



**FACT SHEET FOR HEALTHCARE PROVIDERS ADMINISTERING  
VACCINE (VACCINATION PROVIDERS)  
EMERGENCY USE AUTHORIZATION (EUA) OF  
THE MODERNA COVID-19 VACCINE TO PREVENT CORONAVIRUS DISEASE 2019  
(COVID-19)**

**PRIMARY SERIES PRESENTATION  
6 months through 5 years of age**

The U.S. Food and Drug Administration (FDA) has issued an Emergency Use Authorization (EUA) to permit the emergency use of the unapproved product, Moderna COVID-19 Vaccine, for active immunization to prevent COVID-19 in individuals 6 months of age and older.

This Fact Sheet pertains only to Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border (see image below) which is authorized for use to provide a two-dose primary series to individuals 6 months through 5 years of age. The vaccine is also authorized to provide a third primary series dose to individuals 6 months through 5 years of age who have been determined to have certain kinds of immunocompromise.<sup>1</sup>



 703819	STORE FROZEN between -50° to -15°C (-58° to 5°F). Protect from light. No preservative. After first use, hold at 2° to 25°C (36° to 77°F). Discard after 12 hours. Record date/time of first use: _____	<b>Moderna COVID-19 Vaccine</b> Suspension for Intramuscular Injection For use under Emergency Use Authorization <b>Age 6mo through 5y</b> Vial contains 10 doses of 0.25 mL	NDC 80777-279-05
	 Scan here for FDA-authorized Fact Sheet for dosage and administration, and product expiration dates, or visit <a href="http://www.modernatx.com/covid19vaccine-eua/">www.modernatx.com/covid19vaccine-eua/</a> Mfd. for: Moderna US, Inc., Cambridge, MA 02139		

Moderna COVID-19 Vaccine supplied in multiple-dose vials with a dark blue cap and a label with a magenta border intended for use in individuals 6 months through 5 years of age should not be used in individuals 6 years of age and older because of the potential for

<sup>1</sup> Certain kinds of immunocompromise refers to individuals who have undergone solid organ transplantation, or who are diagnosed with conditions that are considered to have an equivalent level of immunocompromise.

**vaccine administration errors, including dosing errors.<sup>2,3</sup>**

## **SUMMARY OF INSTRUCTIONS FOR COVID-19 VACCINATION PROVIDERS**

Vaccination providers enrolled in the federal COVID-19 Vaccination Program must report all vaccine administration errors, all serious adverse events, cases of myocarditis, cases of pericarditis, cases of Multisystem Inflammatory Syndrome (MIS) in adults and children, and cases of COVID-19 that result in hospitalization or death following administration of the Moderna COVID-19 Vaccine. See “MANDATORY REQUIREMENTS FOR MODERNA COVID-19 VACCINE ADMINISTRATION UNDER EMERGENCY USE AUTHORIZATION” for reporting requirements.

The Moderna COVID-19 Vaccine is a suspension for intramuscular injection.

The Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border is administered as a primary series of two doses (0.25 mL each) 1 month apart to individuals 6 months through 5 years of age.

A third primary series dose (0.25 mL) of the Moderna COVID-19 Vaccine supplied in a multiple dose vial with a dark blue cap and a label with a magenta border is authorized for administration at least 1 month following the second dose to individuals 6 months through 5 years of age with certain kinds of immunocompromise.

See this Fact Sheet for instructions for preparation and administration. This Fact Sheet may have been updated. For the most recent Fact Sheet, please see [www.modernatx.com/covid19vaccine-eua](http://www.modernatx.com/covid19vaccine-eua).

For information on clinical trials that are testing the use of the Moderna COVID-19 Vaccine for active immunization against COVID-19, please see [www.clinicaltrials.gov](http://www.clinicaltrials.gov).

## **DESCRIPTION OF COVID-19**

Coronavirus disease 2019 (COVID-19) is an infectious disease caused by the novel coronavirus, SARS-CoV-2, that appeared in late 2019. It is predominantly a respiratory illness that can affect other organs. People with COVID-19 have reported a wide range of symptoms, ranging from mild symptoms to severe illness. Symptoms may appear 2 to 14 days after exposure to the virus.

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<sup>2</sup> Notwithstanding the age limitations for use of the different presentations of the Moderna COVID-19 Vaccine, individuals who will turn from 5 years to 6 years of age between doses in the primary series may receive, for any dose in the primary series, either: (1) the Moderna COVID-19 Vaccine authorized for use in individuals 6 months through 5 years of age (each 0.25 mL dose containing 25 mcg mRNA) supplied in multiple-dose vials with dark blue caps and labels with a magenta border; (2) the Moderna COVID-19 Vaccine authorized for use in individuals 6 years through 11 years of age (each 0.5 mL dose containing 50 mcg mRNA) supplied in multiple-dose vials with dark blue caps and labels with a purple border stating “BOOSTER DOSES ONLY”; or (3) the Moderna COVID-19 Vaccine authorized for use in individuals 6 years through 11 years of age (each 0.5 mL dose containing 50 mcg mRNA) supplied in multiple-dose vials with dark blue caps and labels with a teal border (currently not available). The multiple-dose vials with dark blue caps and labels with a purple border are authorized to provide 0.5 mL primary series doses for individuals 6 years through 11 years of age.

<sup>3</sup> For primary vaccination of individuals 6 years through 11 years of age and 12 years of age and older, refer to the respective Moderna COVID-19 Vaccine Fact Sheet for Healthcare Providers Administering Vaccine.

Symptoms may include: fever or chills; cough; shortness of breath; fatigue; muscle and body aches; headache; new loss of taste or smell; sore throat; congestion or runny nose; nausea or vomiting; diarrhea.

## **DOSAGE AND ADMINISTRATION**

### **Storage and Handling**

The information in this Fact Sheet supersedes the information on the vial and carton labels.

During storage, minimize exposure to room light, and avoid exposure to direct sunlight and ultraviolet light.

#### Frozen Storage

Store frozen between -50°C to -15°C (-58°F to 5°F).

#### Storage after Thawing

- Storage at 2°C to 8°C (36°F to 46°F):
  - Vials may be stored refrigerated between 2°C to 8°C (36°F to 46°F) for up to 30 days prior to first use.
  - Vials should be discarded 12 hours after the first puncture.
- Storage at 8°C to 25°C (46°F to 77°F):
  - Vials may be stored between 8°C to 25°C (46°F to 77°F) for a total of 24 hours.
  - Vials should be discarded 12 hours after the first puncture.
  - Total storage at 8°C to 25°C (46°F to 77°F) must not exceed 24 hours.

### **Do not refreeze once thawed.**

Thawed vials can be handled in room light conditions.

#### Transportation of Thawed Vials at 2°C to 8°C (36°F to 46°F)

If transport at -50°C to -15°C (-58°F to 5°F) is not feasible, available data support transportation of one or more thawed vials for up to 12 hours at 2°C to 8°C (36°F to 46°F) when shipped using shipping containers which have been qualified to maintain 2°C to 8°C (36°F to 46°F) and under routine road and air transport conditions with shaking and vibration minimized. Once thawed and transported at 2°C to 8°C (36°F to 46°F), vials should not be refrozen and should be stored at 2°C to 8°C (36°F to 46°F) until use.

### **Dose and Schedule**

The Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border is administered as a primary series of two doses (0.25 mL each) 1 month apart to individuals 6 months through 5 years of age.

A third primary series dose (0.25 mL) of the Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border is authorized for

administration at least 1 month following the second dose to individuals 6 months through 5 years of age with certain kinds of immunocompromise.

### Preparation for Administration

- The Moderna COVID-19 Vaccine multiple-dose vial with a dark blue cap and a label with a magenta border is supplied as a frozen suspension that does not contain a preservative and must be thawed prior to administration.
- Verify that the vial of Moderna COVID-19 Vaccine has a dark blue cap and a label with a magenta border.
- Thaw each vial before use following the instructions below.

### Thawing Instructions for Moderna COVID-19 Vaccine Multiple-Dose Vials with Dark Blue Caps and Labels with a Magenta Border

Thaw in Refrigerator	Thaw at Room Temperature
Thaw between 2°C to 8°C (36°F to 46°F) for 2 hours. Let each vial stand at room temperature for 15 minutes before administering.	Alternatively, thaw between 15°C to 25°C (59°F to 77°F) for 45 minutes.

- After thawing, do not refreeze.
- Swirl vial gently after thawing and between each withdrawal. **Do not shake.** Do not dilute the vaccine.
- Parenteral drug products should be inspected visually for particulate matter and discoloration prior to administration, whenever solution and container permit.
- The Moderna COVID-19 Vaccine is a white to off-white suspension. It may contain white or translucent product-related particulates. Do not administer if vaccine is discolored or contains other particulate matter.
- Each multiple-dose vial with a dark blue cap and a label with a magenta border contains 10 primary series doses of 0.25 mL each; low dead-volume syringes and/or needles can be used to extract 10 doses from a single vial. If standard syringes and needles are used, there may not be sufficient volume to extract 10 doses from a single vial. Irrespective of the type of syringe and needle:
  - Each dose must contain 0.25 mL of vaccine;
  - If the amount of vaccine remaining in the vial cannot provide a full dose of 0.25 mL, discard the vial and content;
  - Do not pool excess vaccine from multiple vials.
- After the first 0.25 mL dose has been withdrawn, the vial should be held between 2°C to 25°C (36°F to 77°F). Record the date and time of first use on the Moderna COVID-19 Vaccine vial label. Discard vial after 12 hours. Do not refreeze.

### Administration

Administer the Moderna COVID-19 Vaccine intramuscularly.

### CONTRAINDICATION

Do not administer the Moderna COVID-19 Vaccine to individuals with a known history of a

severe allergic reaction (e.g., anaphylaxis) to any component of the Moderna COVID-19 Vaccine (*see Full EUA Prescribing Information*).

## **WARNINGS**

### **Management of Acute Allergic Reactions**

Appropriate medical treatment to manage immediate allergic reactions must be immediately available in the event an acute anaphylactic reaction occurs following administration of the Moderna COVID-19 Vaccine.

Monitor Moderna COVID-19 Vaccine recipients for the occurrence of immediate adverse reactions according to the Centers for Disease Control and Prevention (CDC) guidelines (<https://www.cdc.gov/vaccines/covid-19/clinical-considerations/managing-anaphylaxis.html>).

### **Myocarditis and Pericarditis**

Postmarketing data demonstrate increased risks of myocarditis and pericarditis, particularly within 7 days following the second dose. The observed risk is highest in males 18 through 24 years of age. Although some cases required intensive care support, available data from short-term follow-up suggest that most individuals have had resolution of symptoms with conservative management. Information is not yet available about potential long-term sequelae. The CDC has published considerations related to myocarditis and pericarditis after vaccination, including for vaccination of individuals with a history of myocarditis or pericarditis (<https://www.cdc.gov/vaccines/covid-19/clinical-considerations/myocarditis.html>).

### **Syncope**

Syncope (fainting) may occur in association with administration of injectable vaccines. Procedures should be in place to avoid injury from fainting.

### **Altered Immunocompetence**

Immunocompromised persons, including individuals receiving immunosuppressant therapy, may have a diminished immune response to the Moderna COVID-19 Vaccine.

### **Limitations of Vaccine Effectiveness**

The Moderna COVID-19 Vaccine may not protect all vaccine recipients.

