

# Vaccination Patterns in Children After Autism Spectrum Disorder Diagnosis and in Their Younger Siblings

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**IMPORTANCE** In recent years, rates of vaccination have been declining. Whether this phenomenon disproportionately affects children with autism spectrum disorder (ASD) or their younger siblings is unknown.

**OBJECTIVES** To investigate if children after receiving an ASD diagnosis obtain their remaining scheduled vaccines according to the Advisory Committee on Immunization Practices (ACIP) recommendations and to compare the vaccination patterns of younger siblings of children with ASD with the vaccination patterns of younger siblings of children without ASD.

**DESIGN, SETTING, AND PARTICIPANTS** This investigation was a retrospective matched cohort study. The setting was 6 integrated health care delivery systems across the United States within the Vaccine Safety Datalink. Participants were children born between January 1, 1995, and September 30, 2010, and their younger siblings born between January 1, 1997, and September 30, 2014. The end of follow-up was September 30, 2015.

**EXPOSURES** Recommended childhood vaccines between ages 1 month and 12 years.

**MAIN OUTCOME AND MEASURE** The proportion of children who received all of their vaccine doses according to ACIP recommendations.

**RESULTS** The study included 3729 children with ASD (676 [18.1%] female), 592 907 children without ASD, and their respective younger siblings. Among children without ASD, 250 193 (42.2%) were female. For vaccines recommended between ages 4 and 6 years, children with ASD were significantly less likely to be fully vaccinated compared with children without ASD (adjusted rate ratio, 0.87; 95% CI, 0.85-0.88). Within each age category, vaccination rates were significantly lower among younger siblings of children with ASD compared with younger siblings of children without ASD. The adjusted rate ratios varied from 0.86 for siblings younger than 1 year to 0.96 for those 11 to 12 years old. Parents who had a child with ASD were more likely to refuse at least 1 recommended vaccine for that child's younger sibling and to limit the number of vaccines administered during the younger sibling's first year of life.

**CONCLUSIONS AND RELEVANCE** Children with ASD and their younger siblings were undervaccinated compared with the general population. The results of this study suggest that children with ASD and their younger siblings are at increased risk of vaccine-preventable diseases.

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Autism spectrum disorder (ASD) is a neurodevelopmental disorder characterized by deficits in communication and social interaction and the exhibiting of stereotyped behaviors, typically occurring before age 3 years. The etiology of ASD is unknown for the vast majority of cases; however, study findings suggest that both genetic<sup>1-3</sup> and environmental<sup>4-6</sup> factors have a role.

Despite numerous scientific studies<sup>7-15</sup> reporting no association between childhood vaccination and ASD, there remain concerns about such a connection for some of the public.<sup>16</sup> In recent years, rates of undervaccination and vaccine refusal have been on the rise in the United States<sup>17-21</sup> and have been associated with vaccine-preventable disease outbreaks.<sup>22-24</sup> Rates of undervaccination among the subpopulation of children with ASD have not been fully investigated. A survey conducted among 98 parents of children with ASD and 65 parents of children without ASD in Canada found that a lower proportion of children with ASD received their measles, mumps, and rubella (MMR) or diphtheria and tetanus toxoids and acellular pertussis and inactivated poliovirus (DTaP-IPV) vaccines compared with children without ASD.<sup>25</sup> Because the first dose of MMR and the first 3 doses of DTaP-IPV are recommended before the age when ASD can be reliably diagnosed (which is at least 2 years), it was not clear from that study if the lower observed vaccination rates among the children with ASD were a consequence of the child's ASD diagnosis. In a recent letter to the editor, Glickman and colleagues<sup>26</sup> reported no significant difference between rates of vaccination of 71 children with ASD and those of 135 children without ASD. However, they found that families with children with ASD were less likely to vaccinate subsequent children. Other studies<sup>25,27-30</sup> also found that parents of children with ASD were more likely to either delay or refuse vaccination for their younger children. In a survey of 197 parents, Bazzano and colleagues<sup>27</sup> found that half of the parents of children with ASD changed vaccination practices for their younger children because of beliefs that vaccines contributed to their child's ASD. After surveying 486 parents of children with ASD, Rosenberg and colleagues<sup>30</sup> found that almost 20% of parents declined or delayed MMR immunization in the younger siblings of children with ASD. Previous studies were limited by small samples, lack of comparable control groups, or restriction to specific vaccines.

