

It seems that unvaccinated children are measurably healthier than vaccinated kids on nine different measures of health.

Whether we're talking about nasal allergies, learning disabilities, or pneumonia, the unvaccinated kids have lower rates of illness.

This study's findings are especially important when we consider that America's children are sicker than ever before, and nobody really knows why.

43% of American kids now suffer from some sort of chronic illness, NOT including obesity.

The study's findings really make you wonder. Here's some of the specifics:

- Vaccinated children were more than three times as likely to be diagnosed on the Autism Spectrum (OR 4.3)
- Vaccinated children were 30-fold more likely to be diagnosed with allergic rhinitis (hay fever) than non-vaccinated children
- Vaccinated children were 22-fold more likely to require an allergy medication than unvaccinated children
- Vaccinated children had more than quadruple the risk of being diagnosed with a learning disability than unvaccinated children (OR 5.2)
- Vaccinated children were 300 percent more likely to be diagnosed with Attention Deficit Hyperactivity Disorder than unvaccinated children (OR 4.3)
- Vaccinated children were 340 percent (OR 4.4) more likely to have been diagnosed with pneumonia than unvaccinated children
- Vaccinated children were 300 percent more likely to be diagnosed with an ear infection than unvaccinated children (OR 4.0)
- Vaccinated children were 700 percent more likely to have surgery to insert ear drainage tubes than unvaccinated children (OR 8.01)
- Vaccinated children were 2.5-fold more likely to be diagnosed with any chronic illness than unvaccinated children

<https://thehomestead.guru/unvaccinated-kids-healthier/>

## Vaccination and Health Outcomes: A Survey of 6- to 12-year-old Vaccinated and Unvaccinated Children based on Mothers' Reports

Anthony R. Mawson<sup>1\*</sup>, Brian D. Ray<sup>2</sup>, Azad R. Bhuiyan<sup>3</sup>, Binu Jacob<sup>4</sup>

<sup>1</sup>Epidemiology and Biostatistics, School of Public Health (Initiative), Jackson State University, USA, <sup>2</sup>National Home Education Research Institute, USA, <sup>3</sup>Epidemiology and Biostatistics, School of Public Health (Initiative), USA, <sup>4</sup>Former Graduate Student, Jackson State University, School of Public Health (Initiative), USA

**Submitted to Journal:**

Frontiers in Public Health

**Specialty Section:**

Child Health and Human Development

**ISSN:**

2296-2565

**Article type:**

Original Research Article

**Received on:**

17 Sep 2016

**Accepted on:**

21 Nov 2016

**Provisional PDF published on:**

21 Nov 2016

**Frontiers website link:**

[www.frontiersin.org](http://www.frontiersin.org)

**Citation:**

Mawson AR, Ray BD, Bhuiyan AR and Jacob B(2016) Vaccination and Health Outcomes: A Survey of 6- to 12-year-old Vaccinated and Unvaccinated Children based on Mothers' Reports. *Front. Public Health* 4:270. doi:10.3389/fpubh.2016.00270

**Copyright statement:**

© 2016 Mawson, Ray, Bhuiyan and Jacob. This is an open-access article distributed under the terms of the [Creative Commons Attribution License \(CC BY\)](https://creativecommons.org/licenses/by/4.0/). The use, distribution and reproduction in other forums is permitted, provided the original author(s) or licensor are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.

Provisional

# **Vaccination and Health Outcomes: A Survey of 6- to 12-year-old Vaccinated and Unvaccinated Children based on Mothers' Reports**

**Anthony R. Mawson<sup>1\*</sup>, Brian D. Ray<sup>2</sup>, Azad R. Bhuiyan<sup>3</sup>, Binu Jacob<sup>4</sup>**

<sup>1</sup>Epidemiology and Biostatistics, School of Public Health (Initiative), Jackson State University, USA

<sup>2</sup>National Home Education Research Institute, USA

<sup>3</sup>Epidemiology and Biostatistics, School of Public Health (Initiative), Jackson State University, USA

<sup>4</sup>Former Graduate Student, School of Public Health (Initiative), Jackson State University, USA

Submitted to Journal:  
Frontiers in Public Health

Specialty Section:  
Child Health and Human Development

Article Type:  
Original Research Article

Manuscript ID:  
231518

Received on:  
17 Sep 2016

Frontiers Website Link:  
[www.frontiersin.org](http://www.frontiersin.org)

### ***Conflict of interest statement***

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest

### ***Author contribution statement***

#### **Author Contributions**

AM designed the study, contributed to data analysis and interpretation, and drafted the paper. BR designed the study, contributed to data collection, and edited the paper. AB contributed to data analyses and edited the paper. BJ contributed to data analyses and editing. All authors read and approved the final version of the paper.

### ***Keywords***

Acute diseases; Chronic diseases; Epidemiology; Evaluation; Health policy; Immunization; Neurodevelopmental disorders; Vaccination, Acute diseases, chronic diseases, Epidemiology, Evaluation, Health Policy, Immunization, Neurodevelopmental disorders

### ***Abstract***

Word count: 343

#### **ABSTRACT**

**Background:** Vaccinations have prevented millions of infectious illnesses, hospitalizations and deaths among US children. Yet the long-term health outcomes of the routine vaccination program remain unknown. Studies have been recommended by the Institute of Medicine to address this question.

**Specific Aims:** To compare vaccinated and unvaccinated children on a broad range of health outcomes, and to determine whether an association found between vaccination and neurodevelopmental disorders (NDD), if any, remains significant after adjustment for other measured factors.

**Design:** A cross-sectional survey of mothers of children educated at home.

**Methods:** Homeschool organizations in four states (Florida, Louisiana, Mississippi, and Oregon) were asked to forward an email to their members, requesting mothers to complete an anonymous online questionnaire on the vaccination status and health outcomes of their biological children 6 to 12 years of age.

**Results:** A total of 415 mothers provided data on 666 children, of whom 261 (39%) were unvaccinated. Vaccinated children were significantly less likely than the unvaccinated to have been diagnosed with chickenpox and pertussis, but significantly more likely to have been diagnosed with pneumonia, otitis media, allergies and NDDs (defined as Autism Spectrum Disorder, Attention Deficit Hyperactivity Disorder, and/or a learning disability). After adjustment, the factors that remained significantly associated with NDD were vaccination (OR 3.1, 95% CI: 1.4, 6.8), male gender (OR 2.3, 95% CI: 1.2, 4.3), and preterm birth (OR 5.0, 95% CI: 2.3, 11.6). In a final adjusted model, vaccination but not preterm birth remained associated with NDD, while the interaction of preterm birth and vaccination was associated with a 6.6-fold increased odds of NDD (95% CI: 2.8, 15.5).

**Conclusions:** In this study based on mothers' reports, the vaccinated had a higher rate of allergies and NDD than the unvaccinated. Vaccination, but not preterm birth, remained significantly associated with NDD after controlling for other factors. However, preterm birth combined with vaccination was associated with an apparent synergistic increase in the odds of NDD. Further research involving larger, independent samples is needed to verify and understand these unexpected findings in order to optimize the impact of vaccines on children's health.

### ***Funding statement***

#### **Funding Sources**

This study was supported by grants from Generation Rescue, Inc., and the Children's Medical Safety Research Institute, charitable organizations that support research on children's health and vaccine safety. The funders had no role or influence on the design and conduct of the research or the preparation of reports.

## **Ethics statement**

(Authors are required to state the ethical considerations of their study in the manuscript including for cases where the study was exempt from ethical approval procedures.)

*Did the study presented in the manuscript involve human or animal subjects:* Yes

*Please state the full name of the ethics committee that approved the study. If the study was exempt from this requirement please state the reason below.*

Institutional Review Board of Jackson State University

*Please detail the consent procedure used for human participants or for animal owners. If not applicable, please state this.* The study was designed as a cross-sectional survey of homeschooling mothers on their vaccinated and unvaccinated biological children ages 6 to 12. As contact information on homeschool families was unavailable, there was no defined population or sampling frame from which response rates could be determined. We proceeded by selecting 4 states (Florida, Louisiana, Mississippi, and Oregon) for the survey (Stage 1). The National Home Education Research Institute (NHERI) compiled a list of statewide and local homeschool organizations, totaling 84 in Florida, 18 in Louisiana, 12 in Mississippi and 17 in Oregon. Initial contacts were made in June 2012. NHERI contacted the leaders of each statewide organization by email to request their support. A second email was then sent, explaining the study purpose and background, which the leaders were asked to forward to their members (Stage 2). A link was provided to an online questionnaire in which no personally identifying information was requested. Families were contacted only indirectly through homeschool organizations. Biological mothers of children ages 6-12 years were asked to serve as respondents in order to standardize data collection and to include data on pregnancy-related factors and birth history that might relate to the children's current health. The age-range of 6 to 12 years was selected because most recommended vaccinations would have been received by then.

### **Recruitment and Informed Consent**

Homeschool leaders were asked to sign Memoranda of Agreement on behalf of their organizations and to provide the number of member families. Non-responders were sent a second notice but few provided the requested information. However, follow-up calls to the leaders suggested that all had contacted their members about the study. Both the letter to families and the survey questions were stated in a neutral way with respect to vaccines. Our letter to parents began:

"Dear Parent, This study concerns a major current health question: namely, whether vaccination is linked in any way to children's long-term health. Vaccination is one of the greatest discoveries in medicine, yet little is known about its long-term impact. The objective of this study is to evaluate the effects of vaccination by comparing vaccinated and unvaccinated children in terms of a number of major health outcomes ..."

Respondents were asked to indicate their consent to participate, to provide their home state and zip code of residence, and to confirm that they had biological children 6 to 12 years of age.

