

End-of-Season Estimates of 2019–20 Seasonal Influenza Vaccine Effectiveness against Medically Attended Influenza from three U.S. Networks

Data from the US Flu VE Network,

New Vaccine Surveillance Network (NVSN), and
Hospitalized Adult Influenza Vaccine Effectiveness Network (HAIVEN)

Advisory Committee on Immunization Practices October 28, 2020

Preliminary results

Three networks to evaluate vaccine effectiveness against laboratory-confirmed influenza in ambulatory and inpatient settings

Ambulatory patients—ages ≥6 months: US Flu VE* Network sites and principal investigators



	US Flu VE
Setting	Ambulatory
Ages	≥6 months
Symptoms	ARI
Symptom duration	≤7

*US Flu VEUS Influenza Vaccine Effectiveness Network

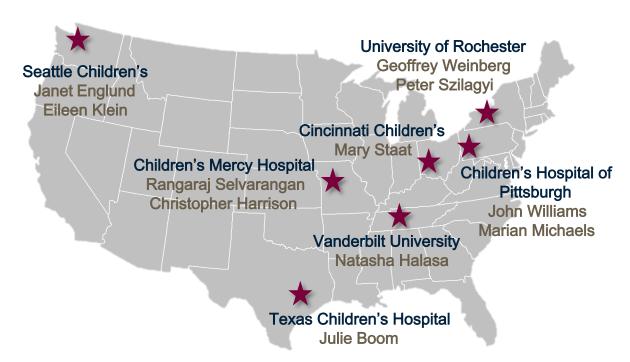
Inpatients—adults aged ≥18 years: HAIVEN* sites and principal investigators



	HAIVEN	
Setting	Inpatient	
Ages	≥18 years	
Symptoms	ARI	
Symptom duration	≤10	

^{*}HAIVEN-Hospitalized Adult Influenza Vaccine Effectiveness Network

Inpatients—children aged 6 months—17 years: NVSN* sites and principal investigators



	NVSN
Setting	Inpatient/ED
Ages	6 months-17 years
Symptoms	ARI
Symptom duration	≤10

*NVSN-New Vaccine Surveillance Network

Methods

Design: Test-negative design

- Comparing odds of influenza vaccination among patients with laboratoryconfirmed influenza vs. patients testing influenza negative by RT-PCR
- Vaccination status: receipt of one dose (≥1 dose for children 6m-8 years) of any 2019–20 seasonal flu vaccine at least 14 days prior to illness onset (if known)
- Sources: Medical records, immunization registries, and/or self-report

Analysis: $VE = (1 - adjusted OR) \times 100\%$

Adjustment for potential confounding variables (i.e. study site, age, calendar time)

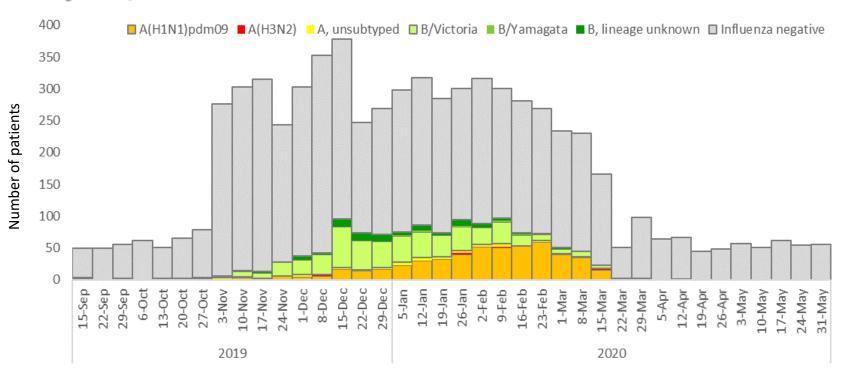
Influenza virus type/subtype distribution by VE network

	US Flu VE	HAIVEN	NVSN-Hosp
Enrollment period	Nov-Mar	Oct-Mar	Sep-May
Enrolled	8,845	3,116	2,029
Influenza pos (%)	2,722(31%)	553 (18%)	335 (17%)