The objectives of this study were 2-fold. First, we investigated if children after receiving an ASD diagnosis obtain all of their remaining scheduled vaccines according to the Advisory Committee on Immunization Practices (ACIP)<sup>31</sup> recommendations. Second, we assessed whether younger siblings of children with ASD receive all recommended vaccines on time compared with younger siblings of children without ASD.

## Methods

### Study Population

The study population included children born between January 1, 1995, and September 30, 2010, and their younger siblings born between January 1, 1997, and September 30, 2014, who were

### Key Points

**Question** After receiving an autism spectrum disorder diagnosis, do children obtain all of their remaining scheduled vaccines, and are the younger siblings of these children vaccinated according to vaccine recommendations?

**Findings** In a matched cohort study of 3729 children with autism spectrum disorder and 592 907 children without autism spectrum disorder, we found that children with autism spectrum disorder were less likely to be fully vaccinated for vaccines recommended between ages 4 and 6 years. The younger siblings were also less likely to be fully vaccinated for vaccines recommended at any age.

**Meaning** Children with autism spectrum disorder and their younger siblings are at increased risk of vaccine-preventable diseases.

members of integrated health care delivery systems (sites) within the Vaccine Safety Datalink (VSD).<sup>32</sup> The VSD is a collaborative project between the Centers for Disease Control and Prevention and 8 sites across the United States and captures comprehensive medical and immunization data for more than 10 million people annually. This study included data from the following 6 VSD sites: Kaiser Permanente Northern California, Kaiser Permanente Colorado, Kaiser Permanente Northwest, Kaiser Permanente Washington, Marshfield Clinic, and Kaiser Permanente Southern California. The study was approved by the institutional review board at each participating VSD site and the Centers for Disease Control and Prevention. Written informed consent was waived by each institutional review board because the study had no direct contact with patients.

### Study Design

In this retrospective matched cohort study, we compared the proportion of vaccination between children with ASD and those without ASD. We also compared the proportion of vaccination of the younger siblings of children with ASD with those of the younger siblings of children without ASD.

### Autism Spectrum Disorder

We defined ASD based on the presence of *International Classification of Diseases, Ninth Revision, Clinical Modification (ICD-9-CM)* codes 299.0, 299.8, and 299.9 in electronic health records on at least 2 occasions from birth until either the sixth birthday or until the end of follow-up (September 30, 2015), whichever was earlier. If the first diagnosis appeared before age 2 years, we required that the second diagnosis be assigned at 2 years or older. A prior medical record review study<sup>33</sup> demonstrated that identifying ASD using at least 2 diagnosis codes predicts valid ASD cases. We matched children without ASD to children with ASD on month and year of birth, sex, and VSD site. Younger siblings of children without ASD were matched to younger siblings of children with ASD on month and year of birth, sex, and VSD site. It is possible that multiple siblings of an individual with or without ASD were included because we did not limit the number of children per family.

### Vaccination Status

To assess the vaccination patterns of children after receiving an ASD diagnosis, we only included children who were at least

7 years old as of September 30, 2015, and in whom ASD was diagnosed at 5 years or younger. We limited this assessment to vaccines routinely recommended between ages 4 and 6 years and ages 11 and 12 years. For the comparison of vaccination patterns of younger siblings of children with ASD and younger siblings of children without ASD, we included children (siblings) who were at least 1 year old as of September 30, 2015, and assessed vaccines routinely recommended by the ACIP at ages 1 to 11 months (vaccine doses given at birth were not assessed), 1 to 2 years, 4 to 6 years, and 11 to 12 years. Children who received all of their vaccine doses within the ACIP-recommended age limits were considered fully vaccinated. For example, we considered a child who received at least 1 dose of DTaP, at least 1 dose of MMR, at least 1 dose of IPV, and at least 1 dose of varicella vaccine between ages 4 and 7 years as fully vaccinated for vaccines recommended at ages 4 to 6 years (Table 1) regardless of vaccination history before age 4 years. We required that children be health plan members during the periods the vaccines were recommended (eg, to assess vaccination status at ages 1-2 years, we required that the children be health plan members between ages 1 and 2 years). For the younger siblings, we assessed vaccination status regardless of ASD diagnosis. Vaccine refusal was identified using ICD-9-CM codes V64.05 and V64.06.

