# Impact of 13-valent Pneumococcal Conjugate Vaccine (PCV13) on Non-bacteremic Pneumococcal Pneumonia (NBPP) among Adults in the United States, 2013-2017

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## Background

- Streptococcus pneumoniae (pneumococcus) is a common etiology of all-cause pneumonia among adults
- True burden of pneumococcal pneumonia is unknown due • to limitations of available diagnostic tests<sup>1</sup>
  - » Blood cultures have low sensitivity
  - » Commercially available pneumococcal urine antigen test (PUAT): 75% sensitivity and not routinely used by all providers<sup>2, 3</sup>
- In 2014, PCV13 recommended for adults aged ≥65 years, in series with 23-valent pneumococcal polysaccharide vaccine (PPSV23) based on demonstrated efficacy against NBPP<sup>4,5</sup>
- PCV13 coverage among adults ≥65 years was 43% in 2017
- We evaluated PCV13 impact on NBPP among adults

# **Methods**

- NBPP defined as clinically or radiographically confirmed pneumonia and a positive PUAT in a hospitalized adult aged  $\geq$ 18 years)
- NBPP cases identified at select hospitals in 10 sites within CDC's Active Bacterial Core surveillance from 2013-2017
- NBPP rates (per 100,000) were estimated using U.S. Census Bureau population denominators
- Rates adjusted for the proportion of pneumonia patients tested by PUAT and the number of pneumonia admissions in the catchment area
- Generalized linear mixed model used to estimate the percent positive PUAT among all cause pneumonia
  - Year and age group fixed effects »
  - Hospitals are treated as random effects
  - Assume PUAT is randomly used within each hospital, age group, » and year
- Multiply the percent positive and test sensitivity (75%) to the all-cause pneumonia count to estimate incidence
- Percent change in incidence rates calculated comparing years 2013 to 2014 and years 2014 to 2017
  - 95% confidence intervals calculated using bootstrap resampling

# Results

- Between 2013 and 2017, 4,435 NBPP cases were identified
- Adults aged  $\geq$  65 years accounted for 49% of cases (Table), with a case fatality ratio of 9%; compared to a case fatality ratio of 4% among adults aged 18-49 years
- From 2013 to 2014, rates of NBPP declined from 162 to 95 (41% reduction, 95%CI 30%, 51%) in ≥ 65 year-olds; 65 to 30 (34% • reduction, 95%CI 22%, 45%) in 50-64 year-olds; and 16 to 10 (37% reduction, 95%CI 25%, 47%) in 18-49 year-olds (Figure)
- From 2014 to 2017, rates of NBPP increased slightly in all ages, but remained below 2013 rates (Figure)

### **Table. Demographics and Clinical Characteristics** of PUAT Positive Case-Patients, 2013–2017

Pre-PCV13, Post-PCV13, 2013-2014 2015-2017 (N= 1,856) (N= 2,579) Demographics n (%) n (%) Age groups, years 18–49 348 (19) 475 (19) 50-64 554 (30) 890 (35) 954 (52) 1214 (47) ≥65 per Median age, years (range) 65 (18-102) 64 (18-105) Male 855 (46) 1196 (46) Hispanic 86 (5) 142 (6) Race: 1,200 (65) 1602 (62) White Black 433 (23) 686 (27) Community onset<sup>1</sup> 1602 (86) 2211 (86) Immunocompromising condition<sup>2</sup> 756 (41) 1495 (42) High risk condition<sup>3</sup> 862 (47) 1190 (46) 61 (3) 32 (1) Received PPSV23 0-3 days before UAT **Clincal Characteristics** Radiographically diagnosed pneumonia 1608 (86) 2275 (88) 812 (32) ICU care 655 (35) Died 119 (7) 152 (6) 5 (0-152) 7 (0 - 120) Median hospitalization (days) (range) Pneumococcal vaccine receipt 259 (14) 230 (9) (current hospitalization)



<sup>1</sup>Not residing in a hospital setting or admitted at least 72 hours before UAT obtained

<sup>2</sup>Immuncompromising conditions defined as those for which PCV13 and PPSV23 are recommended for adults 19-64 years old

<sup>3</sup>High risk conditions defined as those for which PPSV23 is recommended for adults 19–64 years old

<sup>1</sup>Said M.A., et al (2013). Estimating the burden of pneumococcal pneumonia among adults... PloS one. 8(4):e60273. Epub 2013 Apr 2 <sup>2</sup>Horita, N., et al (2013). Sensitivity and specificity of the Streptococcus pneumoniae urinary antigen test... Respirology 18(8): 1177-83. <sup>3</sup>Sinclair, A., et al (2013). Systematic review and meta-analysis of a urine-based pneumococcal antigen test... J Clin Microbiol 51(7): 2303-2310 <sup>4</sup> Bonten MJM, et al (2015). Polysaccharide conjugate vaccine against pneumococcal pneumonia... NEJM. 372:1114-25. <sup>5</sup> McLaughlin, J. M., et al (2018). Effectiveness of PCV13 Against Hospitalization for Community-Acquired Pneumonia in Older US Adults: ... Clin Infect Dis.

# **Figure. Estimated Annual Non-Bacteremic Pneumococcal**

11 (-6, 31)

# **Conclusions**

- NBPP incidence declined among adults
  - » Decrease most dramatic between 2013-2014 (indirect effects of vaccine use in children)
- No additional reductions in NBPP rates among adults since vaccine recommendation for adults aged  $\geq$ 65 years and as vaccine coverage among adults increased
- New PCVs in phase 3 trials covering more serotypes and have potential to further reduce pneumococcal pneumonia among adults through both direct and indirect effects

# Limitations

- Serotype distribution unknown
  - » Unable to determine burden of vaccine-type pneumonia
  - » Unable to determine if increases in non-vaccine type pneumonia minimize overall reductions
- UAT testing practices are likely not at random
- Adjusted incidence based on ICD codes for pneumonia » coding practices may change over time and by hospital/site
- Relatively short time periods for both pre- and post-PCV13 data

# **Affiliates / Partners**

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