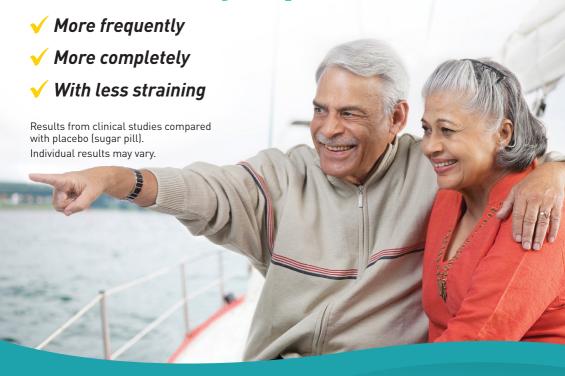


If you're taking opioids for chronic non-cancer pain

Don't stop for constipation Go with Symproic®



What is SYMPROIC?

SYMPROIC is a prescription medicine used to treat constipation that is caused by prescription pain medicines called opioids in adults with long-lasting (chronic) pain that is not caused by active cancer.

It is not known if SYMPROIC is safe and effective in children.

What is opioid-induced constipation?

Opioid-induced constipation (OIC), or constipation due to opioid therapy, is a common and unpleasant effect of opioid treatment.



About 40%-80% of patients receiving opioid therapy report developing OIC.

The most common OIC symptoms include

- · Going less often
- Pushing harder (straining more)
- Not going all the way
- Passing harder stools

Do not take SYMPROIC if you:

- have a bowel blockage (intestinal obstruction) or have a history of bowel blockage.
- are allergic to SYMPROIC or any of the ingredients in SYMPROIC. See the end of this Medication Guide for a complete list of ingredients in SYMPROIC. Tell your healthcare provider or pharmacist before you start or stop any medicines during treatment with SYMPROIC.

How do opioids cause constipation?







It binds to opioid receptors in your brain to reduce or block pain signals, which can help your chronic non-cancer pain.



It binds to opioid receptors found in the digestive system and may cause opioid constipation.

Opioid constipation is different from other types of constipation. Therefore, opioid constipation may last as long as you are on your opioid therapy.*

*Discontinue Symproic if treatment with opioid pain medication is discontinued.

Laxatives do not treat the underlying cause of OIC

Different types of over-the-counter (OTC) laxatives may help relieve constipation in some people. They work by

Coating your intestines to support movement

Stimulating your bowel to move things along

Pulling water into your bowels

However, OTC laxatives may not be enough to treat OIC, especially long term.

Why Symproic®?





More often

Bowel movements occur more frequently

Symproic is a recommended OIC therapy[†]

Symproic is the only OIC therapy with both a **strong recommendation** and **high quality of evidence** from the American Gastroenterological Association.

*For laxative refractory OIC.

SYMPROIC is a prescription medicine used to treat constipation that is caused by prescription pain medicines called opioids, in adults with long-lasting (chronic) pain that is not caused by active cancer. Before you take SYMPROIC, tell your healthcare provider about all of your medical conditions, including if you:

- have any stomach or bowel (intestines) problems, including stomach ulcer, Crohn's disease, diverticulitis, cancer of the stomach or bowel, or Ogilvie's syndrome.
- have liver problems.
- are pregnant or plan to become pregnant. Taking SYMPROIC during pregnancy may cause opioid withdrawal symptoms in your unborn baby. Tell your healthcare provider right away if you become pregnant during treatment with SYMPROIC.



More completely
Bowel movements
feel complete



Symproic° (naldemedine) tablets

- *Results from clinical studies compared with placebo (sugar pill).
- †Individual results may vary.

What are the side effects of Symproic?

SYMPROIC may cause serious side effects, including:

- Tear in your stomach or intestinal wall (perforation). Stomach pain that is severe can be a sign of a serious medical condition. If you get stomach pain that does not go away, stop taking SYMPROIC and get emergency medical help right away.
- Opioid withdrawal. You may have symptoms of opioid withdrawal during treatment with SYMPROIC including sweating, chills, tearing, warm or hot feeling to your face (flush), sneezing, fever, feeling cold, abdominal pain, diarrhea, nausea, and vomiting. Tell your healthcare provider if you have any of these symptoms.

In clinical trials, the most common side effects in patients taking Symproic included:

- Stomach pain/Abdominal pain
- Nausea

Diarrhea

Gastroenteritis

These are not all the possible side effects of Symproic. In clinical studies, up to 3% of patients who took Symproic (and up to 1% of patients who took a placebo/sugar pill) experienced side effects consistent with opioid withdrawal. Tell your healthcare provider if you have any side effects that bother you or that do not go away.

Symproic® fits into your schedule

Taking Symproic won't require you to adjust mealtimes or stop taking laxatives.



Once daily

Take Symproic as prescribed by your doctor (once every day), not only when you feel constipated.



Any time

Choose a time of day that works best for you and continue to take Symproic at approximately the same time every day.



With or without food

Take Symproic before, during, or after a meal. It's totally up to you.



※Symproic®

With or without laxatives

Take Symproic with or without laxatives. It's proven to work either way.

Always remember to ...

