

Bryan was compensated for his time.

WHAT IS COBENFY (co-BEN-fee)?

COBENFY is an oral prescription medicine used to treat schizophrenia in adults.

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

SELECT IMPORTANT SAFETY INFORMATION Do not take COBENFY if you:

- have urinary retention problems that cause your bladder to not empty completely or not empty at all
- · have moderate or severe liver problems (impairment)
- have gastric retention problems that cause your stomach to empty slowly
- are allergic to COBENFY, xanomeline, or trospium chloride, or any of the ingredients in COBENFY
- have an eye problem called untreated narrow-angle glaucoma

COBENFY significantly improved schizophrenia symptoms



In 2 clinical studies,* COBENFY showed...



Proven **significant symptom relief** vs placebo (sugar pill)



Overall improvement across a range of symptoms*

Positive symptoms

Like hallucinations & delusions Negative symptoms

Like lack of interest or emotion

COBENFY was studied in two 5-week clinical studies with 470 adults aged 18 to 65 diagnosed with schizophrenia. Participants either took COBENFY or a sugar pill.

Individual results may vary.

*A schizophrenia rating scale was used to compare meaningful changes in symptoms overall. These symptoms included positive, negative, and general psychopathology symptoms.

In 1 clinical study,† COBENFY...



Reduced the overall severity of schizophrenia

[†]A different schizophrenia rating tool was used to evaluate meaningful overall changes in the severity of schizophrenia.





Interested in learning more about COBENFY? <u>Click here</u> to sign up for more information.

SELECT IMPORTANT SAFETY INFORMATION

Before taking COBENFY, tell your healthcare provider about all of your medical conditions, including if you:

- have an enlarged prostate, problems passing urine, or a blockage in your urinary bladder
- have liver problems
- have or have had gallstones or problems with your bile ducts or pancreas
- have stomach or intestinal problems, including constipation, ulcerative colitis, slow emptying of your stomach, or myasthenia gravis

SELECT IMPORTANT SAFETY INFORMATION

Before taking COBENFY, tell your healthcare provider about all of your medical conditions, including if you (cont'd):

- · have an eye condition called narrow-angle glaucoma
- have kidney problems

COBENEY is not an antipsychotic. It works differently

COBENFY is a different approach to schizophrenia treatment. COBENFY is a unique combination of 2 medicines: xanomeline and trospium chloride.



Xanomeline (Za-NO-muh-leen) is thought to activate muscarinic receptors in your brain to help adjust important chemical messengers related to your schizophrenia symptoms.



Trospium (TRO-spee-um) chloride may help prevent xanomeline from working outside your brain so that other parts of your body are less likely to be impacted.

The exact way COBENFY works is not fully understood.

COBENFY is the first and only FDA-approved schizophrenia medicine that is thought to work by activating specific small proteins in the brain called muscarinic acetylcholine receptors.*

*Receptors are small proteins (or small structures) found on the cells in your brain and body. Certain medicines can interact with receptors to "activate" them. This may change how your cells function. Muscarinic acetylcholine receptors are types of receptors.

SELECT IMPORTANT SAFETY INFORMATION

Before taking COBENFY, tell your healthcare provider about all of your medical conditions, including if you (cont'd):

· are pregnant or plan to become pregnant. It is not known if COBENFY may harm your unborn baby. Tell your healthcare provider if you become pregnant or think you are pregnant during treatment with COBENFY.



about all of your medical conditions, including if you (cont'd):

- There is a pregnancy exposure registry for women who take COBENFY during pregnancy. The purpose of this registry is to collect information about the health of women exposed to COBENFY and their baby. If you become pregnant during treatment with COBENFY, your healthcare provider will register you by calling 1-866-961-2388 or online at https://womensmentalhealth.org/research/ pregnancyregistry/atypicalantipsychotic/
- · are breastfeeding or plan to breastfeed. It is not known if COBENFY passes into your breast milk or if it can harm your baby.

Possible side effects of COBENEY



Most common side effects experienced by 5% or more of people taking COBENFY

Adverse reaction	COBENFY (n=251)	Sugar pill (n=253)
Nausea	19%	4%
Stomach upset or burning (dyspepsia)	18%	5%
Constipation	17%	7 %
Vomiting	15%	1%
High blood pressure	11%	2%
Stomach (abdominal) pain	8%	4%
Diarrhea	6%	2%
Increased heart rate	5%	2%
Dizziness	5%	2%
Heartburn (gastrointestinal reflux disease)	5%	<1%

These are not all of the possible side effects of COBENEY

Your healthcare provider may lower your dose or stop treatment with COBENFY if you get certain side effects.

