Natpara®
(parathyroid hormone)
for Injection
25 • 50 • 75 • 100 mcg per dose strength

NATPARA can help you navigate low blood calcium due to hypoparathyroidism

Look inside for important information about hypoparathyroidism, treatment with NATPARA, and available product support resources.

What is NATPARA® (parathyroid hormone) for Injection?

- NATPARA is a prescription parathyroid hormone used with calcium and vitamin D to control low blood calcium (hypocalcemia) in people with low parathyroid hormone blood levels (hypoparathyroidism).
- NATPARA is only for people who do not respond well to treatment with calcium and active forms of vitamin D alone, because it may increase the possible risk of bone cancer (osteosarcoma).
- NATPARA was not studied in people with hypoparathyroidism caused by calcium-sensing receptor mutations.
- NATPARA was not studied in people who get sudden hypoparathyroidism after surgery.
- It is not known if NATPARA is safe and effective for children 18 years of age and younger. NATPARA should not be used in children and young adults whose bones are still growing.

Important Safety Information

What is the most important information I should know about NATPARA?

Warning: Possible bone cancer (osteosarcoma).

- During animal drug testing, NATPARA caused some rats to develop a bone cancer called osteosarcoma. It is not known if people who take NATPARA will have a higher chance of getting osteosarcoma. Tell your doctor right away if you have pain in any areas of your body that does not go away, or any new or unusual lumps or swelling under your skin that is tender to touch.
- **NATPARA is only available through the NATPARA Risk Evaluation and Mitigation Strategy (REMS) Program.** The purpose of the NATPARA REMS Program is to inform patients about the potential risk of osteosarcoma associated with the use of NATPARA. For more information about this REMS program, call 1-855-NATPARA or go to **www.NATPARAREMS.com**.

Please see Important Safety Information throughout.
Please see full Prescribing Information, Medication Guide, and Instructions for Use for NATPARA.

Learn about hypoparathyroidism and parathyroid hormone

Parathyroid hormone

Most people have 4 parathyroid glands located behind the thyroid gland in the neck. The glands release parathyroid hormone (PTH), which controls the levels of calcium and vitamin D in the blood.

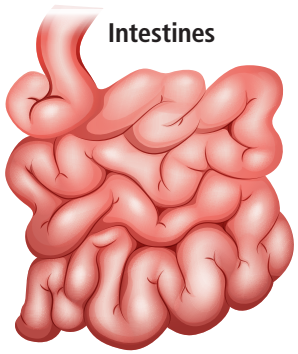
What is hypoparathyroidism?

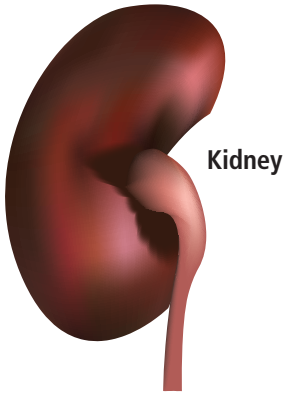
Hypoparathyroidism is a rare disorder that occurs when the parathyroid glands are damaged either due to surgery or a disorder and are not able to produce enough PTH. Not having enough PTH can interfere with important functions of certain organs within the body. In some patients, doctors are not able to tell the cause of the disorder.

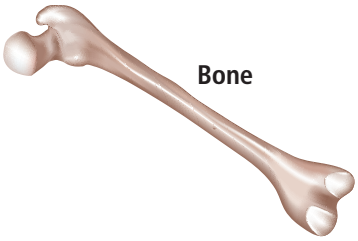
How PTH helps your body get the calcium it needs

PTH works along with the intestines, kidneys, and bones to:

- Activate vitamin D to help the body get calcium from food
- Increase the amount of calcium the kidneys reabsorb
- Release calcium from the bones into the bloodstream







NATPARA works differently in treating low blood calcium in hypoparathyroidism

NATPARA is the first and only FDA-approved prescription parathyroid hormone that is taken along with calcium and vitamin D to control hypocalcemia in adults with hypoparathyroidism.