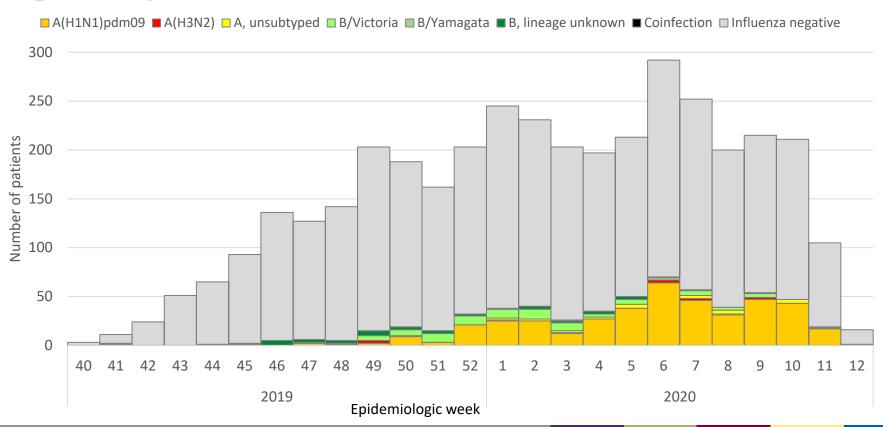
Enrollment period
Enrolled
Influenza pos (%)

NVSN-ED
Sep-May
2,102
671 (32%)

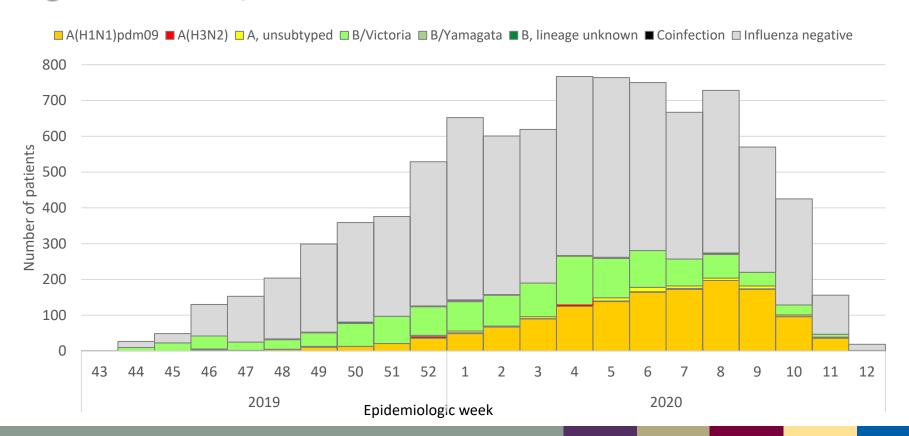
Influenza Virus Infections in NVSN Hospitalized and ED Subjects, 2019–2020



Influenza virus infections in the HAIVEN Network inpatients aged ≥18 years, 2019-2020



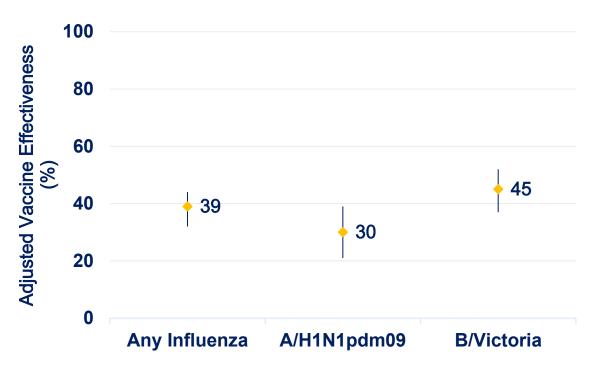
Influenza virus infections in US Flu VE Network outpatients aged ≥6 months, 2019-2020



