

A Guide To **Differential Diagnosis** Selected Rash/Illness In Children

M-M-R® II

Measles, Mumps,
& Rubella Virus
Vaccine Live

VARIVAX®

VARICELLA VIRUS
VACCINE LIVE

ProQuad®

Measles, Mumps,
Rubella and
Varicella Virus
Vaccine Live

measles rubeola¹



Courtesy of CDC.

Courtesy of CDC.

Rash: The measles rash is a maculopapular eruption that usually begins at the hairline, then involves the face and spreads to the trunk and extremities, reaching the hands and feet. Lesions are generally discrete, but may become confluent, particularly on the upper body. The rash usually lasts about 5 to 6 days, and fades first on the areas first involved.²

Other clinical manifestations: The illness usually begins with a prodrome generally lasting 2 to 4 days that is characterized by fever followed by cough, coryza, or conjunctivitis. Fever can reach 105°F. One diagnostic sign is Koplik spots—tiny bluish-white specks on a bright red base—on the mucous membranes of the mouth.²

Indications and Usage

ProQuad is a vaccine indicated for active immunization for the prevention of measles, mumps, rubella, and varicella in children 12 months through 12 years of age.

VARIVAX is a vaccine indicated for active immunization for the prevention of varicella in individuals 12 months of age or older.

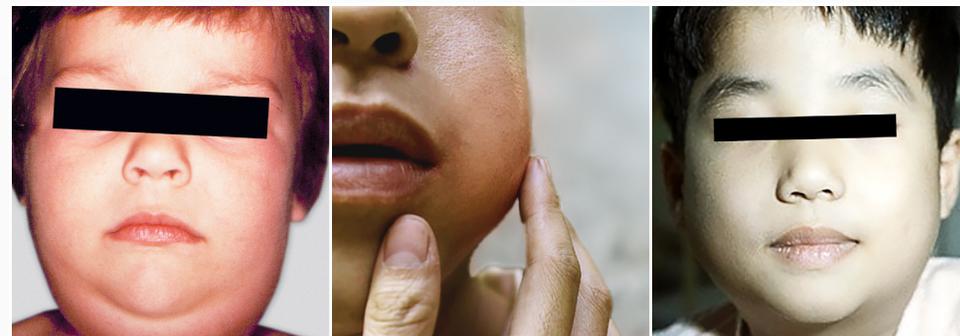
M-M-R® II is indicated for active immunization for the prevention of measles, mumps, and rubella in individuals 12 months of age or older.

Selected Safety Information

• **Hypersensitivity:** **ProQuad**, **M-M-R® II**, and **VARIVAX** are contraindicated in patients with a history of anaphylactic reaction or hypersensitivity to any component of the vaccine (including gelatin or neomycin) or to a prior dose of measles, mumps, rubella, or varicella-containing vaccine. Use caution when administering **ProQuad** and **M-M-R® II** to individuals with anaphylaxis or immediate hypersensitivity to eggs.

- **ProQuad**, **M-M-R® II**, and **VARIVAX** are contraindicated in certain individuals, including those with: immunodeficiency or who are immunosuppressed; an active febrile illness; untreated tuberculosis.
- **Pregnancy:** **ProQuad**, **M-M-R® II**, and **VARIVAX** are contraindicated for use in pregnant women. Do not administer **ProQuad** or **VARIVAX** to individuals who are planning to become pregnant in the next 3 months. Do not administer **M-M-R® II** to individuals who are planning to become pregnant in the next month.
- **Febrile Seizures:** Administration of **ProQuad** (dose 1) to children 12 to 23 months old who have not been previously vaccinated against measles, mumps, rubella, or varicella, nor had a history of the wild-type infections, is associated with higher rates of fever and febrile seizures at 5 to 12 days after vaccination when compared to children vaccinated with a first dose of both **M-M-R® II** and **VARIVAX** administered concomitantly.
- **Febrile Seizures:** Use caution when administering **ProQuad** and **M-M-R® II** to individuals with a history of febrile seizures.

mumps³



Courtesy of CDC.

