Information Regarding the US NATPARA® (parathyroid hormone) for Injection Recall

The latest news about the US NATPARA Recall will be posted here as it becomes available. Takeda is committed to supply integrity, and we are working closely with the FDA to resolve the issue and resume supply as soon as possible.

<table>
<thead>
<tr>
<th>Date</th>
<th>Information</th>
</tr>
</thead>
<tbody>
<tr>
<td>October 9, 2019</td>
<td>Important: FDA classifies NATPARA recall as a Class I recall requiring return of all unused product</td>
</tr>
<tr>
<td>September 25, 2019</td>
<td>Additional details about the NATPARA Special Use Program</td>
</tr>
<tr>
<td>September 24, 2019</td>
<td>Takeda announces NATPARA Special Use Program in the US</td>
</tr>
<tr>
<td>September 13, 2019</td>
<td>Message about the US NATPARA Recall</td>
</tr>
<tr>
<td>September 5, 2019</td>
<td>Takeda Issues US Recall of NATPARA Due to the Potential for Rubber Particulate</td>
</tr>
</tbody>
</table>

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

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S51982 09/19
US Recall of NATPARA® (parathyroid hormone) for Injection

IMPORTANT INFORMATION FOR NATPARA PATIENTS

Dear Valued Patient,

The purpose of this letter is to share some new and important information regarding Takeda’s September 5, 2019 US recall for all doses of NATPARA® (parathyroid hormone) for Injection (25 mcg, 50 mcg, 75 mcg, and 100 mcg).

NATPARA is a parathyroid hormone currently approved in the US as the only adjunctive treatment for adult patients with chronic hypoparathyroidism who cannot be adequately controlled with standard therapy alone (calcium and vitamin D).

On October 2nd, the US Food & Drug Administration (“US FDA”) informed Takeda that, following further review of the NATPARA recall, they are classifying NATPARA recall as Class I due to the potential risk of rubber stopper particles clogging the needle and leading to under-dosing. The Class I recall requires that all patients with product received prior to the recall of September 5, 2019 (“Recalled Product”) return their unused NATPARA to Takeda to prevent the use of Recalled Product. The safety profile of NATPARA remains consistent with the product label.

In parallel, Takeda continues to work closely with the US FDA regarding NATPARA, and values the US FDA’s collaboration and feedback as we work together to resupply NATPARA to patients who need it.

Based on the FDA’s classification, enclosed in this envelope you will find information from Stericycle, a third-party vendor that works with companies such as Takeda to coordinate product recalls. This includes instructions on how to return your unused NATPARA product using the Patient Return Kit.

With patient safety as the company’s main priority, Takeda is communicating directly with healthcare professionals, patients, and specialty pharmacies in the US regarding the actions required as a result of the Class 1 recall. Consistent with the product labelling, Takeda is alerting NATPARA patients and prescribers that discontinuing NATPARA abruptly can cause a sharp decrease in blood calcium levels (severe hypocalcemia) which can result in serious health consequences. It is critically important that patients contact their prescribing healthcare provider to discuss their individual treatment plan and ensure close supervision, including frequent monitoring of blood calcium levels and close titration of active vitamin D and calcium supplements upon stopping NATPARA to avoid low blood calcium (hypocalcemia).

It is critical to note that the Special Use Program is unaffected by the Class I recall and continues to support patients previously prescribed NATPARA who are at extreme risk of life-threatening complications as a result of discontinuation of NATPARA. This means that the
single-use NATPARA cartridges that you have received, or are expected to receive under the Special Use Program, are excluded from the Class I recall and you should continue to use them as prescribed by your physician.

Any patient who believes that they may qualify for the Special Use Program should contact their prescribing healthcare provider. Through this program, healthcare providers will be able to request NATPARA for extraordinary, life-threatening cases. This program requires a signed physician case report confirming the physician’s decision to prescribe NATPARA and their determination that without continued access to NATPARA the patient faces life-threatening health consequences. The prescribing physician and the Medical Review committee will then determine if a patient is at extreme medical risk and should have access to NATPARA. The Medical Review committee is evaluating eligibility on a case-by-case basis for patients who are at most risk for life-threatening complications.

All of us at Takeda understand the impact that this recall has on patients like you, and we will continue to work closely with the FDA until we are able to resolve the issue and resume supply.

