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Transmissibility of Primary Cases Not Completely Vaccinated and Effectiveness of Incomplete Vaccination Against Sars-Cov-2 in Secondary Cases in The Household. Study of 12 Families from February 1 to November 30, 2021 (Before Omicron), in A General Medicine Office in Toledo, Spain: It Is a Very Bad Idea Not to Complete Vaccination

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Abstract

Background: Experts radically changed, during 2021, their opinion on the efficacy of a single dose of vaccine, from being sufficient to being insufficient to reliably prevent infection, but its real impact on the protection of family members is not known.

Objective: Epidemiological evaluation of the transmissibility of SARS-CoV-2 from primary COVID-19 breakthrough infection in not fully vaccinated cases to another family member not fully vaccinated and effectiveness of incomplete vaccination schedule in members of home.

Methodology: An observational, longitudinal and prospective study of families with a primary COVID-19 breakthrough infection in not fully vaccinated people case and at least one secondary case of COVID-19 breakthrough infection in not fully vaccinated family member was conducted, from February 1, 2021, to November 30, 2021 (before omicron), in a general medicine office in Toledo, Spain.

Result: 12 families (with 38 people in total) with a primary case of COVID-19 breakthrough infection in not fully vaccinated people were included. Gross secondary attack rate of covid-19 breakthrough infections in not fully vaccinated people after index cases of covid-19 breakthrough infections in not fully vaccinated people within the family was 100%. Effectiveness of the incomplete vaccine against transmission among household among contacts was 1%.

Conclusion: In the context of general medicine in Toledo (Spain), during 2021 (before omicron), not to complete the vaccination was a very bad idea.

Keywords: COVID-19; SARS-CoV-2; Household contact; Secondary attack rate; Vaccination; Breakthrough infection



Introduction

The COVID-19 pandemic, a public health emergency of international concern, is caused by widespread infection with SARS-CoV-2 and the development of an infectious disease of the respiratory tract [1]. As of December 2020, a single dose of the COVID-19 vaccine was thought to be highly effective (although both doses were advised). So, as the Covid-19 vaccines reached more people, the question arose whether the second dose of the vaccines (from Pfizer, Moderna, Astra) could be delayed allowing more people to get vaccinated more quickly. Experts suggested that people should be calm, since even after a single dose of any of these vaccines, they have very high levels of protection [2]. Before the appearance of the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) variant B.1.617.2 (delta), vaccination reduced the transmission of SARS-CoV-2 from vaccinated persons who became infected, potentially by reducing viral loads. Although vaccination still reduces the risk of infection, similar viral loads in vaccinated and unvaccinated people who are infected with the delta variant call into question the extent to which vaccination prevents transmission [3].

However, since March 2021, the coronavirus has continued to mutate. As of early July 2021, the delta variant had become the most dominant strain of SARS-CoV-2 circulating in the US. Moderna and Pfizer's mRNA vaccines were not specifically designed to prevent the delta variant. While they still generally provide excellent protection after the full two doses [4], new research suggested that a single dose provides less immunity to coronavirus strains than existed in 2021 versus the original strain. Thus, they demonstrated that the emerging Delta variant partially, but remarkably, escaped neutralizing monoclonal antibodies and polyclonal antibodies elicited by previous infection with SARS-CoV-2 or by vaccination [5]. In addition, serum samples from vaccinated individuals neutralized the omicron variant to a much lesser extent than any other variant tested (alpha, beta, or delta) [6].

Consequently, experts modified their advice, suggesting that a single dose of the COVID-19 vaccine from Moderna or Pfizer is not enough to reliably prevent infection [7]. As of December 2021, while in some high-income countries the complete vaccination schedule had already been administered to almost 90% of their population, such as in Spain [8], in low-income countries only slightly more than 6% of the population had received a single vaccine dose [9]. So how effective is a single dose of each of the COVID-19 vaccines? Estimates are controversial and subject to criticism [10] and evidence on the long-term effectiveness of partial vaccination is limited [11]. On the other hand, the transmission of the SARS-CoV-2 virus occurs mainly between people through respiratory droplets expelled through the nose and mouth when coughing, sneezing or talking or when people touch their eyes, nose or mouth after contact with contaminated objects and surfaces [12]. In this context, households are the site of most SARS-CoV-2 transmission globally [13].