### Statistical Analysis

In our primary analysis, we investigated and compared the proportion of fully vaccinated children within each age category for children with ASD, younger siblings of children with ASD, and their respective matched controls. In secondary analyses, we also assessed the proportion of vaccinated children for each individual vaccine or vaccine series for children with ASD, younger siblings of children with ASD, and their respective matched controls. For the younger siblings of children with and without ASD, we compared the proportions of children who received fewer than the recommended vaccinations at each well-child visit (no more than 2 shots per visit or shot limiting in the child's first year of life), as well as the proportion of parents (based on ICD-9-CM codes) who refused to vaccinate their children. We calculated crude and adjusted rate ratios (RRs) using log binomial regression analysis. Because full vaccination is more common than undervaccination, the odds ratio for fully vaccinated would not be a good estimation of the RR. Therefore, we estimated the RRs by using proc GENMOD in SAS (version 9.3; SAS Institute Inc), with the log function as the link with binomial distribution. In multivariable analyses, we included maternal age categories at the child's time of birth ( $\leq 20$ , 21-29, 30-39, or  $\geq 40$  years), maternal self-reported race/ethnicity (Asian, black, white, Hawaiian, Hispanic, Native American, or other), child's sex, and month and year of birth. We used SAS version 9.3 to conduct all analyses. All *P* values were 2 sided, and *P* < .05 was considered statistically significant.

## Results

The study included 3729 children with ASD (676 [18.1%] female), 592 907 children without ASD, and their respective

Table 1. On-Time Recommended Vaccine Doses by Age Group<sup>a</sup>

Vaccine	No. of Doses by Age Group			
	1-11 mo	1-2 y	4-6 y	11-12 y
DTaP	$\geq 3$	$\geq 1$	$\geq 1$	NA
HBV <sup>b</sup>	$\geq 2$	NA	NA	NA
Hib	$\geq 2^c$	$\geq 1$	NA	NA
HPV	NA	NA	NA	$\geq 3$
IPV	$\geq 3$	NA	$\geq 1$	NA
MCV4	NA	NA	NA	$\geq 1$
MMR	NA	$\geq 1$	$\geq 1$	NA
PCV	$\geq 3$	$\geq 1$	NA	NA
Rotavirus	$\geq 2^c$	NA	NA	NA
Tdap	NA	NA	NA	$\geq 1$
VAR	NA	$\geq 1$	$\geq 1$	NA

Abbreviations: DTaP, diphtheria and tetanus toxoids and acellular pertussis; HBV, hepatitis B virus; Hib, *Haemophilus influenzae* type b; HPV, human papillomavirus; IPV, inactivated poliovirus; MCV4, quadrivalent meningococcal conjugate; MMR, measles, mumps, and rubella; NA, not applicable; PCV, pneumococcal conjugate vaccine; Tdap, tetanus, diphtheria, and acellular pertussis; VAR, varicella vaccine.

<sup>a</sup> The Advisory Committee on Immunization Practices recommends seasonal influenza vaccination annually for all persons 6 months or older. Seasonal influenza vaccination was not included in this assessment because these vaccines occur in a defined time frame and may be administered outside of well-child visits or in nonmedical settings.

<sup>b</sup> Limited to 2 or more doses instead of 3 or more doses because the birth dose was not included in the study.

<sup>c</sup> The Advisory Committee on Immunization Practices recommends 2 or more doses or 3 or more doses in this age range depending on which vaccine is used.

younger siblings. Among the children without ASD, 250 193 (42.2%) were female.