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Rash: None, typically.³

Clinical manifestations: Mumps typically presents as parotitis (ie, swelling of the parotid gland) or other salivary gland swelling lasting about 5 days. Parotitis may be unilateral or bilateral, and swelling of any combination of single or multiple salivary glands may be present. Parotitis may first be noted as earache and tenderness on palpation of the angle of the jaw. Prodromal symptoms are nonspecific and can include low-grade fever, headache, myalgia, anorexia, and malaise.³

- **Family History of Immunodeficiency:** Vaccination with **ProQuad**, **M-M-R® II**, and **VARIVAX** should be deferred in individuals with a family history of congenital or hereditary immunodeficiency until the individual's immune status has been evaluated and the individual has been found to be immunocompetent.
- **Thrombocytopenia:** Transient thrombocytopenia has been reported within 4-6 weeks following vaccination with measles, mumps, and rubella vaccine. Carefully evaluate the potential risk and benefit of vaccination in children with thrombocytopenia or in those who experienced thrombocytopenia after vaccination with a previous dose of a measles, mumps, and rubella-containing vaccine.
- **Varicella Transmission and Precautions:** Advise vaccinees administered **ProQuad** or **VARIVAX** to avoid: close contact with high-risk individuals susceptible to varicella for up to 6 weeks following vaccination since transmission of varicella vaccine virus to susceptible contacts has been reported. Varicella vaccine virus transmission may occur between vaccine recipients and contacts susceptible to varicella including healthy individuals.

Selected Safety Information continues on next page.

rubella German measles⁴



Courtesy of CDC.

Courtesy of CDC.

Courtesy of CDC.

Rash: The rubella rash usually first occurs on the face and spreads down the body. It is maculopapular, may be pruritic, and generally lasts about 3 days. The rash is fainter than measles rash and does not coalesce.⁴

Other clinical manifestations: Symptoms in younger children are often mild and prodrome is rare. In older children, there may be a 1- to 5-day prodrome with low-grade fever, malaise, posterior lymphadenopathy, and upper respiratory symptoms preceding the rash. Arthralgia and arthritis are rare in children.⁴

varicella chickenpox⁵



Courtesy of CDC.

Courtesy of CDC.

Courtesy of CDC.

Rash: The rash is generalized and pruritic and progresses rapidly from macules to papules to vesicular lesions before crusting.⁵ Early in the disease, lesions appear at all stages.⁶ The rash usually appears first on the head, or on the trunk, and then the extremities; the highest concentration of lesions is on the trunk. Lesions can occur on mucous membranes throughout the body. Healthy children usually have 250 to 500 lesions in 2 to 4 successive crops.⁵ Crusts completely fall off within 1 to 2 weeks after the onset of infection.⁶

Other clinical manifestations:

The clinical course generally includes malaise, pruritus, and fever up to 102°F for 2 to 3 days. In many children, rash is the first symptom.⁵

CDC, Centers for Disease Control and Prevention.

Selected Safety Information (continued)

- **Immune Globulins and Transfusions:** Immune Globulins and other blood products should not be given concomitantly with **ProQuad**, **M-M-R[®]_{II}**, or **VARIVAX**.
- **Use of Salicylates:** Avoid use of salicylates in children and adolescents administered **ProQuad** or **VARIVAX** for 6 weeks following vaccination due to the association of Reye Syndrome with salicylate therapy and wild-type varicella infection.
- **Adverse Events:** The following adverse events have been reported for both subcutaneous and intramuscular injections of **ProQuad**, **M-M-R[®]_{II}**, and **VARIVAX**: fever, injection-site reactions (pain/tenderness/soreness, erythema, and swelling); and rash on the body or at the injection site. Additionally, irritability has been reported for the subcutaneous injections of **ProQuad**, **M-M-R[®]_{II}**, and **VARIVAX**.
- **ProQuad Systemic Vaccine-Related Adverse Events:** Systemic vaccine-related adverse events that were reported at a significantly

greater rate in recipients of subcutaneous **ProQuad** than in recipients of the component vaccines administered concomitantly were fever and measles-like rash.

- **VARIVAX Dose-related Adverse Events:** In a clinical trial involving children who received 2 doses of **VARIVAX** 3 months apart, the incidence of injection-site clinical complaints observed in the first 4 days following vaccination was slightly higher post-dose 2 (overall incidence 25.4%) than post-dose 1 (overall incidence 21.7%), whereas the incidence of systemic clinical complaints in the 42-day follow-up period was lower post-dose 2 (66.3%) than post-dose 1 (85.8%).
- **Concomitant Vaccines With ProQuad:** **ProQuad** may be administered concomitantly with diphtheria and tetanus toxoids and acellular pertussis vaccine adsorbed (DTaP), *Haemophilus influenzae* type b conjugate (meningococcal protein conjugate) and hepatitis B (recombinant) vaccine. It may also be administered concomitantly

with pneumococcal 7-valent conjugate vaccine and/or hepatitis A vaccine (inactivated) at separate injection sites.

