

# Updated interim influenza vaccine effectiveness estimates by age group and vaccine type for the 2014-15 season: Updates from the US Influenza Vaccine Effectiveness (Flu VE) Network

Brendan Flannery, PhD  
Jessie R. Clippard, MPH  
for the US Flu VE Network  
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# **2014-15 SEASON UPDATE**

# US Flu VE Network Results

- ❑ 4,913 enrolled from Nov 10, 2014–Jan 30, 2015
- ❑ 3,281 (67%) influenza RT-PCR negative
- ❑ 1,632 (33%) influenza RT-PCR positive
  - 1, 537 (94%) influenza A
    - All subtyped influenza A viruses were A(H3N2)
    - 764 genetically characterized by HA pyrosequencing
    - 85% “Low reactor” genetic groups 3C.2a (n=624), 3C.3a (n=25)
    - 15% “Vaccine-like” groups 3C.3, 3C.3b
  - 95 (6%) influenza B
    - 87 (92%) B-Yamagata lineage (determined by RT-PCR)

## Interim adjusted VE against influenza A(H3N2) and B for ≥1 dose of 2014-15 seasonal influenza vaccine

|                               | Influenza-<br>positive | %<br>vaccinated | Influenza-<br>negative | %<br>vaccinated | Adjusted VE | (95% CI)           |
|-------------------------------|------------------------|-----------------|------------------------|-----------------|-------------|--------------------|
| <b>Influenza A (H3N2)</b>     |                        |                 |                        |                 |             |                    |
| <b>All ages</b>               | 1415                   | 53%             | 3281                   | 57%             | <b>18%</b>  | <b>(6 to 29)</b>   |
| <b>Age group (yrs)</b>        |                        |                 |                        |                 |             |                    |
| 6 mos–8                       | 335                    | 46%             | 953                    | 55%             | <b>17%</b>  | <b>(-9 to 37)</b>  |
| 9–17                          | 237                    | 39%             | 414                    | 41%             | <b>12%</b>  | <b>(-27 to 39)</b> |
| 18–49                         | 399                    | 45%             | 980                    | 49%             | <b>13%</b>  | <b>(-13 to 32)</b> |
| 50-64                         | 223                    | 63%             | 529                    | 68%             | <b>25%</b>  | <b>(-7 to 48)</b>  |
| ≥65                           | 221                    | 81%             | 405                    | 86%             | <b>28%</b>  | <b>(-17 to 56)</b> |
| <b>Influenza B (Yamagata)</b> |                        |                 |                        |                 |             |                    |
| <b>All ages</b>               | 87                     | 45%             | 3281                   | 57%             | <b>45%</b>  | <b>(14 to 65)</b>  |

\* Vaccine effectiveness was estimated as  $100\% \times (1 - \text{odds ratio [ratio of odds of vaccination among flu-positive cases to odds of vaccination among flu-negative controls]})$  using logistic regression. Multivariate models adjusted for study site, age, sex, race/Hispanic ethnicity, self-rated health status, days from illness onset to enrollment, and calendar time (biweekly intervals). Models for “all ages” include age as a categorical variable; age-specific models include age in years as a continuous variable.

## Interim adjusted VE against vaccine-like vs drifted A(H3N2) among patients aged ≥6 months, for ≥1 dose of 2014-15 seasonal influenza vaccine

|   | Influenza cases | % vaccinated | Influenza-negative | % vaccinated | Adjusted VE* | (95% CI)   |
|---|-----------------|--------------|--------------------|--------------|--------------|------------|
| <b>Any influenza A (H3N2)</b>                   | 1415            | 53%          | 3281               | 57%          | 18%          | (6 to 29)  |
| <b>A(H3N2), vaccine-like (group 3C.3/3C.3b)</b> | 115             | 39%          | 3281               | 57%          | 49%          | (18 to 69) |
| <b>A(H3N2), low-reactor (group 3C.2a)</b>       | 624             | 55%          | 3281               | 57%          | 15%          | (-5, 30)   |
| <b>A(H3N2), low-reactor (group 3C.3a)</b>       | 25              | 60%          | 3281               | 57%          | -14%         | (-177, 53) |

\* Vaccine effectiveness was estimated as  $100\% \times (1 - \text{odds ratio} [\text{ratio of odds of vaccination among flu-positive cases to odds of vaccination among flu-negative controls}])$  using logistic regression. Multivariate models adjusted for study site, age category, sex, race/Hispanic ethnicity, self-rated health status, days from illness onset to enrollment, and calendar time (biweekly intervals).

