

Initial REMS approval: 01/2017

RISK EVALUATION AND MITIGATION STRATEGY (REMS)

Shared System REMS Program for Sodium Oxybate Oral Solution

I. GOAL:

The goal of the Sodium Oxybate REMS Program is to mitigate the risks of serious adverse outcomes resulting from inappropriate prescribing, misuse, abuse, and diversion of sodium oxybate by:

A. Informing prescribers, pharmacists, and patients of:

1. The risk of significant CNS and respiratory depression associated with sodium oxybate
2. The contraindication of use of sodium oxybate with sedative hypnotics and alcohol
3. The potential for abuse, misuse, and overdose associated with sodium oxybate
4. The safe use, handling, and storage of sodium oxybate

B. Ensuring that pharmacy controls exist prior to filling prescriptions for sodium oxybate that:

1. Screen for concomitant use of sedative hypnotics and other potentially interacting agents
2. Monitor for inappropriate prescribing, misuse, abuse, and diversion of sodium oxybate
3. Notify prescribers when patients are receiving concomitant contraindicated medications or there are signs of potential abuse, misuse, or diversion

II. REMS ELEMENTS:

A. Medication Guide

A Medication Guide will be dispensed with each sodium oxybate prescription in accordance with 21 CFR 208.24

The sodium oxybate Medication Guides are part of the REMS and are available on the Sodium Oxybate REMS Program website

B. Elements to Assure Safe Use

1. Healthcare Providers who prescribe sodium oxybate products are specially certified

- a. Sodium Oxybate sponsors will ensure that healthcare providers who prescribe sodium oxybate are specially certified in the Sodium Oxybate REMS Program.
- b. To become specially certified to prescribe sodium oxybate, each prescriber must complete and submit to the Sodium Oxybate REMS Program the *Sodium Oxybate REMS Program Prescriber Enrollment Form*, which includes the prescriber agreeing to:
 - i. Review the Prescribing Information (PI) and the *Sodium Oxybate REMS Program Prescriber Brochure*
 - ii. Screen each patient for whom sodium oxybate is prescribed for:
 - 1) History of alcohol or substance abuse
 - 2) History of sleep-related breathing disorders
 - 3) History of compromised respiratory function
 - 4) Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
 - 5) History of depression or suicidality
 - iii. Counsel each patient prior to initiating therapy regarding the serious risks and safe use, handling, and storage of sodium oxybate

- iv. Enroll each patient in the Sodium Oxybate REMS Program by completing and submitting the *Sodium Oxybate REMS Program Patient Enrollment Form* to the Sodium Oxybate REMS Program
 - v. Evaluate each patient within the first 3 months of starting sodium oxybate therapy, including an evaluation of the following. It is recommended that patients be re-evaluated every 3 months thereafter while on sodium oxybate therapy:
 - 1) Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
 - 2) Serious adverse events
 - 3) Signs of abuse and misuse, including:
 - a) An increase in dose or frequency of dosing
 - b) Reports of lost, stolen, or spilled medication
 - c) Drug-seeking behavior
 - vi. Report all potential serious adverse events, including CNS depression, respiratory depression, loss of consciousness, coma, death, and any cases of abuse, misuse, or diversion by calling the Sodium Oxybate REMS Program
- b. The prescriber will complete the *Sodium Oxybate REMS Program Prescription Form* for each new prescription and submit the form to one of the specially certified pharmacies. By completing and signing this form, the prescriber acknowledges:
- i. Having an understanding of:
 - 1) The approved indications of sodium oxybate:
 - a) Treatment of cataplexy in narcolepsy
 - b) Treatment of excessive daytime sleepiness in narcolepsy
 - 2) The serious risks associated with sodium oxybate
 - 3) The Prescribing Information (PI) and the *Sodium Oxybate REMS Program Prescriber Brochure*

- ii. Having screened the patient for the following:
 - 1) History of alcohol or substance abuse
 - 2) History of sleep-related breathing disorders
 - 3) History of compromised respiratory function
 - 4) Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
 - 5) History of depression or suicidality
 - iii. Having counseled the patient on:
 - 1) The serious risks associated with sodium oxybate
 - 2) Contraindications (alcohol and sedative hypnotics) and implications of concomitant use of sodium oxybate with other potentially interacting agents
 - 3) Preparation and dosing instructions for sodium oxybate
 - 4) Risk of abuse and misuse associated with sodium oxybate
 - 5) Risk of operating hazardous machinery, including automobiles or airplanes, for the first 6 hours after taking a dose of sodium oxybate
 - 6) Safe use, handling, and storage of sodium oxybate
 - iv. That sodium oxybate is medically appropriate for the patient
 - v. Having listed all known prescription and nonprescription medications and doses on the *Sodium Oxybate REMS Program Prescription Form*
- c. Sodium Oxybate sponsors will:
- i. Ensure that the *Sodium Oxybate REMS Program Prescriber Enrollment Form* can be completed via fax, mail, or the Sodium Oxybate REMS Program website (www.SodiumOxybateREMSProgram.com)
 - ii. Ensure that the *Sodium Oxybate REMS Program Patient Enrollment Form* can be completed via fax, mail, or the Sodium Oxybate REMS Program website (www.SodiumOxybateREMSProgram.com)

- iii. Ensure that the *Sodium Oxybate REMS Program Prescription Form* can be completed via fax
 - iv. Ensure that materials appended to the Sodium Oxybate REMS document will be made available through the Sodium Oxybate REMS Program website (www.SodiumOxybateREMSProgram.com) or by calling the Sodium Oxybate REMS Program at 855-705-2424
 - v. Ensure that a prescriber is specially certified in the Sodium Oxybate REMS Program only after verification that the *Sodium Oxybate REMS Program Prescriber Enrollment Form* is complete and all certification requirements are met
 - vi. Ensure that prescribers are notified when they are successfully specially certified in the Sodium Oxybate REMS Program and are eligible to prescribe sodium oxybate
 - vii. Ensure that secure, validated, separate, and distinct Sodium Oxybate REMS Program databases (patient database, specially certified prescriber database, specially certified pharmacy database and disenrolled prescriber database) are maintained and will only be queried independently through electronic telecommunication verification (see [Section II.C.1.d.](#))
 - viii. Ensure that specially certified prescribers continue to meet the requirements of the Sodium Oxybate REMS Program and can disenroll noncompliant prescribers if the requirements are not met
- d. The following are part of the Sodium Oxybate REMS Program and are appended:
- i. *Sodium Oxybate REMS Program Prescriber Enrollment Form*
 - ii. *Sodium Oxybate REMS Program Prescriber Brochure*
 - iii. *Sodium Oxybate REMS Program Patient Enrollment Form*
 - iv. *Sodium Oxybate REMS Program Prescription Form*
 - v. *Sodium Oxybate REMS Program Patient Quick Start Guide*
 - vi. Sodium Oxybate REMS Program website (www.SodiumOxybateREMSProgram.com)

2. Sodium oxybate will be dispensed only by pharmacies that are specially certified

- a. The Sodium Oxybate REMS Program will certify pharmacies that dispense sodium oxybate. Sodium oxybate will not be stocked in retail pharmacy outlets. To become specially certified in the Sodium Oxybate REMS Program, pharmacies must agree to:
- i. Designate an authorized representative to complete and submit the *Sodium Oxybate REMS Pharmacy Enrollment Form* on behalf of the pharmacy
 - ii. Ensure that the authorized representative oversees implementation and compliance with the Sodium Oxybate REMS Program by the following:
 - 1) Ensure that all pharmacy staff involved in the Sodium Oxybate REMS Program complete the *Sodium Oxybate REMS Program Certified Pharmacy Training Program Module A*
 - 2) Ensure that all pharmacists who dispense sodium oxybate complete the *Sodium Oxybate REMS Program Certified Pharmacy Training Program Modules A and B*
 - iii. Dispense sodium oxybate only to patients enrolled in the Sodium Oxybate REMS Program pursuant to a valid prescription written by a prescriber specially certified in the Sodium Oxybate REMS Program (see [Section II.B.1.a.](#))
 - iv. Dispense only after obtaining a Pre-Dispense Authorization (PDA) for each sodium oxybate prescription by requesting that the Sodium Oxybate REMS Program access the secure, validated, separate, and distinct Sodium Oxybate REMS Program databases (patient database, specially certified prescriber database, specially certified pharmacy database, and disenrolled prescriber database) that will only be queried independently through electronic telecommunication verification to verify the following:
 - 1) Pharmacy is specially certified
 - 2) Prescriber is specially certified
 - 3) Patient is enrolled
 - 4) Patient has no other known active, overlapping prescriptions for sodium oxybate

- v. Recertify in the Sodium Oxybate REMS Program if the pharmacy designates a new authorized representative
 - vi. Provide 24-7 toll-free access to a pharmacist at a Sodium Oxybate REMS Program specially certified pharmacy
 - vii. Ship sodium oxybate directly to each patient or a patient-authorized adult designee, and track and verify receipt of each shipment of sodium oxybate
 - viii. Limit the first shipment for each patient to a one-month supply of sodium oxybate, and subsequent shipments to no more than a three-month supply of sodium oxybate
 - ix. Report all potential adverse events reported by all sources, including any CNS depression, respiratory depression, loss of consciousness, coma, and death, and any cases of suspected abuse, misuse, or diversion to the Sodium Oxybate REMS Program for documentation
- b. Prior to dispensing sodium oxybate, the specially certified pharmacies will:
- i. With every sodium oxybate prescription, ensure that a pharmacist completes the *Sodium Oxybate REMS Program Patient Counseling Checklist* and submits the checklist to the Sodium Oxybate REMS Program
 - ii. Validate each Sodium Oxybate REMS Program prescription by:
 - 1) Verifying that the prescriber is specially certified, the patient is enrolled and the patient has no other active sodium oxybate prescription by entering all prescriptions in the pharmacy management system, including cash payments by obtaining a pre-dispense authorization (PDA) via electronic telecommunication verification
 - 2) Review patient information obtained from the Sodium Oxybate REMS Program website (www.SodiumOxybateREMSProgram.com) and the *Sodium Oxybate REMS Program Prescription Form*, including:
 - a) Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents that either are unknown to the prescriber or pose a high risk of serious interaction with sodium oxybate
 - b) Alerts and *Sodium Oxybate REMS Program Risk Management*

