

## Measles Immunity among Fertile Age Women in Kosovo

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### Abstract

The goal of this research was to examine the immunological status of women of fertile age in Kosovo and to evaluate the immune status of children under one year of age, based on the immune status of mothers.

Immunological status with regard to measles was initially investigated through samples of serum from 433 female participants of fertile age (15-49 years) from the Pristina, Prizren and Gjiilan regions. In the second phase of the research, samples from 84 mothers, and their children below one year of age, were tested for inherited immunity to measles.

The mean age of female participants of fertile age was 29.11±7.91 years. 54.7% of them stated that they had been vaccinated with MMR and 29.6% had been infected with measles. The highest vaccination rate, at 91%, was among women aged between 15 and 19. The lowest immunity rate, at 79%, was found among women born between 1979 and 1983. In the analysis of 84 mothers and their children below one year of age, 42 mothers (50%) confirmed that they had been vaccinated, but their immunological status revealed that 71 mothers (84.5 %) had immunity ( $p < 0.05$ ).

Focusing on and targeting unvaccinated females (before they get pregnant) would ensure passive immunisation of their babies as a protective measure, until they reach the measles vaccination age.

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### Introduction

Measles outbreaks are still part of the public health agenda. Despite WHO programmes and efforts to eradicate measles and rubella in five of the World Health Organization (WHO) regions by 2020, that has not yet been achieved.<sup>1</sup> The European Region has had a variety of problems which have caused measles outbreaks in recent years, including a fall in routine immunisation, lower coverage for certain vulnerable communities, and immunity gaps.<sup>2</sup> A decrease in immunity makes it easier for infected people to get infectious disease.<sup>3</sup> In 2019 alone, more than 13,000 cases of measles were

identified in the 30 EU / EEA Member States, more than 10,000 of which were laboratory-confirmed.<sup>4</sup> During the 2000–2015 period alone, an estimated 20.3 million deaths have been prevented due to measles vaccination.<sup>5</sup>

Furthermore, gaps created by conflict in recent decades, especially in the countries of the former Yugoslavia, have led to persistent cases of measles in Europe. European countries also continue to report cases of measles which may result from contacts in Western Balkan countries.<sup>6</sup> Scandinavian nations, such as Sweden, record cases brought by asylum seekers from various countries, including the former Yugoslavia.<sup>7</sup> Population immunity of 92–95% is considered necessary to interrupt measles transmission.<sup>8</sup> Immunity to measles derives from two sources: active immunisation resulting from infection with wild or attenuated measles virus and passive immunization by transplacentally acquired antibody.<sup>9</sup>

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In Kosovo, 610 infected children died as a result of complications during the measles outbreak of the 1960s and 1970s, of which 426 died before the vaccination period (1960-1970) and 184 died after the vaccination period (1971-1981).<sup>10</sup> Kosovo is not in a good position to respond to cases of measles, despite the fact that it has implemented a programme for the elimination of measles and congenital rubella control for the past two decades. Herd immunity was considered established in various population groups, because vaccination coverage was at 95% in Kosovo after the 1999 war, but this was not the case for MMR1 which was at a lower rate of about 80.5%.<sup>11</sup> A study of cases of measles for the period 2017-2018 by ethnicity found that 263 out of 711 (37.1%) of positive cases occurred in the Roma, Ashkali and Egyptian (RAE) community, and 399 out of 711 (56.1%) occurred in other communities living in Kosovo. For 49 (6.9%) of the patients covered by the study, ethnicity was not identified.<sup>12</sup>

In terms of screening activities for the detection of measles immunity, serological surveillance of the population plays a significant role.<sup>13</sup> To the best of our knowledge, no research has so far been carried out in Kosovo regarding the immunological status of women of fertile age. The goal of this research was to examine the immunological status of women of fertile age in Kosovo and to evaluate the immune status of children below one year of age based on the immune status of mothers.