*Please detail any additional considerations of the study in cases where vulnerable populations were involved, for example minors, persons with disabilities or endangered animal species. If not applicable, please state this.*

Not applicable

# Vaccination and Health Outcomes: A Survey of 6- to 12-year-old Vaccinated and Unvaccinated Children based on Mothers' Reports

Anthony R. Mawson<sup>1\*</sup>, Brian D. Ray<sup>2</sup>, Azad R. Bhuiyan<sup>3</sup>, Binu Jacob<sup>4</sup>

<sup>1</sup>Professor, Department of Epidemiology and Biostatistics, School of Public Health (Initiative), Jackson State University, Jackson, MS 39213, USA

<sup>2</sup>President, National Home Education Research Institute, PO Box 13939, Salem, OR 97309, USA

<sup>3</sup>Associate Professor, School of Public Health (Initiative), Jackson State University, Jackson, MS 39213, USA

<sup>4</sup>Former graduate student, School of Public Health (Initiative), Jackson State University, Jackson, MS 39213, USA

## \*Correspondence:

Anthony R. Mawson

[Anthony.r.mawson@jsums.edu](mailto:Anthony.r.mawson@jsums.edu)

Number of words: 5,589; Number of figures and tables: 9

**Key Words:** Acute diseases; Chronic diseases; Epidemiology; Evaluation; Health policy; Immunization; Neurodevelopmental disorders; Vaccination

## ABSTRACT

**Background:** Vaccinations have prevented millions of infectious illnesses, hospitalizations and deaths among US children. Yet the long-term health outcomes of the routine vaccination program remain unknown. Studies have been recommended by the National Academy of Medicine to address this question.

**Specific Aims:** To compare vaccinated and unvaccinated children on a broad range of health outcomes, and to determine whether an association found between vaccination and neurodevelopmental disorders (NDD), if any, remains significant after adjustment for other measured factors.

**Design:** A cross-sectional survey of mothers of children educated at home.

**Methods:** Homeschool organizations in four states (Florida, Louisiana, Mississippi, and Oregon) were asked to forward an email to their members, requesting mothers to complete an anonymous online questionnaire on the vaccination status and health outcomes of their biological children ages 6 to 12 years.

**Results:** A total of 415 mothers provided data on 666 children, of which 261 (39%) were unvaccinated. Vaccinated children were significantly less likely than the unvaccinated to have been diagnosed with chickenpox and pertussis, but significantly more likely to have been

diagnosed with pneumonia, otitis media, allergies and NDDs (defined as Autism Spectrum Disorder, Attention Deficit Hyperactivity Disorder, and/or a learning disability). After adjustment, the factors that remained significantly associated with NDD were vaccination (OR 3.1, 95% CI: 1.4, 6.8), male gender (OR 2.3, 95% CI: 1.2, 4.3), and preterm birth (OR 5.0, 95% CI: 2.3, 11.6). In a final adjusted model, vaccination but not preterm birth remained associated with NDD, while the interaction of preterm birth and vaccination was associated with a 6.6-fold increased odds of NDD (95% CI: 2.8, 15.5).

**Conclusions:** In this study based on mothers' reports, the vaccinated had a higher rate of allergies and

NDD than the unvaccinated. Vaccination, but not preterm birth, remained significantly associated with

NDD after controlling for other factors. However, preterm birth combined with vaccination was associated with an apparent synergistic increase in the odds of NDD. Further research involving larger, independent samples is needed to verify and understand these unexpected findings in order to optimize the impact of vaccines on children's health.

## INTRODUCTION

Vaccines are among the greatest achievements of biomedical science and one of the most effective public health interventions of the 20th century (1). Among US children born between 1995 and 2013, vaccination is estimated to have prevented 322 million illnesses, 21 million hospitalizations and 732,000 premature deaths, with overall cost savings of \$1.38 trillion (2). About 95% of US children of kindergarten age receive all of the recommended vaccines as a requirement for school and daycare attendance (3, 4) and to prevent the occurrence and spread of infectious disease (5). Advances in biotechnology are contributing to the development of new vaccines for widespread use (6).

Under the currently recommended pediatric vaccination schedule (7), US children receive up to 48 doses of vaccines for 14 diseases from birth to age six years, a figure that has steadily increased since the 1950s, most notably since the Vaccines for Children program was created in 1994. The Vaccines for Children program began with vaccines targeting nine diseases: diphtheria, tetanus, pertussis, polio, *Haemophilus influenzae* type b disease, hepatitis B, measles, mumps, and rubella. Between 1995 and 2013, new vaccines against five other diseases were added for children age 6 and under: varicella, hepatitis A, pneumococcal disease, influenza, and rotavirus vaccine.

Although short-term immunologic and safety testing is performed on vaccines prior to their approval by the Food and Drug Administration, the long-term effects of individual vaccines and of the vaccination program itself remain unknown (8). Vaccines are acknowledged to carry risks of severe acute and chronic adverse effects, such as neurological complications and even death (9), but such risks are considered so rare that the vaccination program is believed to be safe and effective for virtually all children (10).

There are very few randomized trials on any existing vaccine recommended for children in terms of morbidity and mortality, in part because of ethical concerns. One exception, the high-titer measles vaccine, was withdrawn after several randomized trials in west Africa showed that it interacted with the diphtheria-tetanus-pertussis vaccine, resulting in a significant 33% increase in child mortality (11). Evidence of safety from observational



studies includes a limited number of vaccines, e.g., the measles, mumps and rubella vaccine, and hepatitis B vaccine, but none on the vaccination program itself. Knowledge is limited even for vaccines with a long record of safety and protection against contagious diseases (12). The safe levels and long-term effects of vaccine ingredients, such as adjuvants and preservatives, are also unknown (13). Other concerns include the safety and cost-effectiveness of newer vaccines against diseases that are potentially lethal for individuals but have a lesser impact on population health, such as the group B meningococcus vaccine (14).

Knowledge of adverse events following vaccinations is largely based on voluntary reports to the Vaccine Adverse Events Reporting System (VAERS) by physicians and parents. However, the rate of reporting of serious vaccine injuries is estimated to be <1% (15). These considerations led the former Institute of Medicine (now the National Academy of Medicine) in 2005 to recommend the development of a five-year plan for vaccine safety research by the Centers for Disease Control and Prevention (CDC) (16, 17). Yet in its 2011 and 2013 reviews of the adverse effects of vaccines, the Institute of Medicine concluded that few health problems are caused by or associated with vaccines, and found no evidence that the vaccination schedule was unsafe (18, 19). Another systematic review, commissioned by the US Agency for Healthcare Research and Quality to identify gaps in evidence on the safety of the childhood vaccination program, concluded that severe adverse events following vaccinations are extremely rare (20). The Institute of Medicine, however, noted that studies were needed to compare the health outcomes of vaccinated and unvaccinated children, to examine the long-term cumulative effects of vaccines, the timing of vaccination related to the age and condition of the child, the total load or number of vaccines given at one time, the effect of other vaccine ingredients in relation to health outcomes, and the mechanisms of vaccine-associated injury (19).

A complicating factor in evaluating the vaccination program is that vaccines against infectious diseases have complex nonspecific effects on morbidity and mortality that extend beyond prevention of the targeted disease. The existence of such effects challenges the assumption that individual vaccines affect the immune system independently of each other, and have no physiological effect other than protection against the targeted pathogen (21). The nonspecific effects of some vaccines are beneficial, while others appear to increase morbidity and mortality (22, 23). For instance, both the measles and Bacillus Calmette–Guérin vaccine reportedly reduce overall morbidity and mortality (24), whereas the diphtheria-tetanus-pertussis (25) and hepatitis B vaccine (26) have the opposite effect. The mechanisms responsible for these nonspecific effects are unknown but may involve interactions between vaccines and their ingredients, e.g., whether the vaccines are live or inactivated; the most recently administered vaccine; micronutrient supplements such as vitamin A; the sequence in which vaccines are given; and their possible combined and cumulative effects (21).

A major current controversy is the question of whether vaccination plays a role in neurodevelopmental disorders (NDDs), which broadly include learning disabilities, Attention Deficit Hyperactivity Disorder (ADHD) and Autism Spectrum Disorder (ASD). The controversy has been fueled by the fact that the US is experiencing what has been described as a “silent pandemic” of mostly subclinical developmental neurotoxicity, in which 16.6% (1 in 6) children suffers from a learning disability, sensory deficits, and developmental delays (27, 28). In 1996 the estimated prevalence of ASD was 0.42%. By 2010 it had risen to 1.47% (1 in 68), with 1 in 42 boys and 1 in 189 girls affected (29). More recently, based on a CDC

survey of parents in 2011–2014, 2.24% of children (1 in 45) were estimated to have ASD. Rates of other developmental disabilities, however, such as intellectual disability, cerebral palsy, hearing loss, and vision impairments, have declined or remained unchanged (30). Prevalence rates of Attention Deficit Hyperactivity Disorder (ADHD) have also risen markedly in recent decades (31). Earlier increases in the prevalence of learning disability have been followed by declining rates in most states, possibly due to changes in diagnostic criteria (32).

The marked increase in ASD and ADHD in recent decades suggests a role for an environmental factor to which virtually all children are exposed. Agricultural chemicals are currently a major focus of research (33-36). However, in the absence of data on vaccinated and unvaccinated children, the possible role of vaccines in NDD has remained unknown. The need to assess the outcomes of vaccination is supported by the fact that the Vaccine Injury Compensation Program has paid \$3.2 billion in compensation for vaccine injury since its creation in 1986 (37). A study of claims compensated by the Vaccine Injury Compensation Program for vaccine-induced encephalopathy and seizure disorder found 83 claims that were acknowledged as being due to brain damage. In all cases it was noted by the Court of Federal Claims, or indicated in settlement agreements, that the children had autism or ASD (38). On the other hand, numerous epidemiological studies have found no association between receipt of selected vaccines (in particular the combined measles, mumps, and rubella vaccine) and autism (10, 39-44), and there is no accepted mechanism by which vaccines could induce autism (45).

A major challenge in comparing vaccinated and unvaccinated children has been to identify an accessible pool of unvaccinated children, since the vast majority of children in the US are vaccinated.

Children educated at home (“homeschool children”) are suitable for such studies as a higher proportion are unvaccinated compared to public school children (46). Homeschool families have an approximately equal median income to that of married-couple families nationwide, somewhat more years of formal education, and a higher average family size (just over three children) compared to the national average of just over two children (47-49). Homeschooling families are slightly overrepresented in the south, about 23% are nonwhite, and the age distribution of homeschool children in grades K-12 is similar to that of children nationwide (50). About 3% of the school-age population was homeschooled in the 2011-2012 school year (51).