Report (RMR) Forms regarding potential abuse, misuse, or diversion

- 3) Confirming all prescription information, including patient name and two additional identifiers, prescriber name and information, dose, titration information (if applicable), number of refills, dosing directions, total quantity (days' supply), and concomitant medications
 - 4) Contacting the Xyrem REMS Program by phone to:
 - a) Verify that the patient has no other active prescriptions for sodium oxybate that overlap with the current prescription
 - b) Verify that the patient/prescriber has not been disenrolled in the Xyrem REMS Program for suspected abuse, misuse, or diversion
 - c) Report each prescription filled for sodium oxybate
 - 5) Documenting that the call to the Xyrem REMS Program was completed using the *Sodium Oxybate REMS Program Prescription Form*
- c. Sodium Oxybate REMS Program specially certified pharmacies will ship sodium oxybate directly to each patient using an overnight service. In addition, each Sodium Oxybate REMS Program specially certified pharmacy will verify that:
- i. The shipment will be sent to a patient's confirmed shipping address
 - ii. The patient or patient-authorized adult designee will be available to receive the shipment
 - iii. The sodium oxybate Medication Guide is included with each shipment, and a copy of the *Sodium Oxybate REMS Program Patient Quick Start Guide* is provided to a new patient who has not already received it from the prescriber
 - iv. Receipt of each shipment is confirmed and shipment and receipt dates are provided to the Sodium Oxybate REMS Program to be maintained in the patient database

- d. The Sodium Oxybate REMS Program specially certified pharmacies will monitor and report to the Sodium Oxybate REMS Program all instances of patient or prescriber behavior that give rise to a reasonable suspicion of abuse, misuse, or diversion of sodium oxybate
 - i. Pharmacies will document these events, including all requests for early refills by completing and submitting a *Sodium Oxybate REMS Program RMR Form* to the Sodium Oxybate REMS Program
 - ii. Prior to granting an early refill request or if abuse, misuse, or diversion is suspected, the pharmacist will review the patient's RMR history and any alerts obtained from the Sodium Oxybate REMS Program website (www.SodiumOxybateREMSProgram.com), and ensure the request or concern has been discussed with the prescriber prior to shipping sodium oxybate
 - iii. All reports of lost, stolen, destroyed, or spilled drug will be documented in the Sodium Oxybate REMS Program patient database when a specially certified pharmacy completes and submits a *Sodium Oxybate REMS Program RMR Form* to the Sodium Oxybate REMS Program
 - iv. Repeated reports of lost, stolen, destroyed, or spilled drug may be documented as an alert to the patient profile stored in the Sodium Oxybate REMS Program patient database
 - v. Pharmacies and/or prescribers that are specially certified in the Sodium Oxybate REMS Program may direct that a patient be disenrolled from the Sodium Oxybate REMS Program after reviewing or receiving reports of incidents suggestive of abuse, misuse, or diversion by completing and submitting an *Sodium Oxybate REMS Program RMR Form* to the Sodium Oxybate REMS Program
 - vi. Pharmacies may recommend that a prescriber be disenrolled by submitting a *Sodium Oxybate REMS Program RMR Form* to the Sodium Oxybate REMS Program. Sodium Oxybate sponsors will review the information and determine if the prescriber should be disenrolled
- e. Sodium Oxybate Sponsors will:
 - i. Ensure that the *Sodium Oxybate REMS Program Pharmacy Enrollment Form* can be completed via fax, or online at www.SodiumOxybateREMSProgram.com

- ii. Ensure the pharmacy is specially certified in the Sodium Oxybate REMS Program only after verification that the *Sodium Oxybate REMS Program Pharmacy Enrollment Form* is complete and all certification requirements are met
 - iii. Ensure that pharmacies are notified when they are successfully specially certified in the Sodium Oxybate REMS Program
 - iv. Ensure that the secure, validated, separate, and distinct Sodium Oxybate REMS Program databases (patient database, specially certified prescriber database, specially certified pharmacy database, and disenrolled prescriber database) are maintained and will only be queried independently through electronic telecommunication verification (see Section II.C.1.d)
 - v. Ensure that specially certified pharmacies continue to meet the requirements of the Sodium Oxybate REMS Program. Non-compliant pharmacies can be disenrolled if the requirements are not met
- f. The following materials are part of the REMS and are appended:
- i. *Sodium Oxybate REMS Program Certified Pharmacy Training Program*
 - ii. *Sodium Oxybate REMS Program Pharmacy Enrollment Form*
 - iii. *Sodium Oxybate REMS Program Patient Counseling Checklist*
 - iv. *Sodium Oxybate REMS Program Risk Management Report Form*

3. Sodium oxybate will be dispensed and shipped only to patients who are enrolled in the Sodium Oxybate REMS Program with documentation of safe use conditions

- a. Sodium Oxybate sponsors will ensure that sodium oxybate is dispensed only by pharmacies that are specially certified in the Sodium Oxybate REMS Program, by direct shipment, to patients or patient-authorized adult designees enrolled in the Sodium Oxybate REMS Program
- b. Sodium Oxybate sponsors will ensure that patients are enrolled in the Sodium Oxybate REMS Program only if a prescriber specially certified in the Sodium Oxybate REMS Program completes the *Sodium Oxybate REMS Program Patient Enrollment Form* and submits the form to the Sodium Oxybate REMS Program

- c. Sodium Oxybate sponsors will ensure that sodium oxybate is dispensed and shipped only to patients who have signed the *Sodium Oxybate REMS Program Patient Enrollment Form* and acknowledged that:
 - i. He/she has been counseled on the serious risks and safe use of sodium oxybate
 - ii. He/she has asked the prescriber any questions they may have about sodium oxybate
- d. Following enrollment, the patient remains in the Sodium Oxybate REMS Program unless they are disenrolled by the Sodium Oxybate REMS Program at the direction of a specially certified prescriber and/or pharmacy. Specially certified pharmacies and/or prescribers can direct that a patient is disenrolled if the pharmacy and/or prescriber suspect abuse, misuse, or diversion. Reasons for disenrollment include multiple suspicious early refill requests or other information that indicates possible abuse, misuse, or diversion
- e. Following disenrollment, the Sodium Oxybate REMS Program will contact the Xyrem REMS Program to report instances of patient/prescriber disenrollment in Sodium Oxybate REMS Program due to suspected abuse, misuse, or diversion and document that the call was completed in the appropriate database.
- f. A disenrolled patient may be re-enrolled in the Sodium Oxybate REMS Program. In order to re-enroll a patient who had been previously disenrolled for suspicions of abuse, misuse, or diversion, one of the specially certified pharmacies must consult with the specially certified prescriber seeking to re-enroll the patient and will communicate all relevant patient history to the specially certified prescriber, and both the specially certified pharmacy and the requesting specially certified prescriber must agree to re-enroll the patient
- g. A patient may change prescribers if the new prescriber is also specially certified in the Sodium Oxybate REMS Program, and the new prescription does not overlap with another active prescription for sodium oxybate

C. Implementation System

1. The Implementation System for the Sodium Oxybate REMS Program includes the following:

- a. Sodium Oxybate sponsors will ensure that sodium oxybate is distributed only by wholesalers/distributors that have registered with the Sodium Oxybate REMS Program. Registered wholesalers/distributors will only sell/distribute to

pharmacies specially certified in the Sodium Oxybate REMS Program

- b. Sodium Oxybate sponsors will ensure that sodium oxybate is dispensed only by pharmacies that are specially certified in the Sodium Oxybate REMS Program. Sodium oxybate will not be stocked in retail pharmacy outlets
- c. Sodium oxybate will be shipped only to patients enrolled in the Sodium Oxybate REMS Program pursuant to a valid prescription written by a prescriber specially certified in the Sodium Oxybate REMS Program that does not overlap with another active prescription for sodium oxybate
- d. Sodium Oxybate sponsors will ensure that the secure, validated, separate, and distinct Sodium Oxybate REMS Program databases (patient database, specially certified prescriber database, specially certified pharmacy database, and disenrolled prescriber database) are maintained and will only be queried independently through electronic telecommunication verification.
- e. Completed data forms, prescription and distribution data, as well as information related to dosing, concomitant medications, and behavior that raises suspicion of abuse, misuse, or diversion, including complete RMR histories, will be contained only in the appropriate database. The Sodium Oxybate REMS Program will utilize the secure, validated, separate, and distinct databases that will only be queried independently through electronic telecommunication verification, listed below:
 - i. Enrolled patient database
 - ii. Specially certified prescriber database
 - iii. Disenrolled prescriber database
 - iv. Specially certified pharmacy database
 - v. Wholesaler/Distributor database
- f. Sodium Oxybate sponsors will ensure that a sodium oxybate Medication Guide is included with each shipment of sodium oxybate
- g. Sodium Oxybate sponsors will monitor the Sodium Oxybate REMS Program databases for timely reporting to specially certified prescribers and pharmacies of any behavior by enrolled patients or specially certified prescribers in the Sodium Oxybate REMS Program that raises suspicion of abuse, misuse, or diversion

- h. Sodium Oxybate sponsors will monitor the Sodium Oxybate REMS Program databases to ensure compliance with the Sodium Oxybate REMS Program and to evaluate the implementation of the Sodium Oxybate REMS Program. Sodium Oxybate sponsors will ensure that appropriate corrective actions are implemented to address compliance concerns
- i. Sodium Oxybate sponsors must audit the wholesalers/distributors within 90 calendar days after the wholesaler/distributor is authorized to ensure that all processes and procedures are in place and functioning to support the requirements of the Sodium Oxybate REMS Program. Corrective action must be instituted by Sodium Oxybate sponsors if noncompliance is identified
- j. Sodium Oxybate sponsors will audit all specially certified pharmacies after approval of the Sodium Oxybate REMS Program to ensure that each pharmacy implements the Sodium Oxybate REMS Program as directed within 90 calendar days after the pharmacy places its first order of sodium oxybate. Thereafter, Sodium Oxybate sponsors will audit at least 50% of the Sodium Oxybate REMS Program specially certified pharmacy dispensing locations at least annually, identify all issues of noncompliance, and institute appropriate corrective actions, potentially including pharmacy decertification.
- k. The Sodium Oxybate sponsors will monitor the Sodium Oxybate REMS Program for timely reporting of all potential adverse events
- l. Sodium Oxybate sponsors will monitor and evaluate the implementation of the Elements to Assure Safe Use and take reasonable steps to work to improve implementation of these elements

**SODIUM OXYBATE REMS PROGRAM
PRESCRIBER ENROLLMENT FORM**
Sodium oxybate oral solution 500 mg/mL

Sodium Oxybate
REMS Program

Complete this form through www.SodiumOxybateREMSProgram.com.
OR fax the completed form to the Sodium Oxybate REMS Program at 800-353-0987 (toll free),
OR mail to: Sodium Oxybate REMS Program, PO Box XXXXX, City, ST XXXXX-XXXX.
For further information, please call the Sodium Oxybate REMS Program at 855-705-2424.

