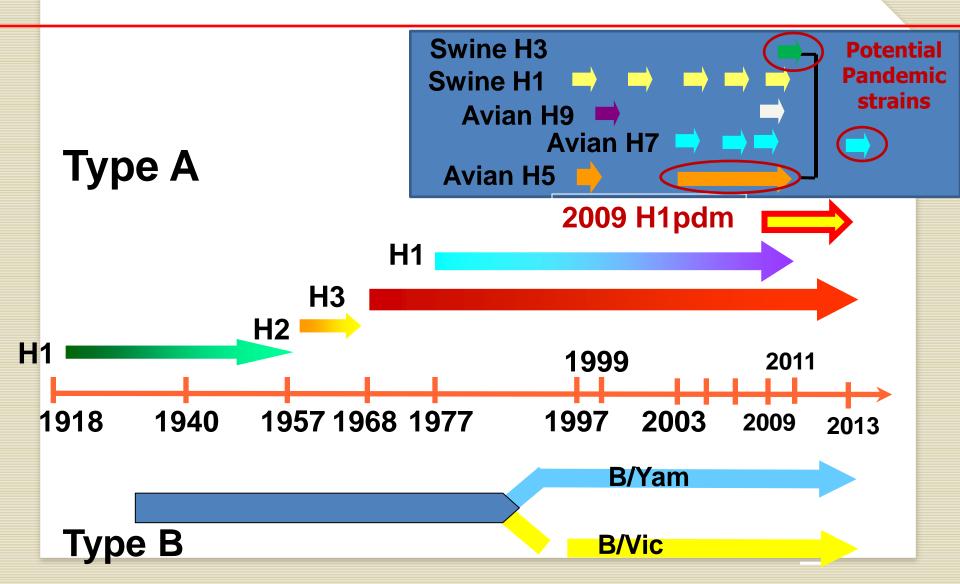








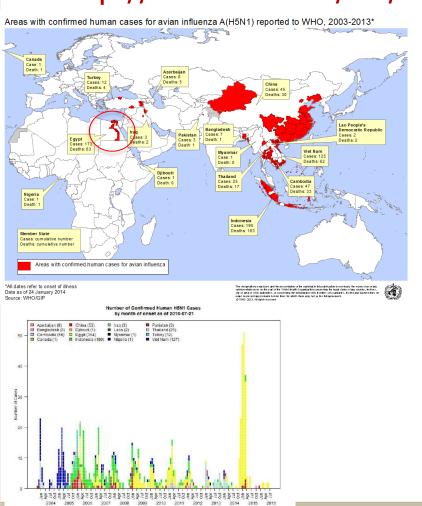
Timeline of Influenza Viruses in Human

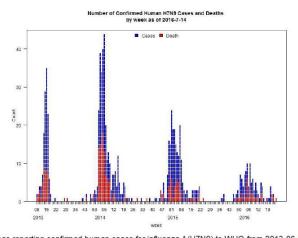


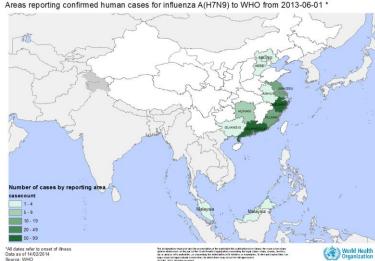
Global Influenza Concerns: A(H5N1) and A(H7N9)



http://www.who.int/csr/disease/avian_influenza/en/







Influenza: Emergence of Novel Flu A Subtypes

Chickens and turkeys take center stage

in 2015

Influenza A

- H1 H17
- N1 N10









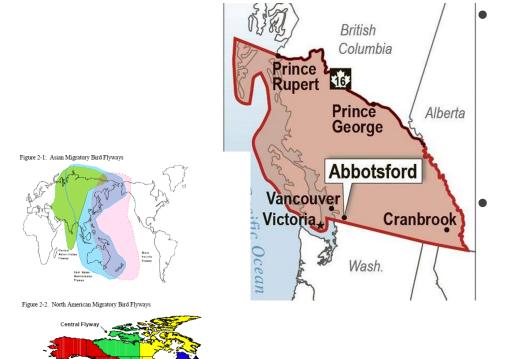






Emergence of Avian Flu (H5Nx) in U.S.