### Materials and methods

The retrospective and prospective epidemiological approaches were used in this study. Using the retrospective approach, the questionnaire designed specifically for this study consists of socio-demographic questions and information on the vaccination status of the participants. Using the prospective approach, the immunological status of measles was investigated through samples of serum from 433 female participants of fertile age (15-49 years) from the Pristina, Prizren and Gjilan regions. In the second phase, samples of 84 mothers and their children below one year of age were tested for inherited immunity to measles. Recruitment of study participants was carried out during their regular visits to primary health care institutions. All participants prior to participating in the

research were informed orally and in writing of the study design and then gave their written consent. Mothers also gave their consent for their children. Samples obtained in the study were analyzed using enzyme-linked immunosorbent assay (ELISA) tests, to determine IgG and IgM.<sup>14</sup> All laboratory analyses of serums were performed at the Department of Microbiology of the National Institute of Public Health of Kosovo.

Continuous variables are summarised by means and standard deviation, while categorical variables are described by frequency and percentage. The Chisquare test was used to compare groups. Statistical analyses were performed with the IBM SPSS v12 software.  $P < 0.05$  was considered significant.

### Ethical approval

The study had been approved by The Ethic Committee of the Faculty of Medicine, University of Prishtina.

### Results

The mean age of female participants of fertile age was  $29.11 \pm 7.91$  years. The distribution of participants by age group is shown in Table 1. Most female participants stated that their occupation was housewife (table 2). 54.7% of them declared that had been vaccinated with MMR, and 29.6% had been infected with measles. The highest vaccination rate, 91%, was among 15-19 year old females, as shown in Table 3. The lowest immunity rate, 79%, was found amongst women born between 1979 and 1983, as shown in Figure 1. In an analysis of 84 mothers (mean age  $27.51 \pm 4.52$  years) and their children below one year of age, 42 (50%) mothers confirmed that they had been vaccinated. Immunological status, however, revealed that 71 out of 84 (84.5 %) had immunity (chi-square;  $p < 0.05$ ) (Table 4).

Age group (years)	Frequency	Percentage (%)
15-19	46	10.6
20-24	88	20.3
25-29	118	27.3
30-34	81	18.7
35-39	46	10.6
40-44	32	7.4
45-49	22	5.1

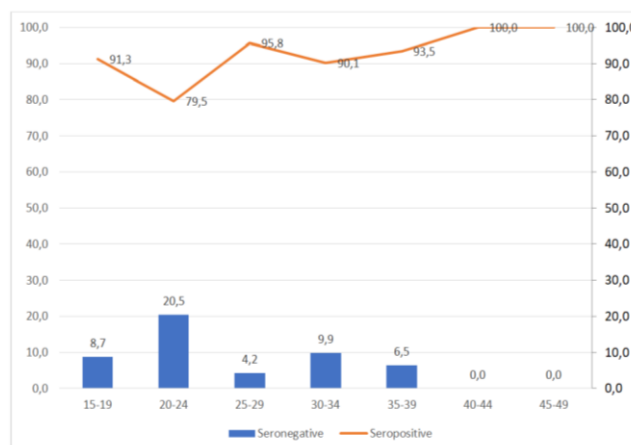
**Table 1.** Fertile age female sample by age groups.

	Frequency	Percentage (%)
Housewife	350	80.8
Economist	4	0.9
Pharmacist	1	0.2
Hairdresser	3	0.7
Nurse	16	3.7
Engineer	1	0.2
Medical doctor	1	0.2
Student	30	6.9
Police officer	2	0.5
Teacher	7	1.6
Lawyer	3	0.7
Technical services	2	0.5
Supervisor/Manager	1	0.2
Sailor	12	2.8

**Table 2.** Occupation of female participats.

Age group (years)		Vaccination Status n(%)	Past infection status n(%)
15-19	Yes	42(91.3)	3(6.5)
	No	4(8.7)	43(93.5)
20-24	Yes	74(84.1)	23(26.1)
	No	14(15.9)	65(73.9)
25-29	Yes	76(64.4)	35(29.7)
	No	42(35.6)	83(70.3)
30-35	Yes	30(3.0)	25(30.9)
	No	51(63.0)	56(69.1)
35-39	Yes	9(19.6)	15(32.6)
	No	37(80.4)	31(67.4)
40-44	Yes	5(15.6)	16(50.0)
	No	27(84.4)	16(50.0)
45-49	Yes	1(4.5)	11(50.0)
	No	21(95.5)	11(50.0)

