OVERVIEW Measles Vaccine
at the Razi Vaccine and Serum Research Institute

[^0]
## Current Products Portfolio

- Measles , Rubella, Mumps, MR, MMR
- OPV (Bopv1,3 and MOPV1)
- DTP, DT, TT
- ppd
- Sera : ATS, ADS,
- Polyvalent Anti Snake \& Scorpion


## Measles Virus

## Paramyxovirus（RNA）

One antigenic type

Hemagglutinin important surface antigen

Rapidly inactivated by heat and light

| Top 10** |  |  |
| ---: | ---: | :--- |
| Country | Cases | Rate |
| Madagascar | 151032 | 6066.87 |
| Ukraine | 78708 | 1771.16 |
| Philippines | 49419 | 478.31 |
| India**** | 36251 | 27.38 |
| Nigeria | 27954 | 150.3 |
| Brazil | 18927 | 91.15 |
| Kazakhstan | 10696 | 594.63 |
| DR Congo | 9245 | 117.42 |
| Yemen | 9156 | 331.93 |
| Thailand | 7738 | 112.37 |


| Other countries with high incidence <br> rates*** |  |  |
| ---: | ---: | ---: |
| Country | Cases | Rate |
| Georgia | 4710 | 1199.88 |
| The Republic of <br> North Macedonia | 1902 | 913.89 |
| Kyrgyzstan | 2839 | 476.68 |
| Israel | 3442 | 420.17 |
| Bosnia and <br> Herzegovina | 1399 | 397.8 |
| New Zealand | 1672 | 358.73 |



| Measles cases from countries with known discrepancies between case-based and aggregate surveillance, as reported by |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: |
| country |  |  |  |



Figure 37.2. Worldwide distribution of measles virus genotypes, March 2014 to February 2015.

Measles Vaccines

1963 Live attenuated and killed vaccines
1965 Live further attenuated vaccine
1967 Killed vaccine withdrawn
1968 Live further attenuated vaccine
(Edmonston-Enders strain)
1971 Licensure of combined measles-mumps-rubella vaccine
1989 Two dose schedule



Figure 37.5. Incidence of fever and rash after natural measles infection and vaccination. The number of susceptibles is included in parentheses. Ig, immunoglobulin. (Modified from Krugman S, Giles JP, Jacobs AM, et al. Studies with a further attenuated live measles-virus vaccine. Pediatrics. 1963;31:919-928.)

The lowest clinical reactions to the AIK-C vaccine were compared
to those of 5 other measles vaccine strain

TABLE 37.3 Measles Vaccine Manufacturers and Vaccine Strains Produced as of 2015

| Manufacturer | Vaccine Strain |
| :---: | :---: |
| Merck (United States) | Moraten |
| SanofiPasteur (France) | Schwarz |
| GlaxoSmithKline (Eelgium) | Sclmwarz |
| Govemment Pharmaceutical Organization (Theriland) | Sclrwarz |
| EioManguinhos (Brazil) | Scltwarz |
| Takeda Pharnarceutical Comparny (Japan) | Schwarz Fess |
| Senumi Institute of India | Edmonston-Zagreb |
| Institute of Immmonology (Croatia) | Edmonston-Zagreb |
| Fesearch Foundation for Microbial Diseases of Osaka University (Japan) | CAM-70 |
| EioFarma (Indonesia) | CAM-70 |
| Kitasato Phammaceutical Industry (Uapan) | AIK-C ${ }^{3}$ |
| The Rlazi State Serum linstitute (Iran) | AK-HDC/Schwarz |
| Moscow Plant of Biological Preparations (Russia) | Leningrad-16 |
| Beiling Tiantan Institute of Eiological Products (China) | Shanghai-191 |

Beijing Minhai Biotechnology Co., Ltd. (China) Chang-47

TABLE 40.10 Incidence of Postvaccine Aseptic Meningitis Following Vaccination With MMR Vaccines Containing the Urabe Am9

