

Trial Information

Simple trial name: A clinical trial to find out if neridronic acid given into a vein eases the pain felt by patients who have severe lasting pain in the hand, wrist, ankle or foot from complex regional pain syndrome (CRPS)

Protocol number: KF7013-02

Trial Sponsor: Grünenthal GmbH

Thank you to the trial patients



If you are a patient who took part in the clinical trial, thank you for your time and commitment.

You made the clinical trial possible.

You helped us on our way to bringing new medicines to patients.

Important note: You should not use this summary to make decisions about the medical treatments you use. You should always see your doctor for advice about medical treatments.

About this summary

This summary is written to share the results of this clinical trial with the public. It is written in a way that should be easy for most people to understand. It describes why the trial was needed, how it was done and the results.

General information about the clinical trial

Why was this trial needed?

Complex regional pain syndrome (CRPS) is a condition that can develop after a minor injury such as a fracture or sprain, usually in the hand, wrist, ankle or foot. People with CRPS feel severe lasting pain in their affected hand, wrist, ankle or foot. The pain felt can be described as a “burning”, “electrical” or “shooting” pain.

Although CRPS treatment options are available, many patients are not treated adequately for their pain.

Researchers are looking for new ways to treat CRPS. In this trial, researchers studied the effect of a trial medicine called neridronic acid in participants with CRPS.

Which medicines were studied?



Neridronic acid

Neridronic acid belongs to a group of medicines that are already being used to treat other conditions like osteoporosis (softening of the bones).



Placebo (Dummy medicine)

A placebo or dummy medicine looks like the trial medicine and is given in the same way but does not have any medicine in it. Researchers sometimes use a placebo or dummy medicine to understand if the changes seen were due to the trial medicine or were caused by other factors.

What were the main objectives of the trial?

The main objectives of the trial were to find out:

- if neridronic acid eased participants’ pain after 12 weeks of treatment.
- if neridronic acid is safe to be given to participants with severe lasting pain in the hand, wrist, ankle or foot from CRPS.

When was the trial?



This trial started on 30 May 2018 and ended on 31 July 2019.

As per the plan, the sponsor measured the initial combined results from this trial and another neridronic acid trial, while they were both underway. The sponsor ended this trial early because the initial results indicated that neridronic acid only had a very small chance of providing pain relief to the participants by the end of the trial.

Where did this trial take place?

The clinical trial took place in the following countries:

European Union (EU) countries

- Germany (1 participant)
- Spain (8 participants)

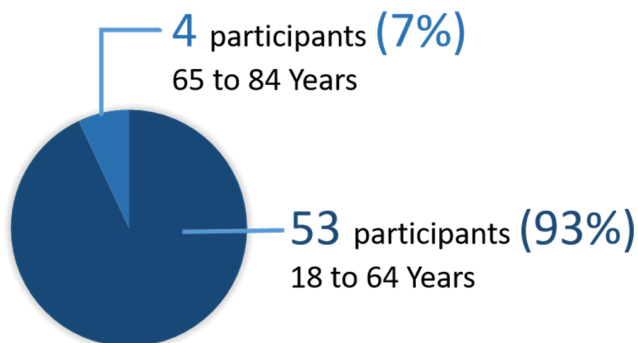
Non-EU countries

- United States (44 participants)
- Australia (1 participant)
- New Zealand (1 participant)
- Republic of Korea (2 participants)

Which participants were included in this trial?