## **ADVERSE REACTIONS**

### *Adverse Reactions in Clinical Trials*

Adverse reactions in individuals 6 months through 23 months of age following administration of the primary series included irritability/crying, pain at the injection site, sleepiness, loss of appetite, fever, swelling at the injection site, erythema at the injection site, and axillary (or groin) swelling/tenderness. (*See Full EUA Prescribing Information*)

Adverse reactions in individuals 24 months through 36 months of age following administration of the primary series included pain at the injection site, irritability/crying, sleepiness, loss of appetite, fever, erythema at the injection site, swelling at the injection site, and axillary (or groin)

swelling/tenderness. *(See Full EUA Prescribing Information)*

Adverse reactions in individuals 37 months through 5 years of age following administration of the primary series included pain at the injection site, fatigue, headache, myalgia, fever, chills, nausea/vomiting, axillary (or groin) swelling/tenderness, arthralgia, erythema at the injection site, and swelling at the injection site. *(See Full EUA Prescribing Information)*

#### *Adverse Reactions in Post-Authorization Experience*

Anaphylaxis and other severe allergic reactions, myocarditis, pericarditis, syncope, and urticaria have been reported following administration of the Moderna COVID-19 Vaccine during post-authorization use.

Additional adverse reactions, some of which may be serious, may become apparent with more widespread use of the Moderna COVID-19 Vaccine.

### **USE WITH OTHER VACCINES**

There is no information on the co-administration of the Moderna COVID-19 Vaccine with other vaccines.

### **INFORMATION TO PROVIDE TO VACCINE RECIPIENTS/CAREGIVERS**

As the vaccination provider, you must communicate to the recipient or their caregiver, information consistent with the “Fact Sheet for Recipients and Caregivers” (and provide a copy or direct the individual to the website [www.modernatx.com/covid19vaccine-eua](http://www.modernatx.com/covid19vaccine-eua) to obtain the Fact Sheet) prior to the individual receiving each dose of the Moderna COVID-19 Vaccine, including:

- FDA has authorized the emergency use of the Moderna COVID-19 Vaccine, which is not an FDA-approved vaccine.
- There is an option to accept or refuse the Moderna COVID-19 Vaccine.
- The significant known and potential risks and benefits of the Moderna COVID-19 Vaccine, and the extent to which such risks and benefits are unknown.
- Information about available alternative vaccines and the risks and benefits of those alternatives.

For information on clinical trials that are evaluating the use of the Moderna COVID-19 Vaccine to prevent COVID-19, please see [www.clinicaltrials.gov](http://www.clinicaltrials.gov).

Provide a vaccination card to the recipient or their caregiver with the date when the recipient needs to return for the second dose of Moderna COVID-19 Vaccine.

Provide the **v-safe** information sheet to vaccine recipients/caregivers and encourage vaccine recipients to participate in **v-safe**. **V-safe** is a voluntary smartphone-based tool that uses text messaging and web surveys to check in with people who have been vaccinated to identify potential side effects after COVID-19 vaccination. **V-safe** asks questions that help CDC monitor the safety of COVID-19 vaccines. **V-safe** also provides dose reminders if needed and live telephone follow-up by CDC if participants report a significant health impact following COVID-19 vaccination. For more information, visit: [www.cdc.gov/vsafe](http://www.cdc.gov/vsafe).



## **MANDATORY REQUIREMENTS FOR MODERNA COVID-19 VACCINE ADMINISTRATION UNDER EMERGENCY USE AUTHORIZATION<sup>4</sup>**

In order to mitigate the risks of using this unapproved product under EUA and to optimize the potential benefit of the Moderna COVID-19 Vaccine, the following items are required. Use of unapproved Moderna COVID-19 Vaccine for active immunization to prevent COVID-19 under this EUA is limited to the following (all requirements **must** be met):

1. The Moderna COVID-19 Vaccine is authorized for use in individuals 6 months of age and older.
2. The vaccination provider must communicate to the individual receiving the Moderna COVID-19 Vaccine or their caregiver information consistent with the “Fact Sheet for Recipients and Caregivers” prior to the individual receiving the Moderna COVID-19 Vaccine.
3. The vaccination provider must include vaccination information in the state/local jurisdiction’s Immunization Information System (IIS) or other designated system.
4. The vaccination provider is responsible for mandatory reporting of the following to the Vaccine Adverse Event Reporting System (VAERS):
  - vaccine administration errors whether or not associated with an adverse event,
  - serious adverse events\* (irrespective of attribution to vaccination),
  - cases of myocarditis,
  - cases of pericarditis,
  - cases of Multisystem Inflammatory Syndrome (MIS) in adults and children, and
  - cases of COVID-19 that result in hospitalization or death.

Complete and submit reports to VAERS online at <https://vaers.hhs.gov/reportevent.html>. For further assistance with reporting to VAERS, call 1-800-822-7967. The reports should include the words “Moderna COVID-19 Vaccine EUA” in the description section of the report.

5. The vaccination provider is responsible for responding to FDA requests for information about vaccine administration errors, adverse events, cases of myocarditis, cases of pericarditis, cases of MIS in adults and children, and cases of COVID-19 that result in hospitalization or death following administration of the Moderna COVID-19 Vaccine to recipients.

\*Serious adverse events are defined as:

- Death;
- A life-threatening adverse event;
- Inpatient hospitalization or prolongation of existing hospitalization;

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<sup>4</sup> Vaccination providers administering SPIKEVAX (COVID-19 Vaccine, mRNA) must adhere to the same reporting requirements.

- A persistent or significant incapacity or substantial disruption of the ability to conduct normal life functions;
- A congenital anomaly/birth defect;
- An important medical event that based on appropriate medical judgement may jeopardize the individual and may require medical or surgical intervention to prevent one of the outcomes listed above.

#### **OTHER ADVERSE EVENT REPORTING TO VAERS AND MODERNATX, INC.**

Vaccination providers may report to VAERS other adverse events that are not required to be reported using the contact information above.


To the extent feasible, report adverse events to ModernaTX, Inc. using the contact information below or by providing a copy of the VAERS form to ModernaTX, Inc.

Email	Fax number	Telephone number
<a href="mailto:ModernaPV@modernatx.com">ModernaPV@modernatx.com</a>	1-866-599-1342	1-866-MODERNA (1-866-663-3762)

#### **ADDITIONAL INFORMATION**

For general questions, visit the website or call the telephone number provided below.

To access the most recent Moderna COVID-19 Vaccine Fact Sheets, please scan the QR code or visit the website provided below.

Website	Telephone number
<a href="http://www.modernatx.com/covid19vaccine-eua">www.modernatx.com/covid19vaccine-eua</a> 	1-866-MODERNA (1-866-663-3762)

#### **AVAILABLE ALTERNATIVES**

There may be clinical trials or availability under EUA of other COVID-19 vaccines.

#### **FEDERAL COVID-19 VACCINATION PROGRAM**

This vaccine is being made available for emergency use exclusively through the CDC COVID-19 Vaccination Program (the Vaccination Program). Healthcare providers must enroll as providers in the Vaccination Program and comply with the provider requirements. Vaccination providers may not charge any fee for the vaccine and may not charge the vaccine recipient any out-of-pocket charge for administration. However, vaccination providers may seek appropriate reimbursement from a program or plan that covers COVID-19 vaccine administration fees for the vaccine recipient (private insurance, Medicare, Medicaid, HRSA COVID-19 Uninsured Program for non-insured recipients). For information regarding provider requirements and



enrollment in the CDC COVID-19 Vaccination Program, see <https://www.cdc.gov/vaccines/covid-19/provider-enrollment.html>.

Individuals becoming aware of any potential violations of the CDC COVID-19 Vaccination Program requirements are encouraged to report them to the Office of the Inspector General, U.S. Department of Health and Human Services, at 1-800-HHS-TIPS or TIPS.HHS.GOV.

### **AUTHORITY FOR ISSUANCE OF THE EUA**

The Secretary of the Department of Health and Human Services (HHS) has declared a public health emergency that justifies the emergency use of drugs and biological products during the COVID-19 Pandemic. In response, the FDA has issued an EUA for the unapproved product, Moderna COVID-19 Vaccine, for active immunization to prevent COVID-19.

FDA issued this EUA, based on ModernaTX, Inc.'s request and submitted data.

For the authorized uses, although limited scientific information is available, based on the totality of the scientific evidence available to date, it is reasonable to believe that the Moderna COVID-19 Vaccine may be effective for the prevention of COVID-19 in individuals as specified in the *Full EUA Prescribing Information*.

This EUA for the Moderna COVID-19 Vaccine will end when the Secretary of HHS determines that the circumstances justifying the EUA no longer exist or when there is a change in the approval status of the product such that an EUA is no longer needed.

For additional information about Emergency Use Authorization, visit FDA at: <https://www.fda.gov/emergency-preparedness-and-response/mcm-legal-regulatory-and-policy-framework/emergency-use-authorization>.

### **COUNTERMEASURES INJURY COMPENSATION PROGRAM**

The Countermeasures Injury Compensation Program (CICP) is a federal program that has been created to help pay for related costs of medical care and other specific expenses to compensate people injured after use of certain medical countermeasures. Medical countermeasures are specific vaccines, medications, devices, or other items used to prevent, diagnose, or treat the public during a public health emergency or a security threat. For more information about CICP regarding the vaccines to prevent COVID-19, visit <http://www.hrsa.gov/cicp>, email [cicp@hrsa.gov](mailto:cicp@hrsa.gov), or call: 1-855-266-2427.

Moderna US, Inc.  
Cambridge, MA 02139

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Patent(s): [www.modernatx.com/patents](http://www.modernatx.com/patents)

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END SHORT VERSION FACT SHEET

Long Version (Full EUA Prescribing Information) Begins On Next Page

# **FULL EMERGENCY USE AUTHORIZATION (EUA) PRESCRIBING INFORMATION**

## **MODERNA COVID-19 VACCINE**

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## **FULL EMERGENCY USE AUTHORIZATION (EUA) PRESCRIBING INFORMATION**

### **1 AUTHORIZED USE**

Moderna COVID-19 Vaccine is authorized for use under an Emergency Use Authorization (EUA) for active immunization to prevent coronavirus disease 2019 (COVID-19) caused by severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) in individuals 6 months of age and older.

This EUA Prescribing Information pertains only to Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border, which is authorized for use in individuals 6 months through 5 years of age.

### **2 DOSAGE AND ADMINISTRATION**

For intramuscular injection only.

The storage, preparation, and administration information in this EUA Prescribing Information apply to the Moderna COVID-19 Vaccine for individuals 6 months through 5 years of age, which is supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border.

## 2.1 Preparation for Administration

- The Moderna COVID-19 Vaccine multiple-dose vial with a dark blue cap and a label with a magenta border is supplied as a frozen suspension that does not contain a preservative and must be thawed prior to administration.
- Verify that the vial of Moderna COVID-19 Vaccine has a dark blue cap and a label with a magenta border.
- Thaw each vial before use following the instructions below.

### Thawing Instructions for Moderna COVID-19 Vaccine Multiple-Dose Vials with Dark Blue Caps and Labels with a Magenta Border

Thaw in Refrigerator	Thaw at Room Temperature
Thaw between 2°C to 8°C (36°F to 46°F) for 2 hours. Let each vial stand at room temperature for 15 minutes before administering.	Alternatively, thaw between 15°C to 25°C (59°F to 77°F) for 45 minutes.