**Table 3.** Vaccination and past infection status of fertile age females, by age groups.



**Figure 1.** Serology status of respondents who participated in the research, by year of birth.

	Child		p	
	IgG positive n(%)	IgG negative n(%)		
Mother	Seronegative	20 (23.8)	7 (8.3)	0.043
	Seropositive	52 (61.9)	5 (6.0)	

**Table 4.** IgG in children below one year of age, according to maternal immune status.

### Discussion

Despite the WHO goal of eradicating measles by 2020, and despite the existence of a highly effective vaccine, measles continues to persist and has increased in prevalence in recent years. In Kosovo, this makes it very important to understand the characterisation trends and to identify any immunological gaps, considering the volatile political past of the country, particularly since 1988.

The results of this study showed that 259 out of 433 subjects (59.8%) had been vaccinated. Of those born between 1974 and 1988, the majority (76.2%) had been vaccinated, while only 24.2 % of those born between 1954 and 1973 had been vaccinated. Considering that the measles vaccine first came into public use in 1963,<sup>5</sup> this is not an unexpected result in terms of the uptake of the measles vaccine during the period 1954-1973. The percentage of vaccine uptake almost doubles for those born between 1974 and 1978 (64.4%) and increases to 84.1% for those born between 1979 and 1983 and then to 91.3% for those born between 1984 and 1988. By comparison, a study conducted by Vyse et al.

reveals that a significant proportion of those born between 1996 and 1999 had no evidence of a specific antibody against measles, mumps, or rubella in England and Wales.<sup>15</sup>

Although, Dine et al. found in their study that even 26 to 33 years after the first or second dose of measles vaccine, more than 90% of respondents had a high titer of antibodies considered protective for measles.<sup>16</sup>

When tested for their immunological status, those born between 1979 and 1983 had the lowest immunological status with regards to measles, at only 75.5%, although their vaccination status showed that 84.1% of them stated that they had received the measles vaccine. By comparison, in their research Tomaskova et al. found that the overall seroprevalence among citizens of the Czech Republic had reached 93.0%.<sup>17</sup>

In our study, 85.7% of children below one year of age who had not received the vaccine were found to be seropositive. Tomaskova et al. show in their results that only 62% of children at the age of one year are seropositive.<sup>17</sup> Wang et al also observed 71.5% measles seropositivity among infants below 7 months of age in their study.<sup>18</sup>

As such, herd immunity was not reached and this creates pockets for possible measles outbreaks, considering what a highly contagious and serious disease measles is.

Considering inherited immunological vulnerability, the mobility scale and the immunological situation in Europe,<sup>5</sup> policy makers in Kosovo need to decide on best practice, and quickly implement all vaccine-related policies, to prevent any possible outbreaks. Catch-up campaigns would help to improve the sustainability of the immunisation programme and would help the government of Kosovo towards achieving measles eradication.

Endemic transmission is defined as continuous transmission of an indigenous measles virus for a period of 12 months or more in a defined geographical area. The diseases are eliminated when the absence of endemic cases for a period of at least 12 months has been documented by a well-performing surveillance system.<sup>19</sup>

A campaign tailored according to the target population should have realistic expectations. The more people we vaccinate against measles, the closer we will come to the

WHO's measles eradication goal. The more future mothers we vaccinate, the more immune babies will be born, giving them passive protection until they reach the vaccination age, and improving their chances of a healthy and long life.

## Conclusions

This situation underlines the need for strong, reliable and effective surveillance systems that are able to detect and prevent possible outbreaks. Focusing on and targeting unvaccinated females (before they get pregnant) would ensure passive immunisation at their babies as a protective measure, until they reach the measles vaccination age.

## Declaration of Interest

The authors report no conflict of interest.

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