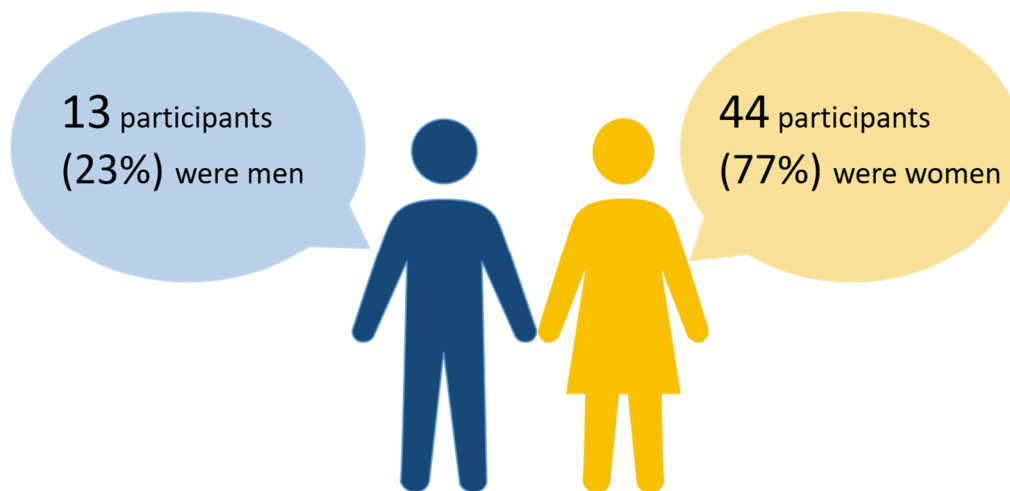
A total of 57 participants were treated with neridronic acid or dummy medicine in this clinical trial.

How old were the participants?



The average age of the participants was 48 years. The youngest participant was 18 years old and the oldest participant was 84 years old.

Were the participants men or women?



Which participants were able to take part in the trial?

Participants were only able to take part in the clinical trial if they met certain criteria. This was important to make sure that it was safe for each participant to take part in the clinical trial, that the results of the clinical trial were valid and that the laws and regulations were followed.

People could take part in this trial if they:

- were at least 18 years of age.
- had severe lasting pain in the hand, wrist, ankle or foot from CRPS for 2 years or less.
- had a pain score of at least 4 on a scale of 0=no pain to 10=worst pain.
- must have failed at least 2 available treatments for severe lasting pain, 1 of which must have been a pain medication.
- must have been on stable treatment for severe lasting pain for at least 1 month before entering the trial.

What happened during this trial?

This was a Phase 3 trial that compared neridronic acid to a dummy medicine. In Phase 3 trials, the trial medicine is given to a large number of participants with the disease condition in order to learn more about the effects of the trial medicine and its safety.

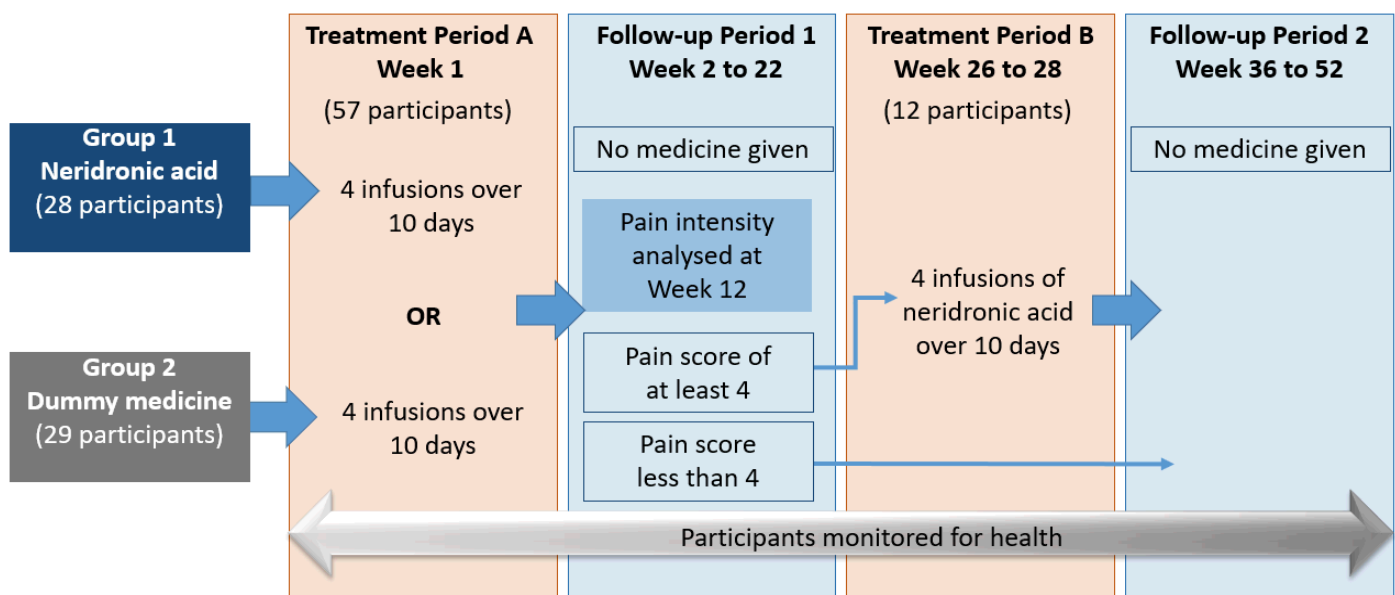
This trial consisted of Treatment Period A with Follow-up Period 1 and Treatment Period B with Follow-up Period 2.

