



# UNIVERSITÀ DEGLI STUDI DI TORINO

# AperTO - Archivio Istituzionale Open Access dell'Università di Torino

# Attitudes towards compulsory vaccination in Italy: Results from the NAVIDAD multicentre study

This is the author's manuscript
Original Citation:
Availability:
This version is available http://hdl.handle.net/2318/1706712 since 2019-07-17T15:32:51Z
Published version:
DOI:10.1016/j.vaccine.2018.04.029
Terms of use:
Open Access
Anyone can freely access the full text of works made available as "Open Access". Works made available under a Creative Commons license can be used according to the terms and conditions of said license. Use of all other works requires consent of the right holder (author or publisher) if not exempted from copyright protection by the applicable law.

(Article begins on next page)

# Title

Attitudes towards compulsory vaccination in Italy: results from the NAVIDAD multicenter study.

# Abstract

BACKGROUND: Vaccine hesitancy is a considerable issue in European countries and leads to low coverage rates. After a long debate, Italy has made vaccination mandatory for admission to its schools. METHODS: For the NAVIDAD study (a cross-sectional multicenter study), a 63-item questionnaire was administered to 1820 pregnant women from 15 Italian cities. The questionnaire assessed interviewee's opinion on mandatory vaccines, as well as their socioeconomic status, sources of information about vaccines, confidence in the Italian National Healthcare Service (NHS), and intention to vaccinate their newborn.

RESULTS: Information sources play a key role in determining the opinion on restoration of mandatory vaccine, in particular women who got information from anti-vaccination movements websites are less likely to accept it (OR: 0.35, 95%CI: 0.21-0.58, p<0.001). Women who had confidence in healthcare professional information agreed more on mandatory vaccination than the other (OR: 2.66, 95%CI:1.62-4.36, p<0.001), who perceive that healthcare professionals have economic interest in child immunization and who declared that healthcare providers inform only on vaccinations benefits not on risks were less likely to agree on compulsory vaccination (OR: 0.66, CI 95%: 0.46-0.96, p=0,03;OR: 0.66, CI 95%: 0.46-0.95, p=0.03).

CONCLUSION: Information sources and confidence towards health professionals are the main determinants of acceptance of mandatory vaccine restoration. In order to increase the acceptability of the restoration and reduce vaccine hesitancy, these aspects need to be strengthened.

# Keywords

Mandatory vaccination; Vaccine hesitancy; attitudes; pregnancy; multicentre survey; Italy

# Highlishts

- Compulsory vaccination is generally welcome
- Mandatory vaccination are not affected by social determinants
- Confidence in health system determinate trustworthiness of mandatory vaccination

# Acknowledgements

The authors declare that no funding or grant was received from other organizations for the conduction

of this study.

### 1 Introduction

2 Vaccination appears to be the most effective and cost-effective intervention to reduce the burden of 3 contagious diseases (1-3). Immunization averts an estimated 2 to 3 million deaths every year; 4 however, an additional 1.5 million deaths could be avoided if global vaccination coverage improves (4). Today, several vaccines are available and differently administered all over the world. Moreover, 5 6 the immunization rates across countries vary considerably and an estimated 19.5 million infants 7 worldwide are still missing out on basic vaccines (5). The Global Vaccine Action Plan (2011-2020) 8 (GVAP) is a framework adopted by all the World Health Organization (WHO) Member States at the 9 Sixty-fifth World Health Assembly in May 2012 to achieve the vision of the Decade of Vaccines 10 (DoV) 2011-2020 of "a world in which all individuals and communities enjoy lives free from 11 vaccine-preventable diseases" (6). The GVAP sets goals, strategic objectives and indicators to achieve 12 the mission, which is "improve health by extending by 2020 and beyond the full benefits of 13 immunization to all people, regardless of where they are born, who they are, or where they live" (6).

The European Vaccine Action Plan (2015-2020) (EVAP) developed by the 53 Member States of the Region with the WHO Regional Office for Europe, immunization partners and stakeholders, contributes directly to the goals of the GVAP and the European Region's overall Health 2020 strategy (7). Despite efforts, the 2015 regional measles and rubella elimination target was missed. The Region's polio-free status was threatened and several countries saw a resurgence of diphtheria and pertussis, which also exposed the unpredictability of vaccine supply in the Region (4).