Ambulatory settings – ages ≥6 months

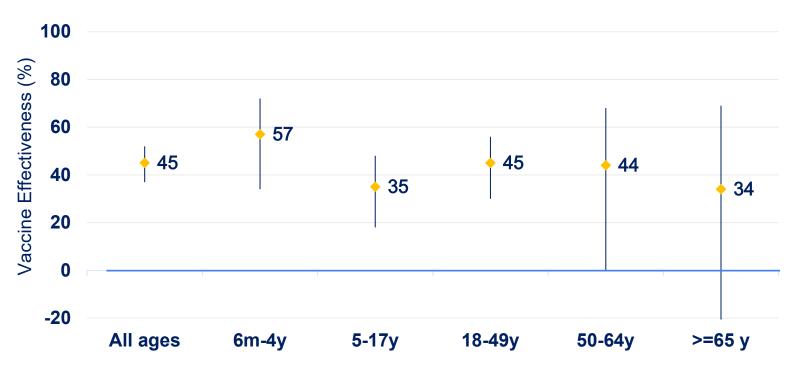
US Flu VE Network

Adjusted vaccine effectiveness against medically attended influenza ages ≥6 months, US Flu VE Network, 2019–20



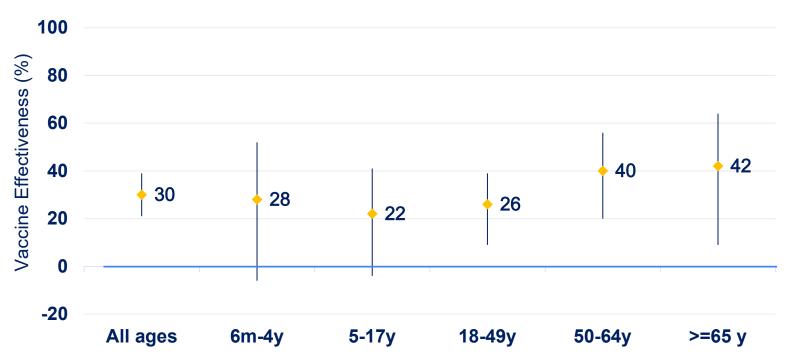
^{*} Multivariable logistic regression models adjusted for site, age, sex, race/ethnicity, self-rated general health status, interval from onset to enrollment, and calendar time.

Adjusted vaccine effectiveness against influenza B/Victoria by age group, US Flu VE Network, 2019–20



^{*} Multivariable logistic regression models adjusted for site, age, sex, race/ethnicity, self-rated general health status, interval from onset to enrollment, and calendar time.

Adjusted vaccine effectiveness against influenza A/H1N1pdm09 by age group, US Flu VE Network, 2019–20

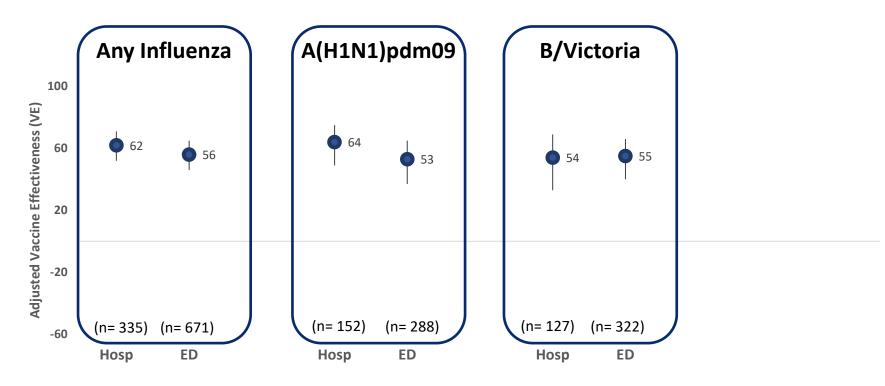


^{*} Multivariable logistic regression models adjusted for site, age, sex, race/ethnicity, self-rated general health status, interval from onset to enrollment, and calendar time.

VE against influenza associated hospitalization and ED visits among patients aged 6 months to 17 years

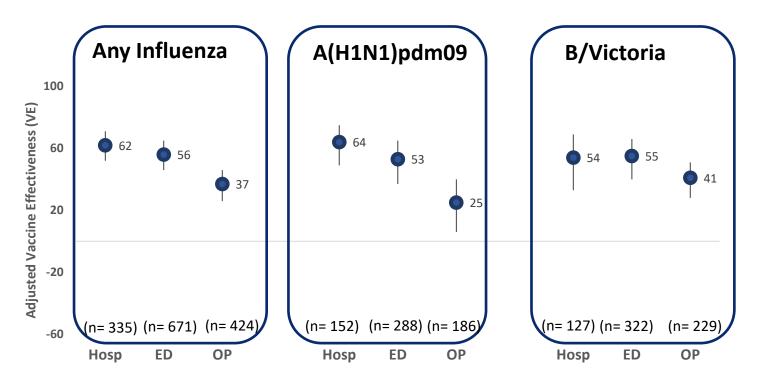
NVSN preliminary results

2019–2020 VE* against influenza hospitalizations, ED visits, by virus type, aged 6m–17 yrs, NVSN