**VE for LAIV and IV in children**

## Interim 2014-15 VE estimates by vaccine type among children and adolescents, US Flu VE Network

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- Analysis restricted to those 2–17 yrs of age at enrollment
- Receipt of  $\geq 1$  dose at least 14 days prior to illness onset considered vaccinated
- Determination of vaccine type
  - Vaccine type determined from medical record if available
  - Accepted vaccine type (shot or nasal spray) from parental-report if no medical record documentation
  - Excluded 12 children with unknown/unreported vaccine type
- VE for LAIV compared to unvaccinated (excludes IIV)
- VE for IIV compared to unvaccinated (excludes LAIV)

## Interim adjusted VE estimates against A(H3N2) for $\geq 1$ dose of 2014-15 seasonal influenza vaccine among children and adolescents, by vaccine type

|                                | Influenza-<br>positive | % vaccinated | Influenza-<br>negative | % vaccinated | Adjusted VE | (95% CI)     |
|--------------------------------|------------------------|--------------|------------------------|--------------|-------------|--------------|
| <b>Any vaccine</b>             |                        |              |                        |              |             |              |
| 2–17 years                     | 519                    | 43%          | 1103                   | 46%          | 7%          | (-16 to 26)  |
| 2–8 yrs                        | 282                    | 45%          | 691                    | 49%          | 6%          | (-27 to 31)  |
| 9–17 yrs                       | 237                    | 39%          | 414                    | 41%          | 12%         | (-27 to 39)  |
| <b>Live-attenuated (LAIV4)</b> |                        |              |                        |              |             |              |
| 2–17 years                     | 383                    | 22%          | 740                    | 20%          | -24%        | (-74 to 11)  |
| 2–8 yrs                        | 205                    | 25%          | 451                    | 23%          | -23%        | (-90 to 21)  |
| 9–17 yrs                       | 178                    | 19%          | 289                    | 15%          | -20%        | (-115 to 33) |
| <b>Inactivated (IIV3/IIV4)</b> |                        |              |                        |              |             |              |
| 2–17 years                     | 434                    | 31%          | 957                    | 38%          | 18%         | (-7 to 37)   |
| 2–8 yrs                        | 231                    | 33%          | 589                    | 41%          | 15%         | (-20 to 40)  |
| 9–17 yrs                       | 203                    | 29%          | 368                    | 33%          | 19%         | (-23 to 46)  |

\* Vaccine effectiveness was estimated as  $100\% \times (1 - \text{odds ratio} [\text{ratio of odds of vaccination among flu-positive cases to odds of vaccination among flu-negative controls}])$  using logistic regression. Multivariate models adjusted for study site, age, sex, race/Hispanic ethnicity, self-rated health status, days from illness onset to enrollment, and calendar time (biweekly intervals). Models for “all ages” include age as a categorical variable; age-specific models include age in years as a continuous variable.



# Adjusted Effectiveness Estimates

|                    | No Vaccination<br>(n=564) |          | IIV<br>(n=283) |          | LAIV<br>(n=141) |          | Effectiveness* (%)<br>[95% confidence interval] |                |                 |
|--------------------|---------------------------|----------|----------------|----------|-----------------|----------|---|----------------|-----------------|
|                    | Cases                     | Controls | Cases          | Controls | Cases           | Controls | Any vaccine                                     | LAIV           | IIV             |
| <b>All strains</b> | 202                       | 362      | 69             | 214      | 44              | 97       | 27<br>[-1;47]                                   | 19<br>[-28;49] | 31<br>[-1;52]   |
| <b>A/H1N1</b>      | 0                         |          | 0              |          | 0               |          | -   | -              | -               |
| <b>A/H3N2</b>      | 193                       |          | 65             |          | 44              |          | 27<br>[-1;47]                                   | 16<br>[-33;47] | 33<br>[1;54]    |
| <b>B</b>           | 9                         |          | 4              |          | 0               |          | 36<br>[-148;83]                                 | 100<br>[NE]    | -5<br>[-328;74] |

\* effectiveness estimates adjusted on site, date of enrollment, age, gender, race, passive smoking, prior vaccination, high risk chronic conditions, days since symptoms onset, health insurance status and number of outpatients visits in the last 6 months.

- For A/H3N2 effectiveness, prior vaccination found to be a significant confounder or effect modifier

# Conclusions and next steps

- ❑ **Low interim VE estimates are consistent with predominance of antigenically drifted A(H3N2) viruses**
  - H3N2 accounted for 95% of influenza-positive cases at US Flu VE Network sites; majority (>80%) were drifted H3N2 viruses
  - Low or no vaccine effectiveness against drifted H3N2 viruses
  - Limited circulation of vaccine-like A(H3N2) and influenza B viruses
  
- ❑ **No evidence of better protection for LAIV4 compared to IIV in 2014-15 season against drifted H3N2 viruses**
  
- ❑ **For final season estimates:**
  - Complete genetic characterization of 2014-15 viruses
  - Update VE estimates for LAIV and IIV

# US Flu VE Network

- *University of Michigan and Henry Ford Health System:* Arnold S. Monto, MD, Joshua G. Petrie, MPH, Suzanne E. Ohmit, DrPH, Emileigh Johnson, Rachel T. Cross, MPH, Casey Martens, RN, EJ McSpadden, MPH, Caroline K. Cheng, MPH, Katherine Reyes, MD, Lois Lamerato, PhD, Heather Lipkovich, MPH;
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- *Baylor Scott and White Health, Texas A&M University Health Science Center College of Medicine:* Manjusha Gaglani, MBBS, Jessica Pruszyński, PhD, Lydia Clipper, Anne Robertson, Kempapura Murthy, MPH, Sophia V James, MS, Teresa Ponder, Deborah Furze, Hope Gonzales, Martha Zayed, Michael Reis, MD, Pedro Piedra, MD, Vasanthi Avadhanula, PhD;
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