- **Concomitant Vaccines With VARIVAX:** **VARIVAX** can be administered with other live viral vaccines. If not given concurrently, at least 1 month should elapse between a dose of a live attenuated measles virus-containing vaccine and a dose of **VARIVAX**. In children, at least 3 months should elapse between administration of 2 doses of a live attenuated varicella virus-containing vaccine.
- **Tuberculin Testing:** If a tuberculin test is to be done with **M-M-R[®]_{II}** and **ProQuad**, it should be administered either any time before, simultaneously with, or at least 4 to 6 weeks after vaccination. With **VARIVAX**, tuberculin testing may be performed before the vaccine is administered or at least 4 weeks following vaccination.

Selected Safety Information continues on next page.



REFERENCES:

1. Centers for Disease Control and Prevention. Measles Symptoms and Complications. Updated May 2024. Accessed May 2024. <https://www.cdc.gov/measles/signs-symptoms/> 2. Centers for Disease Control and Prevention. Epidemiology and Prevention of Vaccine-Preventable Diseases; The Pink Book: Measles. Updated August 2021. Accessed May 2024. <https://www.cdc.gov/vaccines/pubs/pinkbook/meas.html> 3. Centers for Disease Control and Prevention. Epidemiology and Prevention of Vaccine-Preventable Diseases; The Pink Book: Mumps. Updated August 2021. Accessed May 2024. <https://www.cdc.gov/vaccines/pubs/pinkbook/mumps.html> 4. Centers for Disease Control and Prevention. Epidemiology and Prevention of Vaccine-Preventable Diseases; The Pink Book: Rubella. Updated August 2021. Accessed May 2024. <https://www.cdc.gov/vaccines/pubs/pinkbook/rubella.html> 5. Centers for Disease Control and Prevention. Epidemiology and Prevention of Vaccine-Preventable Diseases; The Pink Book: Varicella. Updated September 2021. Accessed May 2024. <https://www.cdc.gov/vaccines/pubs/pinkbook/varicella.html> 6. Whitley RJ. Chickenpox and Herpes Zoster (Varicella-Zoster Virus). In: Bennett JE, Dolin R, Blaser MJ, eds. Mandell, Douglas, and Bennett's Principles and Practice of Infectious Diseases. 9th ed. Philadelphia, PA. Saunders, Elsevier; 2020;2:1849-1856.

Selected Safety Information *(continued)*

- **Additional M-M-R[®] II Precautions:** Additional adverse reactions, which have been reported without regard to causality, include febrile convulsions, arthritis, thrombocytopenia, anaphylaxis, anaphylactoid reactions, arthritis, encephalitis and encephalopathy in their diverse clinical presentations.
- **Additional VARIVAX Precautions:** It is not known if varicella vaccine virus is excreted in human milk. A boost in antibody levels has been observed in vaccinees following exposure to wild-type varicella, which could account for the apparent long-term persistence of antibody levels in studies. The duration of protection from varicella infection after vaccination is unknown.
- **ProQuad/VARIVAX and Herpes Zoster:** The long-term effect of VARIVAX on the incidence of herpes zoster, particularly in those vaccinees exposed to wild-type varicella, is unknown at present.
- **Efficacy:** Vaccination with ProQuad, VARIVAX, or M-M-R[®] II may not result in protection in 100% of vaccinees.

Dosage and Administration

ProQuad:

- Each dose of ProQuad is approximately 0.5 mL and is administered intramuscularly or subcutaneously.
- At least 1 month should elapse between a dose of a measles-containing vaccine such as M-M-R[®] II and a dose of ProQuad. At least 3 months should elapse between a dose of varicella-containing vaccine and ProQuad.

VARIVAX:

- Each dose is approximately 0.5 mL and is administered intramuscularly or subcutaneously.
 - The first dose is administered at 12 to 15 months of age.
 - The second dose is administered at 4 to 6 years of age.
 - There should be a minimum interval of 3 months between doses.
 - 12 months to 12 years of age: If a second dose is administered, there should be a minimum interval of 3 months between doses.
 - Adolescents (≥13 years of age) and Adults: 2 doses, to be administered with a minimum interval of 4 weeks between doses.

M-M-R[®] II:

- The dose for any age is approximately 0.5 mL administered intramuscularly or subcutaneously.
 - The recommended age for primary vaccination is 12 to 15 months and the second dose should be given at 4 to 6 years of age.

Before administering VARIVAX[®] (Varicella Virus Vaccine Live), M-M-R[®] II (Measles, Mumps, and Rubella Virus Vaccine Live), or ProQuad[®] (Measles, Mumps, Rubella and Varicella Virus Vaccine Live), please read the accompanying Prescribing Information. The Patient Information also is available for VARIVAX, M-M-R[®] II, and ProQuad.