The aims of this study were 1) to compare vaccinated and unvaccinated children on a broad range of health outcomes, including acute and chronic conditions, medication and health service utilization, and 2) to determine whether an association found between vaccination and NDDs, if any, remained significant after adjustment for other measured factors.

## **METHODS**

### *Study Planning*

To implement the study, a partnership was formed with the National Home Education Research Institute (NHERI), which has been involved in educational research on homeschooling for many years and has strong and extensive contacts with homeschool organizations throughout the country

([www.nheri.org](http://www.nheri.org)). The study protocol was approved by the Institutional Review Board of Jackson State University.

### *Study Design*

The study was designed as a cross-sectional survey of homeschooling mothers on their vaccinated and unvaccinated biological children ages 6 to 12. As contact information on homeschool families was unavailable, there was no defined population or sampling frame from which response rates could be determined. We proceeded by selecting 4 states (Florida, Louisiana, Mississippi, and Oregon) for the survey (Stage 1). NHERI compiled a list of statewide and local homeschool organizations, totaling 84 in Florida, 18 in Louisiana, 12 in Mississippi and 17 in Oregon. Initial contacts were made in June 2012. NHERI contacted the leaders of each statewide organization by email to request their support. A second email was then sent, explaining the study purpose and background, which the leaders were asked to forward to their members (Stage 2). A link was provided to an online questionnaire in which no personally identifying information was requested. With funding limited to 12 months, we sought to obtain as many responses as possible, contacting families only indirectly through homeschool organizations. Biological mothers of children ages 6-12 years were asked to serve as respondents in order to standardize data collection and to include data on pregnancy-related factors and birth history that might relate to the children's current health. The age-range of 6 to 12 years was selected because most recommended vaccinations would have been received by then.

### *Recruitment and Informed Consent*

Homeschool leaders were asked to sign Memoranda of Agreement on behalf of their organizations and to provide the number of member families. Non-responders were sent a second notice but few provided the requested information. However, follow-up calls to the leaders suggested that all had contacted their members about the study. Both the letter to families and the survey questions were stated in a neutral way with respect to vaccines. Our letter to parents began:

*“Dear Parent, This study concerns a major current health question: namely, whether vaccination is linked in any way to children's long-term health. Vaccination is one of the greatest discoveries in medicine, yet little is known about its long-term impact. The objective of this study is to evaluate the effects of vaccination by comparing vaccinated and unvaccinated children in terms of a number of major health outcomes ...”*

Respondents were asked to indicate their consent to participate, to provide their home state and zip code of residence, and to confirm that they had biological children 6 to 12 years of age. The communications company Qualtrics (<http://qualtrics.com>) hosted the survey website. The questionnaire included only closed-ended questions requiring yes or no responses, with the aim of improving both response and completion rates.

A number of homeschool mothers volunteered to assist NHERI promote the study to their wide circles of homeschool contacts. A number of nationwide organizations also agreed to promote the study in the designated states. The online survey remained open for three months in the summer of 2012. Financial incentives to complete the survey were neither available nor offered.

### *Definitions and Measures*

Vaccination status was classified as unvaccinated (i.e., no previous vaccinations), partially vaccinated (received some but not all recommended vaccinations) and fully vaccinated (received all recommended age-appropriate vaccines), as reported by mothers. These categories were developed on the premise that any long-term effects of vaccines would be more evident in fully-vaccinated than in partially-vaccinated children, and rare or absent in the unvaccinated. Mothers were asked to use their child's vaccination records to indicate the recommended vaccines and doses their child had received. Dates of vaccinations were not requested, in order not to overburden respondents and to reduce the likelihood of inaccurate reporting; nor was information requested on adverse events related to vaccines, as this was not our purpose. We also did not ask about dates of diagnoses because chronic illnesses are often gradual in onset and made long after the appearance of symptoms. Since most vaccinations are given before age 6, vaccination would be expected to precede the recognition and diagnosis of most chronic conditions.

Mothers were asked to indicate on a list of more than 40 acute and chronic illnesses all those for which her child or children had received a diagnosis by a physician. Other questions included the use of health services and procedures, dental check-ups, "sick visits" to physicians, medications used, insertion of ventilation ear tubes, number of days in the hospital, the extent of physical activity (number of hours the child engaged in "vigorous" activities on a typical weekday), number of siblings, family structure (mother and father living in the home, divorced or separated), family income and/or highest level of education of mother or father, and social interaction with children outside the home (i.e., amount of time spent in play or other contact with children outside the household). Questions specifically for the mother included pregnancy-related conditions and birth history, use of medications during pregnancy, and exposure to an adverse environment (defined as living within 1-2 miles of a furniture manufacturing factory, hazardous waste site, or lumber processing factory). NDD, a derived diagnostic category, was defined as having one or more of the following three closely related and overlapping diagnoses: a learning disability, Attention Deficit Hyperactivity Disorder (ADHD) and Autism Spectrum Disorder (ASD) (52).

### *Statistical Methods*

Unadjusted bivariate analyses using chi-square tests were performed initially to test the null hypothesis of no association between vaccination status and health outcomes, i.e., physician-diagnosed acute and chronic illnesses, medications, and the use of health services. In most analyses, partially and fully vaccinated children were grouped together as the "vaccinated" group, with unvaccinated children as the control group. The second aim of the study was to determine whether any association found between vaccination and neurodevelopmental disorders remained significant after controlling for other measured factors. Descriptive statistics on all variables were computed to determine frequencies and percentages for categorical variables and means ( $\pm$  SD) for continuous variables. The strength of associations between vaccination status and health outcomes were tested using odds ratios (OR) and 95% Confidence Intervals (CI). Odds ratios describe the strength of the association between two categorical variables measured simultaneously and are appropriate measures of that relationship in a cross-sectional study (53). Unadjusted and adjusted logistic regression analyses were carried out using SAS (Version 9.3) to determine the factors associated with NDDs.

## RESULTS

### *Socio-Demographic Characteristics of Respondents*

The information contained in 415 questionnaires provided data on 666 homeschool children. Table 1 shows the characteristics of the survey respondents. Mothers averaged about 40 years of age, were typically white, college graduates, with household incomes between \$50,000 to \$100,000, Christian, and married. The reasons for homeschooling for the majority of respondents (80% - 86%) were for a moral environment, better family relationship, or for more contact with their child or children.

---

**Table 1. Characteristics of respondents**

---

The children as a group were similarly mostly white (88%), with a slight preponderance of females (52%), and averaged 9 years of age. With regard to vaccination status, 261 (39%) were unvaccinated, 208 (31%) were partially vaccinated, and 197 (30%) had received all of the recommended vaccinations. All statistical analyses are based on these numbers.

### *Acute Illness*

Vaccinated children (N=405), combining the partially and fully vaccinated, were significantly less likely than the unvaccinated to have had chickenpox (7.9% vs. 25.3%,  $p < 0.001$ ; Odds Ratio = 0.26, 95% Confidence Interval: 0.2, 0.4) and whooping cough (pertussis) (2.5% vs. 8.4%,  $p < 0.001$ ; OR = 0.3, 95% CI: 0.1, 0.6), and less likely, but not significantly so, to have had rubella (0.3% vs. 1.9%,  $p = 0.04$ ; OR = 0.1, 95% CI: 0.01, 1.1). However, the vaccinated were significantly more likely than the unvaccinated to have been diagnosed with otitis media (19.8% vs. 5.8%,  $p < 0.001$ ; OR = 3.8, 95% CI: 2.1, 6.6) and pneumonia (6.4% vs. 1.2%,  $p = 0.001$ ; OR = 5.9, 95% CI: 1.8, 19.7). No significant differences were seen with regard to Hepatitis A or B, high fever in the past 6 months, measles, mumps, meningitis (viral or bacterial), influenza, or rotavirus (Table 2).

---

**Table 2. Vaccination status and health outcomes – acute conditions**

---

### *Chronic Illness*

Vaccinated children were significantly more likely than the unvaccinated to have been diagnosed with the following: allergic rhinitis (10.4% vs. 0.4%,  $p < 0.001$ ; OR = 30.1, 95% CI: 4.1, 219.3), other allergies (22.2% vs. 6.9%,  $p < 0.001$ ; OR = 3.9, 95% CI: 2.3, 6.6), eczema/atopic dermatitis (9.5% vs. 3.6%,  $p = 0.035$ ; OR = 2.9, 95% CI: 1.4, 6.1), a learning disability (5.7% vs. 1.2%,  $p = 0.003$ ; OR = 5.2, 95% CI: 1.6, 17.4), ADHD (4.7% vs. 1.0%,  $p = 0.013$ ; OR = 4.2, 95% CI: 1.2, 14.5), ASD (4.7% vs. 1.0%,  $p = 0.013$ ; OR 4.2, 95% CI: 1.2, 14.5), any neurodevelopmental disorder (i.e., learning disability, ADHD or ASD) (10.5% vs. 3.1%,  $p < 0.001$ ; OR = 3.7, 95% CI: 1.7, 7.9) and any chronic illness (44.0% vs. 25.0%,  $p$



<0.001; OR = 2.4, 95% CI: 1.7, 3.3). No significant differences were observed with regard to cancer, chronic fatigue, conduct disorder, Crohn's disease, depression, Types 1 or 2 diabetes, encephalopathy, epilepsy, hearing loss, high blood pressure, inflammatory bowel disease, juvenile rheumatoid arthritis, obesity, seizures, Tourette's syndrome, or **services received** under the Individuals with Disabilities Education Act (Table 3).

---

**Table 3. Vaccination status and health outcomes – chronic conditions**

---

*Partial versus Full Vaccination*

Partially vaccinated children had an intermediate position **between** the fully vaccinated and unvaccinated in regard to several but not all health outcomes. For instance, as shown in Table 4, the partially vaccinated had an intermediate (apparently detrimental) position in terms of allergic rhinitis, ADHD, eczema, and learning disability.