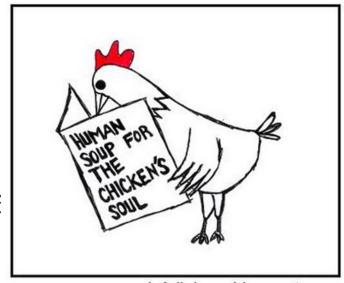
Avian Influenza (H5Nx) emerged in North America (November 2014).

Many flocks in the area were infected by December including those in the US.

Avian Influenza Terminology



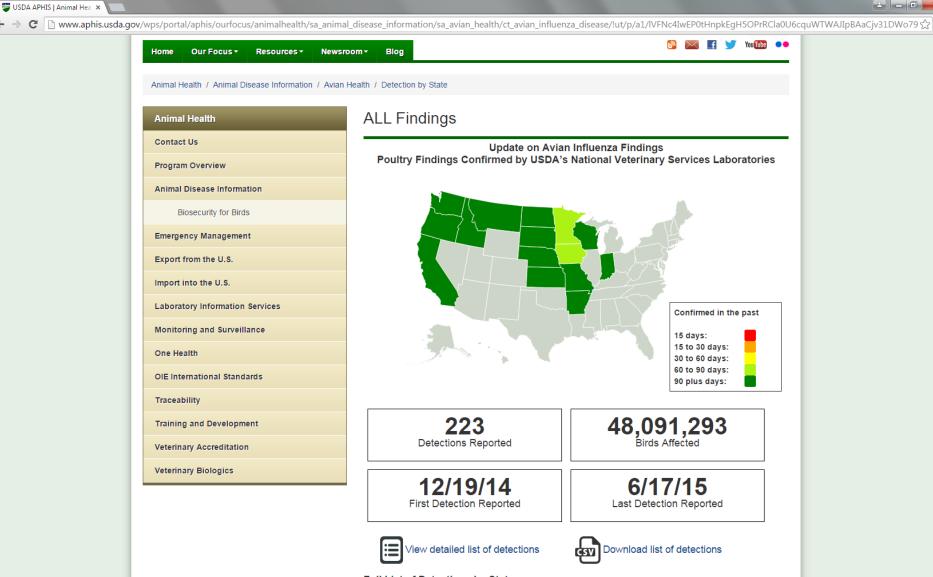
- <u>H</u>ighly <u>P</u>athogenic <u>A</u>vian
 <u>I</u>nfluenza
- Bird flu
- Pathogenicity refers to avian NOT human
- H5N1, H5N2 and H5N8 are collectively referred to as
 H5Nx
- H5N2 and H5N8 have both been detected in the US in 2015.



dcfeliciano.blogspot.com

Current Situation







The question going into the 2015-16 influenza season?



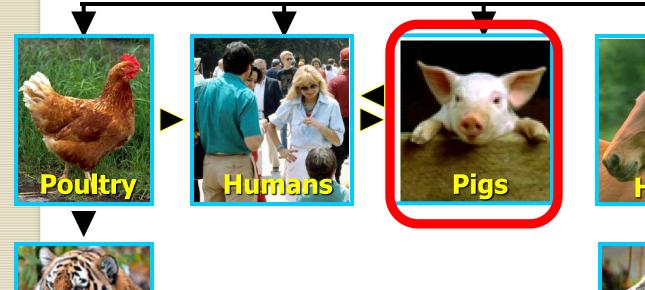
Influenza: Emergence of Novel Flu A Subtypes Don't forget about the little piggies



Influenza A

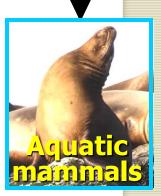
- **H1 H17**
- N1 N10







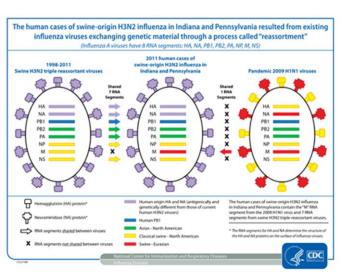


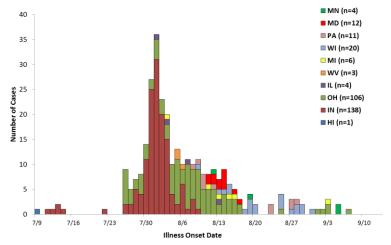




Novel Flu A events happen in the U.S. too!