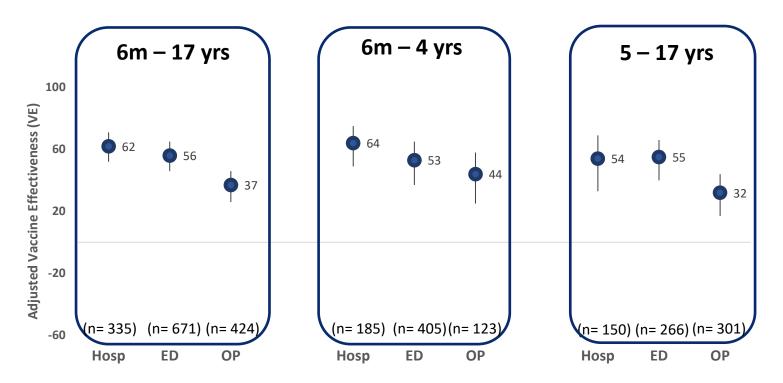
- Final models adjusted for study site, age as a continuous variable and calendar time (monthly intervals)
- n values show the total number of influenza positive subjects in each group

2019–2020 VE* against influenza hospitalizations, ED and outpatient visits by virus type, aged 6m–17 yrs, NVSN/Flu VE



- Final models adjusted for study site, age as a continuous variable and calendar time (monthly intervals)
- n values show the total number of influenza positive subjects in each group

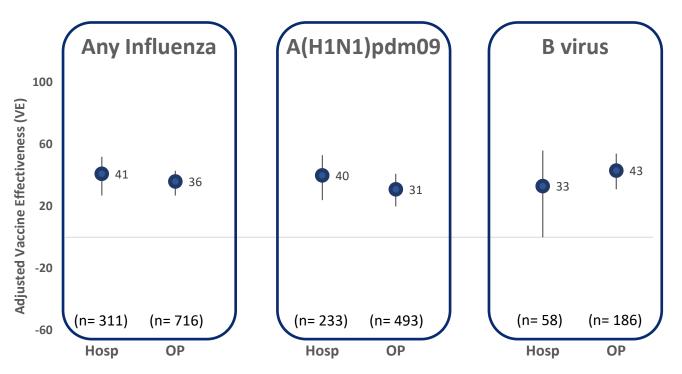
2019–2020 VE* against influenza hospitalizations, ED and outpatient visits aged 6m–17 yrs, by age group, NVSN/Flu VE



- Final models adjusted for study site, age as a continuous variable and calendar time (monthly intervals)
- n values show the total number of influenza positive subjects in each group

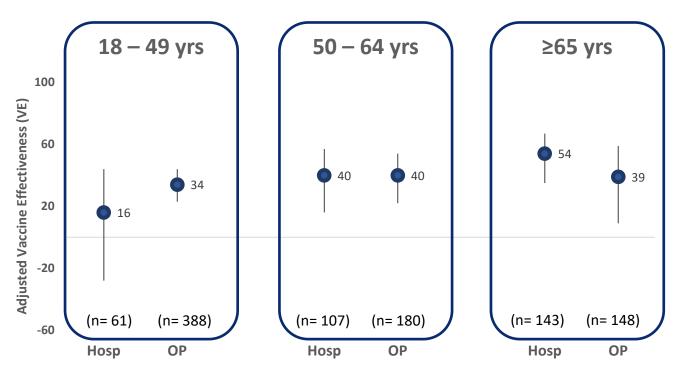
Hospitalized Adult Influenza Vaccine Effectiveness Network (HAIVEN) preliminary results

2019–2020 VE* against influenza hospitalizations and outpatient visits among adults, by virus type, HAIVEN/Flu VE