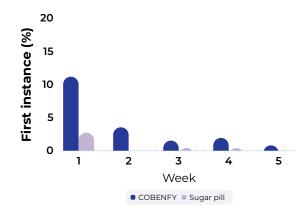
People taking COBENFY had similar rates of uncontrolled facial or body movements (2%) as those taking a sugar pill (<1%).

SELECT IMPORTANT SAFETY INFORMATION

COBENFY may cause serious side effects, including:

Problems with emptying your bladder (urinary retention), risks in people with liver problems, risks in people with bile duct and gallbladder problems (biliary disease), slow emptying of your stomach (decreased gastrointestinal motility), serious allergic reactions (angioedema), an eye problem called narrow-angle glaucoma, increases in heart rate, side effects in people with kidney problems, and central nervous system problems.

First instance of nausea often occurred during the first 2 weeks of treatment and generally decreased over time



If you experience nausea or vomiting, ask your healthcare provider about options that may help.



COBENFY effects on weight Sugar pill – Average change in 5-week studies was 4.96 pounds of weight gain COBENFY – Average change in 5-week studies was 3.04 pounds of weight gain COBENFY – Average change in 1-year studies was 5.64 pounds of weight loss*1 -5.64 lbs

^{*}The results are from 1-year studies where participants and their physicians knew they were taking COBENFY, which may have influenced the results.

[†]On average, participants lost 5.6 lbs in year-long trials. 17.6% of participants had a decrease in body weight of ≥7%. 4.1% of participants had an increase in body weight of ≥7%.

Start and end your day with COBENEY



COBENFY is taken twice a day on an empty stomach



COBENFY is taken twice daily

- Take COBENFY exactly as prescribed. Taking COBENFY twice a day may help keep the medicine at a consistent level in your body throughout the day
- You can take 1 pill when you wake up and 1 pill before bed



Take COBENFY on empty

- Take COBENFY on an empty stomach. Eating food too close to taking COBENFY can make side effects worse
- Take COBENFY at least 1 hour before or at least 2 hours after a meal



Talk to your healthcare provider about how to add COBENFY to your daily routine.

COBENFY is made of xanomeline and trospium chloride. Your healthcare provider will determine which dose is right for you and may change your dose over time.

SELECT IMPORTANT SAFETY INFORMATION

Tell your healthcare provider about all the medicines you take, including prescription and over-the-counter medicines, vitamins, and herbal supplements. Taking COBENFY with certain other medicines may increase your risk of side effects from COBENFY or the other medicine and may affect the way COBENFY or the other medicine works. Know the medicines you take. Keep a list of them to show your healthcare provider and pharmacist when you get a new medicine.

COBENFY comes in different strengths

1. Starting dose for at least 2 days



50 mg/20 mg

When starting COBENFY, the recommended dose is 50 mg/20 mg twice daily for at least 2 days.

2. Increased dose for at least 5 days





100 mg/20 mg

Then your healthcare provider may increase your dose to 100 mg/20 mg twice daily for at least 5 days.

3. Ongoing dose



100 mg/ 20 mg



125 mg/ 30 mg

Your healthcare provider will determine the correct dose.

Pills not shown at actual size. mg/mg is shown as xanomeline/trospium chloride.

Do not change the dose, stop taking COBENFY without talking to your healthcare provider, or open the capsules. If you take too much COBENFY, call your healthcare provider or the Poison Help Line at 1-800-222-1222 or go to the nearest hospital emergency room right away.

Consider your schizophrenia symptoms

It's important to understand the different types of symptoms you may be experiencing so that you can discuss them with your healthcare provider.



