

Post-13-Valent Pneumococcal Conjugate Vaccine Dynamics in Young Children

Corinne Levy, Naim Ouldali, Emmanuelle Varon, Stéphane Béchet, Stéphane Bonacorsi, Robert Cohen

Author affiliations: Association Clinique et Thérapeutique Infantile du Val-de-Marne, Créteil, France (C. Levy, N. Ouldali, S. Béchet, R. Cohen); Groupe de Pathologie Infectieuse Pédiatrique, Paris, France (C. Levy, N. Ouldali, E. Varon, S. Bonacorsi, R. Cohen); Université Paris Est, IMRB-GRC GEMINI, Créteil (C. Levy, E. Varon, R. Cohen); Centre Hospitalier Intercommunal de Créteil, Créteil (C. Levy, E. Varon, R. Cohen); Association Française de Pédiatrie Ambulatoire, Saint-Germain-en-Laye, France (C. Levy, R. Cohen); Assistance Publique-Hôpitaux de Paris, Hôpital Robert Debré, ECEVE INSERM UMR 1123, Paris (N. Ouldali, S. Bonacorsi); Université de Paris, IAME, UMR 1137, INSERM, Paris (S. Bonacorsi)

DOI: <https://doi.org/10.3201/eid2708.210037>

To the Editor: We read with interest the article by Ben-Shimol et al. (1), which described the disproportionate increase of non-13-valent pneumococcal conjugate vaccine (PCV) additional PCV20 serotypes (vaccine type [VT] 20–13) in patients who had respiratory infections or invasive pneumococcal disease (IPD) after PCV13 implementation in Israel. The authors emphasized the higher disease potential of VT20–13 serotypes compared with non-VT20

serotypes. We would like to complement their results with data from France and highlight the similarities and the differences in serotype distribution. Our long-term prospective population-based surveillance comprises pneumococcal isolates from 793 healthy carriers, 4,474 acute otitis media patients, and 441 IPD patients, all of whom were children <24 months of age (2–4). We found that VT13 serotypes accounted for 8%, VT20–13 for 30%, and non-VT20 for 60% of infections in healthy carriers and acute otitis media patients during 2015–2018. Like Ben-Shimol et al. (1), we found that the most common VT20–13 serotypes were 15B/C and 11A, and the most common non-VT20 serotypes were 23B, 15A, and 35B.

From the early PCV13 (2009–2011 in both countries) to late PCV13 period (2015–2017 in Israel and 2015–2018 in France), the prevalence of IPD caused by VT13 serotypes declined by ≈90% in both countries. However, VT20–13 serotypes predominated in Israel, whereas non-VT20 serotypes predominated in France. Although Israel had higher proportions of serotypes 12F (26.9% vs. 0.9%) and 33F (10.6% vs. 4.4%) than France (1), France saw the emergence of the non-VT20 serotype 24F (27.4%) during 2015–2018. This emergent serotype led to a higher proportion of PCV20 serotypes in Israel (62%) than in France (41.6%). The differences in vaccine type distribution between the 2 countries were mainly based on the very high rates of serotypes 12F and 33F in Israel and the emerging serotype 24F in France (5). Apart from these serotypes, the serotype distribution in IPD was very similar (Table).

In conclusion, data from Israel and France show a similar effect of PCV13 on the distribution of VT13 serotypes. The role of emerging non-PCV13 serotypes in carriage was also similar. However, we observed unexpected discrepancies in the serotype replacement pattern, driven by few highly invasive non-PCV13 serotypes. This finding suggests that serotype replacement during the PCV13 era is complex and multifactorial, and has implications for the expected effects of next-generation PCVs. Finally, France and Israel had similar serotype distributions that differed in only the late PCV13 period as a result of the emergence of some invasive specific clones (5–8).

These studies were supported by the Pediatric Infectious Diseases Group of the French Pediatrics Society, Association Clinique et Thérapeutique Infantile du Val de Marne, and Pfizer Investigator-Initiated Research grants. The National Reference Center for Pneumococci was partially funded by the French National Health Agency.

Table. Comparison of pneumococcal serotypes in children <24 mo of age in Israel, 2015–2017, and France, 2015–2018*

Serotypes	Cases of invasive pneumococcal disease, no. (%)	
	Israel	France
Total	216 (100.0)	113 (100.0)
PCV13 serotypes	22 (10.2)	5 (4.4)
22F	8 (3.7)	8 (7.1)
33F	23 (10.6)	5 (4.4)
PCV15 serotypes†	53 (24.5)	18 (15.9)
8	2 (0.9)	2 (1.8)
10A	8 (3.7)	11 (9.7)
11A	3 (1.4)	2 (1.8)
12F	58 (26.9)	1 (0.9)
15B/C	10 (4.6)	13 (11.5)
PCV20 serotypes‡	134 (62.0)	47 (41.6)
Non-VT20	82 (38.0)	66 (58.4)
24F	7 (3.2)	31 (27.4)
VT20–13	112 (51.9)	42 (37.2)
Non-PCV13	194 (89.8)	108 (95.6)

*Non-VT20, serotypes in PCV20; PCV13, 13-valent PCV; PCV15, 15-valent PCV; PCV20, 20-valent PCV; PCV, pneumococcal conjugate vaccine; VT20–13, vaccine types in PCV20 but not PCV13.

†Comprises PCV13 serotypes as well as 22F and 33F.

‡Comprises PCV15 serotypes as well as 8, 10A, 11A, 12F, and 15B/C.