Few data exist on the efficacy or lack of efficacy of a single dose of COVID-19 vaccine, and on the transmissibility of SARS-CoV-2 among partially vaccinated persons. An evaluation of the efficacy of the vaccine in a real-world setting is needed. The lack of data is especially relevant in the family/household. In addition, long-term follow-up to assess the long-term efficacy of a single dose is needed to inform a policy of delaying the second dose [14].

In this scenario, this study aims at the epidemiological evaluation of the transmissibility of SARS-CoV-2 from index cases with COVID-19 progression of infection in people not fully vaccinated to another member of the family did not initially infected (assessing the rate of secondary attack) and the effectiveness of the incomplete and complete vaccination schedule at home.

Material and Method

An observational, longitudinal and prospective study of families with a primary COVID-19 breakthrough infection in not fully vaccinated people case and at least one secondary case of COVID-19 breakthrough infection in not fully vaccinated family member was conducted, from February 1, 2021 to November 30, 2021 (before omicron), in a general medicine office in Toledo, Spain, which has a list of 2,000 patients > 14 years of age (in Spain, the general practitioners [GPs] care for people > 14 years of age, except for exceptions requested by the child's family and accepted by the GP). The GPs in Spain work within the National Health System, which is public in nature, and are the gateway for all patients to the system, and each person is assigned a GP [15].

Outcomes of Interest

1. Assess secondary transmission of SARS-CoV-2 from people vaccinated with an incomplete schedule to other family members also with incomplete vaccination. In this sense, the Secondary attack rate of COVID-19 in the family was determined, which was calculated dividing the number of infection events by the person follow-up time [16].
2. Calculate incomplete vaccine and fully vaccine effectiveness (VE) against infection among household contacts of primary COVID-19 breakthrough infection in not fully vaccinated people cases.

Diagnosis of COVID-19 breakthrough infections in vaccinated people

Because the vaccines require about two weeks reaching their maximum effectiveness, a person is not considered fully vaccinated until two weeks after they completed the recommended number of doses for the vaccine they received [17].

To consider a person as fully vaccinated was required [18]

1. That they have received 2 doses of vaccine separated by a minimum of 19 days if the first dose was BNT162b2 mRNA

vaccine (Comirnaty, Pfizer / BioNTech), 21 days in the case of ChAdOx1 nCoV-19 vaccine (Vaxzevria, Oxford / AstraZeneca) or 25 days in the case of mRNA-1273 vaccine (Spikevax, formerly COVID-19 Vaccine Moderna), and that a minimum period of 7 days has elapsed since the last dose if the last dose was with BNT162b2 mRNA vaccine (Comirnaty), or 14 days if it was with ChAdOx1 nCoV-19 vaccine (Vaxzevria) or mRNA-1273 vaccine (Spikevax). People who received a dose of Janssen vaccine (Johnson & Johnson vaccine) more than 14 days ago were also considered fully vaccinated.

2. Or, that having passed the disease they have received a dose of any of the vaccines, after the minimum period equal to that established for the second doses.

3. In the heterologous regimen in which Vaxzevria (Oxford / AstraZeneca) is used in the first dose and mRNA vaccines in the second, it was considered fully vaccinated after 7 days if the second dose was with Comirnaty, or after 14 days if it was with the Moderna vaccine

To consider a person as COVID-19 partially immunized was required: Having received only 1 doses of vaccine [BNT162b2 mRNA vaccine (Comirnaty, Pfizer / BioNTech), or ChAdOx1 nCoV-19 vaccine (Vaxzevria, Oxford / AstraZeneca) or mRNA-1273 vaccine (Spikevax, formerly COVID-19 Vaccine Moderna), and those 14 days had passed after the first dose; but before the second dose. People who received an only dose of Janssen vaccine (Johnson & Johnson vaccine) were considered fully vaccinated [19].