20 Nowadays, Europe faces many challenges including issues with access to vaccine supply and 21 affordable pricing, sustainable domestic financing and resource mobilization, as well as a growth of 22 anti-vaccination sentiment and visibility (6). Indeed, many Countries and communities are dealing 23 with groups refusing available recommended vaccinations for themselves and/or their children (8-10). 24 The factors underlying these decisions are different and there is no single intervention strategy that 25 can solve the problem (11,12). Vaccines are losing public confidence and several international 26 organizations (WHO, EU, ECDC) warn against the growing phenomenon of vaccine hesitancy and its 27 impact on decreasing vaccine coverage trends (13,14). This has created a need for national 28 immunization programs to find approaches and strategies to address vaccine hesitancy.

In Italy for each vaccine included in the National Immunization Schedule (NIS), it has provided the coverage targets fixed considering herd immunity thresholds needed to break infectious diseases transmission throughout the population. The 24 months coverage target defined in PNPV (Piano

32 Nazionale Prevenzione Vaccinale) was set at  $\geq 95\%$  for the following vaccines: DTPa (Diphtheria, 33 Tetanus, Acellular Pertussis), Hepatitis B, Hib (Haemophilus influenzae type b), first dose of MPR 34 (Measles, Mumps, Rubella), Pneumococcal, Meningococcal C, Chicken pox, Rotavirus (15). The 35 HPV vaccine in females and in males should achieve a coverage  $\geq 70\%$  at 12 years old. In 2016 36 available data on infant vaccines reported that the 24 months of age coverages were all beneath the 37 95% threshold (16). These percentages were a long way off the PNPV published targets and the 38 WHO's recommendations on GVAP. Furthermore, it is important to note that vaccine coverages have 39 been in decline for some years. From 2013, the only coverage that has shown an increase in national 40 data was Meningococcus (17). Negative trend coverage has been reported for all the other vaccines, as 41 Pneumococcal (88.7% in 2015 vs 88.4% in 2016), Measles and Rubella (90.4% in 2013 vs 85.3% in 42 2015 and then look slightly up in 2016 but still far from achieving the coverage needed to eliminate 43 the virus) (17). The general negative trend was also confirmed by the national 36-months vaccination 44 coverage for 2016 (relating to children born in 2013). This data is especially useful for monitoring the 45 share of children who were in default of the previous year's vaccination survey and were recovered. 46 The 36-month coverages showed slightly higher values than those found for the same birth cohort at 47 24 months the previous year. Recuperation is limited and 95% is only achieved for Hib (18).

The national low immunization levels and their negative trend led to the introduction of compulsory vaccination in Italy on 31th July 2017 for ten infectious diseases. Compulsory vaccination has been introduced in order to guarantee public health safeguard and to reach coverage targets of the PNPV (19). Preliminary data from five regions show that, compared with 2016, this strategy lead to an increase in vaccine coverege from June to October, 2017 of 1,0% for the hexavalent vaccine against diphtheria, tetanus, pertussis, poliomyelitis, *H influenzae* type b, and hepatitis B and of 2,9% for the measles, mumps, and rubella vaccine (20).

55 In this context, a study named NAVIDAD (Nozioni e Attitudini sui Vaccini dell'Infanzia nelle Donne 56 in Attesa e loro Decisioni), started in 2016 and lasted for about a year, has been conducted with the 57 aim of analysing the influence of many determinants (21-24) on Italian pregnant women's decision on 58 routine vaccinations of their children (25). This paper focuses on pregnant women's attitude towards 59 compulsory nature of infant vaccinations and its relationship with some vaccine hesitancy 60 determinants: sociodemographic data, information sources, trust on institution, knowledge and 61 perceptions on vaccines and preventable infectious diseases (26,27). The main objective of this paper 62 is to describe pregnant women's attitudes and behaviours towards compulsory nature of paediatric 63 vaccinations, assessing their trustworthiness and acceptability. In particular we wanted to analyse its

64 possible determinants, considering social determinants, source of information and trust in National

65 Health System. In fact, these information can play a role in future public health policies.

66

#### 67 Methods

- 68 A cross-sectional multicenter study was conducted by involving patients in the following Italian cities:
- 69 Bologna, Ferrara, Milan, Parma and Turin that were considered from the North of Italy;
- 70 Ancona, Perugia, Roma and Siena were considered from the Centre of Italy;
- Catania, Chieti-Pescara, L'Aquila, Messina and Naples were considered from the South of
  Italy.