- After thawing, do not refreeze.
- Swirl vial gently after thawing and between each withdrawal. **Do not shake.** Do not dilute the vaccine.
- Parenteral drug products should be inspected visually for particulate matter and discoloration prior to administration, whenever solution and container permit.
- The Moderna COVID-19 Vaccine is a white to off-white suspension. It may contain white or translucent product-related particulates. Do not administer if vaccine is discolored or contains other particulate matter.
- Each multiple-dose vial with a dark blue cap and a label with a magenta border contains 10 primary series doses of 0.25 mL each; low dead-volume syringes and/or needles can be used to extract 10 doses from a single vial. If standard syringes and needles are used, there may not be sufficient volume to extract 10 doses from a single vial. Irrespective of the type of syringe and needle:
  - Each dose must contain 0.25 mL of vaccine;
  - If the amount of vaccine remaining in the vial cannot provide a full dose of 0.25 mL, discard the vial and content;
  - Do not pool excess vaccine from multiple vials.
- After the first 0.25 mL dose has been withdrawn, the vial should be held between 2°C to 25°C (36°F to 77°F). Record the date and time of first use on the Moderna COVID-19 Vaccine vial label. Discard vial after 12 hours. Do not refreeze.

## 2.2 Administration

Administer the Moderna COVID-19 Vaccine intramuscularly.

## 2.3 Dose and Schedule

The Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a

label with a magenta border is administered as a primary series of two doses (0.25 mL each) 1 month apart to individuals 6 months through 5 years of age.

A third primary series dose (0.25 mL) of the Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border is authorized for administration at least 1 month following the second dose to individuals 6 months through 5 years of age with certain kinds of immunocompromise.

### **3 DOSAGE FORMS AND STRENGTHS**

Moderna COVID-19 Vaccine is a suspension for injection. Each dose of the Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border for individuals 6 months through 5 years of age is 0.25 mL.

### **4 CONTRAINDICATIONS**

Do not administer the Moderna COVID-19 Vaccine to individuals with a known history of severe allergic reaction (e.g., anaphylaxis) to any component of the Moderna COVID-19 Vaccine [*see Description (13)*].

### **5 WARNINGS AND PRECAUTIONS**

#### **5.1 Management of Acute Allergic Reactions**

Appropriate medical treatment to manage immediate allergic reactions must be immediately available in the event an acute anaphylactic reaction occurs following administration of the Moderna COVID-19 Vaccine.

Monitor Moderna COVID-19 Vaccine recipients for the occurrence of immediate adverse reactions according to the Centers for Disease Control and Prevention (CDC) guidelines (<https://www.cdc.gov/vaccines/covid-19/clinical-considerations/managing-anaphylaxis.html>).

#### **5.2 Myocarditis and Pericarditis**

Postmarketing data demonstrate increased risks of myocarditis and pericarditis, particularly within 7 days following the second dose. The observed risk is highest in males 18 through 24 years of age. Although some cases required intensive care support, available data from short-term follow-up suggest that most individuals have had resolution of symptoms with conservative management. Information is not yet available about potential long-term sequelae. The CDC has published considerations related to myocarditis and pericarditis after vaccination, including for vaccination of individuals with a history of myocarditis or pericarditis (<https://www.cdc.gov/vaccines/covid-19/clinical-considerations/myocarditis.html>).

#### **5.3 Syncope**

Syncope (fainting) may occur in association with administration of injectable vaccines.

Procedures should be in place to avoid injury from fainting.

#### **5.4 Altered Immunocompetence**

Immunocompromised persons, including individuals receiving immunosuppressive therapy, may have a diminished response to the Moderna COVID-19 Vaccine.

#### **5.5 Limitations of Vaccine Effectiveness**

The Moderna COVID-19 Vaccine may not protect all vaccine recipients.

### **6 OVERALL SAFETY SUMMARY**

**It is MANDATORY for vaccination providers to report to the Vaccine Adverse Event Reporting System (VAERS) all vaccine administration errors, all serious adverse events, cases of myocarditis, cases of pericarditis, cases of Multisystem Inflammatory Syndrome (MIS) in adults and children, and hospitalized or fatal cases of COVID-19 following vaccination with the Moderna COVID-19 Vaccine.<sup>5</sup> To the extent feasible, provide a copy of the VAERS form to ModernaTX, Inc. Please see the REQUIREMENTS AND INSTRUCTIONS FOR REPORTING ADVERSE EVENTS AND VACCINE ADMINISTRATION ERRORS section for details on reporting to VAERS and ModernaTX, Inc.**

In a clinical study, the adverse reactions in participants 6 months through 23 months of age following administration of the primary series were irritability/crying (81.5%), pain at the injection site (56.2%), sleepiness (51.1%), loss of appetite (45.7%), fever (21.8%), swelling at the injection site (18.4%), erythema at the injection site (17.9%), and axillary (or groin) swelling/tenderness (12.2%).

In a clinical study, the adverse reactions in participants 24 months through 36 months of age following administration of the primary series were pain at the injection site (76.8%), irritability/crying (71.0%), sleepiness (49.7%), loss of appetite (42.4%), fever (26.1%), erythema at the injection site (17.9%), swelling at the injection site (15.7%), and axillary (or groin) swelling/tenderness (11.5%).

In a clinical study, the adverse reactions in participants 37 months through 5 years of age following administration of the primary series were pain at the injection site (83.8%), fatigue (61.9%), headache (22.9%), myalgia (22.1%), fever (20.9%), chills (16.8%), nausea/vomiting (15.2%), axillary (or groin) swelling/tenderness (14.3%), arthralgia (12.8%), erythema at the injection site (9.5%), and swelling at the injection site (8.2%).

#### **Post-Authorization Experience**

Anaphylaxis and other severe allergic reactions, myocarditis, pericarditis, syncope, and urticaria

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<sup>5</sup> Vaccination providers administering SPIKEVAX (COVID-19 Vaccine, mRNA) must adhere to the same reporting requirements.

have been reported following administration of the Moderna COVID-19 Vaccine outside of clinical trials.

## **6.1 Clinical Trials Experience**

Because clinical trials are conducted under widely varying conditions, adverse reaction rates observed in the clinical trials of a vaccine cannot be directly compared with rates in the clinical trials of another vaccine and may not reflect the rates observed in practice.

Overall, approximately 39,000 participants aged 6 months and older received at least one dose of Moderna COVID-19 Vaccine in five clinical trials (NCT04283461, NCT04405076, NCT04470427, NCT04649151, and NCT04796896). In a sixth clinical trial (NCT04885907), 60 solid organ transplant recipients received a third dose of Moderna COVID-19 Vaccine.

Study 1 (NCT04470427) is a Phase 3 randomized, placebo-controlled, observer-blind clinical trial conducted in the United States involving 30,346 participants 18 years of age and older who received at least one dose of Moderna COVID-19 Vaccine<sup>6</sup> (n=15,184) or placebo (n=15,162). Study 3 (NCT04649151) is a Phase 2/3 randomized, placebo-controlled, observer-blind, clinical trial conducted in the United States involving 3,726 participants 12 years through 17 years of age who received at least one dose of Moderna COVID-19 Vaccine (n=2,486) or placebo (n=1,240). Study 4 (NCT04796896) includes an ongoing Phase 2/3 randomized, placebo-controlled, observer-blind clinical trial component conducted in the United States and Canada involving 10,390 participants 6 months through 11 years of age who received at least one dose of Moderna COVID-19 Vaccine (n=7,799) or placebo (n=2,591).

### **Participants 6 Months Through 5 Years of Age**

Safety data for Moderna COVID-19 Vaccine from the ongoing Phase 2/3 randomized, placebo-controlled, observer-blind clinical trial conducted in the United States and Canada included data in 6,388 participants 6 months through 5 years of age who received at least one dose of Moderna COVID-19 Vaccine (n=4,792) or placebo (n=1,596) (Study 4). As of the data cutoff date of February 21, 2022, the median duration of blinded follow-up for safety for participants 6 months through 23 months was 68 days after Dose 2. For participants 2 years to 5 years, the median duration of blinded follow-up for safety was 71 days after Dose 2.

For participants 6 months through 23 months, 51.1% were male, 48.9% were female, 13.2% were Hispanic or Latino, 79.0% were White, 3.1% were African American, 4.9% were Asian, 0.2% were American Indian or Alaska Native, 0.0% were Native Hawaiian or Pacific Islander, 1.5% were other races, and 10.6% were Multiracial. For participants 2 years through 5 years, 50.8% were male, 49.2% were female, 14.2% were Hispanic or Latino, 76.5% were White, 4.5% were African American, 6.0% were Asian, 0.4% were American Indian or Alaska Native, 0.3% were Native Hawaiian or Pacific Islander, 1.5% were other races, and 10.4% were Multiracial. Demographic characteristics were similar among participants who received Moderna COVID-19 Vaccine and those who received placebo.

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<sup>6</sup> Moderna COVID-19 Vaccine is marketed as SPIKEVAX (COVID-19 Vaccine, mRNA), which is approved for use in individuals 18 years of age and older.



## Solicited Adverse Reactions

Local and systemic adverse reactions and use of antipyretic medication were solicited in an electronic diary for 7 days following each injection (i.e., day of vaccination and the next 6 days) among participants receiving Moderna COVID-19 Vaccine and participants receiving placebo with at least 1 documented dose (for participants 6 through 23 months, vaccine=1,758, placebo=585; for participants 24 months to 36 months, vaccine=986, placebo=338; for participants 37 months to 5 years, vaccine=2,030, placebo=659). Events that persisted for more than 7 days were followed until resolution.

The reported number and percentage of the solicited local and systemic adverse reactions by dose in Study 4 participants 6 months through 23 months of age are presented in Table 1, participants 24 months through 36 months of age are presented in Table 2, and participants 37 months to 5 years are presented in Table 3.

**Table 1: Number and Percentage of Participants With Solicited Local and Systemic Adverse Reactions Starting Within 7 Days\* After Each Dose in Participants 6 Months Through 23 Months (Solicited Safety Set, Dose 1 and Dose 2)<sup>†</sup>**

	Moderna COVID-19 Vaccine		Placebo <sup>a</sup>	
	Dose 1 (N= 1,746) n (%)	Dose 2 (N=1,596) n (%)	Dose 1 (N= 582) n (%)	Dose 2 (N=526) n (%)
<b>Local Adverse Reactions</b>				
Pain	652 (37.4)	738 (46.2)	175 (30.1)	135 (25.7)
Axillary (or groin) swelling/tenderness	102 (5.9)	148 (9.3)	26 (4.5)	28 (5.3)
Erythema (redness) ≥5 mm	150 (8.6)	216 (13.5)	24 (4.1)	20 (3.8)
Erythema (redness) Grade 3: >50 mm	5 (0.3)	14 (0.9)	2 (0.3)	0 (0)
Swelling (hardness) ≥5 mm	146 (8.4)	244 (15.3)	15 (2.6)	11 (2.1)
Swelling (hardness) Grade 3: >50 mm	5 (0.3)	14 (0.9)	0 (0)	0 (0)
<b>Systemic Adverse Reactions</b>				
Irritability/crying	1,175 (67.6)	1,021 (64.3)	361 (62.1)	307 (58.5)
Irritability/crying, Grade 3 <sup>b</sup>	24 (1.4)	25 (1.6)	6 (1.0)	5 (1.0)
Sleepiness	645 (37.1)	558 (35.1)	217 (37.3)	175 (33.3)
Sleepiness, Grade 3 <sup>c</sup>	4 (0.2)	1 (<0.1)	1 (0.2)	1 (0.2)
Loss of appetite	524 (30.2)	510 (32.1)	152 (26.2)	132 (25.1)

	Moderna COVID-19 Vaccine		Placebo <sup>a</sup>	
	Dose 1 (N= 1,746) n (%)	Dose 2 (N=1,596) n (%)	Dose 1 (N= 582) n (%)	Dose 2 (N=526) n (%)
Loss of appetite, Grade 3 <sup>d</sup>	10 (0.6)	16 (1.0)	1 (0.2)	2 (0.4)
Fever >38.0°C / >100.4°F	191 (11.0)	232 (14.6)	49 (8.4)	44 (8.4)
Fever, Grade 3: 39.6° - 40.0°C / 103.2° - 104.0°F	11 (0.6)	7 (0.4)	3 (0.5)	6 (1.1)
Fever, Grade 4: >40.0°C / >104.0°F	1 (<0.1)	3 (0.2)	1 (0.2)	0 (0)
Use of antipyretic or pain medication	482 (27.6)	543 (34.0)	141 (24.2)	111 (21.1)

N = Included 16 individuals aged 2 years to 4 years randomized in the 6 months through 23 months of age group stratum (13 in the Moderna COVID-19 Vaccine group and 3 in the placebo group).