^{*} Final models adjusted for study site, age, sex, race/ethnicity, days from illness onset to specimen collection, timing of illness onset, ≥1 hospitalizations (versus none) in prior year (HAIVEN)

2019–2020 VE* against influenza hospitalizations and outpatient visits among adults, by age group, HAIVEN/Flu VE



^{*} Final models adjusted for study site, age, sex, race/ethnicity, days from illness onset to specimen collection, timing of illness onset, ≥1 hospitalizations (versus none) in prior year (HAIVEN)

Discussion

Limitations

- VE by vaccine type not included
 - Limited use of LAIV4 among children in NVSN and Flu VE to estimate
 VE for LAIV4 against A(H1N1)pdm09
 - Challenges estimating VE of egg-based vs non-egg based vaccines (recombinant/cell-culture) due to site-specific differences in vaccines offered
 - Among inpatients and outpatients aged ≥65 years, patient characteristics differed by vaccine type (majority received IIV4-HD)
- VE estimates include reported vaccination and partially vaccinated children; results are preliminary pending final vaccination data

Summary—Importance of three VE networks

- 2019–20 influenza vaccination significantly reduced laboratory confirmed medically attended influenza in all VE networks:
 - 39% (95%CI: 32, 44) against outpatient illness aged ≥6 months (Flu VE)
 - 62% (95%CI: 52, 71) against pediatric hospitalizations (NVSN)
 - 56% (95%CI: 46, 65) against pediatric ED visits (NVSN)
 - 41% (95%CI: 27, 52) against adult hospitalizations (HAIVEN)
- Important protection against influenza B virus given severity of 2019-20 season for children and vaccine mismatch
- Outpatient VE trended lower among pediatric outpatient than inpatients/ED visits; outpatient and inpatient VE similar among adults

Acknowledgments

US Flu VE Network Contributors

- Baylor Scott and White Health, Texas A&M University Health Science Center College of Medicine: Manjusha Gaglani, Chandni Raiyani, Madhava Beeram, Kelsey Bounds, Wencong Chen, Lydia Clipper, Renee Day, Amanda Drake, Mary Kylberg, Michael Smith, Kempapura Murthy, Teresa Ponder, Michael Reis, Natalie Settele, Jennifer Thomas, Jamie Walkowiak, Alejandro Arroliga
- University of Pittsburgh Schools of the Health Sciences and UPMC: Richard K. Zimmerman, Mary Patricia Nowalk, Rose Azrak, G.K. Balasubramani, Todd M. Bear, Duane Eisaman, Heather Eng, Andrew Fackler, Edward Garofolo, Robert Hickey, Philip Iozzi, Monika Johnson, Stephanie Kirk, Jason A. Lyons, Donald B. Middleton, Krissy K. Moehling, Jonathan M. Raviotta, Evelyn C. Reis, Bret Rosenblum, Sean Saul, Theresa Sax, Michael Susick, Joe Suyama, Leonard F. Urbanski, Alexandra Weissman, John V. Williams
- Kaiser Permanente Washington Health Research Institute: Michael L. Jackson, Lisa A. Jackson, Zoe Kappelman, Erika Kiniry, Lawrence Madziwa, Matt Nguyen, Suzie Park, C. Hallie Phillips, Stacie Wellwood
- University of Michigan and Henry Ford Health System: Arnold S. Monto, Emily Martin, Allen Achkar, Elizabeth Alleman, Trinh Anh Minh, Habeeb Al-Shohatee, Gabriela Augustinaitis, Sarah Bauer, Danielle Carroll, Caroline K. Cheng, Robert Deblander III, Michelle Groesbeck, Emileigh Johnson, Anne Kaniclides, Armanda Kimberly, Jenna Kiryakos, Marym Kuril, Lois E. Lamerato, Ryan E. Malosh, Maria Matta, E.J. McSpadden, Madeleine Mendelow, Joshua G. Petrie, Niharika Rajesh, Bryan Richardson, Stephanie Robinson, Hannah Segaloff, Caleb Sokolowski, Rachael Swanson, Rachel Truscon
- Marshfield Clinic Research Institute: Edward A. Belongia, Huong Q. McLean, Elizabeth Armagost, Theresa Balinghasay, Tamara Braund, Deanna Cole, Carrie Curtis, Tom Dalcher, Alicia Easley, Terry Foss, Wayne Frome, Hannah Gourdoux, Gregg Greenwald, Sherri Guzinski, Kayla Hanson, Linda Heeren, Lynn Ivacic, Marie Janz, Tara Johnson, Julie Karl, Jennifer King, Tamara Kronenwetter Koepel, Diane Kohnhorst, Sarah Kopitzke, Erik Kronholm, Marcia Lichtenwald, Carrie Marcis, Karen McGreevey, Jennifer Meece, Nidhi Mehta, Vicki Moon, Madalyn Palmquist, Nan Pan, Rebecca Pilsner, DeeAnn Polacek, Martha Presson, Lauren Putnam, Carla Rottscheit, Crystal Sabatke, Jacklyn Salzwedel, Megan Sauer, Julian Savu, Ram Shrestha, Elisha Stefanski, Patrick Stockwell, Sandy Strey
- CDC: Brendan Flannery, Mark Tenforde, Jessie Chung, Sara S. Kim, Rebecca J. Kondor, Juliana DaSilva, Shoshona Le, Thomas Stark, John R. Barnes, David E. Wentworth, Alicia M. Fry, Manish Patel