STEP 1: ALL BOXES BELOW MUST BE CHECKED IN ORDER FOR THE ENROLLMENT PROCESS TO BE COMPLETE AND BEFORE YOU CAN ENROLL PATIENTS AND PRESCRIBE SODIUM OXYBATE

- I understand that sodium oxybate is approved for the treatment of:
- Cataplexy in narcolepsy
 - Excessive daytime sleepiness (EDS) in narcolepsy
- I have read the Prescribing Information (PI) and the *Sodium Oxybate REMS Program Prescriber Brochure* and understand that:
- Sodium oxybate is a Schedule III CNS depressant and can cause obtundation and clinically significant respiratory depression at recommended doses
 - Alcohol and sedative hypnotics are contraindicated in patients who are using sodium oxybate
 - Concurrent use of sodium oxybate with other CNS depressants, including but not limited to opioid analgesics, benzodiazepines, sedating antidepressants or antipsychotics, sedating anti-epileptics, general anesthetics, muscle relaxants, and/or illicit CNS depressants, may increase the risk of respiratory depression, hypotension, profound sedation, syncope, and death
 - Patients who have sleep apnea or compromised respiratory function (e.g., asthma, COPD, etc.) may be at higher risk of developing respiratory depression, loss of consciousness, coma, and death with sodium oxybate use

I agree to:

- Enroll each patient in the Sodium Oxybate REMS Program
- Screen each patient for history of alcohol or substance abuse, sleep-related breathing disorders, compromised respiratory function, depression, suicidality, and concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
- Counsel each patient prior to initiating therapy on the serious risks and safe use, handling, and storage of sodium oxybate
- Evaluate patients within the first 3 months of starting sodium oxybate. It is recommended that patients be re-evaluated every 3 months thereafter while taking sodium oxybate
- Report all potential adverse events, including CNS depression, respiratory depression, loss of consciousness, coma, and death, and any cases of suspected abuse, misuse, or diversion to the Sodium Oxybate REMS Program

STEP 2: TO HELP EXPEDITE THE ENROLLMENT PROCESS, PLEASE PRINT CLEARLY (*denotes required field)

Prescriber Information			
*FIRST NAME:	M.I.:	*LAST NAME:	PROF. DESIGNATION: (MD, DO, PA, NP):
*DEA No.:	*STATE LICENSE No.:		
FACILITY/PRACTICE NAME:			NPI No.:
*STREET ADDRESS:			
*CITY:	*STATE:	*ZIP CODE:	
*PHONE:	*FAX:	EMAIL:	
*PREFERRED METHOD OF CONTACT: <input type="checkbox"/> EMAIL <input type="checkbox"/> FAX			
OFFICE CONTACT:		OFFICE CONTACT PHONE:	

STEP 3: PRESCRIBER SIGNATURE IS REQUIRED BELOW FOR ENROLLMENT IN THE SODIUM OXYBATE REMS PROGRAM

By signing below, I acknowledge the above attestations, and I understand that my personally identifiable information provided above will be shared with the Sodium Oxybate REMS Program, its agents, contractors, and affiliates and entered into a prescriber database for the Sodium Oxybate REMS Program. I agree that I may be contacted in the future by mail, email, fax, and/or telephone concerning sodium oxybate, the Sodium Oxybate REMS Program, and other sodium oxybate programs and services.

*Prescriber Signature: _____ *Date: _____

Report adverse events by contacting the Sodium Oxybate REMS Program at 855-705-2424.

Sodium Oxybate REMS
Program Prescriber Brochure

Dear Prescriber,

Welcome to the Sodium Oxybate REMS Program, which was developed in collaboration with the Food and Drug Administration (FDA) as a Risk Evaluation and Mitigation Strategy (REMS). A REMS is a strategy to manage known or potential serious risks associated with a drug product and is required by the FDA to ensure that the benefits of the drug outweigh its risks.

This brochure provides valuable information about the Sodium Oxybate REMS Program that includes important prescribing information, educational and counseling requirements, and materials necessary for program certification and prescribing sodium oxybate oral solution, including:

- *Sodium Oxybate REMS Program Prescriber Enrollment Form* – a one-time certification is required for all prescribers of sodium oxybate.
- *Sodium Oxybate REMS Program Patient Enrollment Form* – a one-time patient enrollment in the Sodium Oxybate REMS Program is required for each new patient for whom sodium oxybate will be prescribed.
- *Sodium Oxybate REMS Program Prescription Form* – required for prescribing sodium oxybate. This form must be used for new prescriptions and may also be used for refills and renewals of sodium oxybate prescriptions.
- *Sodium Oxybate REMS Program Patient Quick Start Guide* – answers important questions for patients about how to get sodium oxybate, how to use sodium oxybate properly, and how to store it safely. It also gives important information about the risks associated with sodium oxybate.

The *Sodium Oxybate REMS Program Prescriber Enrollment Form* and *Sodium Oxybate REMS Program Patient Enrollment Form* must be completed in full and sent to the Sodium Oxybate REMS Program. The *Sodium Oxybate REMS Program Prescription Form* must be completed in full and sent to one of the certified pharmacies. For your convenience, the *Sodium Oxybate REMS Program Prescriber Enrollment Form* and the *Sodium Oxybate REMS Program Patient Enrollment Form* are available online at www.SodiumOxybateREMSProgram.com and all three forms can be requested by calling the Sodium Oxybate REMS Program toll-free at 855-705-2424. A certified pharmacy in the Sodium Oxybate REMS Program is responsible for processing prescriptions for sodium oxybate.

Continue reading this brochure to learn more about the Sodium Oxybate REMS Program and your responsibilities as a prescriber of sodium oxybate. Please review the Prescribing Information (PI) for sodium oxybate.

Sodium oxybate may be dispensed only to patients enrolled in the Sodium Oxybate REMS Program.

Sodium oxybate is approved for:

- **Treatment of cataplexy in narcolepsy**
- **Treatment of excessive daytime sleepiness (EDS) in narcolepsy**

If you require any additional assistance or information, please call the Sodium Oxybate REMS Program at 855-705-2424 or visit www.SodiumOxybateREMSProgram.com.

Sincerely,

Sodium Oxybate sponsors

IMPORTANT SAFETY INFORMATION

CONTRAINDICATIONS

- Sodium oxybate is contraindicated in patients being treated with sedative hypnotics.
- Patients should not drink alcohol when using sodium oxybate.
- Sodium oxybate is contraindicated in patients with succinic semialdehyde dehydrogenase deficiency.

WARNINGS AND PRECAUTIONS

CNS Depression

- Sodium oxybate is a CNS depressant. Concurrent use of sodium oxybate with other CNS depressants, including but not limited to opioid analgesics; benzodiazepines; sedating antidepressants, antipsychotics, or anti-epileptics; general anesthetics; muscle relaxants; and/or illicit CNS depressants, may increase the risk of respiratory depression, hypotension, profound sedation, syncope, and death.
 - If use of these CNS depressants in combination with sodium oxybate is required, dose reduction or discontinuation of one or more CNS depressants (including sodium oxybate) should be considered.
 - If short-term use of an opioid (e.g., post- or perioperative) is required, interruption of treatment with sodium oxybate should be considered.
- Patients who have sleep apnea or compromised respiratory function may be at a higher risk of developing respiratory depression, loss of consciousness, coma, and death with sodium oxybate use.

Healthcare providers should caution patients about operating hazardous machinery for the first 6 hours after taking a dose of sodium oxybate.

Abuse and Misuse

- Sodium oxybate is a Schedule III controlled substance.
- Sodium oxybate, is the sodium salt of gamma-hydroxybutyrate (GHB), a Schedule I controlled substance. Abuse of illicit GHB, either alone or in combination with other CNS depressants, is associated with CNS adverse events, including seizure, respiratory depression, decreases in the level of consciousness, coma, and death. Illicit GHB has also been associated with drug-facilitated sexual assault.
- The rapid onset of sedation, coupled with the amnesic features of sodium oxybate, particularly when combined with alcohol, has proven to be dangerous for the voluntary and involuntary user (e.g. assault victim).
- You should carefully evaluate patients for a history of drug abuse and follow such patients closely, observing them for signs of misuse or abuse of sodium oxybate (e.g. increase in size or frequency of dosing; reports of lost, stolen, or spilled medication; drug-seeking behavior; feigned cataplexy).

Sodium Oxybate REMS Program

- Sodium oxybate is to be prescribed only to patients enrolled in the Sodium Oxybate REMS Program. Sodium oxybate is available only through a restricted distribution program called the Sodium Oxybate REMS Program. Required components of the Sodium Oxybate REMS Program are:
 - Healthcare providers who prescribe sodium oxybate must be certified. To be certified, prescribers must complete the *Sodium Oxybate REMS Program Prescriber Enrollment Form* and comply with the Sodium Oxybate REMS Program requirements.
 - Sodium oxybate will be dispensed only by pharmacies that are certified.
 - Sodium oxybate will be shipped only to enrolled patients with documentation of safe use conditions. To be enrolled, patients must sign the *Sodium Oxybate REMS Program Patient Enrollment Form* and acknowledge that they have been counseled on the serious risks and safe use of sodium oxybate.

Further information is available at www.SodiumOxybateREMSProgram.com or 855-705-2424.

Depression, Suicidality, and Other Behavioral/Neuropsychiatric Adverse Events

- The emergence of depression in patients treated with sodium oxybate was seen in clinical trials and requires careful and immediate attention. Patients with a previous history of a depressive illness and/or suicide attempt should be monitored especially carefully for the emergence of depressive symptoms while taking sodium oxybate. Sodium oxybate can cause the emergence of neuropsychiatric adverse events (psychosis, paranoia, hallucination, and agitation), loss of consciousness, and sleepwalking. Patients should be instructed to call their healthcare provider if they experience any of these events.
- Anxiety can also occur in patients treated with sodium oxybate.

Use in Patients Sensitive to High Sodium Intake

- Sodium oxybate has a high sodium content.
- Daily sodium intake should be considered in patients on salt-restricted diets or with heart failure, hypertension, or compromised renal function.