### Vaccination Patterns in Children With and Without ASD

For vaccines recommended between ages 4 and 6 years, our analysis included 2855 children with ASD diagnosed by age 5 years matched to 483 961 children without ASD. The proportion of children who received all recommended vaccine doses (Table 1) between ages 4 and 6 years was lower in children with ASD compared with children without ASD (81.6% [2331 of 2855] vs 94.1% [455 435 of 483 961], respectively) (Table 2). The proportion receiving each individual vaccine was also lower among children with ASD compared with children without ASD. For MMR vaccine, 84.0% (2397 of 2855) of those aged 4 to 6 years with ASD were vaccinated compared with 95.9% (464 245 of 483 961) of those without ASD. After adjusting for maternal age at the time of the child's birth and race/ethnicity (which were both associated with ASD in our bivariate analyses) and the matching variables (month and year of birth, sex, and site), children with ASD were significantly less likely to be fully vaccinated (adjusted RR, 0.87; 95% CI, 0.85-0.88) compared with children without ASD. Adjusted RRs were also significant for each individual vaccine.

For vaccines recommended at ages 11 to 12 years, the analysis included 874 children with ASD matched to 218 181 children without ASD. In this age group, the proportions receiving all vaccines (Table 1) and each individual vaccine were similar between children with ASD and children without ASD, and adjusted RRs were not significant (Table 2).

Table 2. On-Time Vaccination Rates and Adjusted RRs Comparing Vaccination Among Children With ASD vs Children Without ASD by Age Group<sup>a</sup>

Vaccine	Recommended Vaccine Doses	No. (%)		Adjusted RR (95% CI)
		ASD	Non-ASD	
<b>Ages 4-6 y</b>				
ASD status, No.		2855	483 961	
DTaP	≥1	2543 (89.1)	467 324 (96.6)	0.92 (0.91-0.93)
IPV	≥1	2497 (87.5)	464 191 (95.9)	0.91 (0.89-0.92)
MMR	≥1	2397 (84.0)	464 245 (95.9)	0.87 (0.86-0.89)
VAR	≥1	2750 (96.3)	478 583 (98.9)	0.97 (0.96-0.98)
Fully vaccinated	All of the above	2331 (81.6)	455 435 (94.1)	0.87 (0.85-0.88)
<b>Ages 11-12 y</b>				
ASD status, No		874	281 181	
HPV	≥3	749 (85.7)	182 119 (83.5)	1.03 (1.00-1.06)
MCV4	≥1	839 (96.0)	212 582 (97.4)	0.98 (0.97-1.00)
Tdap	≥1	795 (91.0)	202 568 (92.8)	0.97 (0.96-1.00)
Fully vaccinated	All of the above	677 (77.5)	167 813 (76.9)	1.00 (0.97-1.04)

Abbreviations: ASD, autism spectrum disorder; DTaP, diphtheria and tetanus toxoids and acellular pertussis; HPV, human papillomavirus; IPV, inactivated poliovirus; MCV4, quadrivalent meningococcal conjugate; MMR, measles, mumps, and rubella; NA, not applicable; RR, rate ratio adjusted for maternal age, maternal race/ethnicity, child's sex, and month and year of birth; Tdap, tetanus, diphtheria, and acellular pertussis; VAR, varicella vaccine.

<sup>a</sup> The Advisory Committee on Immunization Practices recommends seasonal influenza vaccination annually for all persons 6 months or older. Seasonal influenza vaccination was not included in this assessment because these vaccines occur in a defined time frame and may be administered outside of well-child visits or in nonmedical settings.

### Vaccination Patterns in Younger Siblings of Children With and Without ASD

Within each age group, the proportion of children who were fully vaccinated with the recommended vaccines was lower among younger siblings of children with ASD compared with younger siblings of children without ASD. The difference in proportion of fully vaccinated children was greatest in the group aged 1 to 11 months (Table 3). The proportion of children who received each individual vaccine was lower for younger siblings of children with ASD compared with younger siblings of those without ASD within each age group. After adjusting for covariates, younger siblings of children with ASD were significantly less likely to be fully vaccinated than were younger siblings of children without ASD within each age group, except for vaccines recommended between ages 11 and 12 years. Rate ratios of undervaccination comparing younger siblings of children with ASD and those without ASD were lowest in the groups aged 1 to 11 months and 1 to 2 years.