73 The execution of this study was approved by the Ethics Committee of the Hospital "A.O.U. Città della74 Salute e della Scienza di Torino".

75 Study subjects (pregnant women over the age of 18 who were able to understand the protocol 76 information and the questionnaire) were enrolled from September 2016 and May 2017 among patients 77 waiting for a gynecological, ultrasound or hematological examination, in the reference hospitals of the

78 cities involved in the study.

- 79 Informed consent was obtained after the full explanation of the nature and possible consequences of80 the study.
- 81 Trained resident doctors through the submission of a non-self-compiling paper questionnaire
  82 subjected the women to a 25-minutes interview.
- 83 The questionnaire was composed of seven sections for a total of 63-item. Each section was84 investigating:
- 85 1. the socio-economic framing (patient age, qualification, occupation, ...).
- 86 2. whether she intended to vaccinate her child and for which pathologies.
- 87 3. the sources through which the women had sought and obtained information about88 vaccinations
- 89 4. the degree of confidence of the women in healthcare workers
- 5. the perception of the frequency and severity of the major preventable pathologies withvaccinations
- 92 6. an assessment of her vaccine knowledge

93

#### 7. the interviewee's opinion on the restoration of mandatory vaccines

94 This study focuses on the section number 7, interviewee's opinion on the restoration of mandatory

95 vaccines and how it is influenced by socio-economic framing, willingness to vaccinate the newborn,

96 information sources and confidence in healthcare service (section 1, 2, 3 and 4).

97

98 Population and sample size calculation

99 Two different letter were sent to all Italian Public Health Schools' Director. 15 of them agreed to 100 participate in the study. For each city, the sample was defined based on demographic data of the 101 resident population, considering the number of the newborns in the cities included (28).

Considering the MPR vaccine coverage of 86,7% (17), it was possible to provide an estimation of the number of interviews necessary in order to get valid data (29–31). We considered a -10% of MPR vaccine coverage as "Worst Acceptable" for results, in order to find a very conservative value. The confidence level was set at 95% and the power of the study was considered to be 80%. The sample size was then calculated through the statistical software "EPI INFO". To be more conservative, it was required a number of cases in the range between the sample size calculation results and the same increased by 30%.

The final sample size was expected to be in the range between 1764 and 2296 subjects involved, for
each city the number of questionnaire required ranged like follows: Ancona (116-151), Bologna (133172), Catania (132-172), Chieti-Pescara (124- 161), Ferrara (120-160), L'Aquila (111-144), Messina
(129-168), Milan (138-178), Naples (136-177), Parma (128-176), Perugia (125-163), Rome (150180), Siena (99-129) and Turin (123-175).

114

## **115** *Statistical analyses*

116 After the data collection, all the participating centers mailed the original paperwork of the 117 questionnaires filled in anonymous way to the Department of Public Health and Pediatrics at the 118 University of Turin.

119 A total of 1,820 questionnaires were processed by using SPSS 24 Statistical software for Windows.

120 At first, a descriptive analysis of all the variables was conducted. The major outcome was the 121 interviewee's opinion on the restoration of mandatory vaccines. The variables included in the analysis 122 were: the socio-demographic data, any previous pregnancies, the pregnancy quarter, the willingness to vaccinate the newborn, the different ways of obtaining the information and the degree of trust inhealthcare workers.

A logistic regression was conducted to estimate the impact of some variables on the above-described outcomes. The covariates included in the final model have been selected using a stepwise forward selection process, with the criterion of a P value at univariate <0.25 (32). These associations are expressed as Odds Ratios (OR) with 95% Confidence Interval (CI) and the p value ≤0.05 was considered significant for all analyses.</p>

130

# 131 Results

132 A total of 1,820 pregnant women were interviewed. In particular, the number of women that participated for each city was: Ancona 120, Bologna 172, Catania 160, Chieti-Pescara 139, Ferrara 133 134 140, L'Aquila 123, Messina 81, Milan 203, Naples 140, Parma 29, Perugia 125, Rome 127, Siena 90 135 and Turin 171. The mean age of the sample was 32.5 years (SD  $\pm$  5.2). The youngest patient was 18 136 years old, while the oldest 48. Most women declared to be Italian (90.8%), to be married or living with a partner (91.9%), primiparous (63.4%) and at the third trimester of pregnancy (71.9%). Around 137 138 the half of the sample affirmed to be educated at least to university degree level (46.8%). The majority 139 of women declared to be employed (71.8%): 39.9% were office workers or teachers, 14.7% were 140 entrepreneurs or private professionals, 9.3% were labourers or artisans. Moreover, 7.9% of the sample 141 had an occupation in the healthcare field.