These slides contain unpublished data, please DO NOT distribute or reproduce.

H3N2v Cases –September 14, 2012 (n=20*)



U.S. Influenza Surveillance



www.cdc.gov/flu/weekly



Morbidity Surveillance

Virologic Surveillance

CDC CDC

Mortality Surveillance

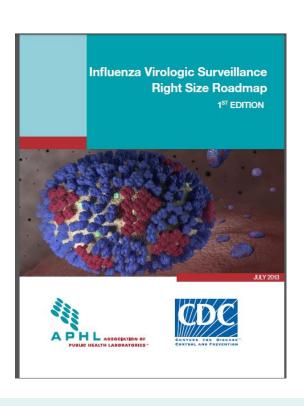


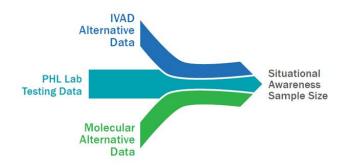
A Weekly Influenza Surveillance Report Prepared by the Influenza Division

State-level data to state surveillance coordinators

"Right-Sizing" Influenza Virologic Surveillance

The Importance of "Alternative Data"



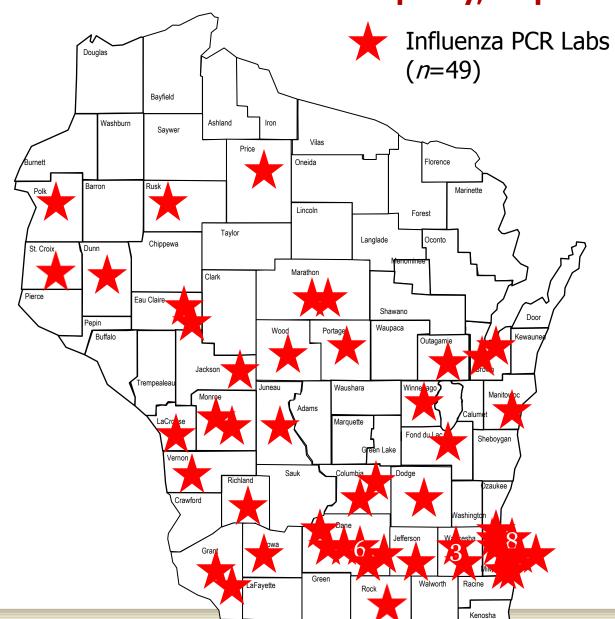


Alternative data is existing virologic data from non-PHL sources that can be used to supplement PHL data for improved situational awareness

Right Size Roadmap

http://www.aphl.org/aphlprograms/infectious/influenza/Pages/Influenza-Virologic-Surveillance-Right-Size-Roadmap.aspx

Wisconsin Labs with Flu PCR Capacity, September 2016



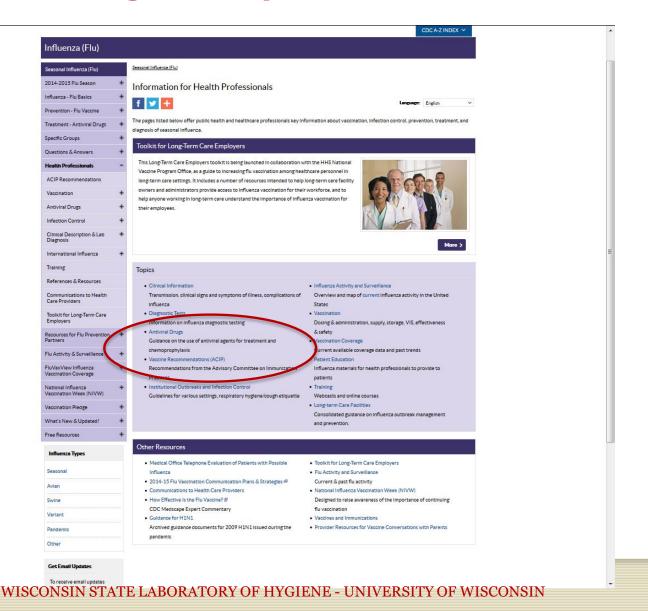


Influenza Virologic Surveillance Increasing Role for the Clinical Lab

- Provide situational awareness
 - Clinical lab testing data Via WSLH or directly CDC
- Detect novel or reassortant viruses
 Inform vaccine strain selection
 Detect and monitor antiviral resistance
 - Specimens/isolates
 → WSLH → CDC from clinical labs