To contact a OnePath Patient Support Manager, please call 1-866-888-0660. OnePath is available Monday through Friday, 8:30am to 8:00pm, Eastern Time.

Sincerely,

Daniel McNamara  
Head of US Patient Services  
Takeda Pharmaceutical Company Limited
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 (628-7272) 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)
  - itching
  - rash
  - hives

Shire-NPS Pharmaceuticals, Inc., part of the Takeda Group
300 Shire Way
Lexington, MA 02421 USA
- fast heartbeat

**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](http://www.fda.gov/medwatch) or call 1-800-FDA-1088.

Please go to [https://www.shirecontent.com/PI/PDFs/Natpara_USA_ENG.pdf](https://www.shirecontent.com/PI/PDFs/Natpara_USA_ENG.pdf) for the Full Prescribing Information and Medication Guide.
September 25, 2019

Dear Valued Patient,

We recognize that the NATPARA® recall in the US has been extremely difficult for patients, their families and caregivers. Since this began on September 5, 2019, our team has been working diligently with the FDA on a number of potential solutions to bring this critical medicine back to patients.

While we are in discussions with the FDA to reintroduce NATPARA to the broader patient community, Takeda and the FDA have developed a Special Use Program for NATPARA. The program is intended to support patients previously prescribed NATPARA who are facing life-threatening complications as a result of discontinuation of NATPARA, despite efforts to actively manage their condition. Through this program, healthcare providers will be able to request NATPARA for patients facing life-threatening complications.

It is anticipated that an extremely small number of patients prescribed NATPARA may be eligible for this limited program. Additional information about the Special Use Program for healthcare providers can be found here: https://www.takeda.com/en-us/newsroom/news-releases/2019/takeda-announces-natpara-special-use-program-in-the-us/. Physicians are also receiving detailed information about this new program.

Patients who have questions about their eligibility under the Special Use Program or about their individual treatment plans should consult with their NATPARA prescribing physician.

While this is an important first step, we realize this short-term and extremely limited program will not address the needs of the vast majority of patients. We have heard your frustration and want to assure you that we are working diligently with the FDA on both short-term and long-term solutions to bring NATPARA back to the broader patient community, which remains our highest priority.

We are dedicated to the hypoparathyroidism community and the resupply of NATPARA for patients. We will continue to work with the FDA on this issue and keep patients and healthcare providers informed as more information becomes available.

As always, our OnePath® team is available to support you. Please reach out to a OnePath team member at 1-866-888-0660, Monday through Friday 8:30 a.m. to 8:00 p.m. ET.

Sincerely,

Dan McNamara
Head of US Patient Services
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 (628-7272) or go to [www.NATPARAREMS.com](http://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.
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.

Please see [Full Prescribing Information](#), [Medication Guide](#), and [Instructions for Use for NATPARA](#).

NATPARA® is a registered trademark of Shire-NPS Pharmaceuticals, Inc., a Takeda company.

OnePath® is a registered trademark of Shire Human Genetic Therapies, Inc., a Takeda company.

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Takeda Announces NATPARA® Special Use Program in the US

Takeda has worked with the FDA to develop a Special Use Program for NATPARA®. The program is intended to support patients previously prescribed NATPARA who are facing life-threatening complications as a result of discontinuation of NATPARA. Through this program, healthcare providers will be able to request NATPARA for these extraordinary, life-threatening cases. It is anticipated that an extremely small number of patients prescribed NATPARA will qualify for this very limited program. Additional information for healthcare providers about the Special Use Program can be found here. Patients who have questions about their eligibility under the Special Use Program or about their individual treatment plans should consult with their healthcare providers.

We realize that while this program is an important first step, it will only help a very small number of patients. We recognize that many more patients are in need of NATPARA to control their hypoparathyroidism. Takeda continues to work with the FDA on both short- and long-term solutions to bring NATPARA back to the broader patient community, which remains our highest priority.

We are committed to the hypoparathyroidism community and the safe supply of NATPARA. We will continue to work urgently on this issue and keep patients and healthcare providers informed of our progress in resuming supply.

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).

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  - Your doctor should check your blood calcium before you start and during your treatment with NATPARA.
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- 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.