Most Common Adverse Events

- In three controlled clinical trials, the most common adverse reactions (incidence 25% and twice the rate seen with placebo) in sodium oxybate -treated patients were nausea (20%), dizziness (15%), vomiting (11%), somnolence (8%), enuresis (7%), and tremor (5%).
- Of the 398 sodium oxybate treated patients with narcolepsy, 10.3% of patients discontinued because of adverse reactions compared with 2.8% of patients receiving placebo. The most common adverse reaction leading to discontinuation was nausea (2.8%). The majority of adverse reactions leading to discontinuation began during the first few weeks of treatment.
- Please see PI for sodium oxybate.

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Prescribing Information and a Medication Guide are also included.

PRESCRIBING SODIUM OXYBATE – A BRIEF GUIDE

The procedure for writing and dispensing prescriptions for sodium oxybate is outlined below.

PRESCRIBERS OF SODIUM OXYBATE

Prescribing sodium oxybate requires a one-time certification.

- If you are prescribing sodium oxybate for the first time, complete the *Sodium Oxybate REMS Program Prescriber Enrollment Form*, found either in this *Sodium Oxybate REMS Program Prescriber Brochure* or online at www.SodiumOxybateREMSProgram.com. If you choose not to complete the *Sodium Oxybate REMS Program Prescriber Enrollment Form* online, please fax it to the Sodium Oxybate REMS Program at 800-353-0987 or mail to Sodium Oxybate REMS Program, PO Box XXXXX, City, ST XXXXX-XXXX.
- On the *Sodium Oxybate REMS Program Prescriber Enrollment Form*, please confirm that:
 - You understand that sodium oxybate is approved for:
 - Treatment of cataplexy in patients with narcolepsy
 - Treatment of excessive daytime sleepiness (EDS) in patients with narcolepsy
 - You have read and understand the PI and this *Sodium Oxybate REMS Program Prescriber Brochure*
 - You agree to screen each patient for:
 - History of alcohol or substance abuse
 - History of sleep-related breathing disorders
 - History of compromised respiratory function
 - Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
 - History of depression or suicidality
 - You agree to counsel your patients on:
 - The serious risks associated with sodium oxybate
 - Contraindications (alcohol and sedative hypnotics)
 - Risks of concomitant use of sodium oxybate with alcohol and/or other CNS depressants
 - Risk of operating hazardous machinery, including automobiles or airplanes, for the first 6 hours after taking a dose of sodium oxybate
 - Preparation and dosing instructions for sodium oxybate
 - Risk of abuse and misuse associated with use of sodium oxybate
 - Safe use, handling, and storage of sodium oxybate
 - You will enroll each patient in the Sodium Oxybate REMS Program by completing the one-time *Sodium Oxybate REMS Program Patient Enrollment Form* and submitting the form to the Sodium Oxybate REMS Program
 - You will evaluate each patient within the first 3 months of starting sodium oxybate, including an evaluation of the following. It is recommended that patients be re-evaluated every 3 months thereafter while on sodium oxybate therapy:
 - Patient's concomitant medications
 - Serious adverse events
 - Signs of abuse and misuse such as an increase in dose or frequency of dosing; reports of lost, stolen, or spilled medication; and/or drug seeking behavior
 - You will report all potential serious adverse events including CNS depression, respiratory depression, loss of consciousness, coma, and death, and any cases of suspected abuse, misuse, or diversion to the Sodium Oxybate REMS Program

PRESCRIBING SODIUM OXYBATE A BRIEF GUIDE (CONT'D)

- On the *Sodium Oxybate REMS Program Patient Enrollment Form*:
 - Verify that you have provided counseling to each patient about the serious risks associated with the use of sodium oxybate and the safe use conditions as described in the *Sodium Oxybate REMS Program Patient Quick Start Guide*
 - Obtain mandatory patient signature acknowledging that he/she has been counseled on the serious risks and safe use conditions of sodium oxybate and has had the opportunity to ask you any questions he/she may have about sodium oxybate, and the patient grants you the authority to release personal information to the Sodium Oxybate REMS Program, other Sodium Oxybate REMS Programs and its business partners and agents, including the certified pharmacy that will fill the prescription
 - Fax the completed *Sodium Oxybate REMS Program Patient Enrollment Form* to the Sodium Oxybate REMS Program at 855-705-2424, complete online at www.SodiumOxybateREMSProgram.com, or mail to Sodium Oxybate REMS Program, PO Box XXXXX, City, ST XXXXX-XXXX

PRESCRIBING REQUIREMENTS

- Write prescriptions for both new and existing patients using the *Sodium Oxybate REMS Program Prescription Form*. If the patient has a lapse in therapy for 6 months or more, a new prescription will be required.
 - Fill out the form completely and clearly to ensure timely fulfillment of your patient's prescription
 - Verify that you have screened your patient for:
 - History of alcohol or substance abuse
 - History of sleep-related breathing disorders
 - History of compromised respiratory function
 - Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
 - History of depression or suicidality
 - Verify that you have counseled the patient regarding:
 - The serious risks associated with sodium oxybate
 - Contraindications (alcohol and sedative hypnotics)
 - The risks of concomitant use of alcohol or other CNS depressants, including sedating antidepressants, antipsychotics, or anti-epileptics; opioids; benzodiazepines; muscle relaxants; and general anesthetics
 - The risks of operating hazardous machinery, including automobiles or airplanes, for the first 6 hours after taking a dose of sodium oxybate
 - Preparation and dosing instructions for sodium oxybate
 - The risk of abuse and misuse associated with sodium oxybate
 - Safe use, handling, and storage of sodium oxybate (refer to pages 13 & 14 of this brochure for Patient Counseling Information)
 - Provide a list of all current prescription and non-prescription medications and dosages that the patient is currently taking, to the best of your knowledge. This can be done by completing the Medications field on the *Sodium Oxybate REMS Program Prescription Form* or by faxing a separate page from the patient's medical history

NOTE: Prior to dispensing each sodium oxybate prescription (including refills), the certified pharmacy responsible to dispense sodium oxybate to the patient will complete a Drug Utilization Review (DUR) and, during the patient counseling process, will ask the patient about the use of other medicines. If the patient's certified pharmacy learns that the patient is taking a previously undisclosed contraindicated medication (sedative hypnotics), an opioid, or more than one CNS depressant, and the prescriber has not indicated awareness of the concomitant medication, the patient's certified pharmacy will contact and inform the prescriber of the concomitant medication use prior to dispensing sodium oxybate. The patient's certified pharmacy may also contact the prescriber about other concomitant medications of concern.

PRESCRIBING SODIUM OXYBATE A BRIEF GUIDE (CONT'D)

- Verify that you have informed the patient that his or her certified pharmacy will send him/her a copy of the sodium oxybate Medication Guide with each prescription fill and a *Sodium Oxybate REMS Program Patient Quick Start Guide* prior to his/her first prescription fill, if you haven't provided one previously. These materials are available through the Sodium Oxybate REMS Program at www.SodiumOxybateREMSProgram.com
- Access www.SodiumOxybateREMSProgram.com to look up the certified pharmacies
- Fax the completed *Sodium Oxybate REMS Program Prescription Form* and all renewal/refill prescriptions to one of the certified pharmacies

Patient Evaluation

- Evaluate each patient within the first 3 months of starting sodium oxybate therapy, including an evaluation of the following. It is recommended that patients be re-evaluated every 3 months thereafter while they are taking sodium oxybate.
 - Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
 - Serious adverse events
 - Signs of abuse and misuse, such as an increase in dose or frequency of dosing; reports of lost, stolen, or spilled medication; and/or drug-seeking behavior

Refill Prescriptions

- One of the certified pharmacies will send you a *Sodium Oxybate REMS Program Prescription Form* in advance of a patient's prescription expiring or running out of refills. Prescription refills and renewals may also be conveyed by phone or fax to the patient's certified pharmacy, and must be documented in the Sodium Oxybate REMS Program.
 - Fill out the form completely and clearly to ensure timely fulfillment of your patient's prescription
 - Access www.SodiumOxybateREMSProgram.com to look up the certified pharmacies
 - Fax the completed *Sodium Oxybate REMS Program Prescription Form* and all subsequent prescriptions to one of the certified pharmacies

RESPONSIBILITIES OF THE SODIUM OXYBATE REMS PROGRAM CERTIFIED PHARMACIES

FOLLOWING RECEIPT OF A PATIENT'S PRESCRIPTION, A CERTIFIED PHARMACY WILL:

- Provide you with confirmation of each new *Sodium Oxybate REMS Program Prescription Form* received from your office
- Contact the patient's insurance provider to verify sodium oxybate prescription benefits
- Prior to the first shipment, contact the patient to:
 - Confirm whether he or she has received a copy of the *Sodium Oxybate REMS Program Patient Quick Start Guide*. The patient's certified pharmacy will send a copy of the *Sodium Oxybate REMS Program Patient Quick Start Guide* to any patient not previously receiving one from his or her prescriber
 - Counsel the patient using the *Sodium Oxybate REMS Program Patient Counseling Checklist* on expectations from sodium oxybate therapy and how to prepare and take sodium oxybate doses safely and effectively
 - Review important sodium oxybate safety information and precautions for sodium oxybate use
 - Review sodium oxybate safe handling and storage procedures
 - Review the adverse events associated with sodium oxybate
 - Review the patient's use of concomitant medications
 - You will be notified of any potential for drug interactions based on patient counseling
 - Ask if the patient has any questions about sodium oxybate and answer the questions and/or refer the patient back to the prescriber, as appropriate
- Provide 24/7 toll-free telephone access to pharmacist support for prescribers, office staff, and patients by answering questions about safety, dosing, and patient care
- Dispense and ship sodium oxybate by overnight service to the patient or his or her authorized adult designee
- Remind patients about monthly refills
- Contact the prescriber if a prescription refill or renewal is required

For your convenience, materials and information regarding the Sodium Oxybate REMS Program are available online at www.SodiumOxybateREMSProgram.com.

Please be sure to review the Prescribing Information prior to prescribing sodium oxybate for your patients.

GUIDELINES FOR DOSING AND TITRATING SODIUM OXYBATE

DOSING SODIUM OXYBATE

Sodium oxybate is a liquid medication taken orally at bedtime. Due to its short half-life, sodium oxybate is taken in 2 equal doses at night, with the first dose taken at bedtime and the second dose taken 2.5 to 4 hours later.