Of the entire sample, 1.9% of women declared they would not vaccinate their next child. As regards to information sources, in the overall sample, only 41.8% of the women obtained information about vaccines from healthcare professionals; the 56.9% declared to have gained information autonomously. The most frequently used information sources were web sites (65.7%), of that a half (50.4%) were non-institutional web sites. The second most used information source was the word of mouth (47.7%). Paediatricians were the third source of information as frequency (37.3%).

Furthermore, the results showed that 92% of our sample had confidence in healthcare professionals and 86.5% declared that they are experienced and knowledgeable; only 18.7% of the interviewed women trusted more private healthcare professionals than the ones engaged by the Italian National Health System. Nevertheless, the 31.6% stated that healthcare providers have economic interest in 152 child immunisation and the 35.5% declared that healthcare professionals inform only on vaccinations 153 benefits not on risks. Moreover, the Italian vaccination schedule was considered too inflexible to adapt 154 to changing parents' needs in 42.8% of questionnaires and the 38% of the women claimed that people 155 who do not vaccine are blamed by healthcare service. Concerning vaccines, 21.7% of the sample 156 thought that they are an imposition and 44.6% that they are more useful for the society than for the 157 individual. Furthermore, the majority of interviewed women (81.6%) was in favour of compulsory 158 vaccination. Most women of the sample (81.6%) declared to be in favour of mandatory vaccinations, 159 13.8% were against them and 4.5% did not answer to the question.

160 Univariate analysis

161 In Table 1 the main demographic, occupational and social features of the sample are resumed, 162 stratified by the propensity towards compulsory vaccinations. Women from the North of Italy had a 163 higher propensity for mandatory vaccination (90.1% of them) comparing to the women from the Centre and the South (83.1% and 82.1% respectively; p<0.001). Women older than 33 agreed more on 164 165 mandatory vaccination than the younger ones (87.2 vs 83.7%; p=0.04), as well as women who were 166 married or lived with a partner when compared with single or divorced women (86.3% vs 76.3%; 167 p=0.04). Moreover, women who wanted to vaccinate their next child were more inclined towards 168 compulsory vaccination (87%) compared with the rest of the sample (19.4%; p<0.001).

169 Table 2 shows the main differences regarding the information sources between women who agreed 170 and disagreed with mandatory vaccination. Women that used institutional information leaflets had a 171 higher propensity for compulsory vaccination than women who did not use this kind of information 172 source (88.5% vs 84.2%; p=0.03), as well as women who went to vaccination clinics to obtain 173 information (88.6% vs 84.6%; p=0.05). Conversely, women who went to a private healthcare 174 professional were less inclined towards compulsory vaccination than the other ones (80.9% vs 86%; 175 p=0.05), as well as people who got information from anti-vaccination movements (69.7% vs 86.8%; 176 p<0.001).

Table 3 describes the association between the trust in health care system and the propensity towards compulsory vaccination. The analysis shows that women who had confidence in healthcare professional information agreed more on mandatory vaccination than the rest of the sample (87.7% vs 55%; p<0.001), as well as women who declared that healthcare professionals are experienced and</p>

181 knowledgeable (87.2% vs 70.2%; p<0.001) and who considered the Italian vaccination schedule</li>
182 flexible (87% vs 82.5%; p=0.02).

183 On the contrary, women who trusted more private healthcare professionals than the ones engaged by the Italian NHS agreed less to mandatory vaccination (79.6% vs 87.1%; p<0.001). The propensity 184 185 towards compulsory vaccination was lower among women who stated that healthcare providers have 186 economic interest in child immunization (77% vs 89.8%; p<0.001) and among the ones who declared 187 that they inform only on vaccinations benefits not on risks (78.3% vs 90.1%; p<0.001). Furthermore, 188 the women that claimed that people who do not vaccine are blamed by healthcare service were less 189 inclined towards mandatory vaccination compared with the rest of the sample (82.9% vs 87.1%, 190 p < 0.001), as well as women who thought that vaccines are an imposition (79% vs 87.6%; p < 0.001).