---

**Table 4. Partial vs. full vaccination and health outcomes**

---

*Gender Differences in Chronic Illness*

Among the vaccinated (combining partially and fully vaccinated children), boys were more likely than girls to be diagnosed with a chronic condition – significantly so in the case of allergic rhinitis (13.9% vs. 7.2%,  $p = 0.03$ ; OR = 2.1, 95% CI: 1.1, 4.1), ASD (7.7% vs. 1.9%,  $p = 0.006$ ; OR = 4.3, 95% CI: 1.4, 13.2), and any neurodevelopmental disorder (14.4% vs. 6.7%,  $p = 0.01$ ; OR = 2.3, 95% CI: 1.2, 4.6) (Table 5).

---

**Table 5. Chronic conditions and gender among vaccinated children**

---

*Use of Medications and Health Services* The vaccinated (combining the partially and fully vaccinated) were significantly more likely than the unvaccinated to use medication for allergies (20.0% vs. 1.2%,  $p < 0.001$ ; OR = 21.5, 95% CI: 6.7, 68.9), to have used antibiotics in the past 12 months (30.8% vs. 15.4%,  $p < 0.001$ ; OR = 2.4, 95% CI: 1.6, 3.6), and to have used fever medications at least once (90.7% vs. 67.8%,  $p < 0.001$ ; OR = 4.6, 95% CI: 3.0, 7.1). The vaccinated were also more likely to have seen a doctor for a routine checkup in the past 12 months (57.6% vs. 37.2%,  $p < 0.001$ ; OR = 2.3, 95% CI: 1.7, 3.2), visited a dentist during the past year (89.4% vs. 80.5%,  $p < 0.001$ ; OR = 2.0, 95% CI: 1.3, 3.2), visited a doctor or clinic due to illness in the past year (36.0% vs. 16.0%,  $p < 0.001$ ; OR = 3.0, 95% CI:

2.0, 4.4), been fitted with ventilation ear tubes (3.0% vs. 0.4%,  $p = 0.018$ ; OR = 8.0, 95% CI: 1.0, 66.1), and spent one or more nights in a hospital (19.8% vs. 12.3%,  $p = 0.012$ ; OR = 1.8, 95% CI: 1.1, 2.7) (Table 6).

---

**Table 6. Vaccination status and the use of medications and health services**

---

### **Factors Associated with Neurodevelopmental Disorders**

The second aim of the study focused on a specific **health** outcome and was designed to determine whether vaccination was associated with neurodevelopmental disorders (NDD) and, if so, whether the association remained significant after adjustment for other measured factors. As noted, because of the relatively small numbers of children with specific diagnoses, NDD was a derived variable combining children with a diagnosis of one or more of ASD, ADHD and a learning disability.

The close association and overlap of these diagnoses in the study is shown in the figure below (Figure 1). The figure shows that the single largest group of diagnoses was learning disability ( $n=15$ ) followed by ASD ( $n=9$ ), and ADHD ( $n=9$ ), with smaller numbers comprising combinations of the three diagnoses.

---

**Figure 1. The overlap and distribution of physician-diagnosed neurodevelopmental disorders, based on mothers' reports**

---

### *Unadjusted Analysis*

Table 7 shows that the factors associated with NDD in unadjusted logistic regression analyses were as follows: vaccination (OR 3.7, 95% CI: 1.7, 7.9); male gender (OR 2.1, 95% CI: 1.1, 3.8); adverse environment, defined as living within 1-2 miles of a furniture manufacturing factory, hazardous waste site, or lumber processing factory (OR 2.9, 95% CI: 1.1, 7.4); maternal use of antibiotics during pregnancy (OR 2.3, 95% CI: 1.1, 4.8); and preterm birth (OR 4.9, 95% CI: 2.4, 10.3). Two factors that almost reached statistical significance were vaccination during pregnancy (OR 2.5, 95% CI: 1.0, 6.3) and three or more fetal ultrasounds (OR 3.2, 95% CI: 0.92, 11.5). Factors that were not associated with NDD in this study included mother's education, household income, and religious affiliation; use of acetaminophen, alcohol, and antacids during pregnancy; gestational diabetes; preeclampsia; Rhogham shot during pregnancy; and breastfeeding (data not shown).

---

**Table 7. Unadjusted analysis of potential risk factors for neurodevelopmental disorders**

---

### *Adjusted Analysis*

After adjustment for all other significant factors, those that remained significantly associated with NDD were: vaccination (OR 3.1, 95% CI: 1.4, 6.8); male gender (OR 2.3, 95% CI: 1.2, 4.3); and preterm birth (OR 5.0, 95% CI: 2.3, 11.6). The apparently strong association between both vaccination and preterm birth and NDD suggested the possibility of an interaction between these factors.

In a final adjusted model designed to test for this possibility, controlling for the interaction of preterm birth and vaccination, the following factors remained significantly associated with NDD: vaccination (2.5, 95% CI: 1.1, 5.6), nonwhite race (2.0, 95% CI: 1.1, 5.5), and male gender (OR 2.3, 95% CI: 1.2, 4.4). Preterm birth itself, however, was not significantly associated with NDD, whereas the combination (interaction) of preterm birth and vaccination was associated with 6.6-fold increased odds of NDD (95% CI: 2.8, 15.5) (Table 8).

---

**Table 8. Adjusted logistic regression analyses of risk factors and NDD**

---

## **DISCUSSION**

Following a recommendation of the Institute of Medicine (19) for studies to compare the health outcomes of vaccinated and unvaccinated children, this study focused on homeschool children ages 6 to 12 years based on mothers' anonymous reports in an online questionnaire. The respondents were mostly white, married and college-educated upper income women who had been contacted and invited to participate in the study by the leaders of their homeschool organizations. Data from the study were also used to determine whether vaccination was associated specifically with NDDs, a derived diagnostic category combining children with the diagnoses of learning disability, ASD and/or ADHD.

With regard to acute and chronic conditions, vaccinated children were significantly less likely than the unvaccinated to have had chickenpox and pertussis. However, contrary to expectation, the vaccinated were significantly more likely to have been diagnosed with otitis media, pneumonia, allergic rhinitis, eczema, ADHD, ASD and a learning disability. The vaccinated were also more likely to have used antibiotics, allergy and fever medications; to have been fitted with ventilation ear tubes; visited a doctor for a health issue in the previous year, and been hospitalized. Although the reason for hospitalization and the age of the child at the time were not determined, the latter finding appears consistent with a study of 38,801 reports to the VAERS of infants who were hospitalized or had died after receiving vaccinations. The study reported a linear relationship between the number of vaccine doses administered at one time and the rate of hospitalization and death; moreover, the younger the infant at the time of vaccination, the higher was the rate of hospitalization and death (54). The hospitalization rate increased from 11% for 2 vaccine doses to 23.5% for 8 doses ( $r^2 = 0.91$ ), while the case fatality rate increased significantly from 3.6% for those receiving from 1-4 doses to 5.4 % for those receiving from 5-8 doses.

In further support of the possibility that the number of vaccinations received could be implicated in risks of associated chronic illness, a comparison of unvaccinated, partially and



fully vaccinated children in the present study showed that the partially vaccinated had increased but intermediate odds of chronic disease, between those of unvaccinated and fully vaccinated children, specifically for allergic rhinitis, ADHD, eczema, a learning disability, and NDD as a whole.

The national rates of ADHD and LD are comparable to those of the study. The US rate of ADHD for ages 4-17 (twice the age range of children than the present study), is 11% (31). The study rate of ADHD for ages 6 to 12 is 3.3%, and 4.7% when only vaccinated children are included. The national LD rate is 5% (32), and the study data show a rate of LD of 3.9% for all groups, and 5.6% when only vaccinated children are included. However, the ASD prevalence of 2.24% from a CDC parent survey is lower than the study rate of 3.3%.

Vaccinated males were significantly more likely than vaccinated females to have been diagnosed with allergic rhinitis, ASD and NDD. The percentage of vaccinated males with an NDD in this study (14.4%) is consistent with national findings based on parental responses to survey questions, indicating that 15% of US children ages 3 to 17 years in the years 2006-2008 had an NDD (28). Boys are also more likely than girls to be diagnosed with an NDD, and ASD in particular (29).

Vaccination was strongly associated with both otitis media and pneumonia, which are among the most common complications of measles infection (55, 56). The odds of otitis media were almost four-fold higher among the vaccinated (OR 3.8, 95% CI: 2.1, 6.6) and the odds of myringotomy with tube placement were eight-fold higher than those of unvaccinated children (OR 8.0, 95% CI: 1.0, 66.1). Acute otitis media (AOM) is a very frequent childhood infection, accounting for up to 30 million physician visits each year in the US, and the most common reason for prescribing antibiotics for children (57, 58). The incidence of AOM peaks at ages 3 to 18 months and 80% of children have experienced at least one episode by 3 years of age. Rates of AOM have increased in recent decades (59). Worldwide, the incidence of AOM is 10.9%, with 709 million cases each year, 51% occurring in children under 5 years of age (60). Pediatric AOM is a significant concern in terms of healthcare utilization in the US, accounting for \$2.88 billion in annual health care costs (61).

Numerous reports of AOM have been filed with VAERS. A search of VAERS for “Cases where age is under 1 and onset interval is 0 or 1 or 2 or 3 or 4 or 5 or 6 or 7 days and Symptom is otitis media” (62) revealed that 438,573 cases were reported between 1990 and 2011, often with fever and other signs and symptoms of inflammation and central nervous system involvement. One study (63) assessed the nasopharyngeal carriage of *S. pneumoniae*, *H. influenzae*, and *M. catarrhalis* during AOM in fully immunized, partly immunized children with 0 or 1 dose of Pneumococcal Conjugate Vaccine-7 (PCV7), and “historical control” children from the pre-PCV-7 era, and found an increased frequency of *M. catarrhalis* colonization in the vaccinated group compared to the partly immunized and control groups (76% vs. 62% and 56%, respectively). A high rate of *M. catarrhalis* colonization is associated with an increased risk of AOM (64).