## Mumps Virus Strain

| Manufacturer | Country (Dates Covered by Study) | Rate (N) | Type of Study and Surveillance |
| :---: | :---: | :---: | :---: |
| SANOFI PASTEUR <br> Rebiere et $\mathrm{al}^{468}$ | France (1991-93) | 1/28,400 (1/65,750) ${ }^{\text {a }}$ | Retrospective: capture-recapture method |
| Joinville-Bera et al ${ }^{469}$ | France (1989-92) | 1/121,951 | Retrospective: passive surveillance; 4 of 54 cases with laboratory confirmation of mumps virus in CSF |
| Dos Santos et $\mathrm{al}^{311}$ | Brazil (1996) | 0/2179 | Randomized double-blind clinical trial in children 6 to 12 years old |
| GLAXOSMITHKLINE Furesz et al ${ }^{470}$ | Canada (1986-90) | 1/62,000 | Retrospective: passive surveillance; 8 cases with virologic confirmation |
| Dourado et $\mathrm{al}^{471}$ | Brazil (1997) | 1/14,00O (452,344) | Prospective: passive surveillance of hospital admissions for meningitis; 32 cases identified |
| Miller et al ${ }^{180}$ | UK (1991-92) | 1/12,400 (49,585) | Retrospective: computerized hospital records for children ages 12-23 months with meningitis; 4 cases identified |
| SANOFI PASTEUR OR Miller et al ${ }^{472}$ | AXOSMITHKLINE ${ }^{\text {D }}$ <br> UK (1990-91) | 1/11,OOO (78,300) | Retrospective: CSF evaluation in children discharged with diagnosis of meningitis within 15 to 35 days after MMR vaccination; 4 of 13 cases virus positive |
| Farrington et $\mathrm{al}^{473}$ | UK (1988-93) | 1/15,000 (77,200) | Retrospective: linkage of vaccination records and hospital discharge for meningitis; 5 cases identified |
| Japan Ministry of Fujinaga et $\mathrm{al}^{474}$ | ALTH AND WELFARE Japan (1989) | BIKEN $1 / 336(11,750)$ | Retrospective: passive surveillance; 35 cases identified |
| Sugiura et $\mathrm{al}^{475}$ | Japan (1989) | 1/6,564 (630,157) | Retrospective: surveillance by physicians; 96 cases with viral confirmation |
| Ueda et $\mathrm{al}^{367}$ | Japan (1990-93) | 1/905 (5430) | Prospective: active surveillance by parents; 6 cases identified |
| Ueda et $\mathrm{al}^{367}$ | Japan (1990-93) | 0/566 (566) | Prospective: active surveillance by parents; physician confirmed |
| MANUFACTURER NOT Maguire et $\mathrm{al}^{476}$ | PECIFIED <br> UK (1990-91) | 1/61,000 (est. 1,650,000) | Prospective: passive surveillance by physicians with active follow up by laboratory or clinical confirmation; 7 definite cases, 17 probable cases |
| Colville et $\mathrm{al}^{477}$ | UK (1988-91) | 1/3,800 doses | Retrospective: 6 virologically confirmed cases in a small population |
| Al Mazrou et al ${ }^{478}$ | Saudi Arabia (2000) | 1/295,000 (est. 2,000,000) | Prospective: active surveillance following booster vaccination in a "catch-up" campaign; 6 cases confirmed |
| ${ }^{\text {a Rate of }} 1 / 28,400$ is based on an estimate of 116 cases of postvaccine aseptic meningitis by the capture-recapture method. The actual number of observed cases was 46 , and no virus isolation was carried out. The rate calculation using the number of observed cases is $1: 65,750$. <br> DAt the time of the study, Urabe Am9 vaccines manufactured by both Sanofi Pasteur (then Aventis Pasteur) and GlaxoSmithKline were licensed, and it is not possible to relate occurrence of postvaccine aseptic meningitis to a specific vaccine. <br> CSF, cerebrospinal fluid; MMR, measles, mumps, rubella; UK, United Kingdom. |  |  |  |

## Measles Vaccine

Time line vaccine In IRAN
development of the live attenuated measles vaccine by Enders and his colleagues in 1960

1. 2. measles vaccination in 1967
1. 2. first local production in 1967 at the Razi Institute using the Sugiyama strain
1. 3. By the end of 1972, approximately 5 million children received this vaccination

By the end of 1975, over 7.5 million children from 9 months to 5 years of age were vaccinated
From 1967 to 1976, the Razi institute produced more than 12,000,000 dose vaccine
Development of a New Measles Vaccine in Iran

1. in 1973 at the Kitasato Institute in Japan with cooperation of IRAN (Razi) marked the beginning of research that made available the new live attenuated measles vaccine (MV), AIK-C. The strain was named AIK to represent the three countriesthat collaborated to develop it.
$90 \%$ of children develop protective antibody levels when given 1 dose of MVat 9 months of
age, whereas $90-95 \%$ respond when vaccinated at 12 monthS
From the time of the vaccine licensure in 1963 until the end of 2020 , over 120 million doses of the measles vaccine were produced at the Razi institute in Iran. annual incidence of measles over the period of 1967-1976 was 150,000-500,000, and the mortality was 5-10\% The successful and high-quality local measles vaccine resulted in < 1 case per million residents during 2004-2009. Recent surveys have revealed a greater than $92 \%$ seropositivity in Iranian children

- 2003 : Iranian campaign : World's largest vaccination operation. Measles and Rubella (MR) vaccine was administered to more than 33 million people aged between 5 and 25 years, in less than one month : Decline in measles incidence due to the vaccination. Ref: Bahmmen Pourabbas eal efficac of meases snd ubbell vaccination one year after the nationwide campaign in Shiraz, Iran; Int.ernational Journal of Infectious Diseases (2008) 12, 43-46.

> MR vaccine was used extensively in mass immunization campaigns in various countries in Latin America, Southern Europe, Central Asia, East Asia and Western Asia

## The elimination of measles in Iran

In 2019, when the Eastern Mediterranean region WHO was experiencing its greatest upsurge in measles cases, Iran received a certificate for measles elimination in October 2019. Iran, with a population of approximately 82 million, is the third country in the Eastern Mediterranean region that has achieved the status of measles elimination, after Oman and Bahrain, both of which have a relatively small population. However, without sustained attention

Iran also received the certification for rubella elimination in May, 2019. The number of rubella cases had dramatically decreased from 1154 cases in 2000, to 33 cases in 2018.

## Impact of Covid-19

## MASS VACCINATION CAMPAIGNS - $\mathbf{3}$ MONTH-

 PAUSE:Campaigns (polio, measles..) put on hold in > 35 countries

At least $\mathbf{8 0}$ million children under one at risk of polio, measles, diphtheria, etc..
POLIO RESOURCES REPURPOSED TO SUPPORT RESPONSE TO COVID-19
PLANNING NOW TO






[^0]:    R. MOR EZZA TAGHIZADEH