Diagnosis of COVID-19

Diagnosis was performed with reverse transcriptase polymerase chain reaction (PCR) or antigen testing. Rapid antigen tests began to be carried out for symptomatic patients with less than 5 days of evolution from december, 2021. The PCR tests were performed both in symptomatic patients and in asymptomatic contacts. The cases included confirmed cases and asymptomatic carriers. Information on COVID-19 patients and their contacts was obtained from the registry systems used by general medical services in the consultation.

A symptomatic confirmed case with active infection was considered to be any person with a clinical picture of sudden onset acute respiratory infection of any severity that occurs, among others, with fever, cough or feeling of shortness of breath. Other symptoms such as odynophagia, anosmia, ageusia, muscle pain, diarrhea, chest pain or headache, among others, were also considered symptoms of suspected SARS-CoV-2 infection according to clinical criteria; and a positive PCR or rapid antigen test positive [20].

Secondary attack rate

Secondary attack rate was defined as the number of new cases divided by the number of people exposed to a primary case. The

existence of second or third generation cases was not assessed. The cases for the determination of the attack rate included symptomatic cases and asymptomatic cases.

Household contacts

Household contacts were defined as people who shared a residence with the COVID-19 index case. We defined family members as those who had lived with primary cases in a house 4 days before and for more than 24 hours after the primary cases developed illness related to COVID-19. Presumed household transmission via an index case in households was cataloged using the definition of secondary transmission from 1 to 14 days [21]. The onset date of a confirmed case was defined as the date of the first appearance of self-reported clinical symptoms [22]. The onset date for an asymptomatic carrier was defined as the date a positive COVID-19 PCR test was obtained [22].

EV calculation against transmission among household contacts

We calculated the vaccine effectiveness against transmission via the secondary attack rate among close contacts of confirmed index cases:

Secondary attack rate in not fully vaccinated people / Secondary attack rate in not vaccinated people) \times 100 [23, 24].

Collected variables

Primary case or secondary case, age, sex, symptoms, chronic diseases (defined as "any alteration or deviation from normal that has one or more of the following characteristics: is permanent, leaves residual impairment, is caused by a non-reversible pathological alteration, requires special training of the patient for rehabilitation, and / or can be expected to require a long period of control, observation or treatment" [25], classified according to the International Statistical Classification of Diseases and Health-Related Problems, CD-10 Version: 2019 [26], symptom duration in days, social-occupancy class (according to the Registrar General's classification of occupations and social status code) [27, 28], if they were Health Care Workers, problems in the family context and low income household based on the genogram and in the experience of the GP for their continuity of care and knowledge of the family (genogram is a schematic model of the structure and processes of a family, which included the family structure, life cycle and family relational patterns. It was understood that "complex" genograms present families with psychosocial problems) [29-32], number of family members, ethnic minority, symptomatic / asymptomatic COVID-19, time in days since last vaccine dose, severity of the disease (mild cases: clinical symptoms are mild and no manifestation of pneumonia can be found on images; moderate cases: with symptoms such as fever and respiratory tract symptoms, and the manifestation of pneumonia can be seen on the imaging tests; and severe cases: respiratory distress, respiratory rate \geq 30 breaths / min; pulse oxygen saturation \leq 93% with room air at rest;

arterial partial pressure of oxygen / oxygen concentration \leq 300 mmHg) [22] (To simplify comparison, moderate and severe cases were counted together), vaccine type: Comirnaty (Pfizer-BioNTech-BNT162b2 mRNA; Pfizer / BioNTech), Moderna-mRNA-1273 mRNA, Vaxzevria, and Janssen / Johnson & Johnson vaccine (Currently, the European Commission has licensed four vaccines: Comirnaty, Pfizer / BioNTech, licensed December 21, 2020; Moderna vaccine, licensed January 6; AstraZeneca vaccine, licensed 29 December and the Janssen / Johnson & Johnson vaccine, authorized on March 11. In Spain, these four vaccines are currently available, all of which have been approved by the European Medicines Agency [33].