Treatment Period A was “double blind”. This means that neither the participants nor the researchers knew who was given which trial medicine. Trials are sometimes done this way to make sure that the trial results are not biased by this information.

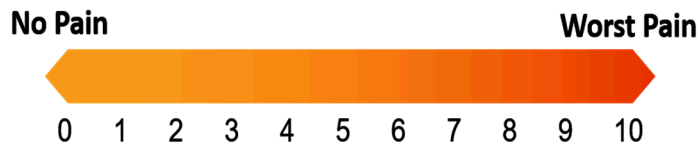
Researchers randomly assigned participants to one of 2 treatment groups using a computer system. This process is called randomisation. It means that each participant could be assigned to any group and it helps to make sure the groups are distributed fairly.

Treatment Period A consisted of 4 infusions of trial medicine over 10 days. Participants in Group 1 received a total dose of 400 milligrams (mg) of neridronic acid into a vein and participants in Group 2 received dummy medicine into a vein. Participants who rated their pain score as at least 4 on a scale of 0=no pain and 10=worst pain could enter **Treatment Period B**, which was “open label”. This means that both the researchers and the participants knew which medicine they took. All participants who entered Treatment Period B received 4 additional infusions of neridronic acid over 10 days.

Participants who did not enter Treatment Period B continued on to Follow-up Period 2. Researchers monitored the health of the participants throughout the trial.



What were the overall results of the trial?



From 1 week before receiving the first dose of trial medicine up until 12 weeks after treatment, every day participants recorded how much pain they felt. Participants measured how strong their pain was using a numerical rating scale from 0=no pain to 10=worst pain.

For this trial, the average pain scores at the end of Week 12 were compared to the average pain scores before the start of the trial treatment. There was a reduction in pain in both neridronic acid and placebo groups. However, researchers could not draw any conclusions on the effect of neridronic acid on easing the pain of the participants in this trial.

During this trial, some participants experienced medical problems that the trial doctor thought could be side effects of the medicine they had taken.