Successful vaccination against pneumococcal infections can lead to replacement of the latter in the nasopharyngeal niche by nonvaccine pneumococcal serotypes and disease (65). Vaccination with PCV-7 has a marked effect on the complete microbiota composition of the upper respiratory tract in children, going beyond shifts in the distribution of pneumococcal serotypes and known potential pathogens and resulting in increased anaerobes, gram-positive

bacteria and gram-negative bacterial species. PCV-7 administration also correlates highly with the emergence and expansion of oropharyngeal types of species. These observations have suggested that eradication of vaccine serotype pneumococci can be followed by colonization of other bacterial species in the vacant nasopharyngeal niche, leading to disequilibria of bacterial composition (dysbiosis) and increased risks of otitis media. Long-term monitoring has been recommended as essential for understanding the full implications of vaccination-induced changes in microbiota structure (66).

The second aim of the paper focused on a specific health outcome and sought to determine whether vaccination remained associated with neurodevelopmental disorders (NDD) after controlling for other measured factors. After adjustment, the factors that remained significantly associated with NDD were vaccination, nonwhite race, male gender, and preterm birth. The apparently strong association between both vaccination and preterm birth and NDD suggested the possibility of an interaction between these factors. This was shown in a final adjusted model with interaction (controlling for the interaction of preterm birth with vaccination). In this model, vaccination, nonwhite race and male gender remained associated with NDD, whereas preterm birth itself was no longer associated with NDD. However, the combination of preterm birth and vaccination was associated with significant 6.6-fold increased odds of NDD.

In summary, vaccination, nonwhite race, and male gender were significantly associated with NDD after controlling for other factors. Preterm birth, although significantly associated with NDD in unadjusted and adjusted analyses, was no longer associated with NDD in the final model with interaction. However, preterm birth combined with vaccination was a strong and synergistic factor for NDD in the final model, more than doubling the odds of NDD compared to vaccination alone. Preterm birth has long been known as a major factor for NDD (67, 68), but since preterm infants are routinely vaccinated, the separate effects of preterm birth and vaccination have not been examined. The present study suggests not only that vaccination may be a contributing factor in the pathogenesis of NDD but that preterm birth alone may have a lesser or even negligible role in NDD, defined here as ASD, ADHD and/or a learning disability. In addition, the results suggest that vaccination coupled with preterm birth may synergistically increase the odds of NDD beyond that of vaccination alone.

### *Strengths and Limitations*

Strengths of the study include the relatively large sample of unvaccinated children (N=261), which made it possible to compare health outcomes across the spectrum of vaccination coverage. The study group also comprised a demographically homogeneous sample of mainly white, higher-income and college-educated homeschooling families in which the children studied were 6 to 12 years of age. Information was provided anonymously by biological mothers who were obviously well-informed about their own children's health, which likely increased the validity of the reports. Recruitment of biological mothers as respondents made it possible to test hypotheses about the role of pregnancy- related factors and birth history as well as vaccination in specific conditions.

Several limitations of the study are acknowledged. Because of the absence of a defined sampling frame, neither the number of homeschool families contacted nor the survey response rate is known or can be validated. To minimize recall bias, respondents were asked to use their child's vaccination records. To enhance reliability, closed-ended questions were

used. To enhance validity, parents were asked to report only physician-diagnosed illnesses and about facts such as whether a child had been hospitalized or fitted with ventilation ear tubes.

Another potential limitation is that self-reported health outcomes were not validated by clinical records or reports. However, self-reports about significant events provide a valid proxy for official records when medical records and administrative data are unavailable (69). Nevertheless, it would be ideal to validate self-reports with official records, hopefully in a future study. A further potential limitation is under-ascertainment of disease in unvaccinated children, who may have seen less often by physicians, thereby reducing their chances of being diagnosed with a disease. Vaccinated children were more likely than the unvaccinated to have seen a doctor for a routine checkup in the past 12 months (57.6% vs. 37.2%,  $p = <0.001$ ; OR = 2.3, 95% CI: 1.7, 3.2). Such visits usually involve vaccinations, which non-vaccinating families would be expected to refuse. However, fewer visits to physicians would not necessarily mean that unvaccinated children are less likely to be seen by a physician if their condition warranted it. In fact, since unvaccinated children were more likely than the vaccinated to be diagnosed with chickenpox and pertussis, which would have involved a visit to the pediatrician, the observed differences in health outcomes are unlikely to be due to under-ascertainment.

## CONCLUSIONS

Assessment of the long-term effects of the current vaccination program on morbidity and mortality has been limited (70). In this study of the health outcomes of vaccinated and unvaccinated homeschool children, reduced odds of chickenpox and whooping cough were found among the vaccinated, as expected, but unexpectedly increased odds were found of otitis media, pneumonia, allergies, neurodevelopmental disorders, medication use, placement of ventilating ear tubes and hospitalizations. Although the cross-sectional design of the study limits causal interpretation, the strength and consistency of the findings, the apparent “dose-response” relationship between vaccination status and several forms of chronic illness, and the significant association between vaccination and NDDs all support the conclusion that some aspect of the current vaccination program may be contributing to risks of childhood morbidity. In particular, vaccination remained significantly associated with NDD after controlling for other factors, whereas preterm birth, long considered a major risk factor for NDD, was not associated with NDD after controlling for vaccination and other significant factors. On the other hand, preterm birth coupled with vaccination was associated with an apparent synergistic increase in the odds of NDD, above that of vaccination alone. Further research involving larger, independent samples is needed to verify and understand these unexpected findings in order to optimize the impact of vaccines on children’s health.

## List of Abbreviations

ADHD, Attention Deficit Hyperactivity Disorder; ASD, Autism Spectrum Disorder; AOM, Acute Otitis Media; CDC, Centers for Disease Control and Prevention; CI, Confidence Interval; NDD, Neurodevelopmental Disorders; NHERI, National Home Education Research Institute; OR, Odds Ratio; PCV-7, Pneumococcal Conjugate Vaccine-7; VAERS, Vaccine Adverse Events Reporting System.

### **Competing Interests**

The authors declare that they have no financial interests that had any bearing on any aspect of the conduct or conclusions of the study and the submitted manuscript.

### **Author Contributions**

AM designed the study, contributed to data analysis and interpretation, and drafted the paper. BR designed the study, contributed to data collection, and edited the paper. AB contributed to data analyses and edited the paper. BJ contributed to data analyses and editing. All authors read and approved the final version of the paper.

### **Funding Sources**

This study was supported by grants from Generation Rescue, Inc., and the Children's Medical Safety Research Institute, charitable organizations that support research on children's health and vaccine safety. The funders had no role or influence on the design and conduct of the research or the preparation of reports.

### **Acknowledgments**

The authors thank those who contributed comments, suggestions and financial support for the project. We also thank the collaborating homeschool organizations and especially the mothers who participated in the survey.

### **Disclaimer**

This study, approved by the Institutional Review Board of Jackson State University, was completed prior to Dr. Mawson's tenure-track appointment at Jackson State University.

### **REFERENCES**

1. Centers for Disease Control and Prevention. Ten great public health achievements -- United States, 1900-1999. *MMWR* (1999) **48**(12):241-243.
2. Whitney CG, Zhou F, Singleton J, Schuchat A. Benefits from immunization during the Vaccines for Children Program era — United States, 1994–2013. *MMMR* (2014) **63**(16):352-355.
3. Stanwyck C, Jain N. Vaccination coverage among children in kindergarten—United States, 2006-2007 school year. *MMWR* (2007) **56**:819-821.
4. CDC. Vaccination coverage among children in kindergarten -- United States, 2012-13 school year. *MMWR* (2013) **62**(30):607-12.
5. CDC April. What Would Happen If We Stopped Vaccinations?  
<http://www.cdc.gov/vaccines/vacgen/whatifstop.htm> (Accessed 19 June 2016)

6. U.S. Department of Health & Human Services. 2010 National Vaccine Plan: Protecting the Nation's Health through Immunization. [http://www.hhs.gov/nvpo/vacc\\_plan/index.html](http://www.hhs.gov/nvpo/vacc_plan/index.html) (Accessed 19 June 2015).
7. CDC. Immunization Schedules. <http://www.cdc.gov/vaccines/schedules/index.html> (Accessed 19 June 2016).
8. Ward BJ. Vaccine adverse events in the new millennium: is there a reason for concern. *Bull WHO* (2000) **78**(2):205-215.
9. Sienkiewicz D, Kulak W, Okurowska-Zawada B, Paszko-Patek G. Neurologic adverse events following vaccination. *Prog Health Sci* (2012) **2**(1):129-141.
10. Pollard AJ. Childhood immunisation: what is the future? *Arch Disease Child* (2007) **92**(5):426-433. doi:10.1136/adc.2006.095760
11. Aaby P, Whittle H, Benn CS. Vaccine programmes must consider their effect on general resistance. *BMJ* (2012) **344**:e3769. doi: 10.1136/bmj.e3769
12. Cunningham AS. Vaccine mandates in the US are doing more harm than good *BMJ* (2015) **351**:h4576. <http://dx.doi.org/10.1136/bmj.h4576> (Accessed 19 June 2016).
13. Dórea JG. Exposure to mercury and aluminum in early life: developmental vulnerability as a modifying factor in neurologic and immunologic effects. *Int J Environ Res Public Health* (2015) **12**(2):1295-313. doi: 10.3390/ijerph120201295
14. Crowcroft NS, Deeks SL, Upshur RE. Do we need a new approach to making vaccine recommendations? *BMJ* (2015) **350**:h308. doi: 10.1136/bmj.h308
15. Kessler D. Introducing MEDWatch: a new approach to reporting medication and device adverse events and product problems. *JAMA* (1993) **269**(21):2765-2768.
16. Institute of Medicine. Vaccine Safety Research, Data Access, and Public Trust. Washington, DC: The National Academies Press (2005). [http://www.nap.edu/catalog.php?record\\_id=11234](http://www.nap.edu/catalog.php?record_id=11234) (Accessed 19 June 2016).
17. Centers for Disease Control and Prevention's Immunization Safety Office. Scientific Agenda. Immunization Safety Office, Division of Healthcare Quality Promotion, National Center for Emerging and Zoonotic Infectious Diseases, February 2011. [http://www.cdc.gov/vaccinesafety/pdf/iso-finalscientific\\_agenda-nov-10.pdf](http://www.cdc.gov/vaccinesafety/pdf/iso-finalscientific_agenda-nov-10.pdf) (Accessed 19 June 2016).
18. Institute of Medicine. Adverse Effects of Vaccines: Evidence and Causality. Washington, DC: The National Academies Press, 2012. <http://www.iom.edu/Reports/2011/Adverse-Effects-of-VaccinesEvidence-and-Causality.aspx> (Accessed 19 June 2016).