Sample

All families in which a primary case of COVID-19 was diagnosed in a not fully vaccinated person, during the study period, and their all-medical information was available, were included.

Statistical analysis

The bivariate comparisons were performed using the Chi

Square test (X²), X² with Yates correction or Fisher Exact Test when necessary, (according to the number the expected cell totals) for percentages, and the Student test for the mean.

Result

12 families (with 38 people in total) with a primary case of COVID-19 breakthrough infection in not fully vaccinated people were included. Family size was 3.1 \pm 1.4 (arithmetic mean \pm standard deviation), with range: 2-7 people. Of these 12 primary cases, 8 were male and 4 females. There were 21 secondary cases (PCR or positive antigen test) (5 men and 16 women), and 5 negative family contacts. Secondary cases vs. primary cases were statistically significantly more women and had a longer mean time from last vaccination dose to COVID-19 diagnosis. Crude secondary attack rate was 21/26 (81%). Of the 21 secondary cases, 4 had the complete vaccination schedule, 8 had a partial schedule, and 9 were not vaccinated. Of the 5 negative family contacts, 1 was not vaccinated and 4 had a complete vaccination schedule (Figure 1) (Tables 1- 4).

Table 1: Description and Comparison of Incomplete Vaccinated Index Cases and Secondary Cases in The Family.

Variables	Primary Cases (Positive index cases with incomplete vaccination) N=12	Positive Secondary Cases N=21	Statistical Significance
Men	8 (67)	5 (24)	X ² Yates correction is 4.2168. p=.040026. Significant at p < .05.
Men-Age (Arithmetic mean and standard deviation; Range)	43.6 \pm 14.5 (Range: 25-71)	28.2 \pm 21.8 (Range: 3-63)	t= 1.40445. p=.093893. NS
Women	4 (33)	16 (76)	
Women-Age (Arithmetic mean and standard deviation; Range)	32.2 \pm 7.8 (Range: 20-40)	36.6 \pm 19.3 (Range: 3-70)	t= -0.42695. p=.337239. NS
\geq 65 years	1 (8)	2 (9)	Fischer exact test= 1. NS
Children and adolescents \leq 22 years	0	4 (19)	Fisher exact test= 0.2713. NS
Total, Age in years (Arithmetic mean and standard deviation; Range)	39.8 \pm 13.7 (Range: 20-71)	34.6 \pm 20.2 (Range: 3-70)	t= 0.76057. p=.226331. NS
Socio-health workers	2 (17)	1 (5)	Fisher exact test= 0.5381. NS
Social-occupancy class of patients (people with some type of labor specialization)	3 (25)	1 (5)	Fisher exact test= 0.125. NS
Complex family	1 (8)	2 (9)	Fisher exact test= 1. NS
Ethnic minority	3 (25)	4 (19)	Fisher exact test= 0.6856. NS
Low income household	1 (8)	2 (9)	Fisher exact test= 1. NS
Asymptomatic	0 (by definition)	9 (43)	NR
Symptomatic	N=12--all symptomatic by definition	12 (57)	NR
Moderate-severe severity	2 (9) (2 pneumonias)	0	Fisher exact test= 0.125. NS
Symptom duration in days (Arithmetic mean and standard deviation; Range)	6.4 \pm 4.7 (Range: 2-20)	(N=12 symptomatic)	t= 1.14345. p=.132572. NS
		4.4 \pm 3.7 (Range: 1-15)	
Incubation period in days (Arithmetic mean and standard deviation; Range)	— No disponible	(N=12)	NA
		4.4 \pm 3.5 (Range: 1-12)	
Previous COVID-19	0	0	NP
Presence of chronic diseases	3 (25)	4 (19)	Fisher exact test= 0.6856. NS

(): Denotes percentages; NS: Not significant; NR: Not relevant; NA: Not available

Table 2: Comparison of Symptoms in Incomplete Vaccinated Index Cases and Secondary Cases in The Family.