Influenza — Prevention and Treatment http://www.cdc.gov/flu/professionals/index.htm







http://www.cdc.gov/flu/professionals/vaccination/ effectivenessga.htm

http://www.cdc.gov/flu/professionals/vaccination/

Table. Adjusted vaccine effectiveness estimates for influenza seasons from 2005-2016

Influenza Season'	Reference	Study Site(s)	No. of Patients:	Adjusted Overall VE (%)	95% CI
2004-05	Belongia 2009	WI	762	10	-36, 40
2005-06	Belongia 2009	WI	346	21	-52, 59
2006-07	Belongia 2009	WI	871	52	22 ,70
2007-08	Belongia 2011	WI	1914	37	22, 49
2009-10	Griffin 2011	WI, MI, NY, TN	6757	56	23, 75
2010-11	Treanor 2011	WI, MI, NY, TN	4757	60	53, 66
2011-12	Ohmit 2014	WI, MI, PA, TX, WA	4771	47	36, 56
2012-13	McLean 2014	WI, MI, PA, TX, WA	6452	49	43, 55
2013-14	Unpublished	WI, MI, PA, TX, WA	5990	51	43, 58
2014-15	ACIP presentation, Flannery	WI, MI, PA, TX, WA	9329	23	14, 31
2015-16*	ACIP presentation, Flannery	WI, MI, PA, TX, WA	563	47*	39, 53*

^{*}Estimate from Nov 2, 2015-April 15, 2016.



Vaccine usage and effectiveness 2015-16

All is not sunshine and roses

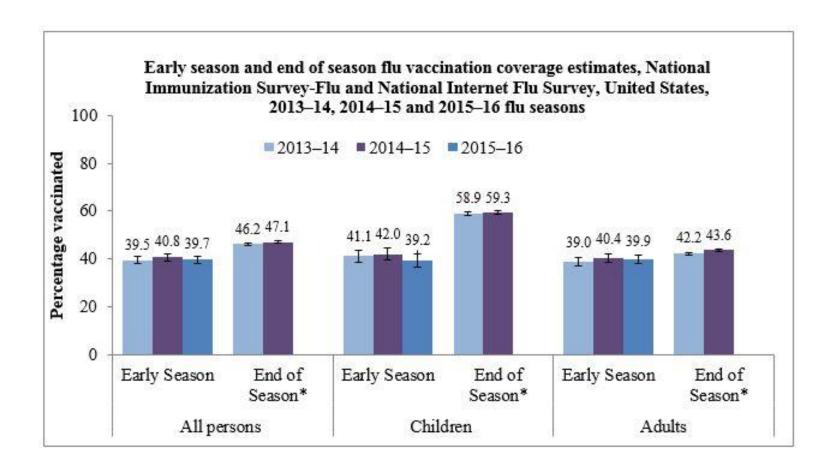
ACIP: Don't Use LAIV During 2016-17 Flu Season

-AAFP News, June 23, 2016-





http://www.cdc.gov/flu/professionals/vaccination/







<u>Multi-element approach</u>

- Rapid Influenza Diagnostic Testing (RIDT)
 Sites
 - >50% of Influenza testing in WI.
 - Confirmatory testing during periods of low prevalence!

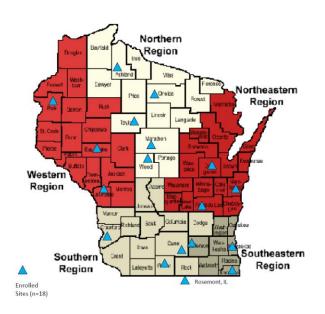
WSLH can provide confirmatory testing for out-of-season positives and the <u>first two positive influenza</u> <u>A and influenza B specimens</u>.