NATPARA helps to control your blood calcium level

NATPARA helps raise low blood calcium levels caused by hypoparathyroidism. It works by:

- Helping your intestines absorb calcium by activating vitamin D
- Increasing the amount of calcium your kidneys reabsorb
- Releasing calcium from your bones

NATPARA was effective in helping to balance blood calcium levels

In a 24-week clinical study of 124 adults with hypoparathyroidism:

- To prove the efficacy of NATPARA, by the end of the study patients had to have their:
  - Dose of active vitamin D reduced by at least 50%
  - Dose of calcium reduced by at least 50%
  - Blood calcium level in a certain target range (between 7.5 mg/dL and 10.6 mg/dL)

55% of patients receiving NATPARA reduced their active vitamin D and oral calcium use by at least half while maintaining calcium levels in the blood within a target range, compared with 3% in the placebo group[a,b]

42% of patients receiving NATPARA were able to **stop all use of active vitamin D** and were able to reduce their oral calcium to no more than 500 mg per day, compared with 3% in the placebo group[a,b]

69% of patients receiving NATPARA reduced their oral calcium use by at least half, compared with 8% in the placebo group[a,b]

For more information about the efficacy of NATPARA, ask your doctor and visit NATPARA.com

[a]At the end of the clinical study, there were no differences in the percentages of patients with serum calcium levels between 7.5 and 10.6 mg/dL in the NATPARA (86.9%) and placebo (87.5%) groups.

[b]Of the 124 adult patients, 84 were taking NATPARA and 40 were taking placebo.

Important Safety Information

What is the most important information I should know about NATPARA? (*cont'd*)

NATPARA may cause other serious side effects, including: (*cont'd*)

High blood calcium (hypercalcemia)

- NATPARA can cause some people to have a higher blood calcium level than normal.
  - Your doctor should check your blood calcium before you start and during your treatment with NATPARA.
  - Tell your doctor if you have nausea, vomiting, constipation, low energy, or muscle weakness. These may be signs that you have too much calcium in your blood.

Please see Important Safety Information throughout.
Please see full Prescribing Information, Medication Guide, and Instructions for Use for NATPARA.

Natpara®
(parathyroid hormone) for Injection
25 • 50 • 75 • 100 mcg per dose strength

What you need to know about using NATPARA

NATPARA is available as an injection (shot). You will need to inject NATPARA once a day directly under the skin of the thigh. Be sure to inject a different thigh every day.

OnePath®, Shire's product support program, will send a registered nurse to your home for training on how to inject NATPARA. See page 6 of this brochure for more details.

(Actual size)

Check with your doctor to get your routine tests

Your doctor will need to monitor how NATPARA is working

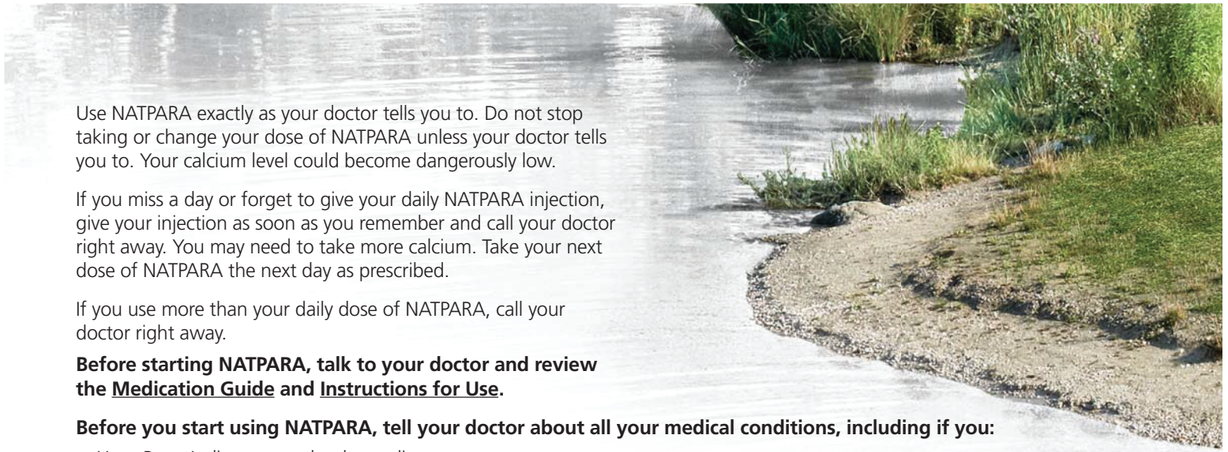
It is important to have routine blood tests and a yearly or twice yearly 24-hour urine collection test once the maintenance dose is achieved, so that your doctor can measure your level of calcium and see how well you are responding to NATPARA.