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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.

Please see Full Prescribing Information, Medication Guide, and Instructions for Use for NATPARA.
Message About the US NATPARA Recall

Cambridge, MA, September 13, 2019 — On September 5, 2019, Takeda issued a US recall for all doses of NATPARA® (parathyroid hormone) for Injection (25 mcg, 50 mcg, 75 mcg, and 100 mcg). The recall is being conducted after discussions with the FDA, which has classified it as a Pharmacy Level recall due to the potential for rubber particulate from the rubber septum component part of the NATPARA cartridge to enter into the drug solution. Patients do not need to return or discard the NATPARA they have on-hand. But, it is imperative that patients immediately see their healthcare provider before stopping NATPARA. Doing so will help patients discontinue the medicine as safely as possible, in a supervised setting, including frequent monitoring of blood calcium levels and close titration of active vitamin D and calcium supplements upon stopping NATPARA to avoid low blood calcium. You can read more information about the NATPARA recall here.

We recognize that the recall has been extremely difficult for our NATPARA patients and their families. Since the recall began on September 5, 2019, our dedicated OnePath team has reached out to the more than 2,000 NATPARA patients in the US with information and support. Our commitment to patients remains our highest priority. We are working urgently with the FDA on a number of potential solutions to bring this critical medicine back to patients as quickly as possible and will continue to keep patients and healthcare providers informed.

If you are a patient with questions, please reach out to our OnePath patient services team at 866-888-0660. Healthcare providers with questions should call 800-828-2088.

We are committed to resolving this critical issue quickly and sincerely regret the impact this recall is having on patients and their families, as well as the broader hypoparathyroidism community.

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.
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- **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 (628-7272) or go to [www.NATPARAREMS.com](http://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 http://www.fda.gov/medwatch or call 1-800-FDA-1088.

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

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Important Notice
For the purposes of this notice, “press release” means this document, any oral presentation, any question and answer session and any written or oral material discussed or distributed by Takeda Pharmaceutical Company Limited (“Takeda”) regarding this press release. This press release (including any oral briefing and any question-and-answer in connection with it) is not intended to, and does not constitute, represent or form part of any offer, invitation or solicitation of any offer to purchase, otherwise acquire, subscribe for, exchange, sell or otherwise dispose of, any securities or the solicitation of any vote or approval in any jurisdiction. No shares or other securities are being offered to the public by means of this press release. No offering of securities shall be made in the United States except pursuant to registration under the U.S. Securities Act of 1933, as amended, or an exemption therefrom. This press release is being given (together with any further information which may be provided to the recipient) on the condition that it is for use by the recipient for information purposes only (and not for the evaluation of any investment, acquisition, disposal or any other transaction). Any failure to comply with these restrictions may constitute a violation of applicable securities laws.

The companies in which Takeda directly and indirectly owns investments are separate entities. In this press release, “Takeda” is sometimes used for convenience where references are made to Takeda and its subsidiaries in general. Likewise, the words “we”, “us” and “our” are also used to refer to subsidiaries in general or to those who work for them. These expressions are also used where no useful purpose is served by identifying the particular company or companies.

Forward-Looking Statements
This press release and any materials distributed in connection with this press release may contain forward-looking statements, beliefs or opinions regarding Takeda’s future business, future position and results of operations, including estimates, forecasts, targets and plans for Takeda. Without limitation, forward looking statements often include the words such as “targets”, “plans”, “believes”, “hopes”,