A higher proportion of parents of children with ASD refused to vaccinate their younger children compared with parents of children without ASD (Figure). These parents were also more likely to limit the number of vaccines administered during well-child visits in the group aged 1 to 11 months (73 of 881 [8.3%] for younger siblings of children with ASD vs 2789 of 189 144 [1.5%] among younger siblings of children without ASD,  $P < .001$ ). For both ASD cases and their siblings, we found no significant differences in the RR of undervaccination over time (eTable in the Supplement).

## Discussion

In this large multisite study, we found that vaccine uptake was high overall. However, after receiving an ASD diagnosis, children

with ASD were subsequently less likely to be vaccinated compared with children without ASD matched on age, sex, and site. We also found that vaccination rates were lower among younger siblings of children with ASD compared with younger siblings of children without ASD. Parents of children with ASD were more likely to refuse vaccinating the children's younger siblings compared with parents of children without ASD. This phenomenon was not observed for vaccines recommended at ages 11 to 12 years for children with ASD and their younger siblings.

Our results are similar to those of a Canadian study,<sup>25</sup> which reported that children with ASD and their younger siblings were undervaccinated for MMR and pertussis-containing vaccines compared with children without ASD. However, there are major differences between our study and the Canadian study, including our larger sample size and more targeted study design. In our comparison of vaccination status between children with ASD and those without ASD, we only assessed vaccines recommended after the child's ASD diagnosis, which enables the inference that the lower vaccination rate in children with ASD was at least in part owing to the ASD diagnosis.

In the present study, within each recommended age category of vaccination before age 10 years, younger siblings of children with ASD were significantly more likely to be undervaccinated compared with younger siblings of children without ASD, suggesting that the ASD diagnosis of the older sibling may have contributed to the undervaccination of the younger children. Knowing that younger siblings of children with ASD are at higher risk for ASD<sup>34-37</sup> may have led some parents of children with ASD to either delay or refuse vaccinations for the younger siblings. It is also possible that health care professionals may have been more likely to assign the code for vaccine refusal for the siblings of children with ASD when they know that the family has a child with ASD. Parents of children with ASD may also delay or alter the vaccination schedule of



**Table 3. On-Time Vaccination Rates and Adjusted RRs Comparing Vaccination Among Younger Siblings of Children With ASD vs Younger Siblings of Children Without ASD by Age Group<sup>a</sup>**