- **The recommended starting dose is 4.5 g/night divided into 2 equal doses of 2.25 g each**
- The effective dose range is 6 g to 9 g/night
- Doses higher than 9 g/night have not been studied and should not ordinarily be administered
- The dose of sodium oxybate should be titrated to effect
 - Sodium oxybate should be titrated in increments of 1.5 g/night at weekly intervals
- An initial sodium oxybate dose reduction of at least 20% is recommended if divalproex sodium is prescribed to patients already taking sodium oxybate. For patients already taking divalproex sodium, it is recommended that prescribers use a lower starting sodium oxybate dose when introducing sodium oxybate. Prescribers are advised to monitor patient response closely and adjust dose accordingly if concomitant use of sodium oxybate and divalproex sodium is warranted
- Improvement may occur during the first weeks of therapy; however, titration to an optimal dose may take longer
- Once a stable dose is established, patients should be evaluated periodically

Note: the patient's first shipment of sodium oxybate will be limited to a 1-month (30-day) supply, and future shipments cannot exceed a 3-month (90-day) supply.

DOSING AND TITRATION			
	1 st Dose	2 nd Dose	Total Nightly Dose
Recommended Starting Dose	2.25 g	2.25 g	4.5 g
	3 g	3 g	6 g
	3.75 g	3.75 g	7.5 g
Maximum Dose	4.5 g	4.5 g	9 g
			Effective Dosing Range

Please see PI for sodium oxybate for additional guidelines for dosing and titration.

PATIENT DOSING INFORMATION:

- Inform patients that all bottles contain concentrated medication ONLY and that water for dilution is not contained in the box. Advise patients to keep sodium oxybate in the provided bottle(s)
- Patients should prepare both nighttime doses at bedtime
 - Instruct patients to make sure that pharmacy vials are empty prior to preparing each dose
 - Each dose of sodium oxybate should be diluted with about 1/4 cup of water
 - Patients should be instructed to store sodium oxybate bottles and prepared nightly doses in a secure place out of the reach of children and pets
- Food significantly reduces the bioavailability of sodium oxybate; therefore, **doses should be taken at least 2 hours after eating**
- Both doses should be taken while in bed
- The first dose should be taken at bedtime and the second dose 2.5 to 4 hours later

ADDITIONAL INFORMATION ABOUT SODIUM OXYBATE

Sodium oxybate has been placed in a bifurcated federal schedule. Sodium oxybate is a Schedule III controlled substance when used for legitimate medical purposes, as prescribed. The active ingredient, sodium oxybate, or gamma-hydroxybutyrate (GHB), is classified as a Schedule I controlled substance when used for any other reason or by anyone other than for whom it was prescribed. Your patients should be informed that federal law prohibits the transfer of sodium oxybate to any persons other than the patient for whom it was prescribed. If you have any questions regarding this, please call the Sodium Oxybate REMS Program toll-free at 855-705-2424.

Illicit use and abuse of GHB have been reported, including drug-facilitated sexual assault. Prescribers should carefully evaluate patients for a history of drug abuse and follow patients closely, observing them for signs of misuse or abuse of GHB (e.g., increase in dose or frequency of dosing, reports of lost, stolen, or spilled medication, drug-seeking behavior).

WHEN PRESCRIBING A CONTROLLED SUBSTANCE:

- Be judicious when deciding to increase a dose. Make sure the appropriate medical indicators for increasing or altering a dose are present
- Be suspicious of a pattern of excuses for additional refills or repeated requests for additional refills on an emergency basis
- Be vigilant. Recognize that there is potential to abuse sodium oxybate

It is important you know that the Sodium Oxybate REMS Program maintains records about who is prescribing sodium oxybate. These records will be made available to any state or federal agency that requests them.

DEPENDENCE AND TOLERANCE

Dependence

- Cases of severe dependence and cravings for GHB have been reported
- There have been case reports of dependence after illicit use of GHB at frequent repeated doses
 - Doses (18 g/day to 250 g/day) were in excess of therapeutic dose range
- Abstinence syndrome has not been reported in clinical trials

Tolerance

- Open-label, long-term (≥6 months) clinical trials did not demonstrate development of tolerance
- There have been some case reports of symptoms of tolerance developing after illicit use at doses far in excess of the recommended sodium oxybate dosage regimen

Discontinuation effects and tolerance of sodium oxybate have not been systematically evaluated in controlled clinical trials.

For your convenience, materials and information regarding the Sodium Oxybate REMS Program are available online at www.SodiumOxybateREMSProgram.com

USE IN SPECIFIC POPULATIONS

PREGNANCY

Teratogenic Effects: Pregnancy Category C.

Nonteratogenic Effects: Because animal reproduction studies are not always predictive of human response, this drug should be used during pregnancy only if clearly needed.

LABOR AND DELIVERY

Sodium oxybate has not been studied in labor or delivery. In obstetric anesthesia using an injectable formulation of sodium oxybate, newborns had stable cardiovascular and respiratory measures but were very sleepy, causing a slight decrease in Apgar scores. There was a fall in the rate of uterine contractions 20 minutes after injection. Placental transfer is rapid, but umbilical vein levels of sodium oxybate were no more than 25% of the maternal concentration. No sodium oxybate was detected in the infant's blood 30 minutes after delivery. Elimination curves of sodium oxybate between a 2-day-old infant and a 15-year-old patient were similar. Subsequent effects of sodium oxybate on later growth, development, and maturation in humans are unknown.

NURSING MOTHERS

It is not known whether sodium oxybate is excreted in human milk. Because many drugs are excreted in human milk, caution should be exercised when sodium oxybate is administered to a nursing woman.

PEDIATRIC USE

Safety and effectiveness in pediatric patients have not been established.

GERIATRIC USE

There is limited experience with sodium oxybate in subjects 65 years and older. In general, dose selection for an elderly patient should be cautious, usually starting at the low end of the dosing range, reflecting the greater frequency of decreased hepatic, renal, or cardiac function, and of concomitant disease and other drug therapy.

RACE AND GENDER EFFECTS

There were too few non-Caucasian patients in the narcolepsy clinical trials to permit evaluation of racial effects on safety or efficacy. More than 90% of the subjects in the clinical trials were Caucasian.

In the narcolepsy clinical trials, with a database that was 58% female, no important differences in safety or efficacy of sodium oxybate were noted between men and women.

Please read accompanying Prescribing Information.

The Sodium Oxybate REMS Program is here to support you, your staff, and your patients.

For assistance, call 855-705-2424 (toll-free).

PATIENT COUNSELING INFORMATION

Prior to initiating therapy, counsel each patient regarding the serious risks and safe use, handling and storage of sodium oxybate using the *Sodium Oxybate REMS Program Patient Quick Start Guide* and encourage all patients to read the sodium oxybate Medication Guide.

- Inform patients that sodium oxybate is available only through certified pharmacies under a restricted distribution program called the Sodium Oxybate REMS Program and provide them with the telephone number and website for more information about sodium oxybate and the Sodium Oxybate REMS Program
- Confirm that patients understand the serious risks and safe use conditions of sodium oxybate and that you have answered any questions the patient has about sodium oxybate by having the patient sign and date the *Sodium Oxybate REMS Program Patient Enrollment Form*. Inform the patient that regular follow-up is recommended

As a component of the Sodium Oxybate REMS Program, the contents of the sodium oxybate Medication Guide are reviewed with every patient by a Sodium Oxybate REMS Program certified pharmacy before initiating treatment with sodium oxybate.

To ensure safe and effective use of sodium oxybate, you should provide your patient with the following guidance:

ALCOHOL OR SEDATIVE HYPNOTICS

Advise patients not to drink alcohol or take other sedative hypnotics if they are taking sodium oxybate.

SEDATION

Inform patients that after taking sodium oxybate they are likely to fall asleep quickly (often within 5 minutes and usually within 15 minutes), but the time it takes to fall asleep can vary from night to night. The sudden onset of sleep, including in a standing position or while rising from bed, has led to falls resulting in injuries, in some cases requiring hospitalization. Instruct patients to remain in bed following ingestion of their first dose, and not to take their second dose until 2.5 to 4 hours later.

FOOD EFFECTS ON SODIUM OXYBATE

Food significantly decreases the bioavailability of sodium oxybate. Inform patients to take the first dose at least 2 hours after eating.

RESPIRATORY DEPRESSION

Inform patients that sodium oxybate can be associated with respiratory depression even at recommended doses and with concurrent use of sodium oxybate with other CNS depressants.

OPERATING HAZARDOUS MACHINERY

Inform patients that until they are reasonably certain that sodium oxybate does not affect them adversely (e.g., impair judgment, thinking, or motor skills) they should not operate hazardous machinery, including automobiles or airplanes.

SUICIDALITY

Instruct patients or families to contact a healthcare provider immediately if the patient develops depressed mood, markedly diminished interest or pleasure in usual activities, significant change in weight and/or appetite, psychomotor agitation or retardation, increased fatigue, feelings of guilt or worthlessness, slowed thinking or impaired concentration, or suicidal ideation.

SLEEPWALKING

Instruct patients and their families that sodium oxybate has been associated with sleepwalking and to contact their healthcare provider if this occurs.

SODIUM INTAKE

Instruct patients who are sensitive to sodium intake (e.g. those with heart failure, hypertension, or renal impairment) that sodium oxybate contains a significant amount of sodium and they should limit their sodium intake.

SAFE USE, HANDLING, STORAGE, AND DISPOSAL

- Discuss safe and proper use of sodium oxybate and dosing information with patients prior to the initiation of treatment
- Instruct patients to store sodium oxybate bottles and sodium oxybate doses in a secure place, out of reach of children and pets
- Patients should be instructed to divide their total nightly dose into 2 separate doses. They should not further divide each of the 2 separate doses
- Patients should be informed that they should be seen by their healthcare provider frequently to review dose titration, symptom response, and adverse reactions
- Instruct patients to store sodium oxybate at room temperature, between 59°F and 86°F. Inform patients that they may safely dispose of sodium oxybate down the sink or toilet drain
- Inform patients that they must report all instances of lost or stolen sodium oxybate to the local police and to the Sodium Oxybate REMS Program

**SODIUM OXYBATE REMS PROGRAM
PATIENT ENROLLMENT FORM**
Sodium oxybate oral solution 500 mg/mL

Sodium Oxybate
REMS Program

Complete this form through www.SodiumOxybateREMSProgram.com,
OR fax completed form to the Sodium Oxybate REMS Program at 800-353-0987 (toll free),
OR mail to: Sodium Oxybate REMS Program, PO Box XXXXX, City, ST XXXXX-XXXX.
For more information, please call the Sodium Oxybate REMS Program at 855-705-2424.