\* 7 days included day of vaccination and the subsequent 6 days. Events and use of antipyretic or pain medication were collected in the electronic diary (e-diary).

† Absence of rows for Grade 3 or Grade 4 adverse reactions indicates no events were reported.

<sup>a</sup> Placebo was a saline solution.

<sup>b</sup> Grade 3 irritability/crying: Defined as lasting >3 hours or inconsolable.

<sup>c</sup> Grade 3 sleepiness: Defined as sleeps most of the time, hard to arouse.

<sup>d</sup> Grade 3 loss of appetite: Defined as missed >2 feeds/meals completely or refuses most feeds/meals.

**Table 2: Number and Percentage of Participants With Solicited Local and Systemic Adverse Reactions Starting Within 7 Days\* After Each Dose in Participants 24 Months Through 36 Months (Solicited Safety Set, Dose 1 and Dose 2)<sup>†</sup>**

	Moderna COVID-19 Vaccine		Placebo <sup>a</sup>	
	Dose 1 (N=944) n (%)	Dose 2 (N=963) n (%)	Dose 1 (N=320) n (%)	Dose 2 (N=330) n (%)
<b>Local Adverse Reactions</b>				
Pain	500 (53.1)	654 (67.9)	119 (37.2)	146 (44.2)
Pain, Grade 3 <sup>b</sup>	3 (0.3)	5 (0.5)	0 (0)	0 (0)
Axillary (or groin) swelling/tenderness	49 (5.2)	84 (8.7)	18 (5.6)	15 (4.5)
Axillary (or groin) swelling/tenderness, Grade 3 <sup>b</sup>	0 (0)	1 (0.1)	0 (0)	0 (0)
Erythema (redness) ≥5 mm	94 (10.0)	117 (12.1)	13 (4.1)	10 (3.0)
Erythema (redness), Grade 3: >50 mm	6 (0.6)	9 (0.9)	2 (0.6)	0 (0)
Swelling (hardness) ≥5 mm	77 (8.2)	111 (11.5)	11 (3.4)	7 (2.1)

	Moderna COVID-19 Vaccine		Placebo <sup>a</sup>	
	Dose 1 (N=944) n (%)	Dose 2 (N=963) n (%)	Dose 1 (N=320) n (%)	Dose 2 (N=330) n (%)
Swelling (hardness), Grade 3: >50 mm	5 (0.5)	8 (0.8)	2 (0.6)	0 (0)
<b>Systemic Adverse Reactions</b>				
Irritability/crying	513 (54.5)	523 (54.3)	163 (51.1)	148 (44.8)
Irritability/crying, Grade 3 <sup>c</sup>	12 (1.3)	10 (1.0)	6 (1.9)	2 (0.6)
Sleepiness	285 (30.3)	347 (36.0)	92 (28.8)	89 (27.0)
Sleepiness, Grade 3 <sup>d</sup>	2 (0.2)	1 (0.1)	0 (0)	0 (0)
Loss of appetite	225 (23.9)	294 (30.5)	71 (22.3)	69 (20.9)
Loss of appetite, Grade 3 <sup>e</sup>	7 (0.7)	8 (0.8)	1 (0.3)	0 (0)
Fever ≥38.0°C / >100.4°F	106 (11.3)	182 (18.9)	25 (7.8)	35 (10.6)
Fever, Grade 3: 39.6° - 40.0°C / 103.2° - 104.0°F	3 (0.3)	12 (1.2)	3 (0.9)	0 (0)
Fever, Grade 4: >40.0°C / >104.0°F	3 (0.3)	3 (0.3)	1 (0.3)	0 (0)
Use of antipyretic or pain medication	193 (20.4)	292 (30.3)	59 (18.4)	62 (18.8)

N = Included 36 individuals younger than 2 years of age randomized in the 2 years through 5 years of age group stratum (24 in the Moderna COVID-19 Vaccine group and 12 in the placebo group). All of these 36 individuals had eDiary for 6 months to ≤36 months age group.

\* 7 days included day of vaccination and the subsequent 6 days. Events and use of antipyretic or pain medication were collected in the electronic diary (e-diary).

† Absence of rows for Grade 3 or Grade 4 adverse reactions indicates no events were reported.

<sup>a</sup> Placebo was a saline solution.

<sup>b</sup> Grade 3 pain, axillary swelling/tenderness: Defined as prevents daily activity.

<sup>c</sup> Grade 3 irritability/crying: Defined as lasting >3 hours or inconsolable.

<sup>d</sup> Grade 3 sleepiness: Defined as sleeps most of the time, hard to arouse.

<sup>e</sup> Grade 3 loss of appetite: Defined as missed >2 feeds/meals completely or refuses most feeds/meals.

**Table 3: Number and Percentage of Participants With Solicited Local and Systemic Adverse Reactions Starting Within 7 Days\* After Each Dose in Participants 37 Months Through 5 Years (Solicited Safety Set, Dose 1 and Dose 2)<sup>†</sup>**

	Moderna COVID-19 Vaccine		Placebo <sup>a</sup>	
	Dose 1 (N=2,013) n (%)	Dose 2 (N= 1,975) n (%)	Dose 1 (N=650) n (%)	Dose 2 (N= 629) n (%)
<b>Local Adverse Reactions</b>				
Pain	1,313 (65.2)	1,445 (73.2)	263 (40.5)	249 (39.6)
Pain, Grade 3 <sup>b</sup>	1 (<0.1)	6 (0.3)	0 (0)	0 (0)
Axillary (or groin) swelling/tenderness	156 (7.7)	183 (9.3)	38 (5.8)	16 (2.5)
Erythema (redness) ≥25 mm	70 (3.5)	143 (7.2)	1 (0.2)	5 (0.8)
Erythema (redness), Grade 3: >100 mm	6 (0.3)	3 (0.2)	1 (0.2)	0 (0)
Swelling (hardness) ≥25 mm	57 (2.8)	129 (6.5)	6 (0.9)	4 (0.6)
Swelling (hardness), Grade 3: >100 mm	5 (0.2)	5 (0.3)	0 (0)	0 (0)
<b>Systemic Adverse Reactions</b>				
Fatigue	807 (40.1)	956 (48.4)	236 (36.3)	185 (29.4)
Fatigue, Grade 3 <sup>c</sup>	21 (1.0)	45 (2.3)	11 (1.7)	8 (1.3)
Headache	232 (11.5)	310 (15.7)	78 (12.0)	51 (8.1)
Headache, Grade 3 <sup>c</sup>	5 (0.2)	8 (0.4)	2 (0.3)	1 (0.2)
Fever ≥38.0°C / >100.4°F	155 (7.7)	316 (16.0)	33 (5.1)	28 (4.5)
Fever, Grade 3: 39.0° - 40.0°C / 102.1° - 104.0°F	23 (1.1)	58 (2.9)	4 (0.6)	2 (0.3)
Fever, Grade 4: >40.0°C / >104.0°F	0 (0)	2 (0.1)	1 (0.2)	0 (0)
Myalgia	200 (9.9)	310 (15.7)	60 (9.2)	47 (7.5)
Myalgia, Grade 3 <sup>c</sup>	5 (0.2)	9 (0.5)	2 (0.3)	3 (0.5)
Chills	129 (6.4)	245 (12.4)	40 (6.2)	31 (4.9)
Chills, Grade 3 <sup>c</sup>	1 (<0.1)	4 (0.2)	0 (0)	2 (0.3)
Nausea/vomiting	137 (6.8)	194 (9.8)	50 (7.7)	30 (4.8)

	Moderna COVID-19 Vaccine		Placebo <sup>a</sup>	
	Dose 1 (N=2,013) n (%)	Dose 2 (N= 1,975) n (%)	Dose 1 (N=650) n (%)	Dose 2 (N= 629) n (%)
Nausea/vomiting, Grade 3 <sup>c</sup>	7 (0.3)	6 (0.3)	2 (0.3)	0 (0)
Arthralgia	124 (6.2)	168 (8.5)	32 (4.9)	28 (4.5)
Arthralgia, Grade 3 <sup>c</sup>	2 (<0.1)	3 (0.2)	1 (0.2)	0 (0)
Use of antipyretic or pain medication	305 (15.2)	508 (25.7)	62 (9.5)	43 (6.8)

\* 7 days included day of vaccination and the subsequent 6 days. Events and use of antipyretic or pain medication were collected in the electronic diary (e-diary).

† Absence of rows for Grade 3 or Grade 4 adverse reactions indicates no events were reported.

<sup>a</sup> Placebo was a saline solution.

<sup>b</sup> Grade 3 pain: Defined as prevents daily activity.

<sup>c</sup> Grade 3 fatigue, headache, myalgia, arthralgia, chills, nausea/vomiting: Defined as prevents daily activity.

Solicited local and systemic adverse reactions reported following administration of Moderna COVID-19 Vaccine had a median duration of 2 to 3 days for participants 6 through 23 months years of age and 2 days for participants 2 through 5 years of age.

An assessment of reactogenicity among participants with evidence of prior SARS-CoV-2 infection (immunologic or virologic evidence of prior SARS-CoV-2 infection [defined as positive RT-PCR test and/or positive Elecsys immunoassay result at Day 1]) compared to those with no evidence of infection at baseline (negative RT-PCR test and negative Elecsys immunoassay result at Day 1) was conducted. In the 6 months through 23 months of age cohort, 6.1% of participants (vaccine=106, placebo=38) had evidence of prior SARS-CoV-2 infection at baseline. In the 2 years through 5 years of age cohort, 8.6% of participants (vaccine=266, placebo=82) had evidence of prior SARS-CoV-2 infection at baseline. In each age cohort, fever (temperature >38°C) was reported in a greater proportion of baseline SARS-CoV-2 positive vaccine participants compared to baseline SARS-CoV-2 negative vaccine participants. There were no notable differences in other reactogenicity events.

### Unsolicited Adverse Events

Participants were monitored for unsolicited adverse events for up to 28 days following each dose and follow-up is ongoing. Serious adverse events and medically attended adverse events will be recorded for the entire study duration.

As of February 21, 2022, among participants 6 months through 23 months of age who had received at least 1 dose of vaccine or placebo (vaccine=1,761, placebo=589), unsolicited adverse events that occurred within 28 days following each vaccination were reported by 49.3% of participants (n=869) who received Moderna COVID-19 Vaccine and 48.2% of participants (n=284) who received placebo. In these analyses, 83.1% of study participants 6 months through 23 months of age had at least 28 days of follow-up after Dose 2. Among participants 2 years through 5 years of age who had received at least 1 dose of vaccine or placebo (vaccine=3,031,

placebo=1,007), unsolicited adverse events that occurred within 28 days following each vaccination were reported by 40.0% of participants (n=1,212) who received Moderna COVID-19 Vaccine and 37.5% of participants (n=378) who received placebo. In these analyses, 89.3% of study participants 2 years through 5 years of age had at least 28 days of follow-up after Dose 2.