## 191 *Multivariate analysis*

192 These results were partially confirmed in the logistic regression model (Table 4): women who want to 193 vaccinate their next child were more inclined towards compulsory vaccination compared with the rest 194 of the sample (OR: 11.83, 95% CI: 3.74-37.45, p<0.001). Moreover, women who went to vaccination 195 clinics to obtain information had a higher propensity for mandatory vaccination (OR: 1.90, 95% CI: 196 1.22-2.95, p=0.01), as opposed to women who got information from anti-vaccination movements 197 comparing to the others (OR: 0.35, 95% CI: 0.21-0.58, p<0.001). Furthermore, women who had 198 confidence in healthcare professional information agreed more on mandatory vaccination than the 199 other women did (OR: 2.66, 95% CI: 1.62-4.36, p<0.001). On the contrary, people who stated that 200 healthcare professionals have economic interest in child immunisation and who declared that 201 healthcare providers inform only on vaccinations benefits not on risks were less likely to agree on 202 compulsory vaccination (OR: 0.66, 95% CI: 0.46-0.96, p=0.03; OR: 0.66, 95% CI: 0.46-0.95, 203 p=0.03).

### 204 Discussion

The implementation of a compulsory vaccination program represents a turning point in vaccination plan in Italy (19). Ethical, political and scientific debate on mandatory vaccinations is an interesting topic. The major aim of this study is to assess pregnant women's attitude toward compulsory nature of paediatric vaccination and identify the main factor influencing it. 209 The results show an interesting association between information sources and attitudes towards 210 compulsory vaccination. In fact, mandatory vaccination is more easily accepted among who seek for 211 information in vaccination clinics. With no surprise, the few women obtaining information from anti-212 vaccine associations are less prone to accept mandatory vaccination plan. A paper published on 213 Vaccine in 2015 by SAGE Working Group on Vaccine Hesitancy proposed the "3Cs" model (12). In 214 this model, a key role in determining the acceptance of vaccination was played by the confidence in 215 the health system and in the reliability and competence of the health services and health professionals. 216 Comparable results are shown in the NAVIDAD study. In fact, women who stated that healthcare 217 professionals have economic interest in child immunisation and who declared that healthcare 218 providers inform only on vaccinations benefits were less likely to agree on compulsory vaccination. 219 To be clear, vaccine hesitancy, defined as a delay or refusal in accepting vaccination (12), is slightly 220 different from attitudes towards compulsory vaccination. Nevertheless, results from NAVIDAD study 221 show how there is a strong association between acceptance of mandatory vaccination and intention to 222 vaccinate the next child. Furthermore, different studies shows how decision about vaccination is made 223 during pregnancy (33,34) and, in particular, first time mother are more vaccine hesitant and undecided 224 about childhood vaccination (35). For these reason, multicomponent and dialogue-based interventions 225 should be considered considering that strategies should be carefully tailored according to the target 226 population, their reasons for hesitancy, and the specific context (36).

A cross-sectional study has some limitation due to the study design. No causality can be proven.Therefore this survey is an assessment tool with some limitations (37).

229 On the other hand, to our knowledge, there is no other study that evaluate which are the factors that 230 can have an influence on the acceptance of compulsory vaccine plan. Furthermore, the results can be 231 easily generalized to Italian population due to the wide sample size and the number of centres 232 involved. It has to be stated that this study has been performed before the Italy's law 119/2017 that 233 makes ten vaccines mandatory for infants. As reported on The Lancet Infectious Disease by Signorelli 234 et al. (20) mandatory vaccine seems to be an effective tool to increase vaccine coverage and the results 235 are encouraging. For this reason, policy makers should consider the results from NAVIDAD study. In 236 particular, the effect on the reliability of health system and professionals should be considered, not 237 only when talking about vaccine.