<u>Multi-element approach</u>

- 2. Enrolled Surveillance Sites
 - 17 labs in 5 public health regions.
 - Provide randomized specimens weekly.



Request to continue to submit the <u>first 3 specimens per</u> <u>week</u> with influenza test requests to WSLH.

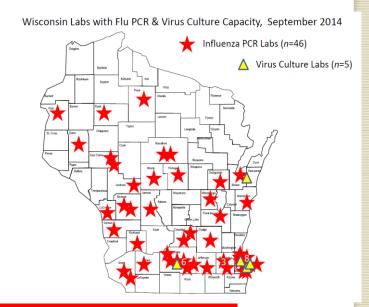
Influenza Surveillance in Wisconsin



<u>Multi-element approach</u>

- 3. PCR Labs
 - "Gold Standard" testing.
 - Provide weekly testing data summary reports.
 - 48 WI PCR labs!





Request to report both the <u>number positive</u> and the <u>number tested</u> weekly.

**Send Flu A unsubtypable specimens when subtyping for both 2009 H1N1 and seasonal H3 were attempted (Ct<35).

Laboratory-based Surveillance



All Clinical Laboratories performing influenza diagnostic testing

All Labs:

- •Send those with international travel histories
- •<u>Sampling</u> of influenza-related hospitalizations
- Unusual presentations/results
- •Contact with swine/ sick or dead poultry
- •Antiviral treatment failure

Other Pathogens of Public Health Importance to Report

- B. pertussis/ parapertussis
- RSV
- Non-influenza respiratory viruses
- Grp A Strep
- VZV
- Rotavirus

NEW! Gastropathogen PCR

Gastrointestinal Pathogens PCR Testing Please report the number of specimens tested and the number positive.

	Number Tested	Number Positive
Aeromonas		
Campylobacter		
Clostridium difficile (Toxin A/B)		
E. coli O157		
Enteroaggregative E. coli (EAEC)		
Enteropathogenic E. coli (EPEC)		
Enterotoxigenic E, coli (ETEC)		
Plesiomonas shigelloides		
Salmonella		
Shiga-like toxin-producing E. coli (STEC)		
Shigella		
Shigella/Enteroinvasive E. coli (EIEC)		
Vibrio		
Vibrio cholerae		
Yersinia enterocolitica		
Adenovirus 40/41		
Astrovirus		
Norovirus GI/GII		
Rotavirus A		
Sapovirus		
Cryptosporidium		
Cyclospora cayetanensis		
Entamoeba histolytica		
Giardia lamblia		



Reporting Lab Results

There are two options.....

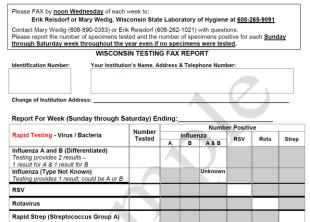
Select the method below to enter data; you must also select "Next".

1. Web-based reporting



Antigen Detection	,,	
O PCR		
O Culture		
	Back Next	

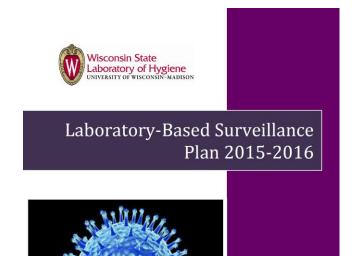
2. FAX reporting





What is the WSLH able to provide to support participating labs?

- Specimen collection supplies.
- Specimen shippers & packaging supplies.
- NO cost specimen transport.
- Influenza confirmatory testing.
- Influenza PCR validation specimen panel.
- Weekly updated surveillance data (B. pertussis, Influenza, RSV & others).
- Laboratory Surveillance Reports



Information, Forms and Instructions

Influenza Surveillance Strategy



WSLH Surveillance Coordinators

Erik Reisdorf
 Virology Lab-Team Lead
 erik.reisdorf@slh.wisc.edu

2. Mary Wedig
Electronic Reporting Coordinator
mary.wedig@slh.wisc.edu

Other Respiratory Pathogens of Public Health Importance.....