New Vaccine Surveillance Network (NVSN)

- Children's Hospital of Pittsburgh: John V. Williams, Marian G. Michaels, Robert W. Hickey, Noreen Jeffrey, Monika Johnson, Samar Musa, Sara Mascaro, Terri Sax, Judy Martin, Heather Eng, Bala Goundappa, Amy Kinzler, Cara Casilli, Andrea Collins, Mike Buklaw
- Children's Mercy Hospital Kansas City: Rangaraj Selvaranjan, Christopher J. Harrison, Jennifer E. Schuster, Barbara A. Pahud, Mary E. Moffatt, Gina Weddle, Paushpala Sen, A Kleinwolterink, Dithi Banerjee, Anjana Sasidharan
- Cincinnati Children's: Mary A. Staat, Monica McNeal, Elizabeth P. Schlaudecker, Marilyn Rice, Sahle Amsalu, Monica Asman, Rose Davis, Maureen Carolin, Skylar Deaton, Carley Dunkman, Nicole Knab, Nicholas Leahy, Julia Majchszak, Chelsea Rohlfs, Tara Keller Kevin Schmidt, Senora Schwab, Meghan Sweeney, Nicole Meyer, Amy Singh, Christina Quigley, Samantha Thomas, Amy Ostrow
- Seattle Children's: Janet A. Englund, Eileen J. Klein, Kirsten Lacombe, Bonnie Strelitz, Ashley Akramoff, Jennifer Baxter, Rachel Buchmeier, Kaitlin Cappetto, Jennifer Jensen, Sarah Korkowski, Katarina Ost, Hanna Schlaack, Hannah Smith, Sarah Steele, Ariundari Tsogoo, Emily Walter
- Texas Children's Hospital: Julie A. Boom, Leila C. Sahni, Flor M. Munoz, Pedro A. Piedra, Vasanthi Avadhanula, Kirti Patel, Karen Gincoo, Lauren Maurer
- University of Rochester: Geoffrey A. Weinberg, Peter G. Szilagyi, Wende Fregoe, Lynne Shelley, Jenelle Putzig, Jennifer Carnahan, Miranda Marchand, Theodore Pristash, Joshua Aldred, Corey Perno
- Vanderbilt University: Natasha B. Halasa, Laura Stewart, Rendie McHenry, Sandy Alvarez Calvillo, Yesenia Romero, Ritu Banerjee, Kristen Ogden, Keipp Talbot, Heidi Rahman, Claudia Guevara Pulido, Erin Yepsen, Derek Williams, Carlos Grijalva, Rebekkah Varjabedia
- Centers for Disease Control and Prevention: Angela P. Campbell, Constance Ogokeh, Manish Patel, Alicia M. Fry, Lauren Beacham, John Barnes, LaShondra Berman, Brian Rha, Joana Y. Lively, Daniel C. Payne, Mary Wiskwo, Sue Gerber, Gayle Langley, Aaron Curns, Idris Sulemana, Yingtao Zhou, Roberto Mejia, Sharad Aggarwal, Umesh Parashar, Jessie Chung