238

# 239 Ethical standards

- 240 The authors assert that all procedures contributing to this work comply with the ethical standards of
- 241 the relevant national and institutional committees on human experimentation and with the Helsinki
- 242 Declaration of 1964 and its later amendments.
- 243

# 244 Conflict of interest

245 The authors declare that they have no conflict of interest.

## 246 References

- 247 [1]. Plotkin S. History of vaccination. Proc Natl Acad Sci U S A. 2014;111(34):12283–7.
- [2]. Ozawa S, Mirelman A, Stack ML, Walker DG, Levine OS. Cost-effectiveness and economic benefits of vaccines in low- and middle-income countries: a systematic review. Vaccine. 2012;31(1):96–108.
- [3] Newall AT, Jit M, Hutubessy R. Are current cost-effectiveness thresholds for low- and middle-income countries useful? Examples from the world of vaccines. PharmacoEconomics. 2014;32(6):525–31.
- [4]. WHO vaccine-preventable disease monitoring system, 2017 global summary. Available at: http://www.who.int/immunization/monitoring\_surveillance/data/gs\_gloprofile.pdf
- [5] 2017 SAGE Assessment Report of the Global Vaccine Action Plan. Available at: http://www.who.int/immunization/web\_2017\_sage\_gvap\_assessment\_report\_en.pdf?ua=1
- [6] WHO | Global Vaccine Action Plan 2011-2020. Available at:
   http://www.who.int/immunization/global\_vaccine\_action\_plan/GVAP\_doc\_2011\_2020/en/
- [7]. The Regional Office for Europe of WHO. European Vaccine Action Plan (2015 2020). Available at: http://www.euro.who.int/\_\_data/assets/pdf\_file/0007/255679/WHO\_EVAP\_UK\_v30\_WEBx.pdf
- [8] Nowak GJ, Gellin BG, MacDonald NE, Butler R, SAGE Working Group on Vaccine Hesitancy. Addressing
   vaccine hesitancy: The potential value of commercial and social marketing principles and practices. Vaccine.
   2015;33(34):4204–11.
- [9] Larson HJ, Jarrett C, Eckersberger E, Smith DMD, Paterson P. Understanding vaccine hesitancy around vaccines and vaccination from a global perspective: a systematic review of published literature, 2007-2012. Vaccine.
   2014;32(19):2150–9.
- [10] Dubé E, Vivion M, MacDonald NE. Vaccine hesitancy, vaccine refusal and the anti-vaccine movement: influence, impact and implications. Expert Rev Vaccines. 2015;14(1):99–117.
- [11] Dubé E, Gagnon D, Nickels E, Jeram S, Schuster M. Mapping vaccine hesitancy—Country-specific characteristics of a global phenomenon. Vaccine. 2014;32(49):6649–54.
- [12] MacDonald NE, SAGE Working Group on Vaccine Hesitancy. Vaccine hesitancy: Definition, scope and determinants. Vaccine. 2015;33(34):4161–4.
- [13] Hickler B, Guirguis S, Obregon R. Vaccine Special Issue on Vaccine Hesitancy. Vaccine. 2015;33(34):4155–6.
- [14] Odone A, Signorelli C. When vaccine hesitancy makes headlines. Vaccine. 2017;35(9):1209–10.
- [15] Ministero della Salute. Piano Nazionale Prevenzione Vaccinale (PNPV) 2016-2018. Available at: http://www.adnkronos.com/r/Pub/AdnKronos/Assets/PDF/piano\_prevenzione\_vaccini.pdf
- [16] Ministero della Salute, Ufficio 5. [Internet] Coperture vaccinali a 24 mesi (per 100 abitanti). Anno 2016 (Coorte 2014) [Cited 2018 March 5]. Available at:
- http://www.salute.gov.it/imgs/C\_17\_tavole\_20\_allegati\_iitemAllegati\_0\_fileAllegati\_itemFile\_5\_file.pdf
- [17] Istituto Superiore Sanità, Epicentro [Internet] Copertura vaccinale in Italia. [Cited 2018 March 5] Available at: http://www.epicentro.iss.it/temi/vaccinazioni/dati\_Ita.asp
- [18]. Ministero della Salute [Internet] Vaccinazioni dell'età pediatrica e dell'adolescente Coperture vaccinali [Cited 2018 March 5]. Available
- $at: http://www.salute.gov.it/portale/documentazione/p6\_2\_8\_3\_1.jsp?lingua=italiano&id=20$
- [19] Urgent disposition conerning vaccine prevention. Public Law No.119/17. Available at: http://www.gazzettaufficiale.it/eli/id/2017/08/5/17G00132/sg