Symptoms *According to Who, Icd-10 Groups	Primary Cases (Positive index cases with incomplete vaccination) N=12	Positive Secondary Cases N=21	Statistical Significance
General (discomfort, asthenia, myalgia, fever, artralgiias)	13 (48)	15 (68)	X2= 1.9867. p= .158689. NS
Respiratory (cough, dyspnea, chest pain)	4 (15)	4 (18)	X2 with Yates correction= 0.0051. p= .943108. NS
ENT (Anosmia / ageusia,odynophagia, rhinorrhea, pharyngeal dryness-mucus, epixtasis)	7 (26)	3 (14)	X2 with Yates correction= 0.4975. p= .480583. NS
Digestive (anorexia, nausea / vomiting, diarrhea, abdominal pain)	2 (7)	0	Fisher exact test= 0.4949. NS
Neurological (headache, dizziness, mental confusion -brain fog)	1 (4)	0	Fisher exact test= 1. NS
Psychiatric (Anxiety, insomnia)	0	0	Fisher exact test= 1. NS
Skin (chilblains, flictenas, rash)	0	0	Fisher exact test= 1. NS
Total, symptoms*	27 (100)	22 (100)	--

(): Denotes percentages; NS: Not significant; * Patients could have more than one symptom. The percentages are over the total of symptoms

Table 3: Comparison of Chronic Diseases in Incomplete Vaccinated Index Cases and Secondary Cases in The Family.

Chronic Diseases* According To Who, Icd-10 Groups	Primary Cases (Positive index cases with incomplete vaccination) N=12	Positive Secondary Cases N=21	Statistical Significance
-I Infectious	0	0	Fisher exact test= 1. NS
-II Neoplasms	1 (11)	0	Fisher exact test= 1. NS
-III Diseases of the blood	1 (12)	0	Fisher exact test= 1. NS
-IV Endocrine	2 (22)	1 (11)	Fisher exact test= 1. NS
-V Mental	1 (11)	1 (11)	Fisher exact test= 1. NS
-VI-VIII Nervous and Senses	1 (11)	1 (11)	Fisher exact test= 1. NS
-IX Circulatory system	0	0	Fisher exact test= 1. NS
-X Respiratory system	0	0	Fisher exact test= 1. NS
-XI Digestive system	2 (22)	1 (11)	Fisher exact test= 1. NS
-XII Diseases of the skin	0	0	Fisher exact test= 1. NS
-XIII Musculo-skeletal	0	2 (22)	Fisher exact test= 0.4706. NS
-XIV Genitourinary	1 (11)	3 (34)	Fisher exact test= 0.5765. NS
-XVII Congenital malformations	0	0	Fisher exact test= 1. NS
-XIX Injury, poisoning and certain other consequences of external causes	0	0	Fisher exact test= 1. NS
-XXI Factors influencing health status	0	0	Fisher exact test= 1. NS
TOTAL, chronic diseases*	9 (100)	9 (100)	---

(): Denotes percentages; * Patients could have more than one chronic disease. The percentages are over the total of chronic diseases; NS: Not significant

Table 4: Comparison of Vaccination Data in Incomplete Vaccinated Index Cases and Secondary Cases in The Family.