During the 28-day follow-up period following any dose, lymphadenopathy-related events were reported by 1.5% of vaccine recipients and 0.2% of placebo recipients who were 6 months through 23 months of age and 0.9% of vaccine recipients and <0.1% of placebo recipients who were 2 years through 5 years of age. These events included lymphadenopathy, injection-site lymphadenopathy, and vaccination-site lymphadenopathy which were plausibly related to vaccination. This imbalance is consistent with the imbalance observed for solicited axillary (or groin) swelling/tenderness in the injected limb.

During the 28-day follow-up period following any dose, hypersensitivity adverse events were reported in 3.9% of vaccine recipients and 5.3% of placebo recipients who were 6 months through 23 months of age and 3.5% of vaccine recipients and 2.5% of placebo recipients who were 2 years through 5 years of age. Hypersensitivity events in the vaccine group included injection site rash and injection site urticaria, which are likely related to vaccination. Delayed injection site reactions that began >7 days after vaccination were reported in 1.2% of vaccine recipients and no placebo recipients who were 6 months through 23 months of age and 1.4% of vaccine recipients and <0.1% of placebo recipients who were 2 years through 5 years of age. Delayed injection site reactions included pain, erythema, and swelling and are likely related to vaccination.

During the 28-day follow-up period following any dose, events of abdominal pain (including abdominal pain, abdominal pain upper, and abdominal discomfort) were reported by 0.7% of vaccine recipients and 0.4% of placebo recipients who were 2 years through 5 years of age. Currently available information is insufficient to determine a causal relationship with the vaccine.

There were no other notable patterns or numerical imbalances between treatment groups for specific categories of adverse events that would suggest a causal relationship to Moderna COVID-19 Vaccine.

### Serious Adverse Events

As of February 21, 2022, serious adverse events were reported by 0.9% (n=15) of participants who received vaccine and 0.2% (n=1) of participants who received placebo who were 6 months through 23 months of age and 0.3% (n=9) of participants who received Moderna COVID-19 Vaccine and 0.2% (n=2) of participants who received placebo who were 2 years through 5 years of age. In these analyses, 83.1% of study participants 6 months through 23 months of age had at least 28 days of follow-up after Dose 2, and the median follow-up time for all participants was 68 days after Dose 2. In these analyses, 89.3% of study participants 2 years through 5 years of age had at least 28 days of follow-up after Dose 2, and the median follow-up time for all participants was 71 days after Dose 2.

In participants 6 months through 23 months of age who received the vaccine, a 1-year-old female experienced serious adverse events of a Grade 3 fever 6 hours after Dose 1 and a febrile convulsion 1 day after Dose 1. These events were considered related to vaccination. In participants 2 years through 5 years of age who received Moderna COVID-19 Vaccine, none of the events were considered related to vaccine.

### **Participants 6 Years Through 11 Years of Age**

Safety data for Moderna COVID-19 Vaccine from Study 4 included data in 4,002 participants 6 years through 11 years of age who received at least one dose of Moderna COVID-19 Vaccine (n=3,007) or placebo (n=995). As of the data cutoff date of November 10, 2021, the median duration of blinded follow-up for safety was 51 days after Dose 2, and 1,284 participants had been followed for at least 2 months after Dose 2 (vaccine=1,006, placebo=218).

Demographic characteristics in Study 4 were similar among participants who received Moderna COVID-19 Vaccine and those who received placebo. Overall, 50.8% were male, 49.2% were female, 18.5% were Hispanic or Latino, 65.6% were White, 10.0% were African American, 9.9% were Asian, 0.4% were American Indian or Alaska Native, <0.1% were Native Hawaiian or Pacific Islander, 2.1% were other races, and 10.6% were Multiracial.

### **Unsolicited Adverse Events**

Participants were monitored for unsolicited adverse events for up to 28 days following each dose. Serious adverse events and medically attended adverse events will be recorded for the entire study duration. As of November 10, 2021, among participants who had received at least 1 dose of vaccine or placebo (vaccine=3,007, placebo=995), unsolicited adverse events that occurred within 28 days following each vaccination were reported by 29.6% of participants (n=891) who received Moderna COVID-19 Vaccine and 25.1% of participants (n=250) who received placebo. In these analyses, 98.6% of study participants had at least 28 days of follow-up after Dose 2.

During the 28-day follow-up period following any dose, lymphadenopathy-related events were reported by 1.8% of vaccine recipients and 0.6% of placebo recipients. These events included lymphadenopathy, lymph node pain, injection-site lymphadenopathy, and vaccination-site lymphadenopathy which were plausibly related to vaccination.

During the 28-day follow-up period following any dose, hypersensitivity adverse events were reported in 4.3% of vaccine recipients and 2.1% of placebo recipients. Hypersensitivity events in the vaccine group included injection site rash and injection site urticaria, which are likely related to vaccination. Delayed injection site reactions that began >7 days after vaccination were reported in 2.7% of vaccine recipients and in 0.2% of placebo recipients. Delayed injection site reactions included pain, erythema, and swelling and are likely related to vaccination.

During the 28-day follow-up period following any dose, events of abdominal pain (including abdominal pain, abdominal pain upper, and abdominal pain lower) were reported by 1.1% of vaccine recipients and 0.6% of placebo recipients. Currently available information is insufficient



to determine a causal relationship with the vaccine.

There were no other notable patterns or numerical imbalances between treatment groups for specific categories of adverse events that would suggest a causal relationship to Moderna COVID-19 Vaccine.

### Serious Adverse Events

As of November 10, 2021, serious adverse events were reported by 0.2% (n=6) of participants who received Moderna COVID-19 Vaccine and 0.2% (n=2) participants who received placebo. None of the events in the Moderna COVID-19 Vaccine group were considered related to vaccine. In these analyses, 98.6% of study participants had at least 28 days of follow-up after Dose 2, and the median follow-up time for all participants was 51 days after Dose 2.

There were no notable patterns or imbalances between treatment groups for specific categories of serious adverse events that would suggest a causal relationship to Moderna COVID-19 Vaccine.

### Additional Safety Analyses

Participants 6 years through 11 years in Study 4 started to enter an open-label, observational phase after November 1, 2021. A long-term safety analysis was conducted in participants 6 years through 11 years from Study 4 who received Moderna COVID-19 Vaccine (n=3,007) with a cut-off date of February 21, 2022. In these analyses, the median duration of follow-up including both the blinded and open-label phases was 158 days after Dose 2. Through the cut-off date, there were no serious adverse events causally related to the vaccine.

### **Participants 12 Years Through 17 Years of Age**

Safety data for Moderna COVID-19 Vaccine in adolescents were collected in an ongoing Phase 2/3 randomized, placebo-controlled, observer-blind, clinical trial conducted in the United States involving 3,726 participants 12 years through 17 years of age who received at least one dose of Moderna COVID-19 Vaccine (n=2,486) or placebo (n=1,240) (Study 3, NCT04649151). Overall, 51.4% were male, 48.6% were female, 11.6% were Hispanic or Latino, 83.9% were White, 3.4% were African American, 5.9% were Asian, 0.5% were American Indian or Alaska Native, <0.1% were Native Hawaiian or Pacific Islander, 1.0% were other races, and 4.5% were Multiracial. Demographic characteristics were similar among participants who received Moderna COVID-19 Vaccine and those who received placebo.

### Unsolicited Adverse Events

Participants were monitored for unsolicited adverse events for up to 28 days following each dose. Serious adverse events and medically attended adverse events will be recorded for the entire study duration. As of May 8, 2021, among participants who had received at least 1 dose of vaccine or placebo (vaccine=2,486, placebo=1,240), unsolicited adverse events that occurred within 28 days following each vaccination were reported by 20.5% of participants (n=510) who received Moderna COVID-19 Vaccine and 15.9% of participants (n=197) who received placebo.

In these analyses, 97.3% of study participants had at least 28 days of follow-up after Dose 2.

A 14-year-old male experienced probable myocarditis with onset of symptoms 1 day after Dose 2 of Moderna COVID-19 Vaccine. Symptoms resolved after 8 days and no sequelae were observed at 5 months. There were no cases of myocarditis among placebo recipients.

During the 28-day follow-up period following any dose, lymphadenopathy-related events that were not necessarily captured in the 7-day e-diary were reported by 5.0% of vaccine recipients and 0.5% of placebo recipients. These events included lymphadenopathy, vaccination-site lymphadenopathy and injection-site lymphadenopathy which were plausibly related to vaccination.

During the 28-day follow-up period following any dose, hypersensitivity adverse events were reported in 1.8% of vaccine recipients and 0.6% of placebo recipients. Hypersensitivity events in the vaccine group included injection site rash and injection site urticaria, which are likely related to vaccination. Delayed injection site reactions that began >7 days after vaccination were reported in 0.9% of vaccine recipients and in no placebo recipients. Delayed injection site reactions included pain, erythema, and swelling and are likely related to vaccination.

There were no other notable patterns or numerical imbalances between treatment groups for specific categories of adverse events that would suggest a causal relationship to Moderna COVID-19 Vaccine.

### Serious Adverse Events

As of May 8, 2021, serious adverse events were reported by 0.2% (n=6) of participants who received Moderna COVID-19 Vaccine and 0.2% (n=2) of participants who received placebo. In these analyses, 97.3% of study participants had at least 28 days of follow-up after Dose 2, and the median follow-up time for all participants was 53 days after Dose 2.

There were no notable patterns or imbalances between treatment groups for specific categories of serious adverse events that would suggest a causal relationship to Moderna COVID-19 Vaccine.

### Additional Safety Analyses

Study 3 participants started to enter an open-label, observational phase after May 10, 2021. A long-term safety analysis was conducted in participants from Study 3 who received Moderna COVID-19 Vaccine (n=2,486) with a cut-off date of January 31, 2022. In these analyses, the median duration of follow-up including both the blinded and open-label phases was 312 days after Dose 2 and 95.6% of study participants have had at least 6 months of follow-up after Dose 2. Through the cut-off date, there were no serious adverse events causally related to the vaccine.

### **Participants 18 Years of Age and Older**

The safety of Moderna COVID-19 Vaccine was evaluated in an ongoing Phase 3 randomized, placebo-controlled, observer-blind clinical trial conducted in the United States involving 30,346

participants 18 years of age and older who received at least one dose of Moderna COVID-19 Vaccine (n=15,184) or placebo (n=15,162) (Study 1, NCT04470427). Upon issuance of the Emergency Use Authorization (December 18, 2020) for Moderna COVID-19 Vaccine, participants were unblinded in a phased manner over a period of months to offer placebo participants Moderna COVID-19 Vaccine. The median duration of follow up for safety after the second injection during the blinded phase was 4 months. The median duration of follow up for safety after the second injection including both the blinded phase and the open-label phase was 6 months.

In Study 1, the median age of the population was 52 years (range 18-95); 22,826 (75.2%) participants were 18 to 64 years of age and 7,520 (24.8%) participants were 65 years of age and older. Overall, 52.6% of the participants were male, 47.4% were female, 20.5% were Hispanic or Latino, 79.2% were White, 10.2% were African American, 4.6% were Asian, 0.8% were American Indian or Alaska Native, 0.2% were Native Hawaiian or Pacific Islander, 2.0% were other races, and 2.1% were Multiracial. Demographic characteristics were similar between participants who received Moderna COVID-19 Vaccine and those who received placebo.