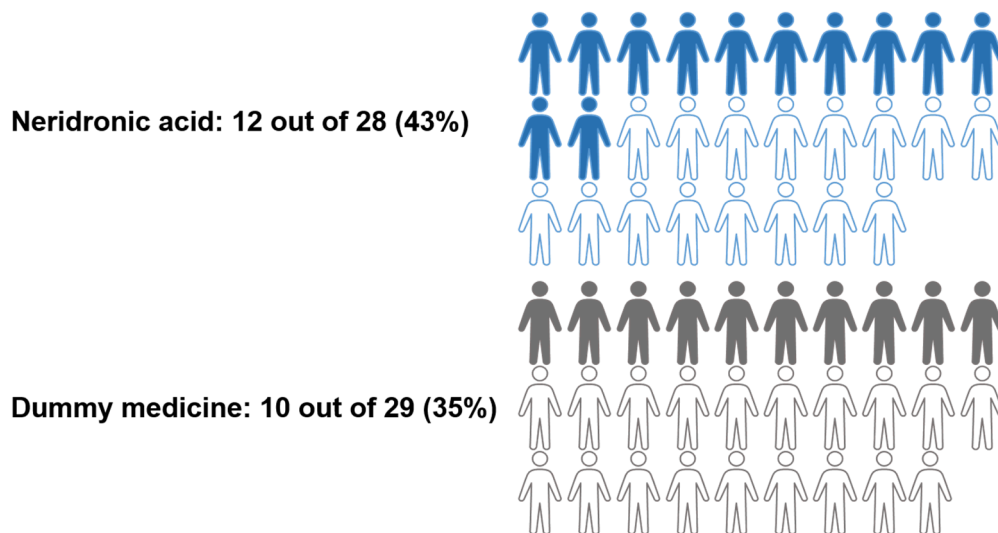
Number of participants with side effects by treatment group

Before Week 26

Serious side effects: Serious side effects are those that may cause death, disability, lasting problems, life-threatening conditions or hospitalisation.

Serious side effects were not reported in either the neridronic acid or dummy medicine group.

Most common side effects:



The most common side effects in participants before Week 26 were non-specific diseases at the infusion site, stomach problems, diseases of muscles and connective tissues and nerve problems.

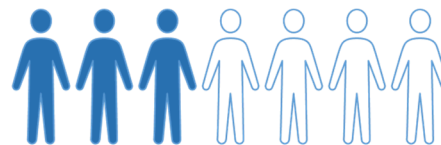
After Week 26

Serious side effects:

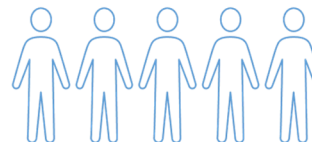
Serious side effects were not reported.

Most common side effects:

**Placebo (Treatment Period A) + Neridronic acid
(Treatment Period B): 3 out of 7 (43%)**



**Neridronic acid (Treatment Period A) + Neridronic acid
(Treatment Period B): 0 out of 5 (0%)**



The most common side effects in participants after Week 26 were related to non-specific diseases at the infusion site. The remaining 22 participants in the placebo group and the 23 participants in the neridronic acid group who did not receive neridronic acid in Treatment Period B, did not report any side effects after Week 26.

How was this trial useful for patients and researchers?

This trial was planned to help researchers learn more about the effects of neridronic acid in participants with severe pain in the hand, wrist, ankle or foot from CRPS lasting for 2 years or less.

The sponsor ended this trial early because the combined initial results from this trial and another neridronic acid trial, indicated that neridronic acid only had a very small chance of providing pain relief to the participants by the end of the trial. Because the trial ended early, researchers could not draw any conclusions on the effect of neridronic acid on easing the pain of the participants in this trial.

Findings from this trial may be used in other trials with neridronic acid. There are no ongoing trials for neridronic acid as at 23-June-2020 and the sponsor does not plan to conduct any more trials for neridronic acid in the future.

The results described in this report are for one trial. The findings of other trials might be different. How neridronic acid works and how safe it is to use must not be judged on the results of one clinical trial alone.

If you have questions, please contact your trial doctor.

Where can I learn more about this trial?

You can find more information about this trial on the following websites:

www.clinicaltrials.gov

Use the NCT identifier
NCT03530345 in the search field.

<https://www.clinicaltrialsregister.eu/ctr-search/search>

Use the EudraCT identifier
2016-003833-91 in the search field.

Full trial name: Randomised, double-blind, placebo-controlled trial investigating the efficacy and safety of intravenous neridronic acid in subjects with complex regional pain syndrome (CRPS)

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