19. Institute of Medicine. The childhood immunization schedule and safety: Stakeholder concerns, scientific evidence, and future studies: Washington, DC: The National Academies Press, 2013.  
[http://www.iom.edu/~media/Files/Report%20Files/2013/Childhood-Immunization-Schedule/ChildhoodImmunizationScheduleandSafety\\_RB.pdf](http://www.iom.edu/~media/Files/Report%20Files/2013/Childhood-Immunization-Schedule/ChildhoodImmunizationScheduleandSafety_RB.pdf) (Accessed 19 June 2016).
20. Maglione MA, Das L, Raaen L, Smith A, Chari R, Newberry S, et al. Safety of vaccines used for routine immunization of US children: a systematic review. *Pediatrics* (2014) **134**(2):325-37. doi: 10.1542/peds.2014-1079
21. Siegrist CA. "Vaccine Immunology,". In: Plotkin SA, Orenstein WA, Offit PA, editors. *Vaccines*. edit 5. Saunders Elsevier (2008).  
[http://www.who.int/immunization/documents/Elsevier\\_Vaccine\\_immunology.pdf](http://www.who.int/immunization/documents/Elsevier_Vaccine_immunology.pdf) (Accessed 19 June 2016).
22. Benn CS, Netea MG, Selin LK, Aaby P. A small jab - a big effect: nonspecific immunomodulation by vaccines. *Trends Immunol* (2013) **34**(9):431-9. doi: 10.1016/j.it.2013.04.004
23. Jensen KJ, Benn CS, van Crevel R. Unravelling the nature of non-specific effects of vaccines - A challenge for innate immunologists. *Semin Immunol* (2016) **28**(4):377-83. doi: 10.1016/j.smim.2016.05.005
24. Sørup S, Benn CS, Poulsen A, Krause TG, Aaby P, Ravn H. Live vaccine against measles, mumps, and rubella and the risk of hospital admissions for nontargeted infections. *JAMA* (2014) **311**(8):826-35. doi: 10.1001/jama.2014.470
25. Aaby P, Benn C, Nielsen J, Lisse IM, Rodrigues A, Ravn H. Testing the hypothesis that diphtheria-tetanus-pertussis vaccine has negative non-specific and sex-differential effects on child survival in high-mortality countries. *BMJ Open* (2012) **2**(3):e000707.
26. Garly ML, Jensen H, Martins CL, et al. Hepatitis B vaccination associated with higher female than male mortality in Guinea-Bissau: an observational study. *Pediatr Infect Dis J* (2004) **23**(12):1086-1092.
27. Grandjean P, Landrigan PJ. Developmental neurotoxicity of industrial chemicals. *Lancet* (2006) **368**(9553):2167-78.
28. Boyle CA, Boulet S, Schieve LA, Cohen RA, Blumberg SJ, Yeargin-Allsopp M, et al. Trends in the prevalence of developmental disabilities in US Children, 1997-2008. *Pediatrics* (2011) **127**:1034-1042. doi:10.1542/peds.2012-2989
29. Baio J. Prevalence of Autism Spectrum Disorder among children aged 8 years — Autism and Developmental Disabilities Monitoring Network, 11 Sites, United States, 2010 Surveillance Summaries. *MMWR* (2014) **63**(SS02):1-21.  
[http://www.cdc.gov/mmwr/preview/mmwrhtml/ss6302a1.htm?s\\_cid=ss6302a1\\_w](http://www.cdc.gov/mmwr/preview/mmwrhtml/ss6302a1.htm?s_cid=ss6302a1_w) (Accessed 19 June 2016).



30. Zablotsky B, Black LI, Maenner MJ, Schieve LA, Blumberg SJ. Estimated prevalence of autism and other developmental disabilities following questionnaire changes in the 2014 National Health Interview Survey. *Natl Health Stat Report* (2015) **13**(87):1-20. <http://www.cdc.gov/nchs/nhis/index.htm> (Accessed 19 June 2016).
31. Visser SN, Danielson ML, Bitsko RH, Holbrook JR, Kogan MD, Ghandour RM, et al. Trends in the parent-report of health care provider-diagnosed and medicated attention-deficit/hyperactivity disorder: United States, 2003-2011. *J Am Acad Child Adolesc Psychiatry* (2014) **53**(1):34-46.e2. doi: 10.1016/j.jaac.2013.09.001
32. Cortiella C, Horowitz SH. *The State of Learning Disabilities: Facts, Trends and Emerging Issues*. New York: National Center for Learning Disabilities (2014). <https://www.nclld.org/wpcontent/uploads/2014/11/2014-State-of-LD.pdf> (Accessed 19 June 2016).
33. Landrigan P. What causes autism? Exploring the environmental contribution. *Curr Opin Pediatr* (2010) **22**(2):219-25. doi: 10.1097/MOP.0b013e328336eb9a
34. Nevison CD. A comparison of temporal trends in United States autism prevalence to trends in suspected environmental factors. *Environmental Health* (2014) **13**:73. doi:10.1186/1476-069X-13-73
35. Shaw CA, Seneff S, Kette SD, Tomljenovic L, Oller JW Jr, Davidson RM. Aluminum-induced entropy in biological systems: implications for neurological disease. *J Toxicol* (2014) **2014**:491316. doi: 10.1155/2014/491316
36. Sealey LA, Hughes BW, Sriskanda AN, Guest JR, Gibson AD, Johnson-Williams L, et al. Environmental factors in the development of autism spectrum disorders. *Environ Int* (2016) **88**:288-98. doi: 10.1016/j.envint.2015.12.021
37. U.S. Department of Health and Human Services. Health Resources and Services Administration, HRSA. <http://www.hrsa.gov/vaccinecompensation/data.html> (Accessed 20 June 2016).
38. Holland M, Conte L, Krakow R, Colin L. Unanswered questions from the Vaccine Injury Compensation Program: A review of compensated cases of vaccine-induced brain injury. *Pace Envtl L Rev* (2011) **28**(2):480. Available at: <http://digitalcommons.pace.edu/pelr/vol28/iss2/6> (Accessed 20 June 2016).
39. Doja A, Roberts W. Immunizations and autism: A review of the literature. *Can J Neurol Sci* (2000) **33**(4):341-346.
40. Price CS, Thompson WW, Goodson B, Weintraub ES, Croen LA, Hinrichsen VL, et al. Prenatal and infant exposure to thimerosal from vaccines and immunoglobulins and risk of autism. *Pediatrics* (2010) **126**(4):656-64. doi: 10.1542/peds.2010-0309

41. DeStefano F, Price CS, Weintraub ES. Increasing exposure to antibody-stimulating proteins and polysaccharides in vaccines is not associated with risk of autism. *J Pediatr* (2013) **163**(2):561-7. doi: 10.1016/j.jpeds.2013.02.001
42. McNeil MM, Gee J, Weintraub ES, Belongia EA, Lee GM, Glanz JM, et al. The Vaccine Safety Datalink: successes and challenges monitoring vaccine safety. *Vaccine* (2014) **32**(42):5390-8. doi: 10.1016/j.vaccine.2014.07.073
43. Taylor LE, Swerdfeger AL, Eslick GD. Vaccines are not associated with autism: an evidence-based meta-analysis of case-control and cohort studies. *Vaccine* (2014) **32**(29):3623-9. doi: 10.1016/j.vaccine.2014.04.085
44. Jain A, Marshall J, Buikema A, Bancroft T, Kelly JP, Newschaffer CJ. Autism occurrence by MMR vaccine status among US children with older siblings with and without autism. *JAMA* (2015) **313**(15):1534-40. doi: 10.1001/jama.2015.3077. Erratum in: *JAMA* (2016) **315**(2):204.
45. Gerber JS, Offit PA. Vaccines and autism: a tale of shifting hypotheses. *Clin Infect Dis* (2009) **48**(4):456-61. doi: 10.1086/596476
46. Choi BK, Manning ML. The immunization status of home-schooled children in America. *J Pediatr Health Care* (2010) **24**:42-47.
47. Ray BD. Academic achievement and demographic traits of homeschool students: a nationwide study. *J Acad Leadership* (2010) **8**(1). <http://www.nheri.org/research/> (Accessed 19 August 2016).
48. United States Census Bureau. Statistical Abstracts Series. Statistical Abstract of the United States: 2009. [https://www.census.gov/library/publications/time-series/statistical\\_abstracts.html](https://www.census.gov/library/publications/time-series/statistical_abstracts.html) (Accessed 19 August 2016).
49. United States Department of Education. Elementary/secondary education, Table A-6-1: Number and percentage distribution of all school-age children who were homeschooled and homeschooling rate, by selected characteristics: 1999, 2003, and 2007. Published 2009. <http://files.eric.ed.gov/fulltext/ED505409.pdf> (Accessed 22 August 2016).
50. Princiotta D, Bielick S, Chapman C. Homeschooling in the United States: 2003 statistical analysis report [NCES 2006-042]. Washington, DC: U.S. Department of Education, Institute of Education Sciences. <http://nces.ed.gov/pubs2006/2006042.pdf> (Accessed 22 August 2016).
51. Noel A, Stark P, Redford J. Parent and family involvement in education, from the National Household Education Surveys Program of 2012 (NCES 2013-028). Washington, DC: *National Center for Education Statistics*, Institute of Education Sciences, U.S. Department of Education (2013). <http://eric.ed.gov/?id=ED544174> (Accessed 22 August 2016).