#### Unsolicited Adverse Events

Participants were monitored for unsolicited adverse events for 28 days following each dose. Serious adverse events and medically attended adverse events will be recorded for the entire study duration (2 years). Among the 30,346 participants who had received at least 1 dose of vaccine (N=15,184) or placebo (N=15,162), unsolicited adverse events that occurred within 28 days following any vaccination were reported by 31.3% of participants (n=4,752) who received Moderna COVID-19 Vaccine and 28.6% of participants (n=4,338) who received placebo.

During the 28-day follow-up period following any dose, lymphadenopathy-related events were reported by 1.7% of vaccine recipients and 0.8% of placebo recipients. These events included lymphadenopathy, lymphadenitis, lymph node pain, vaccination-site lymphadenopathy, injection-site lymphadenopathy, and axillary mass.

During the 7-day follow-up period of any vaccination, hypersensitivity events of injection site rash or injection site urticaria, likely related to vaccination, were reported by 6 participants in the Moderna COVID-19 Vaccine group and none in the placebo group. Delayed injection site reactions that began >7 days after vaccination were reported in 1.4% of vaccine recipients and 0.7% of placebo recipients. Delayed injection site reactions included pain, erythema, and swelling and are likely related to vaccination.

In the blinded portion of the study, there were 8 reports of facial paralysis (including Bell's palsy) in the Moderna COVID-19 Vaccine group, and 3 in the placebo group. In the 28-day follow-up period there were two cases of facial paralysis in the Moderna COVID-19 Vaccine group, which occurred on 8 and 22 days, respectively, after vaccination, and one in the placebo group, which occurred 17 days after vaccination. Currently available information on facial paralysis is insufficient to determine a causal relationship with the vaccine.

In the blinded portion of the study, there were 50 reports of herpes zoster in the Moderna

COVID-19 Vaccine group, and 23 in the placebo group. In the 28-day period after any vaccination, there were 22 cases of herpes zoster in the Moderna COVID-19 Vaccine group, and 15 in the placebo group. Currently available information on herpes zoster infection is insufficient to determine a causal relationship with the vaccine.

There were no other notable patterns or numerical imbalances between treatment groups for specific categories of adverse events (including other neurologic, neuro-inflammatory, and thrombotic events) that would suggest a causal relationship to Moderna COVID-19 Vaccine.

### Serious Adverse Events

During the blinded phase of the study, serious adverse events were reported by 1.8% (n=268) of participants who received Moderna COVID-19 Vaccine and 1.9% (n=292) of participants who received placebo.

There were three serious adverse events of angioedema/facial swelling in the vaccine group in recipients with a history of injection of dermatological fillers. The onset of swelling was reported 1-2 days after the second dose and was likely related to vaccination.

There were no other notable patterns or imbalances between treatment groups for specific categories of serious adverse events (including neurologic, neuro-inflammatory, and thrombotic events) that would suggest a causal relationship to Moderna COVID-19 Vaccine.

## **6.2 Post-Authorization Experience**

The following adverse reactions have been identified during post-authorization use of the Moderna COVID-19 Vaccine. Because these reactions are reported voluntarily, it is not always possible to reliably estimate their frequency or establish a causal relationship to vaccine exposure.

Cardiac Disorders: myocarditis, pericarditis  
Immune System Disorders: anaphylaxis, urticaria  
Nervous System Disorders: syncope

## **8 REQUIREMENTS AND INSTRUCTIONS FOR REPORTING ADVERSE EVENTS AND VACCINE ADMINISTRATION ERRORS<sup>7</sup>**

See Overall Safety Summary (Section 6) for additional information.

The vaccination provider enrolled in the federal COVID-19 Vaccination Program is responsible for the MANDATORY reporting of the listed events following Moderna COVID-19 Vaccine to the Vaccine Adverse Event Reporting System (VAERS)

- Vaccine administration errors whether or not associated with an adverse event
- Serious adverse events\* (irrespective of attribution to vaccination)

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<sup>7</sup> Vaccination providers administering SPIKEVAX (COVID-19 Vaccine, mRNA) must adhere to the same reporting requirements.

- Cases of myocarditis
- Cases of pericarditis
- Cases of Multisystem Inflammatory Syndrome (MIS) in adults and children
- Cases of COVID-19 that results in hospitalization or death

\*Serious Adverse Events are defined as:

- Death;
- A life-threatening adverse event;
- Inpatient hospitalization or prolongation of existing hospitalization;
- A persistent or significant incapacity or substantial disruption of the ability to conduct normal life functions;
- A congenital anomaly/birth defect;
- An important medical event that based on appropriate medical judgement may jeopardize the individual and may require medical or surgical intervention to prevent one of the outcomes listed above.

### Instructions for Reporting to VAERS

The vaccination provider enrolled in the federal COVID-19 Vaccination Program should complete and submit a VAERS form to FDA using one of the following methods:

- Complete and submit the report online: <https://vaers.hhs.gov/reportevent.html>, or
- If you are unable to submit this form electronically, you may fax it to VAERS at 1-877-721-0366. If you need additional help submitting a report, you may call the VAERS toll-free information line at 1-800-822-7967 or send an email to [info@vaers.org](mailto:info@vaers.org).

**IMPORTANT: When reporting adverse events or vaccine administration errors to VAERS, please complete the entire form with detailed information. It is important that the information reported to FDA be as detailed and complete as possible. Information to include:**

- Patient demographics (e.g., patient name, date of birth)
- Pertinent medical history
- Pertinent details regarding admission and course of illness
- Concomitant medications
- Timing of adverse event(s) in relationship to administration of Moderna COVID-19 Vaccine
- Pertinent laboratory and virology information
- Outcome of the event and any additional follow-up information if it is available at the time of the VAERS report. Subsequent reporting of follow-up information should be completed if additional details become available.

The following steps are highlighted to provide the necessary information for safety tracking:

1. In Box 17, provide information on Moderna COVID-19 Vaccine and any other vaccines administered on the same day; and in Box 22, provide information on any other vaccines received within one month prior.
2. In Box 18, description of the event:

- a. Write “Moderna COVID-19 Vaccine EUA” as the first line
  - b. Provide a detailed report of vaccine administration error and/or adverse event. It is important to provide detailed information regarding the patient and adverse event/medication error for ongoing safety evaluation of this unapproved vaccine. Please see information to include listed above.
3. Contact information:
  - a. In Box 13, provide the name and contact information of the prescribing healthcare provider or institutional designee who is responsible for the report.
  - b. In Box 14, provide the name and contact information of the best doctor/healthcare professional to contact about the adverse event.
  - c. In Box 15, provide the address of the facility where vaccine was given (NOT the healthcare provider’s office address).

### Other Reporting Instructions

Vaccination providers may report to VAERS other adverse events that are not required to be reported using the contact information above.

To the extent feasible, report adverse events to ModernaTX, Inc. using the contact information below or by providing a copy of the VAERS form to ModernaTX, Inc.

Email	Fax number	Telephone number
<a href="mailto:ModernaPV@modernatx.com">ModernaPV@modernatx.com</a>	1-866-599-1342	1-866-MODERNA (1-866-663-3762)

## **10 DRUG INTERACTIONS**

There are no data to assess the concomitant administration of the Moderna COVID-19 Vaccine with other vaccines.

## **11 USE IN SPECIFIC POPULATIONS**

### **11.3 Pediatric Use**

Moderna COVID-19 Vaccine is authorized for use in individuals 6 months through 17 years of age. This authorization is based on safety and effectiveness data in this age group and adults.

Moderna COVID-19 Vaccine is not authorized for use in individuals younger than 6 months of age.

### **11.4 Use in Immunocompromised Individuals**

Safety and effectiveness of the Moderna COVID-19 Vaccine in individuals 6 months through 17 years of age with immunocompromise have been extrapolated from adult data. In an independent study (*Hall VG, Ferreira VH, Ku T et al. Randomized Trial of a Third Dose of mRNA-1273*

*Vaccine in Transplant Recipients. N Engl J Med 2021 DOI: 10.1056/NEJMc2111462; NCT04885907*), safety and effectiveness of a third primary series dose of the Moderna COVID-19 Vaccine have been evaluated in participants who received solid organ transplants. In this study, in 60 adult participants who had undergone various solid organ transplant procedures (heart, kidney, kidney-pancreas, liver, lung, pancreas) a median of 3.57 years previously (range 1.99-6.75 years) who received a third vaccine dose, the adverse event profile was similar to that after the second dose and no Grade 3 or Grade 4 events were reported. The administration of a third primary series vaccine dose appears to be only moderately effective in increasing antibody titers. Patients should be counseled to maintain physical precautions to help prevent COVID-19. In addition, close contacts of immunocompromised persons should be vaccinated, as appropriate for their health status.

### **13 DESCRIPTION**

Moderna COVID-19 Vaccine is provided as a white to off-white suspension for intramuscular injection.

Each 0.25 mL primary series dose of Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border contains 25 mcg of nucleoside-modified messenger RNA (mRNA) encoding the pre-fusion stabilized Spike glycoprotein (S) of the SARS-CoV-2 Wuhan-Hu-1 strain. Each 0.25 mL dose of the Moderna COVID-19 Vaccine supplied in a multiple-dose vial with a dark blue cap and a label with a magenta border contains the following ingredients: a total lipid content of 0.5 mg (SM-102, polyethylene glycol [PEG] 2000 dimyristoyl glycerol [DMG], cholesterol, and 1,2-distearoyl-sn-glycero-3-phosphocholine [DSPC]), 0.13 mg tromethamine, 0.62 mg tromethamine hydrochloride, 0.011 mg acetic acid, 0.049 mg sodium acetate trihydrate, and 21.8 mg sucrose.

Moderna COVID-19 Vaccine does not contain a preservative.

The vial stoppers are not made with natural rubber latex.

### **14 CLINICAL PHARMACOLOGY**

#### **14.1 Mechanism of Action**

The nucleoside-modified mRNA in the Moderna COVID-19 Vaccine is formulated in lipid particles, which enable delivery of the nucleoside-modified mRNA into host cells to allow expression of the SARS-CoV-2 S antigen. The vaccine elicits an immune response to the S antigen, which protects against COVID-19.

### **18 CLINICAL TRIAL RESULTS AND SUPPORTING DATA FOR EUA**

#### **18.1 Efficacy of Two-Dose Primary Series in Participants 18 Years of Age and Older**

Study 1 is an ongoing Phase 3 randomized, placebo-controlled, observer-blind clinical trial to evaluate the efficacy, safety, and immunogenicity of the Moderna COVID-19 Vaccine in



participants 18 years of age and older in the United States (NCT04470427). Randomization was stratified by age and health risk: 18 to <65 years of age without comorbidities (not at risk for progression to severe COVID-19), 18 to <65 years of age with comorbidities (at risk for progression to severe COVID-19), and 65 years of age and older with or without comorbidities. Participants who were immunocompromised and those with a known history of SARS-CoV-2 infection were excluded from the study. Participants with no known history of SARS-CoV-2 infection but with positive laboratory results indicative of infection at study entry were included. The study allowed for the inclusion of participants with stable pre-existing medical conditions, defined as disease not requiring significant change in therapy or hospitalization for worsening disease during the 3 months before enrollment, as well as participants with stable human immunodeficiency virus (HIV) infection. A total of 30,420 participants were randomized equally to receive 2 doses of the Moderna COVID-19 Vaccine or saline placebo 1 month apart. Participants will be followed for efficacy and safety until 24 months after the second dose.