New viral pathogens discovered since

- Human metapneumovirus (2001)
- SARS-coronavirus (2003)
- Coronavirus NL63 (2004)
- Coronavirus HKU1 (2005)
- Human bocavirus (2005)
- MERS- coronavirus (2012)

Other "not so new" ones impacting public health.....

• Enterovirus D68 (2014)

Rapid Molecular Platforms









Flu A/B GAS



GAS

GAS Flu A/B Flu A/B & RSV



Benefits of NIRV Testing

- Establishes situational awareness
- Establishes etiology when influenza is not detected
- NIRV have similar clinical presentations
- Antibiotic stewardship
- Broaden capacity for outbreak investigations
- Define etiologies with severe acute respiratory illness (SARI)
- Understand burden of co-infections, emerging pathogens

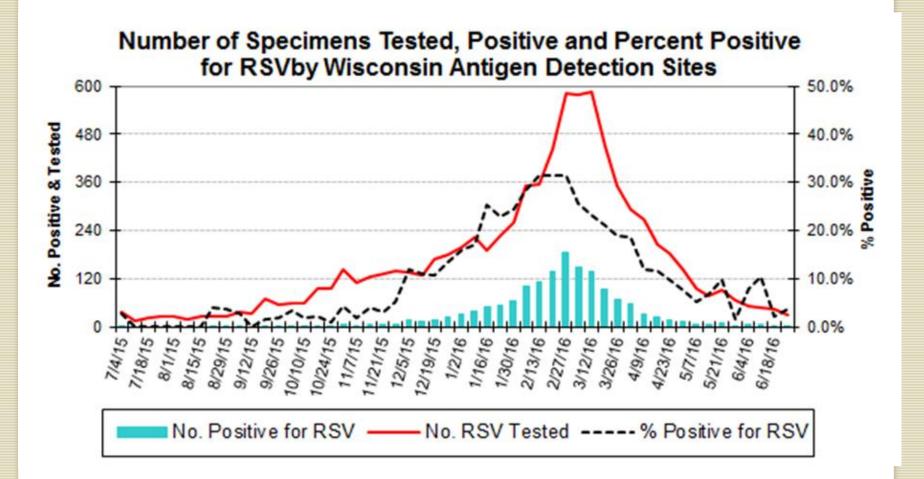


The Major Players- RSV

- Types A and B
- Leading cause of severe disease in infants and young children
- Bronchiolitis, hospitalizations (1-2% infants)
- Burden on elderly
- Seasonality varies with geography
- Infections occur throughout life
- Prophylaxis for vulnerable population--\$\$
- Vaccines in development



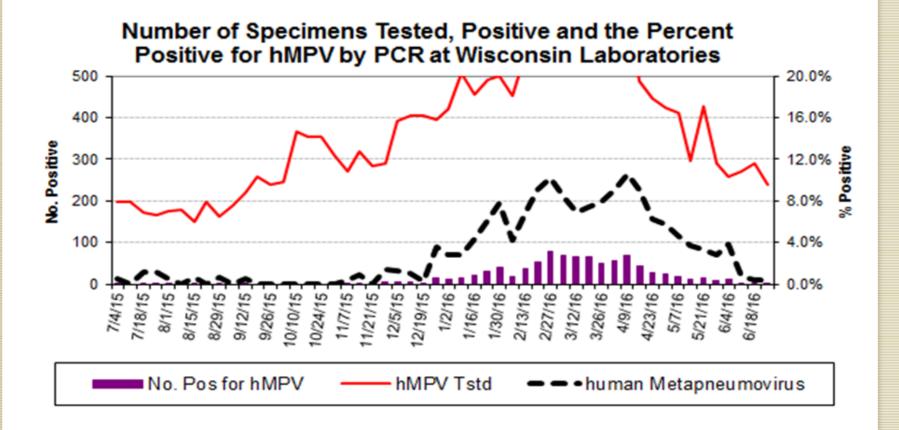
RSV Seasonality-Wisconsin



The Major Players- Metapneumovirus

- Burden primarily on children, elderly immunocompromised, COPD
- Occurs every year
- Symptoms indistinguishable with RSV
- No antivirals, treatments, or vaccines
- 51% hMPV patients prescribed antibiotics
 (Williams J et al. J Infect Dis 2006, 193: 387-95)