Please Print (*denotes required field)

Patient Information			
*FIRST NAME:	M.I.:	*LAST NAME:	*PRIMARY PHONE:
*DATE OF BIRTH (MM/DD/YYYY):	*GENDER:	<input type="checkbox"/> M <input type="checkbox"/> F	CELL PHONE:
*ADDRESS:			WORK PHONE:
*CITY:	*STATE:	*ZIP CODE:	EMAIL:
*MEDICATIONS: (list all known current prescription and non-prescription medications and dosages or submit as a separate page) <input type="checkbox"/> Check box if separate page attached			
Insurance Information			
Does Patient Have Prescription Coverage? <input type="checkbox"/> Yes (Please provide photocopy of both sides of insurance identification Card with this form) <input type="checkbox"/> No			
POLICY HOLDER'S NAME:		POLICY HOLDER'S DATE OF BIRTH:	
INSURANCE COMPANY NAME:		RELATIONSHIP TO PATIENT:	
INSURANCE PHONE:	RxID No:	RxGrp No:	
RxBIN No:	RxPCN No:		
Prescriber Information			
*FIRST NAME:	M.I.:	*LAST NAME:	*DEA No.:
*STREET ADDRESS:			*PHONE:
*CITY:	*STATE:	*ZIP CODE:	*FAX:
OFFICE CONTACT:	OFFICE CONTACT PHONE:	*NPI No.:	

PATIENT: FORM MUST BE SIGNED BEFORE ENROLLMENT CAN BE PROCESSED

By signing below, I acknowledge that:

- My doctor/prescriber has counseled me on the serious risks and safe use of sodium oxybate
- I have asked my doctor/prescriber any questions I have about sodium oxybate
- I understand that my personally identifiable information provided above will be shared with the Sodium Oxybate REMS Program, its agents, contractors, and affiliates, and entered into a patient database for the Sodium Oxybate REMS Program
- I understand that my personally identifiable information provided above will be shared with other sodium oxybate REMS programs, its agents, contractors, and affiliates

*Patient/Guardian Signature: _____ *Date: _____

*Printed Guardian Name (if applicable): _____

PRESCRIBER: FORM MUST BE SIGNED BEFORE ENROLLMENT CAN BE PROCESSED

By signing below, I acknowledge that:

- I have counseled the patient about the serious risks associated with the use of sodium oxybate and the safe use conditions as described in the *Sodium Oxybate REMS Program Patient Quick Start Guide*
 I have provided the patient with the *Sodium Oxybate REMS Program Patient Quick Start Guide* (optional)

*Prescriber Signature: _____ *Date: _____

**SODIUM OXYBATE REMS PROGRAM
PRESCRIPTION FORM**

Sodium oxybate oral solution 500 mg/mL

Sodium Oxybate
REMS Program

Fax the completed *Sodium Oxybate REMS Program Prescription Form* to one of the certified pharmacies for the patient.
You can look up certified pharmacies on www.SodiumOxybateREMSProgram.com,
or call the Sodium Oxybate REMS Program at 855-705-2424.
For more information, please call the Sodium Oxybate REMS Program at 855-705-2424.

Please Print (*denotes required field)

Prescriber Information			
*FIRST NAME:	M.I.:	*LAST NAME:	*DEA No.:
*STREET ADDRESS:			*PHONE:
*CITY:	*STATE	*ZIP CODE:	*FAX:
OFFICE CONTACT:	OFFICE CONTACT PHONE:		*NPI No.:
Patient Information			
*FIRST NAME:	M.I. (opt):	*LAST NAME	*PRIMARY PHONE:
*DATE OF BIRTH (MM/DD/YYYY):	GENDER: <input type="checkbox"/> M <input type="checkbox"/> F		CELL PHONE:
*ADDRESS:			WORK PHONE:
*CITY:	*STATE:	*ZIP CODE:	EMAIL:
*MEDICATIONS: (list all known current prescription and non-prescription medications and dosages or submit as a separate page) <input type="checkbox"/> Check box if separate page attached			

Please complete either the fixed dosing or titrated dosing section.

Fixed Sodium Oxybate Dosing

Dose: First dose (bedtime): _____ g + Second dose (2.5 to 4 hours later): _____ g = _____ g Total Nightly Dose

Titrated Sodium Oxybate Dosing (First dose is at bedtime; second dose is taken 2.5 to 4 hours later)

Starting Dose	First dose: _____ g +	Second dose: _____ g	=	Total Nightly Dose for _____ days
1 st Titration:	First dose: _____ g +	Second dose: _____ g	_____ g	Total Nightly Dose for _____ days
2 nd Titration:	First dose: _____ g +	Second dose: _____ g	=	Total Nightly Dose for _____ days
3 rd Titration:	First dose: _____ g +	Second dose: _____ g	_____ g	Total Nightly Dose for _____ days

Dispensing Instructions

Total Quantity: 1 2 3 month(s) supply (circle one) (initial prescription fill cannot exceed 1 month of therapy; refills cannot exceed 3 months).	Refills: 0 1 2 3 4 5 (circle one)
Directions: Take first dose p.o., diluted in ¼ cup of water at bedtime. Take second dose p.o., diluted in ¼ cup of water 2.5 to 4 hours later. Note: Prepare both doses at the same time prior to bedtime. The sodium oxybate shipment does not include water for dilution.	
Special Instructions:	

Prescriber Verification – My signature below signifies that: I understand the statements and agree to the Sodium Oxybate REMS Program requirements which are found on the back of this form; sodium oxybate is medically appropriate for this patient; and, I have informed the patient that the Sodium Oxybate REMS Program will send him or her a copy of the sodium oxybate Medication Guide with each prescription fill and a *Sodium Oxybate REMS Program Patient Quick Start Guide* prior to his or her first prescription fill, if I have not previously provided one.

*Prescriber Signature: _____ *Date: _____
Supervising Physician Signature: _____ Date: _____
(if required by state law for prescriptions written by NPs or PAs)

Note: This form may not satisfy all legal requirements for prescribing sodium oxybate in your state. Please submit all prescriptions in accordance with applicable state laws.

Prescriber: Signature verification is required on the **front** page of this *Sodium Oxybate REMS Program Prescription Form* as acknowledgment that you have an understanding of and/or agree to the following:

I understand that sodium oxybate is approved for:

- Treatment of cataplexy in narcolepsy
- Treatment of excessive daytime sleepiness (EDS) in narcolepsy

I understand that:

- Sodium oxybate is a CNS depressant and can cause obtundation and clinically significant respiratory depression at recommended doses
- Alcohol and sedative hypnotics are contraindicated in patients who are using sodium oxybate
- Concurrent use of sodium oxybate with other CNS depressants, including but not limited to opioid analgesics, benzodiazepines, sedating antidepressants or antipsychotics, sedating anti-epileptics, general anesthetics, muscle relaxants, and/or illicit CNS depressants, may increase the risk of respiratory depression, hypotension, profound sedation, syncope, and death
 - If use of these CNS depressants in combination with sodium oxybate is required, dose reduction or discontinuation of one or more CNS depressants (including sodium oxybate) should be considered
 - If short-term use of an opioid (e.g., post- or perioperative) is required, interruption of treatment with sodium oxybate should be considered
- Patients who have sleep apnea or compromised respiratory function (e.g., asthma, COPD, etc.) may be at higher risk of developing respiratory depression, loss of consciousness, coma, and death with sodium oxybate use
- Sodium oxybate is a Schedule III controlled substance with potential for abuse and misuse
- Safe use, handling and storage by patients is important in order to prevent abuse/misuse and accidental exposure to family/friends including children
- Sodium oxybate is to be prescribed only to patients enrolled in the Sodium Oxybate REMS Program

I have read and understand the Prescribing Information (PI) and *Sodium Oxybate REMS Program Prescriber Brochure*.

I have screened this patient for:

- History of alcohol or substance abuse
- History of sleep-related breathing disorders
- History of compromised respiratory function
- Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
- History of depression or suicidality

I have counseled this patient on:

- The serious risks associated with sodium oxybate
- Contraindications (alcohol and sedative hypnotics)
- Risk of concomitant use of sodium oxybate with alcohol, other CNS depressants, or other potentially interacting agents
- Preparation and dosing instructions for sodium oxybate
- Risk of abuse and misuse associated with use of sodium oxybate
- Risk of operating hazardous machinery, including automobiles or airplanes, for the first 6 hours after taking a dose of sodium oxybate
- Preparation and dosing instructions for sodium oxybate
- Safe use, handling, and storage of sodium oxybate

Pharmacy Use Only – My signature below signifies that: I have contacted the Xyrem REMS Program to:

- Verify that the patient has no other active prescriptions for sodium oxybate that overlap with the current prescription
- Verify the patient/prescriber has not been disenrolled in the Xyrem REMS Program for suspected abuse, misuse, or diversion
- Report this prescription filled for sodium oxybate

*Pharmacist Name (please print): _____ *Phone: _____

*Pharmacist Signature: _____ *Date: _____

Read this **Quick Start Guide** and the sodium oxybate Medication Guide carefully before you start taking sodium oxybate.

YOUR DOCTOR HAS PRESCRIBED
SODIUM OXYBATE ORAL SOLUTION

Frequently asked questions about the safe use and handling of sodium oxybate

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Dear Patient,

Welcome to the Sodium Oxybate REMS Program. You are receiving these materials because your healthcare provider has prescribed sodium oxybate oral solution for you. Sodium oxybate is a medicine used to treat excessive daytime sleepiness and/or cataplexy in patients with narcolepsy.

Because of the serious risks associated with sodium oxybate, the Food and Drug Administration (FDA) has required a special program called a Risk Evaluation and Mitigation Strategy (REMS) for sodium oxybate. The purpose of the Sodium Oxybate REMS Program is to make sure the benefits of sodium oxybate outweigh the risks. All patients must be enrolled in the Sodium Oxybate REMS Program to receive sodium oxybate. This *Quick Start Guide* and the sodium oxybate Medication Guide contain information you need to know about sodium oxybate and will help you to use sodium oxybate correctly. Read this *Quick Start Guide* and the sodium oxybate Medication Guide before you start taking sodium oxybate.