HAIVEN Contributors

- Baylor Scott & White Health, Texas A&M University Health Science Center College of Medicine: Manjusha Gaglani, Kempapura Murthy, Wencong Chen, Tresa McNeal, Shekhar Ghamande, Kevin Chang, Heath White, Chaitanya Reddy, John Manning, Nathan Drew, Michael Ladogana, Jadvinder Goraya, Gonzalo Salgado Gonzales, Joseph Fernandez, Robert Stewart, March Eleison, Karen Patterson, Robert Fader, Manohar Mutnal, Arundhati Rao, Lydia Clipper, Victor Escobedo, Kelsey Bounds, Tommie Bell, Michael Smith, Chandni Raiyani, Martha Zayed, Mary Kylberg, Justin Paradeza, Natalie Settele, Jennifer Thomas, Deborah Price, Jeremy Ray, Renee Day, Jaime Walkowiak, Madhava Beeram, Michael Reis, Wright Bates, Jay Fox, Jim Morrison, Glenn Robinson, John Erwing, Alejandro Arroliga
- University of Michigan, Henry Ford Health System, Saint Joseph Mercy Health System: Arnold S. Monto, Emily Toth Martin, Joshua G. Petrie, Lois E. Lamerato, Anurag Malani, Adam Lauring, Ryan E. Malosh, EJ McSpadden, Hannah Segaloff, Caroline Cheng, Rachel Truscon, Emileigh Johnson, Armanda Kimberly, Anne Kaniclides, Emily Nichols, Elizabeth Alleman, Sarah Bauer, Michelle Groesbeck, Amy Burghardt, Jenna Russell, Lisa Mayer, Caleb Ward, Chinwendu Uzosike, Asmaa Ibrahim, Hope Wheeler, Hanna Wilhelm, Danielle Kassa, Emily Wade, Stephanie Otto
- University of Pittsburgh Schools of the Health Sciences and UPMC: Richard K. Zimmerman, Mary Patricia Nowalk, G.K. Balasubramani, Donald Middleton, Silveira Fernanda, Lori Stiefel, John Williams, Heather Eng, Sean Saul, Michael Susick, Lydia Taylor, Melissa Saul, Kailey Hughes, Joseph Penzelik, Terrie Sax, Monika Johnson, Charles Rinaldo, Tracey Conti MD, Mohammed Yassin MD
- Vanderbilt University and St. Thomas Health System: H. Keipp Talbot, Dayna Wyatt, Yuwei Zhu, Zhouwen Liu, Stephanie Longmire, Parker Hodge, Jennifer Patrick, Emily Sedillo, Chris Trabue, Jessica Freeman, Alina Simion, Karen Speer, Peiman Nebi, Hollie Horton, Bethany Alicie, Briana Krantz, Donna Carillo, Laura Adams, Amelia Drennan, Jan Orga and Lynn Peterson, Natasha Halasa, Rendi McHenry, Claudia Guevara Pulido
- Centers for Disease Control and Prevention: Jill M. Ferdinands, Manish Patel, Emily Smith, Mark Tenforde, Samantha Olson, Alicia M. Fry