R.C., C.L., and E.V. received personal fees and nonfinancial support from Pfizer. R.C. reports personal fees from AstraZeneca GSK, Merck Sharp & Dohme, and Sanofi outside of the context of the submitted work.

References

1. Ben-Shimol S, Givon-Lavi N, Kotler L, Adriaan van der Beek B, Greenberg D, Dagan R. Post-13-valent pneumococcal conjugate vaccine dynamics in young children of serotypes included in candidate extended-spectrum conjugate vaccines. *Emerg Infect Dis.* 2021;27:150–60. <https://doi.org/10.3201/eid2701.201178>
2. Cohen R, Levy C, Ouldali N, Goldrey M, Béchet S, Bonacorsi S, et al. Invasive disease potential of pneumococcal serotypes in children after PCV13 implementation. *Clin Infect Dis.* 2021;72:1453–6. <https://doi.org/10.1093/cid/ciaa917>
3. Levy C, Vie le Sage F, Varon E, Chalumeau M, Grimprel E, Cohen R. Pediatric ambulatory and hospital networks for surveillance and clinical epidemiology of community-acquired infections. *J Pediatr.* 2018;194:269–270.e2. <https://doi.org/10.1016/j.jpeds.2017.11.050>
4. Ouldali N, Cohen R, Levy C, Gelbert-Baudino N, Seror E, Corrad F, et al. Pneumococcal susceptibility to antibiotics in carriage: a 17 year time series analysis of the adaptive evolution of non-vaccine emerging serotypes to a new selective pressure environment. *J Antimicrob Chemother.* 2019;74:3077–86. <https://doi.org/10.1093/jac/dkz281>
5. Rokney A, Ben-Shimol S, Korenman Z, Porat N, Gorodnitzky Z, Givon-Lavi N, et al. Emergence of *Streptococcus pneumoniae* serotype 12F after sequential introduction of 7- and 13-valent vaccines, Israel. *Emerg Infect Dis.* 2018;24:453–61. <https://doi.org/10.3201/eid2403.170769>
6. Janoir C, Lepoutre A, Gutmann L, Varon E. Insight into resistance phenotypes of emergent non 13-valent pneumococcal conjugate vaccine type pneumococci isolated from invasive disease after 13-valent pneumococcal conjugate vaccine implementation in France. *Open Forum Infect Dis.* 2016;3:ofw020. <https://doi.org/10.1093/ofid/ofw020>
7. Lo SW, Gladstone RA, van Tonder AJ, Lees JA, du Plessis M, Benisty R, et al.; Global Pneumococcal Sequencing Consortium. Pneumococcal lineages associated with serotype replacement and antibiotic resistance in childhood invasive pneumococcal disease in the post-PCV13 era: an international whole-genome sequencing study. *Lancet Infect Dis.* 2019;19:759–69. [https://doi.org/10.1016/S1473-3099\(19\)30297-X](https://doi.org/10.1016/S1473-3099(19)30297-X)
8. Ouldali N, Levy C, Varon E, Bonacorsi S, Béchet S, Cohen R, et al.; French Pediatric Meningitis Network. Incidence of paediatric pneumococcal meningitis and emergence of new serotypes: a time-series analysis of a 16-year French national survey. *Lancet Infect Dis.* 2018;18:983–91. [https://doi.org/10.1016/S1473-3099\(18\)30349-9](https://doi.org/10.1016/S1473-3099(18)30349-9)

Address for correspondence: Corinne Levy, ACTIV, 31, rue Le Corbusier, 94000 Créteil, France; email: corinne.levy@activ-france.fr

Estimate of Burden and Direct Healthcare Cost of Infectious Waterborne Disease in the United States

Stephanie DeFlorio-Barker, Abhilasha Shrestha, Samuel Dorevitch

Author affiliations: US Environmental Protection Agency, Research Triangle Park, North Carolina, USA (S. DeFlorio-Barker); University of Illinois–Chicago School of Public Health, Chicago, Illinois, USA (A. Shrestha, S. Dorevitch)

DOI: <https://doi.org/10.3201/eid2708.210242>

To the Editors: We read with interest an article by S.A. Collier et al. (1) estimating the economic burden of waterborne illnesses in the United States. Although we found the study noteworthy, the burden estimates differ greatly from those in our 2018 study (2) of the economic burden from recreational waterborne illness in the United States. The studies estimated very different numbers of cases: Collier et al. estimated ≈ 7.1 million total waterborne illnesses, but we estimated ≈ 90 million recreational waterborne illnesses in untreated water. Collier et al. estimated \$3.3 billion in total direct costs from all waterborne illness caused by 17 pathogens, but we estimated \$2.9 billion from recreational waterborne illness alone. Both studies used similar methods to address underreporting and underdiagnosis of illness. Key differences between studies include that Collier et al. summarized healthcare costs associated with infections caused by 17 pathogens that might be waterborne, then relied heavily on expert judgment (3) to estimate the proportion attributable to water exposure. In contrast, our study used data from large cohort studies of water recreation to estimate the burden from mild and moderate illnesses and outbreak data to estimate the burden from severe illnesses from water recreation. Collier et al. estimated the direct costs of illness, whereas our study estimated both direct and indirect costs (e.g., workplace absence). Enteric pathogens responsible for gastrointestinal symptoms after water recreation are generally not identified in clinical testing (4); because Collier et al. used economic burden estimates from waterborne illness based only on 17 pathogens, the study substantially underestimates the overall number of cases and associated economic burden. Future research should consider cohort and outbreak data for treated and untreated recreational water and drinking water to estimate the total economic burden from waterborne illness.