Human Metapneumovirus-Seasonality



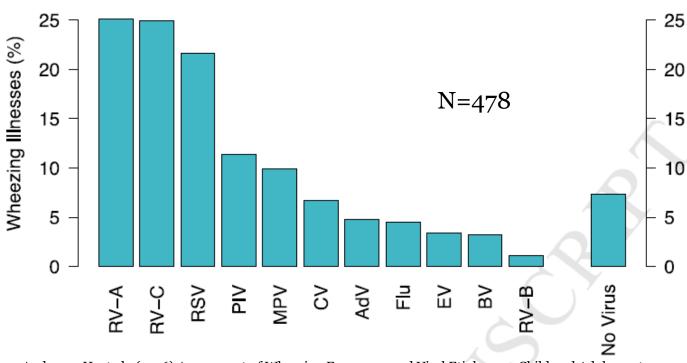


The Major Players-Rhino/Enterovirus

- "Common cold", URI sinusitis but.....LRI
- Exacerbations of asthma, wheezing
- Occurs every year, year-round
- Re-infections common
- Many types co-circulate, diversity
- Some types associated with more severe disease (e.g. paralysis, myocarditis, encephalitis); Others are mild

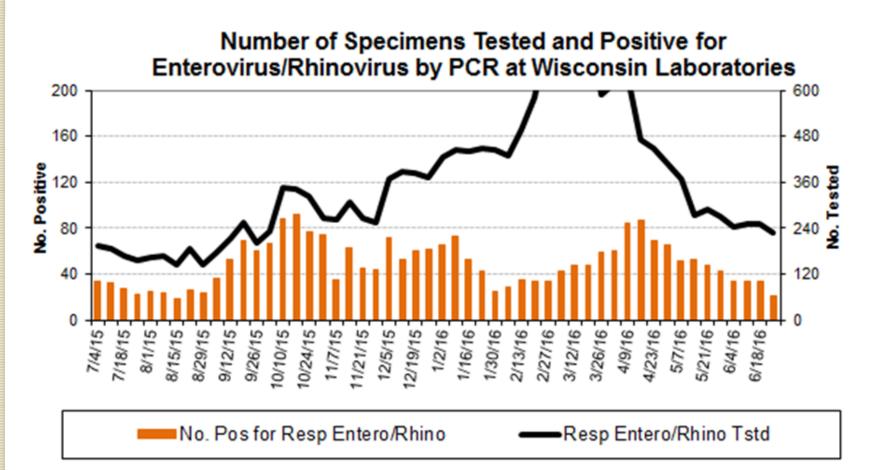


Rhinovirus and Wheezing



Anderson H, et al., (2016) Assessment of Wheezing Frequency and Viral Etiology on Child and Adolescent Asthma Risk. J Allergy and Clin Immun 26 July 2016







WI Pediatric SARI Patients (2016)

	Infants (n=51)	Toddlers (n=47)	School-Aged Children (n=37)
Bacterial growth	4 (8%)	3 (6%)	3 (8%)
Coinfection	11 (22%)	11 (23%)	5 (14%)
Virus Detected	45 (88%)	42 (89%)	27 (73%)
RSV	25 (62%)	6 (14%)	2 (7%)
Influenza	2 (4%)	1 (2%)	5 (19%)
Coronavirus	4 (9%)	4 (10%)	5 (19%)
Adenovirus	5 (11%)	7 (17%)	3 (11%)
Bocavirus	3 (7%)	3 (7%)	1 (4%)
Metapneumovirus	1 (2%)	5 (12%)	0 (0%)
Rhinovirus	14 (31%)	19 (45%)	10 (37%)
Parinfluenza	1 (2%)	3 (7%)	2 (7%)

Preliminary data courtesy of Lina Elbadawi, MD (CDC) and the WI Div. of Public Health (unpublished)





Your participation in the Wisconsin surveillance system is **vital** to monitor for emerging novel strains with pandemic potential and other pathogens that impact community health.