“continues”, “expects”, “aims”, “intends”, “ensures”, “will”, “may”, “should”, “would”, “could” “anticipates”, “estimates”, “projects” or words or terms of similar substance or the negative thereof. Any forward-looking statements in this document are based on the current assumptions and beliefs of Takeda in light of the information currently available to it. Such forward-looking statements do not represent any guarantee by Takeda or its management of future performance and involve known and unknown risks, uncertainties and other factors, including but not limited to: the economic circumstances surrounding Takeda’s business, including general economic conditions in Japan, the United States and worldwide; competitive pressures and developments; applicable laws and regulations; the success of or failure of product development programs; decisions of regulatory authorities and the timing thereof; changes in exchange rates; claims or concerns regarding the safety or efficacy of marketed products or products candidates; and post-merger integration with acquired companies, any of which may cause Takeda’s actual results, performance, achievements or financial position to be materially different from any future results, performance, achievements or financial position expressed or implied by such forward-looking statements. For more information on these and other factors which may affect Takeda’s results, performance, achievements, or financial position, see “Item 3. Key Information—D. Risk Factors” in Takeda’s Annual Report on Form 20-F filed with the U.S. Securities and Exchange Commission, available on Takeda’s website at: https://www.takeda.com/investors/reports/sec-filings/ or at www.sec.gov. Neither Takeda nor its management gives any assurances that the expectations expressed in these forward-looking statements will turn out to be correct, and actual results, performance or achievements could materially differ from expectations. Persons receiving this press release should not place undue reliance on forward looking statements. Takeda undertakes no obligation to update any of the forward-looking statements contained in this press release or any other forward-looking statements it may make, except as required by law or stock exchange rule. Past performance is not an indicator of future results and the results of Takeda in this press release may not be indicative of, and are not an estimate, forecast or projection of Takeda’s future results.

About Takeda Pharmaceutical Company Limited

Takeda Pharmaceutical Company Limited (TSE:4502/NYSE:TAK) is a global, values-based, R&D-driven biopharmaceutical leader headquartered in Japan, committed to bringing Better Health and a Brighter Future to patients by translating science into highly-innovative medicines. Takeda focuses its R&D efforts on four therapeutic areas: Oncology, Gastroenterology (GI), Rare Diseases and Neuroscience. We also make targeted R&D investments in Plasma-Derived Therapies and Vaccines. We are focusing on developing highly innovative medicines that contribute to making a difference in people's lives by advancing the frontier of new treatment options and leveraging our enhanced collaborative R&D engine and capabilities to create a robust, modality-diverse pipeline. Our employees are committed to improving quality of life for patients and to working with our partners in health care in approximately 80 countries and regions.

For more information, visit https://www.takeda.com.
September 6, 2019

US Recall of NATPARA® (parathyroid hormone) for Injection

IMPORTANT INFORMATION REQUIRING ACTION

Dear Valued Patient,

The purpose of this letter is to inform you that Takeda has issued a US recall for all doses of NATPARA® (parathyroid hormone) for Injection (25 mcg, 50 mcg, 75 mcg, and 100 mcg). The recall is effective immediately. The safety profile of NATPARA remains consistent with the product label. As of the date of this recall, Takeda is not aware of any adverse events directly related to this matter. After discussions with the US FDA, the company is issuing this recall as a precaution.

NATPARA is a parathyroid hormone indicated as an adjunct to calcium and vitamin D to control hypocalcemia in patients with hypoparathyroidism.

This recall is due to a potential issue related to rubber particulates originating from the rubber septum of the NATPARA cartridge. During the 14-day NATPARA treatment period, the septum is punctured by a needle each day to obtain the daily dosage of medicine solution. When the septum is repeatedly punctured, it is possible that small rubber fragments may detach into the cartridge.

It is critically important that you contact your prescribing healthcare provider, who has also received this important information. As part of this communication, and consistent with the product label, we are alerting NATPARA patients and prescribers that stopping NATPARA abruptly can cause a sharp decrease in blood calcium levels (severe hypocalcemia) which can have serious health consequences. You need to be under close supervision by your healthcare provider to discuss your individual treatment plan, including frequent monitoring of your blood calcium levels and close titration of active vitamin D and calcium supplements upon stopping NATPARA to avoid low blood calcium (hypocalcemia).
In addition, a OnePath® Patient Support Manager will be contacting you to answer any questions you may have about the content of this letter.

Takeda is committed to supply integrity, and we are working closely with the FDA to resolve the issue and resume supply as soon as possible.

We sincerely regret the impact this interruption has on you.

To contact a OnePath Patient Support Manager, please call 1-866-888-0660. OnePath is available Monday through Friday, 8:30am to 8:00pm, Eastern Time.

Sincerely,

Daniel McNamara
Head of US Patient Services
Takeda Pharmaceutical Company Limited

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 (628-7272) 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.

1. Your doctor should check your blood calcium before you start and during your treatment with NATPARA.

2. 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.

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.

Please see Full Prescribing Information, including Boxed Warning for potential risk of osteosarcoma.