The primary efficacy analysis population (referred to as the Per-Protocol Set) included 28,207 participants who received two doses (at 0 and 1 month) of either Moderna COVID-19 Vaccine (n=14,134) or placebo (n=14,073) and had a negative baseline SARS-CoV-2 status. In the Per-Protocol Set, 47.4% were female, 19.7% were Hispanic or Latino; 79.5% were White, 9.7% were African American, 4.6% were Asian, and 2.1% other races. The median age of participants was 53 years (range 18-95) and 25.3% of participants were 65 years of age and older. Of the study participants in the Per-Protocol Set, 18.5% were at increased risk of severe COVID-19 due to at least one pre-existing medical condition (chronic lung disease, significant cardiac disease, severe obesity, diabetes, liver disease, or HIV infection) regardless of age. Between participants who received Moderna COVID-19 Vaccine and those who received placebo, there were no notable differences in demographics or pre-existing medical conditions.

### Efficacy Against COVID-19

COVID-19 was defined based on the following criteria: The participant must have experienced at least two of the following systemic symptoms: fever ( $\geq 38^{\circ}\text{C}$  /  $\geq 100.4^{\circ}\text{F}$ ), chills, myalgia, headache, sore throat, new olfactory and taste disorder(s); or the participant must have experienced at least one of the following respiratory signs/symptoms: cough, shortness of breath or difficulty breathing, or clinical or radiographical evidence of pneumonia; and the participant must have at least one NP swab, nasal swab, or saliva sample (or respiratory sample, if hospitalized) positive for SARS-CoV-2 by RT-PCR. COVID-19 cases were adjudicated by a Clinical Adjudication Committee.

The median length of follow-up for efficacy for participants in the study was 9 weeks post Dose 2. There were 11 COVID-19 cases in the Moderna COVID-19 Vaccine group and 185 cases in the placebo group, with a vaccine efficacy of 94.1% (95% confidence interval of 89.3% to 96.8%).

**Table 4: Primary Efficacy Analysis: COVID-19\* in Participants 18 Years of Age and Older Starting 14 Days After Dose 2 per Adjudication Committee Assessments – Per-Protocol Set**

Moderna COVID-19 Vaccine			Placebo			% Vaccine Efficacy (95% CI)†
Participants (N)	COVID-19 Cases (n)	Incidence Rate of COVID-19 per 1,000 Person-Years	Participants (N)	COVID-19 Cases (n)	Incidence Rate of COVID-19 per 1,000 Person-Years	
14,134	11	3.328	14,073	185	56.510	94.1 (89.3, 96.8)

\* COVID-19: symptomatic COVID-19 requiring positive RT-PCR result and at least two systemic symptoms or one respiratory symptom. Cases starting 14 days after Dose 2.

† VE and 95% CI from the stratified Cox proportional hazard model.

The subgroup analyses of vaccine efficacy are presented in Table 5.

**Table 5: Subgroup Analyses of Vaccine Efficacy: COVID-19\* Cases Starting 14 Days After Dose 2 per Adjudication Committee Assessments – Per-Protocol Set**

Age Subgroup (Years)	Moderna COVID-19 Vaccine			Placebo			% Vaccine Efficacy (95% CI)†
	Participants (N)	COVID-19 Cases (n)	Incidence Rate of COVID-19 per 1,000 Person-Years	Participants (N)	COVID-19 Cases (n)	Incidence Rate of COVID-19 per 1,000 Person-Years	
18 to <65	10,551	7	2.875	10,521	156	64.625	95.6 (90.6, 97.9)
≥65	3,583	4	4.595	3,552	29	33.728	86.4 (61.4, 95.2)

\* COVID-19: symptomatic COVID-19 requiring positive RT-PCR result and at least two systemic symptoms or one respiratory symptom. Cases starting 14 days after Dose 2.

† VE and 95% CI from the stratified Cox proportional hazard model.

Severe COVID-19 was defined based on confirmed COVID-19 as per the primary efficacy endpoint case definition, plus any of the following: Clinical signs indicative of severe systemic illness, respiratory rate  $\geq 30$  per minute, heart rate  $\geq 125$  beats per minute, SpO2  $\leq 93\%$  on room air at sea level or PaO2/FIO2  $< 300$  mm Hg; or respiratory failure or ARDS (defined as needing high-flow oxygen, non-invasive or mechanical ventilation, or ECMO), evidence of shock (systolic blood pressure  $< 90$  mmHg, diastolic BP  $< 60$  mmHg or requiring vasopressors); or significant acute renal, hepatic, or neurologic dysfunction; or admission to an intensive care unit or death.

Among all participants in the Per-Protocol Set analysis, which included COVID-19 cases confirmed by an adjudication committee, no cases of severe COVID-19 were reported in the

Moderna COVID-19 Vaccine group compared with 30 cases reported in the placebo group (incidence rate 9.138 per 1,000 person-years). One PCR-positive case of severe COVID-19 in a vaccine recipient was awaiting adjudication at the time of the analysis.

## 18.2 Effectiveness of Two-Dose Primary Series in Participants 6 Months Through 5 Years of Age

Study 4 includes an ongoing Phase 2/3 randomized, placebo-controlled, observer-blind, clinical trial component to evaluate the safety, reactogenicity, and effectiveness of the Moderna COVID-19 Vaccine in individuals ages 6 months through 5 years in the United States and Canada (NCT04796896). Participants with a known history of SARS-CoV-2 infection within 2 weeks of study vaccination were excluded from the study. A total of 6,403 participants were randomized 3:1 to receive 2 doses of the Moderna COVID-19 Vaccine or saline placebo 1 month apart. Participants will be followed for occurrence of COVID-19 and safety until 1 year after the last dose.

Effectiveness in individuals 6 months through 5 years of age is based on a comparison of immune responses in this age group to adults 18 years through 25 years of age.

In Study 4, an analysis was conducted of SARS-CoV-2 neutralizing antibody concentrations and seroresponse rates 28 days after Dose 2 in a subset of individuals 6 months through 5 years of age in Study 4 and participants 18 years through 25 years of age in Study 1 who had no immunologic or virologic evidence of prior SARS-CoV-2 at baseline. Noninferior immune responses as assessed by neutralizing antibody concentrations in arbitrary units (AU)/mL and seroresponse rates were demonstrated in a comparison of individuals 6 months through 23 months of age to participants 18 years through 25 years of age (Table 6) and 2 years through 5 years of age to participants 18 years through 25 years of age (Table 7).

**Table 6: Summary of Geometric Mean Concentration Ratio and Seroresponse Rate – Comparison of Individuals 6 Months Through 23 Months of Age to Participants 18 Years Through 25 Years of Age – Per-Protocol Immunogenicity Set**

		Moderna COVID-19 Vaccine			
		6 Months Through 23 Months n=230	18 Years Through 25 Years n=291	6 Months Through 23 Months/ 18 Years Through 25 Years	
Assay	Time Point	GMC (95% CI)*	GMC (95% CI)*	GMC Ratio (95% CI) <sup>a</sup>	Met Noninferiority Objective (Y/N) <sup>b</sup>
SARS-CoV-2 neutralization assay <sup>c</sup>	28 days after Dose 2	1780.7 (1606.4, 1973.8)	1390.8 (1269.1, 1524.2)	1.3 (1.1, 1.5)	Y
		Seroresponse % (95% CI) <sup>d</sup>	Seroresponse % (95% CI) <sup>d</sup>	Difference in Seroresponse Rate % (95% CI) <sup>e</sup>	
		100 (98.4, 100)	99.3 (97.5, 99.9)	0.7 (-1.0, 2.5)	

GMC = Geometric mean concentration

n = number of participants with non-missing data at baseline and at Day 57

\* Antibody values reported as below the lower limit of quantification (LLOQ) are replaced by 0.5 x LLOQ. Values greater than the upper limit of quantification (ULOQ) are replaced by the ULOQ if actual values are not available.

<sup>a</sup> The log-transformed antibody levels are analyzed using an analysis of covariance (ANCOVA) model with the group variable (individuals in Study 4 and young adults in Study 1) as fixed effect. The resulted LS means, difference of LS means, and 95% CI are back transformed to the original scale for presentation.

<sup>b</sup> Noninferiority is declared if the lower bound of the 2-sided 95% CI for the GMC ratio is greater than 0.67, with a point estimate of >0.8 and the lower bound of the 2-sided 95% CI for difference in seroresponse rate is greater than -10%, with a point estimate of >-5%.

<sup>c</sup> Final geometric mean antibody concentrations (GMC) in AU/mL were determined using SARS-CoV-2 microneutralization assay. The SARS CoV-2 MN is a cell-based assay that is designed to determine the ability of SARS-CoV-2 neutralizing antibodies to inhibit the infection of 293T-ACE2 cells by SARS-CoV-2 Reporter Virus Particles (RVP) which express green fluorescent protein (GFP). A given serum sample is pre-incubated with a known quantity of SARS-CoV-2-GFP for 60 (±5) minutes prior to infection of 293T-ACE2 cells. COVID-19 infection is monitored 48 (±4) hours following infection by counting the number of green fluorescent cells using the Cytation 5 cell imaging reader.

<sup>d</sup> Seroresponse due to vaccination specific to SARS-CoV-2 RVP neutralizing antibody concentration at a subject level is defined in protocol as a change from below LLOQ to equal or above 4 x LLOQ, or at least a 4-fold rise if baseline is equal to or above LLOQ. Seroresponse 95% CI is calculated using the Clopper-Pearson method.

<sup>e</sup> Difference in seroresponse rate 95% CI is calculated using the Miettinen-Nurminen (score) confidence limits.

**Table 7: Summary of Geometric Mean Concentration Ratio and Seroresponse Rate – Comparison of Individuals 2 Years Through 5 Years of Age to Participants 18 Years Through 25 Years of Age – Per-Protocol Immunogenicity Set**

		Moderna COVID-19 Vaccine			
		2 Years Through 5 Years n=264	18 Years Through 25 Years n=291	2 Years Through 5 Years/ 18 Years Through 25 Years	
Assay	Time Point	GMC (95% CI)*	GMC (95% CI)*	GMC Ratio (95% CI) <sup>a</sup>	Met Noninferiority Objective (Y/N) <sup>b</sup>
SARS-CoV-2 neutralization assay <sup>c</sup>	28 days after Dose 2	1410.0 (1273.8, 1560.8)	1390.8 (1262.5, 1532.1)	1.0 (0.9, 1.2)	Y
		Seroresponse % (95% CI) <sup>d</sup>	Seroresponse % (95% CI) <sup>d</sup>	Difference in Seroresponse Rate % (95% CI) <sup>e</sup>	
		98.9 (96.7, 99.8)	99.3 (97.5, 99.9)	-0.4 (-2.7, 1.5)	

GMC = Geometric mean concentration

n = number of participants with non-missing data at baseline and at Day 57

\* Antibody values reported as below the lower limit of quantification (LLOQ) are replaced by 0.5 x LLOQ. Values greater than the upper limit of quantification (ULOQ) are replaced by the ULOQ if actual values are not available.

<sup>a</sup> The log-transformed antibody levels are analyzed using an analysis of covariance (ANCOVA) model with the group variable (individuals in Study 4 and young adults in Study 1) as fixed effect. The resulted LS means, difference of LS means, and 95% CI are back transformed to the original scale for presentation.

<sup>b</sup> Noninferiority is declared if the lower bound of the 2-sided 95% CI for the GMC ratio is greater than 0.67, with a point estimate of >0.8 and the lower bound of the 2-sided 95% CI for difference in seroresponse rate is greater than -10%, with a point estimate of >-5%.

<sup>c</sup> Final geometric mean antibody concentrations (GMC) in AU/mL were determined using SARS-CoV-2 microneutralization assay. The SARS CoV-2 MN is a cell-based assay that is designed to determine the ability of SARS-CoV-2 neutralizing antibodies to inhibit the infection of 293T-ACE2 cells by SARS-CoV-2 Reporter Virus Particles (RVP) which express green fluorescent protein (GFP). A given serum sample is pre-incubated with a known quantity of SARS-CoV-2-GFP for 60 ( $\pm$ 5) minutes prior to infection of 293T-ACE2 cells. COVID-19 infection is monitored 48 ( $\pm$ 4) hours following infection by counting the number of green fluorescent cells using the Cytation 5 cell imaging reader.