Vaccination Data	PRIMARY CASES (Positive Index Cases With Incomplete Vaccination) N=12	Positive Secondary Cases N=21	Statistical Significance
Fully vaccinated	0 (100) [by definition]	21-Apr	NR
Vaccinated one dose or without fulfilling adequate time from last dose to positive test	12 (100) [by definition]	21-Aug	NR
Completely vaccinated, or with a dose or without complying with adequate time from last dose to positive test	12 (100) [by definition]	12/21 (57)	NR
Not vaccinated	0 (100) [by definition]	9/21 (43)	NR

Time in days from last vaccine dose to positive COVID-19 test	14.9+-17.0 (Range: 1-67)	(N=12 positive vaccinated)	t= -2.02819. p= .027409. The result is significant at p < .05.
(Arithmetic mean and standard deviation; Range)		40.4 +- 38.0 (Range: 1-120)	
VACCINE TYPES			
BNT162b2 mRNA vaccine (Comirnaty, Pfizer / BioNTech)	7 (58)	6 (50)	X2= 0.1678. p= .682046. NS
mRNA-1273 vaccine (Spikevax, formerly COVID-19 Vaccine Moderna)	1 (8)	1 (8)	Fisher exact test= 1. NS
2ChAdOx1 nCoV-19 vaccine (Vaxzevria, Oxford / AstraZeneca)	3 (25)	4 (33)	X2 with Yates correction= 0. p= 1. NS
Janssen (Johnson & Johnson vaccine) una sola dosis se consideró completa!	1 (8)	1 (8)	Fisher exact test= 1. NS
TOTAL	12 (100)	12 (100)	---

(): Denotes percentages; NS: Not significant; NR: Not relevant

Table 5: Secondary Attack Rate Among Contacts in The Family of Confirmed Index Cases with Covid-19 Breakthrough Infections in Partially (Not Fully) Vaccinated People and Vaccine Effectiveness Against Infection.

Secondary Cases Of Covid-19 Breakthrough Infections In Fully Vaccinated People After Index Cases Of Covid-19 Breakthrough Infections In Not Fully Vaccinated People Within The Family	Gross Secondary Attack Rate	Secondary Cases Of Covid-19 Breakthrough Infections In Partially Vaccinated People After Index Cases Of Covid-19 Breakthrough Infections In Not Fully Vaccinated People Within The Family	Gross Secondary Attack Rate	Secondary Cases Of Covid-19 Breakthrough Infections In Partially Vaccinated People Or No Vaccinated After Index Cases Of Covid-19 Breakthrough Infections In Not Fully Vaccinated People Within The Family	Gross Secondary Attack Rate
4/8	50%	8/8	100%	17/18	94%

Vaccine effectiveness against transmission among household contacts not fully vaccinated = (Secondary attack rate in not fully vaccinated people/ Secondary attack rate in unvaccinated people) × 100 = 100 / 94 (1%)

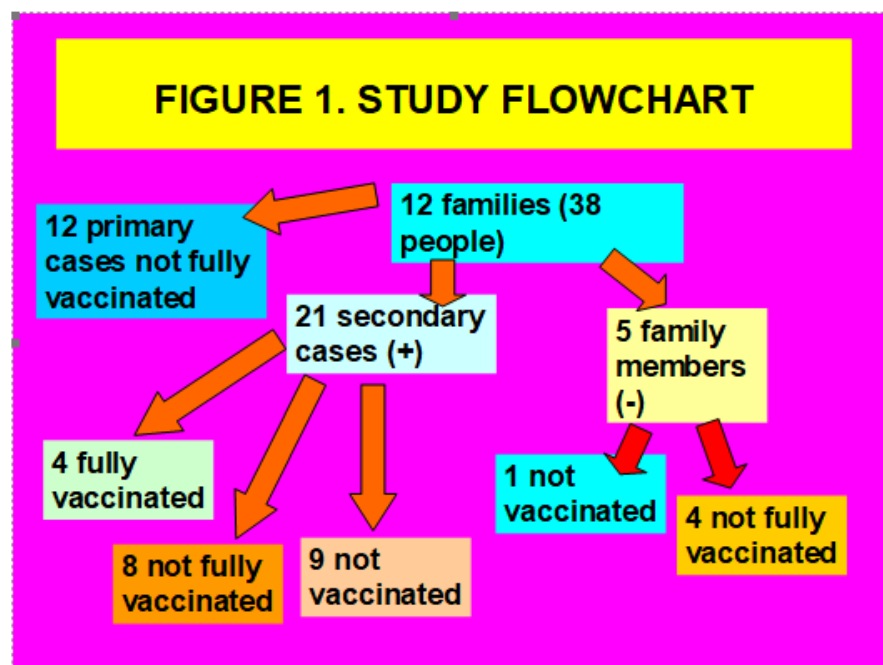


Figure 1: Study Flowchart.