After your healthcare provider sends your enrollment form to the Sodium Oxybate REMS Program and first prescription for sodium oxybate to your certified pharmacy, you will receive a call from your certified pharmacy of the Sodium Oxybate REMS Program to tell you how the Sodium Oxybate REMS Program helps you get started with taking sodium oxybate and to answer any questions you may have about sodium oxybate.

You will also speak with appropriate staff at a certified pharmacy, who will go over your insurance information with you. Before you can receive your first shipment of sodium oxybate, a pharmacist at a certified pharmacy must confirm whether you have read and understood this *Quick Start Guide*, ask you about your medical history and other medications you may be taking, and give you advice on how to prepare and take your sodium oxybate and how to store it safely. **You must take this call before you can get your sodium oxybate.**

Please call your healthcare provider if you have questions about sodium oxybate, or you can contact the Sodium Oxybate REMS Program toll free at 855-705-2424. You can reach your certified pharmacy through this number 24 hours a day, 7 days a week with any questions.

We hope you find this information and the Sodium Oxybate REMS Program services helpful.

Sincerely,

Sodium Oxybate sponsors

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

**WARNING: Sodium oxybate can cause
serious side effects.**

Do not drink alcohol or take other medicines that make you sleepy.

Sodium oxybate is a prescription medicine used to treat patients with narcolepsy to reduce too much daytime sleepiness and to reduce cataplexy (suddenly weak or paralyzed muscles).

Important information about sodium oxybate includes the following:

- When taking sodium oxybate, do not drink alcohol or take other medicines that slow your breathing or mental activity or make you sleepy. You could have serious side effects
- Sodium oxybate can cause serious side effects, including trouble breathing while asleep, confusion, unusual or disturbing thoughts, depression, and passing out, even at recommended doses. Tell your healthcare provider if you have any of these problems while taking sodium oxybate
- Abuse of sodium oxybate can lead to dependence (a physical need to take the drug), craving for the medicine, and severe withdrawal symptoms (symptoms that start when the drug is stopped, especially when it is stopped suddenly)

(continued on next page)

- Patients usually fall asleep in about 5 to 15 minutes, although some patients have reported falling asleep more quickly (without first feeling drowsy) and others take more time. The time that it takes to fall asleep might be different from night to night. You should take each dose of sodium oxybate while in bed. Take the first dose at bedtime and the second 2 ½ to 4 hours later. You may need to set an alarm to awaken for the second dose
- Do not drive a car, use heavy machinery, fly an airplane, or do anything that is dangerous or that requires you to be alert for the first 6 hours after taking sodium oxybate. When you first start taking sodium oxybate, be careful until you know how sodium oxybate affects you
- Keep sodium oxybate out of the reach of children and pets. Get emergency medical help right away if a child drinks your sodium oxybate
- Report all side effects to your healthcare provider

Any questions? Please call the Sodium Oxybate REMS Program at 855-705-2424.

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more detailed information about sodium oxybate.

What will you find in this booklet?

This booklet answers important questions about how to get your sodium oxybate, how to use sodium oxybate properly, and how to store it safely. It also gives you important information about sodium oxybate.

What is the Sodium Oxybate REMS Program?

Because of the serious risks associated with sodium oxybate, the FDA has required a special program called REMS for sodium oxybate. Enrollment in the Sodium Oxybate REMS Program by prescribers, pharmacies, and patients is required by the FDA to ensure the benefits of sodium oxybate outweigh the risks associated with sodium oxybate. You are enrolled in the program when your healthcare provider sends the enrollment form you signed in his or her office to the Sodium Oxybate REMS Program. At that time, your healthcare provider also sent your prescription for sodium oxybate to a certified pharmacy.

Certified pharmacy staff will review important information about sodium oxybate with you. They will also answer any questions you may have about sodium oxybate.

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Any questions? Please call the Sodium Oxybate REMS Program at 855-705-2424.

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more detailed information about sodium oxybate.

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Any questions? Please call the Sodium Oxybate REMS Program at 855-705-2424.

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more detailed information about sodium oxybate.

ENROLLING IN THE SODIUM OXYBATE REMS PROGRAM

What am I required to do in this program?

As a patient, your responsibility is to discuss the safe use of sodium oxybate with your healthcare provider and to read this *Sodium Oxybate REMS Program Patient Quick Start Guide* before receiving your first sodium oxybate prescription. Be sure to let your healthcare provider know if you are taking other medications or if you have any conditions that might affect your breathing.

You must also read the sodium oxybate Medication Guide that you will receive with each prescription from your certified pharmacy.

Do I have to enroll in this program?

You will be required to sign an enrollment form at your healthcare provider's office in order to receive sodium oxybate. You must verify that you have been counseled by your healthcare provider on the serious risks and safe use of sodium oxybate and that you were able to ask your healthcare provider any questions you have about sodium oxybate.

FILLING YOUR SODIUM OXYBATE PRESCRIPTION

How is my prescription filled?

All sodium oxybate prescriptions are filled only by pharmacies certified in the Sodium Oxybate REMS Program.

What does a certified pharmacy do?

Your healthcare provider sends your sodium oxybate prescription directly to a certified pharmacy.

After your healthcare provider sends in your first prescription of sodium oxybate, you will receive a call from your certified pharmacy to tell you how the Sodium Oxybate REMS Program helps you get started with taking sodium oxybate and to answer any questions you may have about sodium oxybate. A staff member from your certified pharmacy will call you to complete a *Patient Counseling Checklist*. The *Patient Counseling Checklist* will include information about other medications that you are taking and other medical conditions that might increase your risk of serious side effects. Your certified pharmacy will go over the information about how to use sodium oxybate safely and provide a copy of the Medication Guide with each sodium oxybate shipment.

Your certified pharmacy will always ask you where and when you would like your sodium oxybate delivered and who will sign for the shipment. Sodium oxybate will be shipped by an overnight service. When the courier arrives, you or an adult you name must sign for your sodium oxybate.

Any questions? Please call the Sodium Oxybate REMS Program at 855-705-2424.

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more detailed information about sodium oxybate.

FILLING YOUR SODIUM OXYBATE PRESCRIPTION (cont'd)

What will I get with my sodium oxybate prescription?

With each prescription, you will get 1 or more bottles of sodium oxybate (each bottle, whether full or partial, has the concentrated medicine), a sodium oxybate-specific dosing syringe for drawing up your sodium oxybate dose, 2 empty pharmacy containers with child-resistant caps, and a printed Medication Guide.

How do I get my sodium oxybate refills?

Your certified pharmacy will contact you when it is close to your refill time. You may also call your certified pharmacy to schedule your refills.

Can my local pharmacy provide sodium oxybate?

No. You can get your sodium oxybate only from a Sodium Oxybate REMS Program certified pharmacy. You may be able to have your sodium oxybate shipped to your place of work or to a local overnight carrier hub for pickup. Saturday deliveries may also be an option for you. Your certified pharmacy will work with you on the best options available.

INSURANCE COVERAGE

Will insurance pay for my sodium oxybate?

In most cases, YES. A staff member from your certified pharmacy will call and work with your insurance company to help you get coverage for sodium oxybate.

What is the pharmacy's role with my insurance?

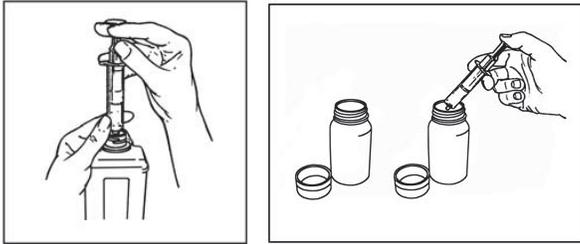
An experienced staff member will:

- Go over your prescription benefits and coverage
- Tell you what your co-pay is, if applicable
- Work with your healthcare provider on prior authorizations, if required by your insurance company

Your certified pharmacy's attempt to get coverage from a third-party payer does not guarantee that you will get coverage.

HOW DO I TAKE MY SODIUM OXYBATE?

Take sodium oxybate only as your healthcare provider tells you to take it.



*For illustration purposes only. Your product may look different.

How do I prepare my doses?

Before going to bed, draw up each of your sodium oxybate doses with the syringe that comes in your shipment. Add each sodium oxybate dose into 1 of the 2 empty pharmacy containers by pushing down on the plunger. Be sure each pharmacy container is empty before adding sodium oxybate into it.

HOW DO I TAKE MY SODIUM OXYBATE? (cont'd)

How do I prepare my doses? (cont'd)

Any questions? Please call the Sodium Oxybate REMS Program at 855-705-2424.

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more detailed information about sodium oxybate.

Add about $\frac{1}{4}$ cup of water to each dose of sodium oxybate. Then place the child-resistant caps onto the pharmacy containers and turn each cap clockwise (to the right) until it clicks and locks in its child-resistant position.

Then put the 2 prepared doses in a safe place by your bed, out of the reach of children and pets.

Place the cap back on the sodium oxybate bottle and store it in a safe and secure place (locked up if needed), out of the reach of children and pets.

Sodium oxybate should always be stored in the bottle provided. Rinse out the syringe and pharmacy containers with water after each use.

How do I take my doses?

Food will lower the amount of sodium oxybate that passes into your body. You should allow at least 2 hours after a meal before taking your first dose of sodium oxybate.

Sodium oxybate is a medicine that can make you sleepy quickly; therefore, take your doses while you are in bed. Take the first dose at bedtime and the second dose 2 ½ to 4 hours later. As with any medicine that causes sleepiness, if you continue evening activities after taking your dose, such as watching television or walking around, you may experience light-headedness, dizziness, nausea, confusion, or other unpleasant feelings.

Any questions? Please call the Sodium Oxybate REMS Program at 855-705-2424.

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more detailed information about sodium oxybate.

HOW DO I TAKE MY SODIUM OXYBATE? (cont'd)

What should I do if I miss a sodium oxybate dose?

- It is very important to take both doses of sodium oxybate each night, as prescribed. If you miss the second dose, skip that dose
 - Do not take sodium oxybate again until the next night
 - Never take both sodium oxybate doses at once
- Any unused sodium oxybate doses that you prepared but didn't take must be thrown away within 24 hours from the time you first prepared your doses

How soon will I see a change in my symptoms?

After starting sodium oxybate, it may take a few weeks or longer to see your symptoms improve. It may also take time to find the right dose that works for you. It is important that you talk with your healthcare provider often when you first start taking sodium oxybate.

Tell your healthcare provider if you don't feel any improvements while taking sodium oxybate. Sodium oxybate may not be right for you.

What are the side effects of sodium oxybate?