<sup>d</sup> Seroresponse due to vaccination specific to SARS-CoV-2 RVP neutralizing antibody concentration at a subject level is defined in protocol as a change from below LLOQ to equal or above 4 x LLOQ, or at least a 4-fold rise if baseline is equal to or above LLOQ. Seroresponse 95% CI is calculated using the Clopper-Pearson method.

<sup>e</sup> Difference in seroresponse rate 95% CI is calculated using the Miettinen-Nurminen (score) confidence limits.

A descriptive efficacy analysis evaluating confirmed COVID-19 cases accrued up to the data cutoff date February 21, 2022, was performed in 5,476 participants 6 months through 5 years of age who received two doses (at 0 and 1 month) of either Moderna COVID-19 Vaccine or placebo and had a negative baseline SARS-CoV-2 status (referred to as the Per-Protocol Set for Efficacy) (for participants 6 months through 23 months, vaccine=1,511, placebo=513; for participants 2 years through 5 years, vaccine=2,594, placebo=858). For participants 6 months through 23 months in the Per-Protocol Set for Efficacy, 51.2% were male, 48.8% were female, 12.7% were Hispanic or Latino; 78.9% were White, 3.1% were African American, 4.6% were Asian, 0.2% were American Indian or Alaska Native, 0.0% were Native Hawaiian or Pacific Islander, 1.8% were other races, and 10.7% were Multiracial. For participants 2 years through 5 years, 50.7% were male, 49.3% were female, 14.0% were Hispanic or Latino, 76.8% were White, 4.1% were African American, 6.1% were Asian, 0.4% were American Indian or Alaska Native, 0.3% were Native Hawaiian or Pacific Islander, 1.6% were other races, and 10.3% were Multiracial. Between participants who received Moderna COVID-19 Vaccine and those who received placebo, there were no notable differences in demographics.

The median length of follow-up for efficacy post-Dose 2 was 68 days for participants 6 months through 23 months of age and 71 days for participants 2 years through 5 years of age.

Vaccine efficacy among individuals 6 months through 5 years in Study 4 was evaluated during the period when the B.1.1.529 (Omicron) variant was the predominant variant in circulation.

The efficacy information in individuals 6 months through 23 months of age and 2 years through 5 years of age are presented in Table 8 and Table 9, respectively.

**Table 8: Efficacy Analyses: COVID-19 in Participants 6 Months Through 23 Months of Age Starting 14 Days After Dose 2 – Per Protocol Set for Efficacy**

	Moderna COVID-19 Vaccine N=1,511		Placebo N=513		% Vaccine Efficacy (95% CI)*
	Cases (n)	Incidence Rate of COVID-19 per 1,000 Person-Years	Cases (n)	Incidence Rate of COVID-19 per 1,000 Person-Years	
<b>COVID-19 Cases - Definition 1<sup>a</sup></b>	37	99.981	18	146.042	31.5 (-27.7, 62.0)
<b>COVID-19 Cases - Definition 2<sup>b</sup></b>	51	138.239	34	279.822	50.6 (21.4, 68.6)

N = Included 15 individuals aged 2 years to 4 years randomized in the 6 months through 23 months of age group stratum (12 in the Moderna COVID-19 Vaccine group and 3 in the placebo group), and none of them had a COVID-19 case starting 14 days after Dose 2.

\* Vaccine efficacy defined as 1 — ratio of incidence rate (Moderna COVID-19 Vaccine vs. placebo). The 95% CI of the ratio is calculated using the exact method conditional upon the total number of cases, adjusting for person-years.

<sup>a</sup> Participant must have experienced at least two of the following systemic symptoms: fever ( $\geq 38^{\circ}\text{C}$  /  $\geq 100.4^{\circ}\text{F}$ ), chills, myalgia, headache, sore throat, new olfactory and taste disorder(s); or the participant must have experienced at least one of the following respiratory signs/symptoms: cough, shortness of breath or difficulty breathing, or clinical or radiographical evidence of pneumonia; and the participant must have at least one NP swab, nasal swab, or saliva sample (or respiratory sample, if hospitalized) positive for SARS-CoV-2 by RT-PCR.

<sup>b</sup> Presence of at least one symptom from a list of COVID-19 symptoms and a positive NP swab or saliva sample for SARS-CoV-2 by RT-PCR. Listed symptoms were fever (temperature  $> 38^{\circ}\text{C}$  /  $\geq 100.4^{\circ}\text{F}$ ), or chills, cough, shortness of breath or difficulty breathing, fatigue, muscle aches, or body aches, headache, new loss of taste or smell, sore throat, congestion or runny nose, nausea, abdominal pain, poor appetite/poor feeding, or vomiting or diarrhea.

**Table 9: Efficacy Analyses: COVID-19 in Participants 2 Years Through 5 Years of Age Starting 14 Days After Dose 2 – Per-Protocol Set for Efficacy**

	Moderna COVID-19 Vaccine N=2,594		Placebo N=858		% Vaccine Efficacy (95% CI)*
	Cases (n)	Incidence Rate of COVID-19 per 1,000 Person-Years	Cases (n)	Incidence Rate of COVID-19 per 1,000 Person-Years	
<b>COVID-19 Cases - Definition 1<sup>a</sup></b>	71	103.761	43	193.528	46.4 (19.8, 63.8)
<b>COVID-19 Cases - Definition 2<sup>b</sup></b>	119	175.023	61	276.980	36.8 (12.5, 54.0)

N = Included 25 individuals younger than 2 years of age randomized in the 2 years through 5 years of age group stratum (18 in the Moderna COVID-19 Vaccine group and 7 in the placebo group), and one in each treatment group had a COVID-19 case starting 14 days after Dose 2.

\* Vaccine efficacy defined as 1 — ratio of incidence rate (Moderna COVID-19 Vaccine vs. placebo). The 95% CI of the ratio is calculated using the exact method conditional upon the total number of cases, adjusting for person-years.

<sup>a</sup> Participant must have experienced at least two of the following systemic symptoms: fever ( $\geq 38^{\circ}\text{C}$  /  $\geq 100.4^{\circ}\text{F}$ ), chills, myalgia, headache, sore throat, new olfactory and taste disorder(s); or the participant must have experienced at least one of the following respiratory signs/symptoms: cough, shortness of breath or difficulty breathing, or clinical or radiographical evidence of pneumonia; and the participant must have at least one NP swab, nasal swab, or saliva sample (or respiratory sample, if hospitalized) positive for SARS-CoV-2 by RT-PCR.

<sup>b</sup> Presence of at least one symptom from a list of COVID-19 symptoms and a positive NP swab or saliva sample for SARS-CoV-2 by RT-PCR. Listed symptoms were fever (temperature  $>38^{\circ}\text{C}$  /  $\geq 100.4^{\circ}\text{F}$ ), or chills, cough, shortness of breath or difficulty breathing, fatigue, muscle aches, or body aches, headache, new loss of taste or smell, sore throat, congestion or runny nose, nausea, abdominal pain, poor appetite/poor feeding, or vomiting or diarrhea.

### 18.3 Immunogenicity in Solid Organ Transplant Recipients

Immunogenicity of the Moderna COVID-19 Vaccine in children with immunocompromise has been extrapolated from adult data. An independent randomized-controlled study has been conducted in 120 adult participants who had undergone various solid organ transplant procedures (heart, kidney, kidney-pancreas, liver, lung, pancreas) a median of 3.57 years previously (range 1.99-6.75 years). A third primary series dose of the Moderna COVID-19 Vaccine was administered to 60 participants approximately 2 months after they had received a second dose; saline placebo was given to 60 individuals for comparison. Significant increases in levels of SARS-CoV-2 antibodies occurred four weeks after the third dose in 55.0% of participants in the Moderna COVID-19 Vaccine group (33 of 60) and 17.5% of participants in the placebo group (10 of 57).

## 19 HOW SUPPLIED/STORAGE AND HANDLING

The information in this section applies to the Moderna COVID-19 Vaccine that is supplied in multiple-dose vials with dark blue caps and labels with a magenta border containing a volume of 2.5 mL. These multiple-dose vials are supplied as follows:

NDC 80777-279-99 Carton of 10 multiple-dose vials, each containing 10 doses of 0.25 mL

During storage, minimize exposure to room light, and avoid exposure to direct sunlight and ultraviolet light.

#### Frozen Storage

Store frozen between  $-50^{\circ}\text{C}$  to  $-15^{\circ}\text{C}$  ( $-58^{\circ}\text{F}$  to  $5^{\circ}\text{F}$ ).

#### Storage after Thawing

- Storage at  $2^{\circ}\text{C}$  to  $8^{\circ}\text{C}$  ( $36^{\circ}\text{F}$  to  $46^{\circ}\text{F}$ ):
  - Vials may be stored refrigerated between  $2^{\circ}\text{C}$  to  $8^{\circ}\text{C}$  ( $36^{\circ}\text{F}$  to  $46^{\circ}\text{F}$ ) for up to 30 days prior to first use.
  - Vials should be discarded 12 hours after the first puncture.
- Storage at  $8^{\circ}\text{C}$  to  $25^{\circ}\text{C}$  ( $46^{\circ}\text{F}$  to  $77^{\circ}\text{F}$ ):
  - Vials may be stored between  $8^{\circ}\text{C}$  to  $25^{\circ}\text{C}$  ( $46^{\circ}\text{F}$  to  $77^{\circ}\text{F}$ ) for a total of 24 hours.
  - Vials should be discarded 12 hours after the first puncture.
  - Total storage at  $8^{\circ}\text{C}$  to  $25^{\circ}\text{C}$  ( $46^{\circ}\text{F}$  to  $77^{\circ}\text{F}$ ) must not exceed 24 hours.



**Do not refreeze once thawed.**

Thawed vials can be handled in room light conditions.

Transportation of Thawed Vials at 2°C to 8°C (36°F to 46°F)

If transport at -50°C to -15°C (-58°F to 5°F) is not feasible, available data support transportation of one or more thawed vials for up to 12 hours at 2°C to 8°C (36°F to 46°F) when shipped using shipping containers which have been qualified to maintain 2°C to 8°C (36°F to 46°F) and under routine road and air transport conditions with shaking and vibration minimized. Once thawed and transported at 2°C to 8°C (36°F to 46°F), vials should not be refrozen and should be stored at 2°C to 8°C (36°F to 46°F) until use.

## **20 PATIENT COUNSELING INFORMATION**

Advise the recipient or caregiver to read the Fact Sheet for Recipients and Caregivers.

The vaccination provider must include vaccination information in the state/local jurisdiction's Immunization Information System (IIS) or other designated system. Advise recipient or caregiver that more information about IISs can be found at:

<https://www.cdc.gov/vaccines/programs/iis/about.html>.

## **21 CONTACT INFORMATION**

For general questions, send an email or call the telephone number provided below.

<b>Email</b>	<b>Telephone number</b>
<a href="mailto:medinfo@modernatx.com">medinfo@modernatx.com</a>	1-866-MODERNA (1-866-663-3762)

This EUA Prescribing Information may have been updated. For the most recent Full EUA Prescribing Information, please visit [www.modernatx.com/covid19vaccine-eua](http://www.modernatx.com/covid19vaccine-eua).

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Patent(s): [www.modernatx.com/patents](http://www.modernatx.com/patents)

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