52. Surén P, Bakken IJ, Aase H, Chin R, Gunnes N, Lie KK, et al. Autism spectrum disorder, ADHD, epilepsy, and cerebral palsy in Norwegian children. *Pediatrics* (2012) **130**(1):e152-8. doi: 10.1542/peds.2011-3217
53. Zocchetti C, Consonni D, Bertazzi PA. Relationship between prevalence rate ratios and odds ratios in cross-sectional studies. *Int J Epidemiol* (1997) **26**(1):220-3.
54. Goldman GS, Miller NZ. Relative trends in hospitalizations and mortality among infants by the number of vaccine doses and age, based on the Vaccine Adverse Event Reporting System (VAERS), 1990-2010. *Hum Exp Toxicol* (2012) **31**(10):1012-1021.  
<http://het.sagepub.com/content/31/10/1012.full> (Accessed February 14, 2016).
55. Orenstein WA, Perry RT, Halsey NA. The clinical significance of measles: a review. *J Infect Dis* (2004) **189**:S4-S16. doi:10.1086/377712
56. CDC. Prevention of measles, rubella, congenital rubella syndrome, and mumps, 2013: Summary Recommendations of the Advisory Committee on Immunization Practices (ACIP). Recommendations and Reports. *MMWR* (2013) **62**(RR04):1-34.
57. Dhooze IJ. Risk factors for the development of otitis media. *Current Allergy Asthma Rep* (2003) **3**:321–325.
58. Siegel RM. Acute otitis media guidelines, antibiotic use, and shared medical decision-making. *Pediatrics* (2010) **125**(2):384-386. doi: 10.1542/peds.2009-3208
59. Casselbrant ML, Mandel EM. Epidemiology. In: Rosenfeld RM, Bluestone CD, editors. *Evidence-based otitis media*. Hamilton, ON, Canada: BC Decker (2003). p. 147–162.
60. Monasta L, Ronfani L, Marchetti F, Montico M, Vecchi Brumatti L, Bavcar A, et al. Burden of disease caused by otitis media: systematic review and global estimates. *PLoS ONE* (2012) **7**(4): e36226. doi:10.1371/journal.pone.0036226
61. Ahmed S, Shapiro NL, Bhattacharyya N. Incremental health care utilization and costs for acute otitis media in children. *Laryngoscope* (2014) **124**(1):301-5. doi: 0.1002/lary.24190
62. National Vaccine Information Center. 21525 Ridgetop Circle, Suite 100, Sterling, VA 20166. Available:  
[http://www.medalerts.org/vaersdb/findfield.php?TABLE=ON&GROUP1=AGE&EVENTS=ON&SYMPTOMS\[\]=Otitis+media+%2810033078%29&NUMDAYS\[\]=0&NUMDAYS\[\]=1&NUMDAYS\[\]=2&NUMDAYS\[\]=3&NUMDAYS\[\]=4&NUMDAYS\[\]=5&NUMDAYS\[\]=6&NUMDAYS\[\]=7&WhichAge=range&LOWAGE=0.0&HIGHAGE=1.0](http://www.medalerts.org/vaersdb/findfield.php?TABLE=ON&GROUP1=AGE&EVENTS=ON&SYMPTOMS[]=Otitis+media+%2810033078%29&NUMDAYS[]=0&NUMDAYS[]=1&NUMDAYS[]=2&NUMDAYS[]=3&NUMDAYS[]=4&NUMDAYS[]=5&NUMDAYS[]=6&NUMDAYS[]=7&WhichAge=range&LOWAGE=0.0&HIGHAGE=1.0) (Accessed 25 August, 2016).
63. Revai K, McCormick DP, Patel J, Grady JJ, Saeed K, Chonmaitree T. Effect of pneumococcal conjugate vaccine on nasopharyngeal bacterial colonization during acute otitis media. *Pediatrics* (2006) **117**(5):1823–9.

64. Faden H, Harabuchi Y, Hong JJ. Epidemiology of *Moraxella catarrhalis* in children during the first 2 years of life: relationship to otitis media. *J Infect Dis* (1994) **169**(6):1312-7.
65. Weinberger DM, Malley R, Lipsitch M. Serotype replacement in disease after pneumococcal vaccination. *Lancet* (2011) **378**(9807):1962-73. doi: 10.1016/S0140-6736(10)62225-8
66. Biesbroek G, Wang X, Keijser BJ, Eijkemans RM, Trzciński K, Rots NY, et al. Seven-valent pneumococcal conjugate vaccine and nasopharyngeal microbiota in healthy children. *Emerg Infect Dis* (2014) **20**(2):201-10. doi: 10.3201/eid2002.131220
67. Goldin RL, Matson JL. Premature birth as a risk factor for autism spectrum disorder. *Dev Neurorehabil* (2016) **19**(3):203-6. doi: 10.3109/17518423.2015.1044132
68. Padilla N, Eklöf E, Mårtensson GE, Bölte S, Lagercrantz H, Ådén U. Poor brain growth in extremely preterm neonates long before the onset of autism spectrum disorder symptoms. *Cereb Cortex* (2015) pii: bhv300.
69. Short ME, Goetzel RZ, Pei X, Tabrizi MJ, Ozminkowski RJ, Gibson TB, et al. How accurate are self-reports? Analysis of self-reported health care utilization and absence when compared with administrative data. *J Occup Environ Med* (2009) **51**(7):786-96. doi: 10.1097/JOM.0b013e3181a86671
70. Fisker AB, Hornshøj L, Rodrigues A, Balde I, Fernandes M, Benn CS, et al. Effects of the introduction of new vaccines in Guinea-Bissau on vaccine coverage, vaccine timeliness, and child survival: an observational study. *Lancet Glob Health* (2014) **2**(8):e478-87. doi: 10.1016/S2214109X(14)70274-8

**Table 1. Characteristics of the respondents <sup>a</sup>**

	<b>Mean (SD) <sup>a</sup></b>
<b>Age (n=407)</b>	40.59 (6.7)
	<b>Number (%) <sup>a</sup></b>
<b>Race</b>	
White	382 (92.5%)
Non-White	21 (7.6%)
Total	413
<b>Education</b>	
High School Graduate or Less	35 (8.5%)
Some College	114 (27.5%)
College Graduate	187 (45.2%)
Post-Graduates	78 (18.5%)
Total	414
<b>Total Gross Household Income</b>	
< \$49,999	123 (30.8%)
\$50,000-100,000	182 (45.5%)
> \$100,000	95 (23.8%)
Total	400
<b>Religious Affiliation</b>	
Christianity	375 (91.2%)
Non-Christianity	36 (8.8%)
Total	411
<b>Marital Status</b>	
Married	386 (93.7%)
Not Married	26 (6.3%)
Total	412

<sup>a</sup> Missing observations are excluded.

**Table 2. Vaccination status and health outcomes – Acute Conditions**

	<b>Vaccinated (n=405)</b>	<b>Unvaccinated (n=261)</b>	<b>Total (n=666)</b>	<b>Chi- square</b>	<b>P-value</b>	<b>Odds Ratio (95% CI)</b>
Chickenpox						
Yes	32 (7.9%)	66 (25.3%)	98 (14.7%)	38.229	< 0.001	0.26 (0.2 - 0.4)
No	373 (92.1%)	195 (74.7%)	568 (85.3%)			
Otitis media						
Yes	80 (19.8%)	16(5.8%)	96 (14.4%)	26.643	< 0.001	3.8 (2.1 - 6.6)
No	325 (80.2%)	245 (94.2%)	507 (85.6%)			
Pneumonia						
Yes	26 (6.4%)	3 (1.2%)	29 (4.4%)	10.585	< 0.001	5.9 (1.8 - 19.7)
No	379 (93.6%)	258 (98.8%)	637 (95.6%)			
Whooping cough						
Yes	10 (2.5%)	22 (8.4%)	32 (4.8%)	12.326	< 0.001	0.3 (0.1 - 0.6)
No	395 (97.5%)	239 (91.6%)	634 (95.2%)			
Rubella						
Yes	1 (0.3%)	5 (1.9%)	6 (0.9%)	4.951	0.037	0.1 (0.01 - 1.1)
No	404 (99.6%)	256 (98.1%)	660 (99.1%)			

Provisional

**Table 3. Vaccination status and health outcomes – Chronic Conditions**

<b>Chronic Disease</b>	<b>Vaccinated (n=405)</b>	<b>Unvaccinated (n=261)</b>	<b>Chi- square</b>	<b>P-value</b>	<b>Odds Ratio (95% CI)</b>
Allergic rhinitis					
Yes	42 (10.4%)	1 (0.4%)	26.21	< 0.001	30.1 (4.1 - 219.3)
No	363 (89.6%)	260 (99.6%)			
Allergies					
Yes	90 (22.2%)	18 (6.9%)	29.44	< 0.001	3.9 (2.3 - 6.6)
No	315 (77.9%)	243 (93.1%)			
ADHD					
Yes	19 (4.7%)	3 (1.0%)	6.23	0.013	4.2 (1.2 - 14.5)
No	386 (95.3%)	258 (99.0%)			
ASD					
Yes	19 (4.7%)	3 (1.0%)	6.23	0.013	4.2 (1.2 - 14.5)
No	386 (95.3%)	258 (99.0%)			
Eczema (atopic dermatitis)					
Yes	38 (9.5%)	9 (3.6%)	8.522	0.035	2.9 (1.4 - 6.1)
No	367 (90.5%)	252 (96.4%)			
Learning Disability					
Yes	23 (5.7%)	3 (1.2%)	8.6803	0.003	5.2 (1.6 - 17.4)
No	382 (94.3%)	258 (98.9%)			
Neurodevelopment Disorder					
Yes	42 (10.5%)	8 (3.1%)	12.198	< 0.001	3.7 (1.7 - 7.9)
No	313 (89.5%)	253 (96.9%)			
Any Chronic Condition					
Yes	178 (44.0%)	65 (24.9%)	24.8456	< 0.001	2.4 (1.7 - 3.3)
No	227 (56.0%)	196 (75.1%)			