- Stop taking Symproic when you stop taking your opioid pain medicine
- Talk to your doctor before you stop taking Symproic

Before you take SYMPROIC, tell your healthcare provider about all of your medical conditions, including if you (cont):

 are breastfeeding or plan to breastfeed. It is not known if SYMPROIC passes into your breast milk. You should not breastfeed during treatment with SYMPROIC and for 3 days after your last dose. Taking SYMPROIC while you are breastfeeding may cause opioid withdrawal symptoms in your baby. You and your healthcare provider should decide if you will take SYMPROIC or breastfeed. You should not do both.

Tell your healthcare provider about all the medicines you take, including prescription and over-the-counter medicines, vitamins, and herbal supplements. Other medicines may affect the way SYMPROIC works.

How should I take SYMPROIC?

- Take SYMPROIC exactly as your healthcare provider tells you to take it.
- Take your prescribed dose of SYMPROIC 1 time each day.
- SYMPROIC can be taken with or without food.
- SYMPROIC has been shown to be effective in people who have taken opioid pain medicines for at least 4 weeks.
- Tell your healthcare provider if you stop taking your opioid pain medicine. If you stop taking your opioid pain medicine, you should also stop taking SYMPROIC.

Talk to your doctor

Pay as little as \$0* for Symproic®



OIC is different from other forms of constipation and may require a treatment that blocks the effects opioids have on the gut. Talk to your doctor about how Symproic may be the right treatment for your opioid-induced constipation.

List of Potential Questions to Discuss with Your Doctor at Your Next Appointment

Since starting your opioid treatment, about how often do you have bowel movements?

Daily

Every other day

Every few days

Once a week or less

How often do you need to use an OTC laxative to help you have a bowel movement?

Always

Most of the time

About half of the time

Never

How often do you physically strain to have a bowel movement?

Daily

Every other day

Every few days

Once a week

How often do you have a bowel movement that feels incomplete?

Every time I go

Most of the time

About half of the time

Sometimes

Never

Commercially insured patients may be eligible to save on their prescription.

How to Save on Symproic

(eligibility requirements apply*)





Visit symproic.com

to download your savings card anytime

Present the Savings Card

to your pharmacist with your Symproic prescription

*For commercially insured patients only. Visit symproic.com for a complete list of terms and conditions



*Maximum savings limit applies; patient out-of-pocket expense may vary.

MEDICATION GUIDE SYMPROIC® (sim proe' ik)

(naldemedine) tablets, for oral use

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

SYMPROIC may cause serious side effects, including:

- **Tear in your stomach or intestinal wall (perforation).** Stomach pain that is severe can be a sign of a serious medical condition. If you get stomach pain that does not go away, stop taking SYMPROIC and get emergency medical help right away.
- **Opioid withdrawal.** You may have symptoms of opioid withdrawal during treatment with SYMPROIC including sweating, chills, tearing, warm or hot feeling to your face (flush), sneezing, fever, feeling cold, abdominal pain, diarrhea, nausea, and vomiting. Tell your healthcare provider if you have any of these symptoms.

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Do not take SYMPROIC if you:

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- SYMPROIC has been shown to be effective in people who have taken opioid pain medicines for at least 4 weeks.
- Tell your healthcare provider if you stop taking your opioid pain medicine. If you stop taking your opioid pain medicine, you should also stop taking SYMPROIC.

What are the possible side effects of SYMPROIC?

See "What is the most important information I should know about SYMPROIC?" The most common side effects of SYMPROIC include stomach (abdomen) pain, diarrhea, nausea and vomiting (gastroenteritis).

Tell your healthcare provider if you have any side effect that bothers you or that does not go away. These are not all the possible side effects of SYMPROIC. Call your doctor for medical advice about side effects. You may report side effects to FDA at 1-800-FDA-1088.

How should I store SYMPROIC?

- Store SYMPROIC at room temperature between 68°F to 77°F (20°C to 25°C).
- Keep SYMPROIC in the bottle that it comes in.

Keep SYMPROIC and all medicines out of the reach of children.

General information about the safe and effective use of SYMPROIC.

Medicines are sometimes prescribed for purposes other than those in a Medication Guide. Do not take SYMPROIC for a condition for which it was not prescribed. Do not give SYMPROIC to other people, even if they have the same symptoms that you have. It may harm them. You can ask your pharmacist or healthcare provider for information about SYMPROIC that is written for health professionals.

What are the ingredients in SYMPROIC?

Active Ingredient: naldemedine tosylate

Inactive ingredients: D-mannitol, croscarmellose sodium, magnesium stearate, hypromellose, talc, and yellow ferric oxide.

Manufactured for: Collegium Pharmaceutical, Inc. Stoughton, MA 02072

Symproic® may help you go*,†



More often

Bowel movements
occur more frequently



More completely Bowel movements feel complete



With less straining
Bowel movements
are easier to pass

Visit symproic.com for resources, potential savings, and more



^{*}Results from clinical studies compared with placebo (sugar pill).

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Please see Medication Guide and full Prescribing Information accompanying this piece, or at symproic.com/PI. Speak to your healthcare provider if you have questions about Symproic.



[†]Individual results may vary.