- [20] Signorelli C, Iannazzo S, Odone A. The imperative of vaccination put into practice. Lancet Infect Dis.2018;18(1):26–7.
- [21] Forbes TA, McMinn A, Crawford N, Leask J, Danchin M. Vaccination uptake by vaccine-hesitant parents
   attending a specialist immunization clinic in Australia. Hum Vaccines Immunother. 2015;11(12):2895–903.
- [22] Brown KF, Kroll JS, Hudson MJ, Ramsay M, Green J, Long SJ, et al. Factors underlying parental decisions about combination childhood vaccinations including MMR: a systematic review. Vaccine. 2010;28(26):4235–48.
- [23] Weiner JL, Fisher AM, Nowak GJ, Basket MM, Gellin BG. Childhood Immunizations: First-Time Expectant
   Mothers' Knowledge, Beliefs, Intentions, and Behaviors. Am J Prev Med. dicembre 2015;49(6 Suppl 4):S426 434.
- [24] Mollema L, Wijers N, Hahné SJM, van der Klis FRM, Boshuizen HC, de Melker HE. Participation in and attitude towards the national immunization program in the Netherlands: data from population-based questionnaires. BMC Public Health. 2012;12:57.
- [25] Signorelli C, Odone A, Cella P, Iannazzo S, D'Ancona F, Guerra R. Infant immunization coverage in Italy (2000-2016). Ann Ist Super Sanita. 2017;53(3):231–7.
- Scaioli G, Bert F, Galis V, Brusaferro S, De Vito E, La Torre G, et al. Pregnancy and internet: sociodemographic and geographic differences in e-health practice. Results from an Italian multicenter study. Public Health.
   2015;129(9):1258–66.
- 303 [27] Bert F, Passi S, Scaioli G, Gualano MR, Siliquini R. There comes a baby! What should I do? Smartphones'
   304 pregnancy-related applications: A web-based overview. Health Informatics J. 2016;22(3):608–17.
- 305 [28] Tuttitalia.it [Internet] Popolazione per classi di Età Scolastica 2016 Italia. Fonte ISTAT. [Cited 2018 March 5]
   306 Available at: https://www.tuttitalia.it/statistiche/popolazione-eta-scolastica-2016/
- [29] Pourhoseingholi MA, Vahedi M, Rahimzadeh M. Sample size calculation in medical studies. Gastroenterol
   Hepatol Bed Bench. 2013;6(1):14–7.
- 309 [30] Habib A, Johargy A, Mahmood K. Design And Determination Of The Sample Size In Medical Research. IOSR
   310 Journal of Dental and Medical Sciences. 2014;13 (5):21-31.
- [31] Charan J, Biswas T. How to calculate sample size for different study designs in medical research? Indian J
   Psychol Med. aprile 2013;35(2):121–6.
- 313 [32] Hosmer DW, Lemeshow S. Applied Logistic Regression. John Wiley & Sons, Inc.; 2000
- [33] Glanz JM, Wagner NM, Narwaney KJ, Shoup JA, McClure DL, McCormick EV, et al. A mixed methods study of
   parental vaccine decision making and parent-provider trust. Acad Pediatr. 2013;13(5):481–8.
- [34]. Benin AL, Wisler-Scher DJ, Colson E, Shapiro ED, Holmboe ES. Qualitative analysis of mothers' decision making about vaccines for infants: The importance of trust. Pediatrics. 2006;117(5):1532–41.
- [35]. Danchin MH, Costa-Pinto J, Attwell K, Willaby H, Wiley K, Hoq M, et al. Vaccine decision-making begins in pregnancy: Correlation between vaccine concerns, intentions and maternal vaccination with subsequent childhood vaccine uptake. Vaccine. 2017 [Preprint]; Available at: https://doi.org/10.1016/j.vaccine.2017.08.003
- [36] Jarrett C, Wilson R, O'Leary M, Eckersberger E, Larson HJ, SAGE Working Group on Vaccine Hesitancy.
   Strategies for addressing vaccine hesitancy A systematic review. Vaccine. 2015;33(34):4180–90.
- 323 [37] Gualano MR, Bontempi C, Saulle R, Ricciardi W, La Torre G. Validation of the Global Health Professions
   324 Students Survey questionnaire in Italy. Ital J Public Health. 2011;8(4):392–8.
- 325

326