With 4 dosage strengths of NATPARA, your doctor can find the dose that is right for you

NATPARA offers dosing options to help your doctor find the dose that is best for you

The starting dose for NATPARA is 50 mcg once a day as a subcutaneous injection in the thigh (alternate thigh every day). Your doctor will adjust the doses of NATPARA, calcium, and vitamin D you take based on what you need. If necessary, your doctor will increase the dose of NATPARA.

A majority of patients in the NATPARA clinical study required a maximum dose of 100 mcg. The goal of treatment is to have you take the lowest dose of NATPARA that will achieve a calcium level within the lower half of the normal range without the need for active forms of vitamin D and with enough calcium to meet your body's daily needs.



Use NATPARA exactly as your doctor tells you to. Do not stop taking or change your dose of NATPARA unless your doctor tells you to. Your calcium level could become dangerously low.

If you miss a day or forget to give your daily NATPARA injection, give your injection as soon as you remember and call your doctor right away. You may need to take more calcium. Take your next dose of NATPARA the next day as prescribed.

If you use more than your daily dose of NATPARA, call your doctor right away.

Before starting NATPARA, talk to your doctor and review the Medication Guide and Instructions for Use.

Before you start using NATPARA, tell your doctor about all your medical conditions, including if you:

- Have Paget's disease or other bone disease
- Have or have had cancer in your bones
- Have or have had radiation therapy
- Have or have had too much calcium in your blood
- Have or have had high blood levels of certain electrolytes (for example, alkaline phosphatase)
- Are pregnant or plan to become pregnant, or are breastfeeding or plan to breastfeed. It is not known if NATPARA will harm your unborn baby or if NATPARA passes into your breast milk. You and your doctor should decide if you will use NATPARA or breastfeed. You should not do both.

Tell your doctor about all the medicines you take, including prescription and over-the-counter medicines, vitamins, and herbal supplements.

NATPARA and other medicines may affect each other, causing side effects. Especially tell your doctor if you are taking medicines that contain digoxin, alendronate, calcium supplements, or food products that contain calcium or active vitamin D.

Know the medicines you take. Keep a list of them to show your doctor or pharmacist each time you get a new medicine.

Important Safety Information

What is the most important information I should know about NATPARA? (*cont'd*)

NATPARA may cause other serious side effects, including: (*cont'd*)

Low blood calcium (hypocalcemia)

- People who stop using or miss a dose of NATPARA may have an increased risk of severe low blood calcium levels.
- Tell your doctor if you have tingling of your lips, tongue, fingers and feet, twitching of face muscles, cramping of feet and hands, seizures, depression, or have problems thinking or remembering.

Tell your doctor right away if you have any of these signs and symptoms of **high or low blood calcium** levels.

Who should not use NATPARA?

- **Do not use NATPARA** if you are allergic to parathyroid hormone or any of the ingredients in NATPARA.

What should I tell my healthcare provider before using NATPARA?

- **Before you start using NATPARA, tell your doctor about all of your medical conditions. Tell your doctor about all the medicines you take,** including prescription and over-the-counter medicines, vitamins, and herbal supplements.

Please see Important Safety Information throughout.

Please see full Prescribing Information, Medication Guide, and Instructions for Use for NATPARA.

Natpara®
(parathyroid hormone)
for Injection
25 • 50 • 75 • 100 mcg per dose strength

A product support program to help you get started on NATPARA

Shire is committed to supporting patients with hypoparathyroidism on their treatment journey.

OnePath is a support program for patients like you.

How OnePath can help

OnePath helps eligible adults with hypoparathyroidism who have been prescribed NATPARA gain access to treatment. OnePath® is a product support program that can help eligible patients with many aspects of their Shire therapy, from insurance and financial options to training and prescription fulfillment. For more information, contact your OnePath Patient Support Manager.