Vaccine	Recommended Vaccine Doses	No. (%)		Adjusted RR (95% CI)
		Siblings of ASD	Siblings of non-ASD	
<b>Ages 1-11 mo</b>				
Sibling status, No.	NA	881	189 144	NA
DTaP	≥3	745 (84.6)	175 414 (92.7)	0.91 (0.88-0.94)
HBV <sup>b</sup>	≥2	751 (85.2)	182 907 (96.7)	0.88 (0.86-0.91)
Hib	≥2	791 (89.8)	182 807 (96.6)	0.93 (0.91-0.95)
IPV	≥3	708 (80.4)	172 878 (91.4)	0.88 (0.85-0.91)
PCV	≥3	NA	NA	NA
Rotavirus	≥2 <sup>c</sup>	741 (84.1)	173 680 (91.8)	0.91 (0.89-0.94)
Fully vaccinated	All of the above	645 (73.2)	160 773 (85.0)	0.86 (0.82-0.89)
<b>Ages 1-2 y</b>				
Sibling status, No.	NA	816	196 309	NA
DTaP	≥1	696 (85.3)	182 926 (93.2)	0.91 (0.88-0.94)
Hib	≥1	718 (88.0)	185 005 (94.2)	0.93 (0.91-0.95)
MMR	≥1	626 (76.7)	183 544 (93.5)	0.82 (0.79-0.85)
PCV	≥1	685 (83.9)	179 804 (91.6)	0.91 (0.88-0.94)
VAR	≥1	664 (81.4)	182 514 (93.0)	0.87 (0.85-0.90)
Fully vaccinated	All of the above	491 (60.2)	138 751 (70.7)	0.84 (0.79-0.89)
<b>Ages 4-6 y</b>				
Sibling status, No.	NA	315	86 878	NA
DTaP	≥1	288 (91.4)	84 163 (96.9)	0.94 (0.91-0.97)
IPV	≥1	281 (89.2)	83 781 (96.4)	0.92 (0.89-0.96)
MMR	≥1	270 (85.7)	83 499 (96.1)	0.89 (0.85-0.93)
VAR	≥1	290 (92.1)	84 884 (97.7)	0.94 (0.91-0.97)
Fully vaccinated	All of the above	262 (83.2)	82 157 (94.6)	0.88 (0.84-0.92)
<b>Ages 11-12 y</b>				
Sibling status, No.	NA	19	3170	NA
HPV	≥3	12 (63.2)	2170 (68.5)	0.93 (0.66-1.30)
MCV4	≥1	18 (94.7)	2962 (93.4)	1.01 (0.90-1.13)
Tdap	≥1	18 (94.7)	3084 (97.3)	0.98 (0.87-1.09)
Fully vaccinated	All of the above	12 (63.2)	2077 (65.5)	0.96 (0.68-1.36)

Abbreviations: ASD, autism spectrum disorder; DTaP, diphtheria and tetanus toxoids and acellular pertussis; HBV, hepatitis B virus; Hib, *Haemophilus influenzae* type b; HPV, human papillomavirus; IPV, inactivated poliovirus; MCV4, quadrivalent meningococcal conjugate; MMR, measles, mumps, and rubella; NA, not applicable; PCV, pneumococcal conjugate vaccine; RR, rate ratio adjusted for maternal age, maternal race/ethnicity, child's sex, and month and year of birth; Tdap, tetanus, diphtheria, and acellular pertussis; VAR, varicella vaccine.

<sup>a</sup> The Advisory Committee on Immunization Practices recommends seasonal influenza vaccination annually for all persons 6 months or older. Seasonal influenza vaccination was not included in this assessment because these vaccines occur in a defined time frame and may be administered outside of well-child visits or in nonmedical settings.

<sup>b</sup> Limited to 2 or more doses instead of 3 or more doses because the birth dose was not included in this study.

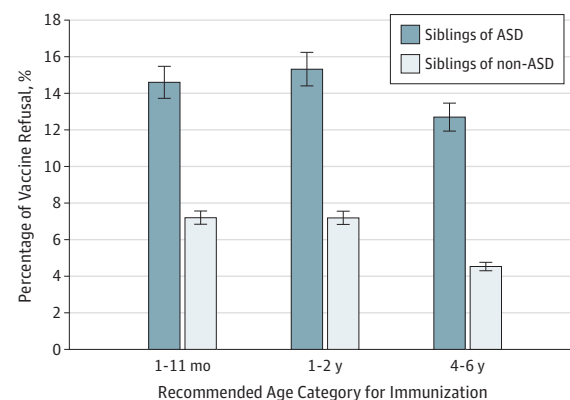
<sup>c</sup> The Advisory Committee on Immunization Practices recommends 2 or more doses or 3 or more doses in this age range depending on which vaccine is used.

their younger children because of concerns that vaccines may have had a role in causing the ASD of the older siblings<sup>27,30</sup>; this is despite considerable scientific evidence that vaccines do not cause autism. As previously reported, most parents vaccinate their children according to the ACIP-recommended schedule.<sup>18</sup> However, an increasing number of parents, especially parents of children with ASD, also appear to limit the number of vaccines their children receive during their child's first year of life; the safety of such alternative vaccination schedules is unknown, but this practice increases the chances for contracting a vaccine-preventable disease.