Positive symptoms are additional experiences that your brain creates

- ☐ Hallucinations Seeing, hearing, feeling, tasting, or smelling something that is not there
- Delusions Holding beliefs that are not true or based in reality
- Difficulty thinking or speaking



Negative symptoms are like missing pieces of your usual self

- Difficulty staying motivated to complete tasks
- Decreased sense of joy or happiness
- Showing little emotion, not smiling or using facial expressions
- Social withdrawal

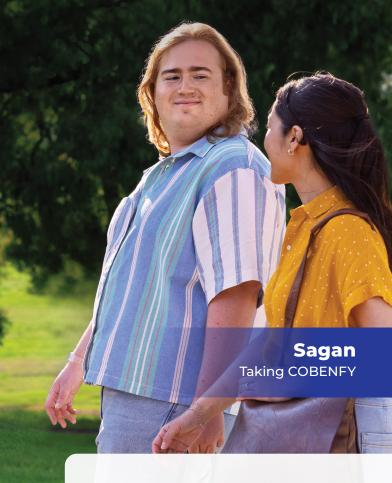


Cognitive symptoms affect how you think and process information

- Trouble with memory
- Difficulty understanding and making decisions
- Difficulty paying attention

This is not a complete list of symptoms, and everyone living with schizophrenia experiences symptoms differently.





<u>Click here</u> to download a daily symptom tracker.

Sagan was compensated for his time.

SELECT IMPORTANT SAFETY INFORMATION

What should I avoid while taking COBENFY?

Do not drive, operate heavy machinery, or do other dangerous activities until you know how COBENFY affects you. COBENFY may cause dizziness, confusion, seeing or hearing things that are not real (hallucinations), and sleepiness.



Support starts with COBENFY Cares™



Access free, 1:1 support 24/7 with a COBENFY prescription

Our team of COBENFY Cares Champions* is available 365 days a year and may be able to help you:



Answer questions about COBENFY



Get ongoing support with regularly scheduled check-ins



Build routines for taking your medication as prescribed



Understand insurance coverage and sign up for a COBENFY Co-Pay Card[†]

*COBENFY Cares Champions are available to provide support to patients who have been prescribed COBENFY. Champions do not provide medical advice or care. COBENFY Cares Champions are provided as a service by Bristol Myers Squibb. Patients should discuss any questions about their medical conditions and treatment options with their healthcare providers.

†COBENFY Cares does not guarantee insurance coverage for COBENFY. Coverage is determined by your insurance provider. Please see COBENFY Terms and Conditions in pocket.

Sign up for our Co-Pay Card

Eligible participants may pay as little as \$0

If you have private or commercial health insurance, you may be eligible to pay as little as \$0 per month through a co-pay savings offer. Certain terms and conditions apply.





Want tips for remembering to take your medicine regularly? COBENFY Cares has tools available.

<u>Click here</u> or call 1-877-COBENFY (1-877-262-3639) to enroll in COBENFY Cares or sign up for more information.

Important Safety Information (cont'd)



COBENFY may cause serious side effects, including:

- Problems with emptying your bladder (urinary retention). See "Do not take COBENFY if you:"

 COBENFY may cause your bladder to not empty completely or not empty at all. You are at increased risk for urinary retention if you are elderly, have a blockage in your bladder, have an enlarged prostate called benign prostatic hyperplasia (BPH), have bladder emptying problems from diabetes, or are taking higher doses of COBENFY. Urinary retention may increase your risk for getting a urinary tract infection. Call your healthcare provider or get emergency help right away if you get any signs or symptoms of urinary retention during treatment with COBENFY, including:
 - difficulty urinating
 - urination in a weak stream or drips
 - urinating frequently
- full bladder and difficulty emptying your bladder
- pain when you urinate
- Risks in people with liver problems. See "Do not take COBENFY if you:" It is not recommended that people with mild liver problems (impairment) take COBENFY because they have an increased risk of getting side effects from COBENFY. Your healthcare provider will check the liver enzyme levels in your blood before starting treatment and as needed during treatment with COBENFY. Tell your healthcare provider if you get any signs or symptoms of liver problems during treatment with COBENFY, including:
 - yellowing of your skin or the white part of your eyes
 - dark urine
 - pain and swelling in the upper right part of your stomach (abdomen)
 - stomach pain that spreads to your back or to below your right shoulder

- itching
- nausea or vomiting
- loss of appetite
- fever
- chills
- light-colored stools
- tiredness

• Risks in people with bile duct and gallbladder problems (biliary disease). COBENFY may cause a blockage in your bile ducts that could lead to gallstones, pancreatitis, and increases in your liver enzymes. Your healthcare provider will check your liver enzyme and bilirubin levels in your blood before starting treatment and as needed during treatment with COBENFY. Tell your healthcare provider if you get any signs or symptoms of biliary disorders during treatment with COBENFY, including:

- stomach upset or
- vomiting
- burning (dyspepsia)
 nausea
- pain in the upper right part of your stomach
- Slow emptying of your stomach (decreased gastrointestinal motility). See "Do not take COBENFY if you:" You are at increased risk for getting decreased gastrointestinal motility if you have ulcerative colitis, already have problems with slow stomach emptying, and have myasthenia gravis. Tell your healthcare provider if you get any signs and symptoms of decreased gastrointestinal motility during treatment with COBENFY, including:
 - constipation
 - vomiting
 - nausea
 - stomach
 - (abdominal) bloating
- stomach (abdominal) pain
- a feeling of fullness after eating just a few bites
- acid reflux

· Serious allergic reactions (angioedema).

Angioedema may happen during treatment with COBENFY and can be life threatening. Stop taking COBENFY and call your healthcare provider or get emergency help right away if you get any of the following signs or symptoms of a serious allergic reaction during treatment with COBENFY, including:

- hives
- swelling of your face, lips, mouth, or tongue
- swelling of your throat
- hoarseness or difficulty speaking
- breathing problems

Important Safety Information (cont'd)

COBENFY may cause serious side effects, including (cont'd):

- · An eye problem called narrow-angle glaucoma. See "Do not take COBENFY if you:" If you already have narrow angles in your eyes, COBENFY may cause a sudden attack (acute angle closure) of glaucoma. Tell your healthcare provider if you get any signs or symptoms of narrow-angle glaucoma during treatment with COBENFY, including:
 - red eyes - blurred vision
 - discomfort - seeing halos or bright - nausea or vomiting colors around lights - severe headache

- eve pain or

- · Increases in heart rate. COBENFY may increase your heart (pulse) rate. Your healthcare provider should check your heart rate before you start treatment and during treatment as needed. Tell your healthcare provider if you get a racing or pounding feeling in your chest during treatment with COBENFY.
- · Side effects in people with kidney problems. People with kidney problems may have an increased risk of getting dry mouth, constipation, stomach upset or burning, urinary tract infection, and urinary retention during treatment with COBENFY.
- · Central nervous system problems. See "What should I avoid while taking COBENFY?"

The most common side effects of COBENFY include:

- nausea
- · stomach (abdominal) pain
- stomach upset or
- diarrhea
- burning (dyspepsia)
- · increased heart rate
- constipation
- dizziness
- vomitina
- · heartburn (gastrointestinal
- high blood pressure reflux disease)

Your healthcare provider may lower your dose or stop treatment with COBENFY if you get certain side effects.

These are not all of the possible side effects of COBENFY.

Call your doctor for medical advice about side effects. You may report side effects to FDA at 1-800-FDA-1088.

Please see Important Safety Information throughout and Full Prescribing Information, including Patient Information.



Notes:		

Click here to hear real stories from people taking COBENFY and others living with schizophrenia.

If you decide to try COBENFY, we'd love to hear your story. Click here to share your experience with us.

Reasons to **BE**lieve in CO**BE**NFY



The first and only FDA-approved muscarinic agonist* for schizophrenia



Proven symptom relief across a range of symptoms, including positive and negative symptoms[†]



It is not an antipsychotic—
it works differently*

*COBENFY is a muscarinic agonist (xanomeline) and muscarinic antagonist (trospium chloride). The exact way COBENFY works is not fully understood.

[†]COBENFY was studied in two 5-week clinical studies with 470 adults aged 18 to 65 diagnosed with schizophrenia. Participants either took COBENFY or a sugar pill. A schizophrenia rating scale was used to compare meaningful changes in symptoms overall. These symptoms included positive, negative, and general psychopathology symptoms. Individual results may vary.

Ask your healthcare provider if COBENFY may be right for you.

<u>Click here</u> to download a guide to help start the conversation.

SELECT IMPORTANT SAFETY INFORMATION

The most common side effects of COBENFY include nausea, stomach upset or burning (dyspepsia), constipation, vomiting, high blood pressure, stomach (abdominal) pain, diarrhea, increased heart rate, dizziness, and heartburn (gastrointestinal reflux disease).

Please see Important Safety Information throughout and <u>Full Prescribing Information</u>, including <u>Patient Information</u>.

COBENFY. (xanomeline and trospium chloride) capsules 50mg/20mg, 100mg/20mg, 125mg/30mg



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