**Table 4. Partial versus full vaccination and chronic health conditions**

	Unvaccinated (n=261)	Partially Vaccinated (n=208)	Fully Vaccinated (n=197)	Total (n=666)	Chi-Square	P-value
<b>Chronic Conditions</b>						
Allergic rhinitis						
Yes	1 (0.4%)	17 (8.2%)	25 (12.7%)	43 (6.5%)	29.6306	< 0.001
No	260 (99.6%)	191 (91.8%)	172 (87.3%)	623 (93.5%)		
Allergies						
Yes	18 (6.9%)	47 (22.6%)	43 (21.8%)	108 (16.2%)	27.4819	< 0.001
No	243 (93.1%)	161 (77.4%)	154 (78.2%)	558 (83.8%)		
ADHD						
Yes	3 (1.2%)	8 (3.9%)	11 (5.6%)	22 (3.3%)	7.1900	0.075
No	258 (98.8%)	200 (96.1%)	186 (94.4%)	644 (96.7%)		
ASD						
Yes	3 (1.2%)	11 (5.3%)	8 (4.6%)	22 (3.3%)	6.7109	0.034
No	258 (98.8%)	197 (94.7%)	189 (95.4%)	644 (96.7%)		
Eczema (atopic dermatitis)						
Yes	9 (3.5%)	18 (8.7%)	20 (10.2%)	47 (7.1%)	8.8683	0.012
No	252 (96.5%)	190 (91.3%)	177 (89.8%)	619 (92.9%)		
Learning Disability						
Yes	3 (1.2%)	11 (5.3%)	12 (6.1%)	26 (3.9%)	8.8541	0.012
No	258 (98.8%)	197 (94.7%)	185 (93.9%)	640 (96.1%)		
NDD						
Yes	8 (3.1%)	21 (10.1%)	21 (10.7%)	50 (7.5%)	12.2443	0.002
No	253 (96.9%)	187 (89.9%)	176 (89.3%)	616 (92.5%)		
Any Chronic Condition						
Yes	65 (24.9%)	94 (45.2%)	84 (42.6%)	243 (36.5%)	25.1301	< 0.001
No	196 (75.1%)	114 (54.8%)	113 (57.4%)	423 (63.5%)		

**Table 5. Chronic conditions and gender among vaccinated children**

	Male (n=194)	Female (n=209)	Total (n=403)	Chi- square	P-value	Odds Ratio (95% CI)
Allergic rhinitis						
Yes	27 (13.9%)	15 (7.2%)	42 (10.4%)	4.8964	0.0269	2.1 (1.1 - 4.1)
No	167 (86.1%)	194 (92.8%)	361 (90.0%)			
Allergies						
Yes	50 (25.8%)	40 (19.1%)	90 (22.3%)	2.5531	0.1101	1.5 (0.91 - 2.4)
No	144 (74.2%)	168 (80.9%)	313 (77.7%)			
ADHD						
Yes	13 (6.7%)	6 (2.9%)	19 (4.7%)	3.2856	0.0699	2.4 (0.90 - 6.5)
No	181 (93.3%)	203 (97.1%)	384 (95.3%)			
ASD						
Yes	15 (7.7%)	4 (1.9%)	19 (4.7%)	7.5810	0.0059	4.3 (1.4 - 13.2)
No	178 (92.3%)	205 (98.1%)	384 (95.3%)			
Eczema						
Yes	19 (9.89%)	19 (9.1%)	38 (9.4%)	0.0582	0.8094	1.1 (0.6 - 2.1)
No	175 (90.2%)	190 (90.9%)	365 (90.6%)			
Learning Disability						
Yes	14 (7.2%)	9 (4.3%)	23 (5.7%)	1.5835	0.2083	1.7 (0.7 - 4.1)
No	180 (92.8%)	200 (95.7%)	380 (94.3%)			
NDD						
Yes	28 (14.4%)	14 (6.7%)	42 (10.4%)	6.4469	0.0111	2.3 (1.2 - 4.6)
No	166 (85.6%)	195 (93.3%)	361 (89.6%)			
Any Chronic Condition						
Yes	94 (48.5%)	83 (39.7%)	177 (43.9%)	3.1208	0.0773	1.4 (1.0 - 2.1)
No	100 (51.5%)	126 (60.3%)	226 (56.1%)			

**Table 6. Vaccination status, medication use and health services utilization**

	Vaccinated (n=405)	Unvaccinated (n=261)	Total (n=666)	Chi- square	P-value	Odds Ratio (95% CI)
<b>Medication Use</b>						
Medication for Allergy						
Yes	81 (20.0%)	3 (1.2%)	84 (12.6%)	51.170	< 0.001	21.5 (6.7 - 68.9)
No	324 (80.0%)	258 (98.8%)	582 (87.4%)			
Used antibiotics in the past 12 months						
Yes	124 (30.8%)	40 (15.4%)	164 (24.7%)	20.092	< 0.001	2.4 (1.6 - 3.6)
No	279 (69.2%)	220 (84.6%)	499 (75.3%)			
Used fever medication 1+ times						
Yes	350 (90.7%)	173 (67.8%)	523 (81.6%)	53.288	< 0.001	4.6 (3.0 - 7.1)
No	36 (9.3%)	82 (32.2%)	118 (18.4%)			
Using fitted ear drainage tubes						
Yes	12 (3.0%)	1 (0.4%)	13 (2.0%)	5.592	0.018	8.0 (1.0 - 66.1)
No	389 (97.0%)	260 (99.6%)	649 (98.0%)			
Used medication for ADHD						
Yes	7 (1.7%)	3 (1.2%)	10 (1.5%)	0.346	0.556	-
No	398 (98.3%)	256 (98.8%)	654 (98.5%)			
Used medication for Seizures						
Yes	4 (1.0%)	1 (0.4%)	5 (0.8%)	0.769	0.653	-
No	400 (99.0%)	258 (99.6%)	658 (99.2%)			
<b>Health Services Utilization</b>						
Emergency Department visit in the past 12 months						
Yes	38 (9.5%)	23 (9.0%)	61 (9.3%)	0.047	0.828	-
No	364 (90.5%)	234 (91.0%)	598 (90.7%)			
Sick visit to doctor in the past year						
Yes	145 (36.0%)	41 (16.0%)	186 (28.2%)	31.096	< 0.001	3.0 (2.0 - 4.4)
No	258 (64.0%)	216 (84.0%)	474 (71.8%)			
Ever spent one or more nights in the hospital						
Yes	80 (19.8%)	32 (12.3%)	112 (16.8%)	6.267	0.012	1.8 (1.1 - 2.7)
No	325 (80.2%)	228 (87.7%)	553 (83.2%)			
Seen doctor for checkup in past 12 months						
Yes	233 (57.6%)	97 (37.2%)	330 (49.6%)	26.336	< 0.001	2.3 (1.7 - 3.2)
No	172 (42.4%)	164 (62.8%)	336 (50.4%)			
Seen dentist in the past 12 months						
Yes	362 (89.4%)	210 (80.5%)	572 (85.9%)	10.424	< 0.001	2.0 (1.3 - 3.2)
No	43 (10.6%)	51 (19.5%)	94 (14.1%)			



**Table 7. Unadjusted analysis of potential risk factors for neurodevelopmental disorders**

	NDD					
Vaccination Status	Yes (N=50)	No (N=616)	Total* (N=666)	Chi-Square	P-value	OR (95% CI)**
Vaccinated	42	363	405	12.198	<0.001	3.7 (1.7 - 7.9)
Not Vaccinated	8	253	261			Ref
Race						
Non-White	9	71	80	1.8208	0.177	1.7 (0.7 - 3.6)
White	41	544	585			Ref
Child's Gender						
Male	32	283	315	5.9471	0.015	2.1 (1.1 - 3.8)
Female	18	331	349			Ref
Adverse Environment						
Yes	6	27	33	5.8706	0.053	2.9 (1.1 - 7.4)
No	40	523	563			Ref
Do not know	4	66	70			0.8 (0.3 - 2.3)
Medication during Pregnancy - Antibiotics						
Yes	10	61	71	4.950	0.026	2.3 (1.1 - 4.8)
No	40	555	595			Ref
Medication during Pregnancy - Vaccinated						
Yes	6	32	38	3.965	0.057	2.5 (1.0 - 6.3)
No	44	583	627			Ref
Preterm birth						
Yes	12	37	49	22.910	< 0.001	4.9 (2.4 - 10.3)
No	38	578	616			Ref
Ultrasound						
None	3	71	74	5.898	0.052	Ref
1-3 times	30	419	449			1.7 (0.5 - 5.7)
> 3 times	17	124	141			3.2 (0.92 - 11.5)

\*Numbers may not add to column totals due to missing or incomplete data.

\*\*Note that Odds Ratios are the cross-product ratios of the entries in the 2-by-2 tables, and are an estimate of the relative incidence (or risk) of the outcome associated with the exposure factor.

**Table 8. Adjusted logistic regression analyses of risk factors and NDD\***

	Adjusted Model (Model 1)	Adjusted Model with Interaction (Model 2)
<b>Vaccination Status</b>		
<b>Vaccinated</b>	3.1 (1.4 - 6.8)	2.5 (1.1 - 5.6)
<b>Not Vaccinated</b>	Ref	Ref
<b>Race</b>		
<b>Non-White</b>	2.3 (1.0 - 5.2)	2.0 (1.1 - 5.5)
<b>White</b>	Ref	Ref
<b>Child's Gender</b>		
<b>Male</b>	2.3 (1.2 - 4.3)	2.3 (1.2 - 4.4)
<b>Female</b>	Ref	Ref
<b>Pre-term birth</b>		NS
<b>Yes</b>	5.0 (2.3 - 11.6)	
<b>No</b>	Ref	
<b>Pre-term birth and Vaccination interaction</b>		
<b>No interaction</b>	Not in the model	Ref
<b>Pre-term and Vaccinated</b>		6.6 (2.8 - 15.5)

\*Number of observation read 666, number of observations used 629. NDD=47, Not NDD = 582

**Figure 1. The overlap and distribution of physician-diagnosed neurodevelopmental disorders based on mothers' reports**