The highest rates of undervaccination in this study were among siblings of children with ASD who were in the groups aged 1 to 11 months and 1 to 2 years. This suggests that some parents consider the potential risks of ASD associated with vaccination to be greatest at these younger ages at which more vaccines are recommended. However, as these children grow older, these parents may be more willing to vaccinate.

### Limitations and Strengths

Our study has some limitations. Autism spectrum disorder status was determined using specific diagnostic codes from automated data, and medical record reviews were not con-

**Figure. Parental Vaccine Refusal of Any Vaccine Dose for Younger Siblings by Age Category and by Child Autism Spectrum Disorder (ASD) Status of Older Siblings**

Vaccine refusal was identified based on *International Classification of Diseases, Ninth Revision, Clinical Modification* codes. The error bars represent 95% CIs.

ducted. However, our case definition, which required at least 2 codes for ASD on different days, has been shown to identify true cases of ASD with high accuracy.<sup>33</sup> Although the overall

sample size of the study was large, the analysis comparing vaccination status of the younger siblings for the group aged 11 to 12 years was small, which limited the validity and power to observe differences in vaccine patterns in this group. We required that children in the study be a health plan member during the periods we assessed their vaccination status; while it is possible that some children may have received some of their vaccines outside of the VSD sites, this is considered unlikely because vaccines are offered free of charge at all of the participating sites. For missing vaccine doses to affect our results, the data missing would have to be differential for families with and without children with ASD. Our rates of under-vaccination among children with ASD may reflect an under-estimation of the true rates because we did not assess vaccination status before the ASD diagnosis. We did not conduct medical record reviews to validate the codes for vaccine refusal. Rates of vaccine refusal in this study likely underestimate the true rates of vaccine refusal because vaccine refusal is not always coded in the medical record consistently by physicians. Our rates could be biased toward or away from the null value because we cannot determine if ASD status is always associated with better or worse documentation of vaccine refusal. For some vaccines, different numbers of doses are recommended depending on which vaccine is used; we did not examine specific vaccine formulations. We were not able to assess the birth dose of hepatitis B vaccine because not all children included in our study were born at the health care organizations included in the study. Finally, we cannot attribute all of the under-vaccination findings for the younger siblings of children with ASD to the ASD diagnosis of the older sibling because it is possible that some parents may have modified their younger children's vaccine schedule without knowing the ASD status of the older children or for other reasons, including health care professional recommendations or some unknown factors.

This study's strengths include our large racially/ethnically and socioeconomically diverse population<sup>38</sup> from 6 different geographic areas, suggesting that the findings may be broadly generalizable to other populations. We also used vaccination data validated from the medical record instead of a parental report, which could be subject to recall bias or incomplete information. Furthermore, we had extensive vaccination data over many years and were thus able to assess the vaccination rate for recommended childhood vaccines between ages 1 month and 12 years for the younger siblings of children with ASD. In addition, we were able to match a large comparison group with the children with ASD and identify the younger siblings of both children with and without ASD. By matching on month and year of birth, the study minimized the possibility of a difference in the interpregnancy interval between cases and controls or between siblings of cases and siblings of controls.

## Conclusions

This large multisite study found that children with ASD and their younger siblings were under-vaccinated compared with the general population, suggesting that they are at increased risk of vaccine-preventable diseases. Although we do not know all factors contributing to under-vaccination among children with ASD, the results of our study suggest that parental vaccine refusal could have a role. Previous studies reported that a large proportion of parents of children with ASD consider that vaccines contributed to their child's ASD, and consequently they either changed or discontinued vaccination, suggesting that current strategies to address vaccine hesitancy have not been effective for parents of children with ASD. New strategies, including establishing or promoting a better dialogue among parents, health care professionals, and public health authorities, may be needed to increase vaccine uptake in populations with low uptake.

### ARTICLE INFORMATION

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**Study concept and design:** Zerbo, Goddard, Lewis, Daley, Irving, Jackson, DeStefano, Klein.

**Acquisition, analysis, or interpretation of data:** All authors.

**Drafting of the manuscript:** Zerbo.

**Critical revision of the manuscript for important intellectual content:** All authors.

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