Updates and Work Group Considerations

Lisa A. Grohskopf, MD, MPH
Influenza Division
CDC/NCIRD

Advisory Committee on Immunization Practices October 28, 2020

Acknowledgements

Influenza Division

Elif Alyanak

Noreen Alabi

Lenee Blanton

Lynnette Brammer

Alicia Budd

Jessie Chung

Vivien Dugan

Scott Epperson

Jill Ferdinands

Brendan Flannery

Alicia Fry

Krista Kniss

Manish Patel

Melissa Rolfes

Jerry Tokars

Tim Uyeki

Immunization Safety Office

Karen Broder

Frank Destefano

Penina Haber

Tom Shimabukuro

Patricia Wodi

Immunization Services Division

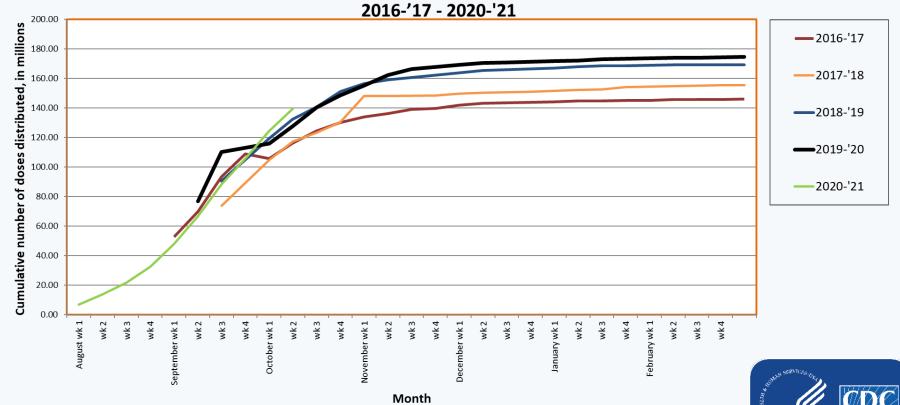
Sam Graitcer

Andrew Kroger

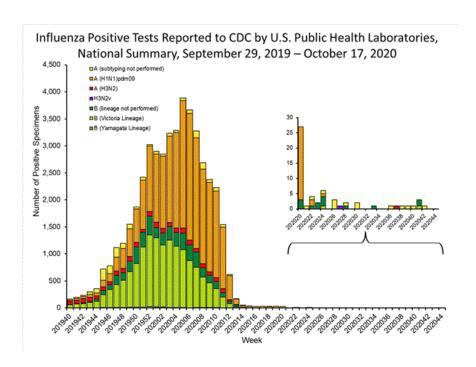
Amy Parker Fiebelkorn

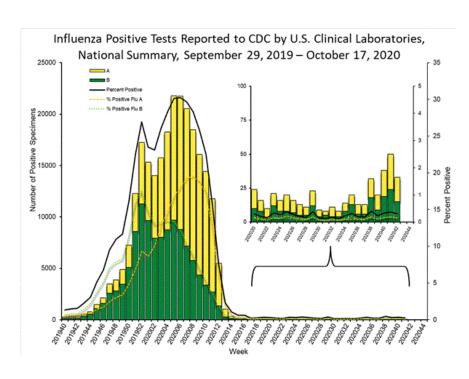
Jeanne Santoli

CDC's FluFinder Distribution Tracking Program: Cumulative doses of influenza vaccines distributed by month, by season:

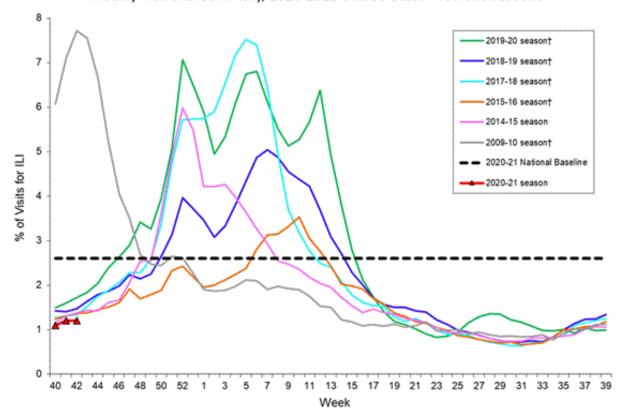


FluView, Week 42 (ending October 17, 2020): Virologic Surveillance





Percentage of Visits for Influenza-like Illness (ILI) Reported by the U.S. Outpatient Influenza-like Illness Surveillance Network (ILINet), Weekly National Summary, 2020-2021 and Selected Previous Seasons



FluView, Week 42 (ending October 17, 2020):

Influenza-Like
Illness Surveillance

†These seasons did not have a week 53, so the week 53 value is an average of week 52 and week 1

Work Group Discussion and Considerations

- Influenza WG heard and discussed presentation of trial of Flucelvax Quadrivalent for children aged ≥2 through <18 years.</p>
- Some clarifying questions concerning study design and study population.
- No specific concerns raised.

For more information, contact CDC 1-800-CDC-INFO (232-4636) TTY: 1-888-232-6348 www.cdc.gov

The findings and conclusions in this report are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention.