OnePath Programs and Services

Options for financial assistance	• OnePath may be able to help you find options for financial support • OnePath Co-pay Assistance Program – For patients enrolled in OnePath and with commercial insurance – 100% coverage for eligible co-pay expenses up to the program maximum regardless of financial status – Eligible co-pay expenses may also be billed as deductibles and coinsurance
At-home injection training	• Your Patient Support Manager (PSM) will connect you with a specially trained nurse educator who will provide injection training in your own home • You will receive: – 2 at-home visits from your dedicated nurse on days 1 and 15 of treatment – 3 follow-up calls from a nurse educator at OnePath on days 3, 8, and 30 • You can call OnePath at 1-866-888-0660 to connect with a registered nurse, Monday through Friday, from 8:30 AM to 8 PM ET
Patient Support Manager (PSM) and Patient Access Manager (PAM) product support	• Once enrolled in OnePath, you will be connected with your dedicated PSM • Your PSM will: – Facilitate insurance benefits verification – Work with a specialty pharmacy to facilitate treatment access – Serve as a point of contact throughout your treatment • A PAM is also available in your local area to help address your insurance and treatment access-related needs

Enrolling in OnePath

To enroll in OnePath, you and your doctor will first need to fill out a OnePath Start Form. Your doctor's office will then submit the completed form to OnePath. Additional forms may be required.

Connect with dedicated and knowledgeable patient ambassadors

The NATPARA Shire Patient Ambassador Program enables patients to connect directly with real hypoparathyroidism patients to share their experiences through:

- **Live Events**
  - Live meetings in your area will feature healthcare experts and Patient Ambassadors discussing hypoparathyroidism and treatment with NATPARA
  - These gatherings allow patients to receive education from NATPARA Patient Ambassadors in a positive environment

To sign up for the program, including live events, visit NATPARA.com

Cece is in her 50s and is a community leader.
Actual NATPARA Patient Ambassador

Important Safety Information

What are the possible side effects of NATPARA?

- **NATPARA may cause serious side effects like allergic (hypersensitivity) reaction, including anaphylaxis.** Tell your healthcare provider or get emergency medical help right away if you have any of the following symptoms of an allergic reaction:
  - swelling of your face, lips, mouth, or tongue
  - breathing problems
  - fainting, dizziness, feeling lightheaded (low blood pressure)
  - fast heartbeat
  - itching
  - rash
  - hives
- **The most common side effects of NATPARA include:** tingling, tickling or burning feeling of the skin, low or high blood calcium, headache, nausea, reduced sense of touch or sensation, diarrhea, vomiting, pain in joints, too much calcium in urine, and pain in limbs.

These are not all the possible side effects of NATPARA. For more information, talk with your doctor. You are encouraged to report negative side effects of prescription drugs to the FDA. Visit: **www.fda.gov/medwatch** or call **1-800-FDA-1088**.

Natpara®
(parathyroid hormone)
for Injection
25 • 50 • 75 • 100 mcg per dose strength

Please see Important Safety Information throughout.
Please see full Prescribing Information, Medication Guide, and Instructions for Use for NATPARA.

Things you may be wondering about

Here is some important information about NATPARA you should know before starting treatment.

Q. Will my insurance cover NATPARA?

A. Every person's insurance coverage is different. OnePath may be able to help. OnePath can help facilitate an insurance benefits verification to determine if your insurance covers NATPARA.

Q. Why does NATPARA have a warning about bone cancer?

A. During animal drug testing, NATPARA caused some rats to develop a bone cancer called osteosarcoma. It is not known if people who take NATPARA will have a higher chance of getting osteosarcoma. Tell your doctor right away if you have pain in any areas of your body that does not go away, or any new or unusual lumps or swelling under your skin that is tender to touch. These are some of the signs and symptoms of osteosarcoma and your doctor may need to do further tests.

Q. Do I need to have any blood tests before I begin taking NATPARA?

A. Yes. Before you can begin treatment, you need to have a blood test to check the levels of calcium and vitamin D in your blood.

Q. How will I get NATPARA?

A. NATPARA is only available through the NATPARA Risk Evaluation and Mitigation Strategy (REMS) Program. The purpose of the NATPARA REMS Program is to inform patients about the potential risk of osteosarcoma associated with the use of NATPARA.

The NATPARA REMS Program Coordinating Center will call you with the name and phone number of the certified pharmacy that will fill your NATPARA prescription.

The certified pharmacy will call you to arrange the date to ship NATPARA to you. For more information about this REMS program, call 1-855-NATPARA or go to www.NATPARAREMS.com.

For more information on treatment with NATPARA, ask your doctor and visit NATPARA.com

Important Safety Information

What is the most important information I should know about NATPARA?

Warning: Possible bone cancer (osteosarcoma).