Gross secondary attack rate of COVID-19 breakthrough infections in fully vaccinated people after index cases of covid-19 breakthrough infections in not fully vaccinated people within the family was 50%. Gross secondary attack rate of COVID-19 breakthrough infections in not fully vaccinated people after index cases of COVID-19 breakthrough infections in not fully vaccinated people within the family was 100%. Gross secondary attack rate of covid-19 breakthrough infections in no vaccinated people after index cases of COVID-19 breakthrough infections in not fully vaccinated people within the family was 94%. VE against transmission among household contacts not fully vaccinated was $[(\text{Secondary attack rate in not fully vaccinated people} / \text{Secondary attack rate in not vaccinated people}) \times 100] 100/94 = 1\%$ (Table 5).

Discussion

Real-world evidence from vaccine rollout programs, through the end of 2021, has shown that full COVID-19 vaccinations are highly effective against severe illness, hospitalization, and death from SARS-CoV-2, including B.1.1.7 (alfa) and B.1.617.2 (delta), and reduce both asymptomatic infection and intra-household transmission by reducing the progressive spread of people who have become infected despite vaccination [3, 34].

How second booster doses of vaccines work

When the immune system first encounters a vaccine, it activates two important types of white blood cells. First of all there are the plasmatic B cells, which are mainly focused on the production of antibodies. Unfortunately, this type of cell has a short life, without the second injection, a rapid decline occurs. Then there are T cells that can stay in the body for decades until they hit their target, meaning immunity to vaccinations or infections can sometimes last a lifetime; but usually not many of these cells are produced until the second dose. The booster dose is a way of re-exposing the body to antigens that activate the immune system, to initiate the second part of the response. On the second exposure to the same vaccine, the B cells can divide rapidly and create a second spike in the amount of circulating antibodies. The second dose also kickstarts the B-cell maturation process, which involves selecting the immature ones with the best receptors to bind to a particular pathogen. Meanwhile, memory T cells also proliferate rapidly [10].

Duration in time of protection after the first dose

There are no clear data showing that protection after the first dose is maintained after 21 days. It is possible that the protection that people seem to have suddenly disappears after that point. Reliably calculating how long protection from a single dose might last is further complicated by the fact that all currently approved Covid-19 vaccines use entirely new technology [10]. In our study, secondary cases vs. primary cases had a statistically significant longer mean (arithmetic mean) time from last vaccination dose to COVID-19 diagnosis (40 vs. 14 days).

VE of partial immunization (14 days after the first dose but before the second dose)

The story of Colin Horseman, 85, one of the first people in the world to receive the initial dose of a Covid-19 vaccine (Pfizer-BioNTech) in the UK, who died of COVID-19 less than three weeks later, has been published. He received that first dose; he must have received the second dose two days before his death. In fact, most vaccines require booster doses to work: around 40% of people who have received a single dose of the MMR (measles, mumps and rubella) vaccine are not protected from all three viruses, compared to 4 % of those who have received the second; People in the first group are four times more likely to contract measles than those in the second, and there have been outbreaks in places where a high proportion of people have not completed the full MMR vaccination schedule.

The minimum acceptable level of efficacy of a given vaccine is 50% in preventing COVID-19 as indicated by the World Health Organization [35] and the US Food and Drug Administration (36). In this sense, the figures communicated in the different published data regarding the VE of a single dose of COVID-19 vaccine are confusing and variable according to the definition of effectiveness (number of people who tested positive for the virus, prevention of symptomatic disease, prevention of severe disease, hospitalization), population, type of vaccine and predominant variant of the virus, but in general they are between 36% and 80% [10, 14, 37,45]. We found a gross secondary attack rate of COVID-19 breakthrough infections in not fully vaccinated people after index cases of COVID-19 breakthrough infections in not fully vaccinated people within the family of 100%.