- During animal drug testing, NATPARA caused some rats to develop a bone cancer called osteosarcoma. It is not known if people who take NATPARA will have a higher chance of getting osteosarcoma. Tell your doctor right away if you have pain in any areas of your body that does not go away, or any new or unusual lumps or swelling under your skin that is tender to touch.
- **NATPARA is only available through the NATPARA Risk Evaluation and Mitigation Strategy (REMS) Program.** The purpose of the NATPARA REMS program is to inform patients about the potential risk of osteosarcoma associated with the use of NATPARA. For more information about this REMS program, call 1-800-828-2088 or go to **www.NATPARAREMS.com**.

NATPARA may cause other serious side effects, including:

High blood calcium (hypercalcemia)

- NATPARA can cause some people to have a higher blood calcium level than normal.
  - Your doctor should check your blood calcium before you start and during your treatment with NATPARA.
  - Tell your doctor if you have nausea, vomiting, constipation, low energy, or muscle weakness. These may be signs that you have too much calcium in your blood.

Low blood calcium (hypocalcemia)

- People who stop using or miss a dose of NATPARA may have an increased risk of severe low blood calcium levels.
- Tell your doctor if you have tingling of your lips, tongue, fingers and feet, twitching of face muscles, cramping of feet and hands, seizures, depression, or have problems thinking or remembering.

Tell your doctor right away if you have any of these signs and symptoms of **high or low blood calcium** levels.

Who should not use NATPARA?

- **Do not use NATPARA** if you are allergic to parathyroid hormone or any of the ingredients in NATPARA.

What should I tell my healthcare provider before using NATPARA?

- **Before you start using NATPARA, tell your doctor about all of your medical conditions. Tell your doctor about all the medicines you take,** including prescription and over-the-counter medicines, vitamins, and herbal supplements.

What are the possible side effects of NATPARA?

- **NATPARA may cause serious side effects like allergic (hypersensitivity) reaction, including anaphylaxis.** Tell your healthcare provider or get emergency medical help right away if you have any of the following symptoms of an allergic reaction:
  - swelling of your face, lips, mouth, or tongue
  - breathing problems
  - fainting, dizziness, feeling lightheaded (low blood pressure)
  - fast Heartbeat
  - itching
  - rash
  - hives
- **The most common side effects of NATPARA include:** tingling, tickling or burning feeling of the skin, low or high blood calcium, headache, nausea, reduced sense of touch or sensation, diarrhea, vomiting, pain in joints, too much calcium in urine, and pain in limbs.

These are not all the possible side effects of NATPARA. For more information, talk with your doctor. You are encouraged to report negative side effects of prescription drugs to the FDA. Visit: **www.fda.gov/medwatch** or call **1-800-FDA-1088**.

Natpara®
(parathyroid hormone)
for Injection
25 • 50 • 75 • 100 mcg per dose strength

Please see full Prescribing Information, Medication Guide, and Instructions for Use for NATPARA.

Talk to your doctor to learn more about NATPARA and see if NATPARA is right for you

OnePath®

Shire's product support program for eligible patients prescribed NATPARA

For more information on OnePath and product support resources, visit **NATPARA.com** or call **1-866-888-0660**.

Patient Support Managers are available to speak with you Monday through Friday, 8:30am to 8:00pm EST

To learn more about hypoparathyroidism and treatment with NATPARA, visit www.NATPARA.com

Important Safety Information

What is the most important information I should know about NATPARA?

Warning: Possible bone cancer (osteosarcoma).

- During animal drug testing, NATPARA caused some rats to develop a bone cancer called osteosarcoma. It is not known if people who take NATPARA will have a higher chance of getting osteosarcoma. Tell your doctor right away if you have pain in any areas of your body that does not go away, or any new or unusual lumps or swelling under your skin that is tender to touch.
- **NATPARA is only available through the NATPARA Risk Evaluation and Mitigation Strategy (REMS) Program.** The purpose of the NATPARA REMS Program is to inform patients about the potential risk of osteosarcoma associated with the use of NATPARA. For more information about this REMS program, call 1-855-NATPARA or go to **www.NATPARAREMS.com**.

Please see Important Safety Information throughout.
Please see full Prescribing Information, Medication Guide, and Instructions for Use for NATPARA.

Natpara®
(parathyroid hormone) for Injection
25 • 50 • 75 • 100 mcg per dose strength

Shire