Prevention of transmission of SARS-CoV-2

EV was largely tested by looking at whether they prevented people from developing symptoms, not whether they prevented them from becoming infected with the virus. And we know that it is possible to have an asymptomatic infection. There is no evidence yet that one dose, or even two, of the existing vaccines will stop people from transmitting the virus to other people [3,10].

After a single dose, the SARS-CoV-2 vaccines Vaxzevria (AstraZeneca) and BNT162b2 mRNA vaccine (Comirnaty, Pfizer / BioNTech) impact differently on the peripheral immune compartment. Although both vaccines elicited the induction of spike-specific effector cells and spike binding antibodies, the different compilation of these immunological features suggests that the strategy for spike delivery impacts on how and to what extent immune priming against the main SARS-CoV-2 antigen proceeds [46]. There is no evidence that any of the current COVID-19 vaccines can completely prevent people from becoming infected, and this has implications for our prospects of achieving herd immunity.

If vaccines don't completely stop transmission, it will increase the number of people we need to vaccinate to truly cross herd immunity thresholds and bring cases down to a level close to zero.

For example, one calculation suggests that for a vaccine that totally eliminates transmission, 60-72% of the population would need to have it to achieve full herd immunity. But if the vaccine was 80% effective, 75-90% of people would need it. However, most of scientists do not expect eliminate virus; for the time being, the goal is to reduce its transmission as much as possible [10]. In our study, VE against transmission among household contacts not fully vaccinated $[(\text{Secondary attack rate in not fully vaccinated people} / \text{Secondary attack rate in not fully vaccinated people}) \times 100]$ was 1%.

Could some vaccines drive the evolution of more virulent pathogens?

Conventional wisdom is that natural selection will eliminate highly lethal pathogens if host death greatly reduces transmission. Vaccines that keep hosts alive but still allow transmission could allow highly virulent strains to circulate in a population. When vaccines prevent transmission, as is the case with almost all vaccines used in humans, this type of evolution toward greater virulence is blocked. But when vaccines leak, allowing at least some transmission of pathogens, they could create the ecological conditions that allow hot strains to emerge and persist. Therefore, the use of leaky vaccines may facilitate the evolution of pathogen strains that put unvaccinated hosts at increased risk of severe disease [47]. In any case, what is certain is that the appearance of variants that avoid vaccines, will be more likely through viral propagation and replication [48].

Can the behavior change after receiving a single dose?

The fact of receiving a single dose can make people think that they will be sufficiently protected, and encourage risky behavior. It should be very clear with the message to be given, which should include two key concepts: You are not going to be protected; and there is no evidence that having received the vaccine prevents you from contracting the virus and transmitting it [10].

Limitations and strengths of the study

1. It must be taken into account that the changes in community transmission during the study period may also imply changes in one direction or another in cautious and personal protection behaviors, which could be an element of confusion in interpreting the results
2. The study may have missed asymptomatic cases that did not attend in GP consultation, as no surveillance or systematic screening was done.
3. There is potential for misclassification of household transmission if the secondary case infection was actually acquired outside the household or if the true household index case was not assessed.

Conclusion

In the context of general medicine in Toledo (Spain), gross secondary attack rate of COVID-19 breakthrough infections in not

fully vaccinated people after index cases of COVID-19 breakthrough infections in not fully vaccinated people within the family was 100%. Por otra parte, VE against transmission among not fully vaccinated household contacts $[(\text{Secondary attack rate in not fully vaccinated people} / \text{Secondary attack rate in not vaccinated people}) \times 100]$ was 1%. Consequently it is a very bad idea not to complete the vaccination.

Acknowledgement

None.

Conflict of Interest

No conflict of interest.

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