Sodium oxybate can cause serious side effects, including breathing problems (slower breathing, trouble breathing, and short periods of no breathing while asleep), mental health problems (confusion, seeing or hearing things that are not real, unusual or disturbing thoughts, feeling anxious or upset, depression, thoughts of suicide), and sleepwalking. If you have any of these side effects, call your healthcare provider right away.

The most common side effects with sodium oxybate are nausea, dizziness, throwing up, bedwetting, and diarrhea. Side effects may increase with higher doses.

These are not the only possible side effects with sodium oxybate. If you are worried about any possible side effects with sodium oxybate, talk with your healthcare provider or the pharmacist at one of the certified pharmacies. You should report all side effects by contacting your healthcare provider, the Sodium Oxybate REMS Program at 855-705-2424, or the FDA at 1-800-FDA-1088.

**Any questions? Please call the
Sodium Oxybate REMS
Program at 855-705-2424.**

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more detailed information about sodium oxybate.

HOW DO I TAKE MY SODIUM OXYBATE? (cont'd)

Are there any precautions I should take while on sodium oxybate?

- While taking sodium oxybate, do not drink alcohol or take medicines that cause sleepiness
- Do not drive a car, use heavy machinery, or do anything that is dangerous or requires you to be alert, for the first 6 hours after taking sodium oxybate. When you first start taking sodium oxybate, be careful until you know how it will affect you
- Before starting sodium oxybate, tell your healthcare provider if you are pregnant, or plan to become pregnant, or if you are breastfeeding. It is not known whether sodium oxybate can pass through your breast milk
- Keep your sodium oxybate in a safe place, out of the reach of children
- Take sodium oxybate while in bed

Tell your healthcare provider and pharmacist about any other medicines you are taking, including prescription and non-prescription medicines, vitamins, and supplements.

It is also important to tell other healthcare providers, including pharmacists, that you are taking sodium oxybate before you start or change any medications.

How often should my healthcare provider check my progress with sodium oxybate?

When you first start taking sodium oxybate, you may need to talk to your healthcare provider often until he or she has determined the best dose for you. You can expect that your dose may need to be adjusted. After your dose has been established, your healthcare provider should check on you every 3 months while you are taking sodium oxybate.

Any questions? Please call the Sodium Oxybate REMS Program at 855-705-2424.

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more detailed information about sodium oxybate.

STORAGE AND SAFETY TIPS AT HOME

How do I store sodium oxybate?

- Always store sodium oxybate in its original bottle
- Store sodium oxybate at room temperature. Do not refrigerate sodium oxybate
- Keep sodium oxybate in a safe place, out of the reach of children and pets. Get emergency medical help (call 911) right away if a child drinks your sodium oxybate

How do I properly dispose of sodium oxybate?

When you have finished a bottle, pour any unused sodium oxybate down the sink or toilet drain. Mark out over the prescription label with a marker to protect your confidentiality before putting the empty bottle in the trash.

If you misplace, lose, or damage your sodium oxybate dosing syringe, contact your certified pharmacy to have it replaced. Do not use a different syringe or try to guess the correct dose.

What if I have concerns about having sodium oxybate in my home?

- If your sodium oxybate is lost or stolen, report the incident right away to the local police and to your certified pharmacy
- Use sodium oxybate only as your healthcare provider tells you. Remember that use of your sodium oxybate by others is illegal
- If you have any questions or concerns, or if you need advice about sodium oxybate, call your healthcare provider or your certified pharmacy

Any questions? Please call the Sodium Oxybate REMS Program at 855-705-2424.

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more detailed information about sodium oxybate.

Where can I get more information about sodium oxybate?

For more information about sodium oxybate, contact the Sodium Oxybate REMS Program

- **Phone:** 855-705-2424
- **Fax:** 800-353-0987 (toll free)
- **Outside the US:** +1-855-705-2424
- **Website:** www.SodiumOxybateREMSProgram.com

----- **Keep this booklet as a helpful reminder** -----

If you have questions or need information,
contact the Sodium Oxybate REMS Program

**Any questions? Please call the
Sodium Oxybate REMS
Program at 855-705-2424.**

SODIUM OXYBATE REMS PROGRAM
Sodium oxybate oral solution
For Patients

Please see the Medication Guide for more
detailed information about sodium oxybate.

SODIUM OXYBATE REMS PROGRAM PHARMACY ENROLLMENT FORM

Sodium oxybate oral solution 500 mg/mL

Sodium Oxybate
REMS Program

For immediate enrollment, please go to www.SodiumOxybateREMSprogram.com.

To submit this form via fax, please complete all required fields below and fax to 800-353-0987. You will receive a confirmation via the contact preference you list below.

Pharmacies must be specially certified in the Sodium Oxybate REMS Program to dispense sodium oxybate. Sodium oxybate will not be stocked in retail pharmacy outlets. To become certified, every pharmacy must designate an authorized representative to:

1. Complete certification using this *Sodium Oxybate REMS Pharmacy Enrollment Form* and submit the completed form to the Sodium Oxybate REMS Program.
2. Provide relevant training to the pharmacy staff and pharmacists in each pharmacy and maintain a record of the training.
3. Ensure the pharmacy enables its Pharmacy Management System (PMS) to support electronic communication with the Sodium Oxybate REMS Program system using established telecommunication standards.

Authorized Representative Responsibilities

By signing this form, I attest that my pharmacy has put processes and procedures in place to:

1. Dispense sodium oxybate only to patients enrolled in the Sodium Oxybate REMS Program pursuant to a valid prescription written by a prescriber certified in the Sodium Oxybate REMS Program.
2. Ensure that all pharmacy staff involved in the Sodium Oxybate REMS Program complete the *Sodium Oxybate REMS Program Certified Pharmacy Training Program* and maintain a record of the training.
3. Ensure that all pharmacists that dispense sodium oxybate complete the pharmacist training in the *Sodium Oxybate REMS Program Certified Pharmacy Training Program* and maintain a record of the training.
4. Obtain a Pre-Dispense Authorization (PDA) for each sodium oxybate prescription by entering all prescriptions in the pharmacy management system, including cash payments.
5. Provide 24-7 toll-free access to a pharmacist at a Sodium Oxybate REMS Program specially certified pharmacy.
6. Recertify in the Sodium Oxybate REMS Program if the pharmacy designates a new authorized representative.
7. Ship sodium oxybate directly to each patient or a patient-authorized adult designee using an overnight service, track and verify receipt of each shipment of sodium oxybate, and provide shipment and receipt dates to the Sodium Oxybate REMS program for documentation.
8. Limit the first shipment for each patient to a one-month supply of sodium oxybate, and subsequent shipments to no more than a three-month supply of sodium oxybate.
9. Include a sodium oxybate Medication Guide with each shipment and provide a copy of the *Sodium Oxybate REMS Program Patient Quick Start Guide* to each new patient.
10. Document and report all potential serious adverse events reported by all sources, including any CNS depression, respiratory depression, loss of consciousness, coma, and death, and any instances of patient or prescriber behavior that give rise to a reasonable suspicion of abuse, misuse, or diversion to the Sodium Oxybate REMS Program.
11. Report all potential adverse events related to suspected abuse, misuse, or diversion, by completing and submitting the *Sodium Oxybate REMS Program Risk Management Report (RMR)* to the Sodium Oxybate REMS Program.
12. Maintain documentation that all processes and procedures are in place and are being followed for the Sodium Oxybate REMS Program and provide upon request to the Sodium Oxybate Sponsors, FDA, or a third party acting on behalf of the Sodium Oxybate Sponsors or FDA.
13. Comply with audits by the Sodium Oxybate Sponsors, FDA, or a third party acting on behalf of the Sodium Oxybate Sponsors or FDA to ensure that all processes and procedures are in place and are being followed for the Sodium Oxybate REMS Program.
14. Prohibit the sale, loan, or transfer of any sodium oxybate inventory to any other pharmacy institution, distributor, or prescriber.

Prior to dispensing sodium oxybate, my pharmacy will:

15. Complete the *Sodium Oxybate REMS Program Patient Counseling Checklist* and its requirements each time sodium oxybate is dispensed and submit the completed checklist to the Sodium Oxybate REMS Program.
16. Validate each sodium oxybate prescription by:
 - a. Verifying that the prescriber is certified, the patient is enrolled and the patient has no other active sodium oxybate prescription by entering all prescriptions in the pharmacy management system, including cash payments by obtaining a PDA via electronic telecommunication verification.
 - b. Confirming all prescription information, including patient name and two additional identifiers, prescriber name and information, dose, titration information (if applicable), number of refills, dosing directions, totally quantity (days' supply), and concomitant medications.
 - c. Contacting the Xyrem REMS Program by phone to verify that the patient has no other active prescriptions for sodium oxybate, the patient/prescriber has not been disenrolled in the Xyrem REMS program for suspected abuse, misuse, or diversion, and to report all prescriptions filled for sodium oxybate.
 - d. Documenting that the call to the Xyrem REMS Program was completed using the *Sodium Oxybate REMS Program Prescription Form*.
17. Review the relevant patient information obtained from the Sodium Oxybate REMS Program website (www.SodiumOxybateREMSProgram.com) and the *Sodium Oxybate REMS Program Prescription Form*, including:
 - a. Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents that either are unknown to the prescriber or pose a high risk of serious interaction with sodium oxybate.
 - b. Alerts and *Sodium Oxybate REMS Program RMR* regarding potential abuse, misuse, or diversion.

Pharmacy Information (All fields required)

Pharmacy Name:

Address:

City:

State:

Zip Code:

NCPDP:

NPI:

DEA:

Authorized Representative Information (All fields required)

First Name:

Last Name:

Phone:

Fax:

Email:

Preferred Contact Method: Phone Fax Email

By signing below, I acknowledge that I will comply with the Authorized Representative Responsibilities outlined on this form.

Authorized Representative Signature:

Date:

Next Steps

1. After completing and signing this form, please fax to 800-353-0987.
2. Once this form is processed, you will receive instructions on submitting test transaction to the Sodium Oxybate REMS Program to ensure that your pharmacy management system has been successfully configured/updated to communicate with the Sodium Oxybate REMS Program.
3. After successful completion of the test transactions, you will receive a pharmacy certification confirmation. Upon receipt, your pharmacy is certified and your pharmacy staff is now eligible to complete their training.

Sodium Oxybate REMS Program

Certified Pharmacy Training Modules A and B

All Sodium Oxybate REMS Program Certified Pharmacy staff and pharmacists must complete **Module A** and the Module A Knowledge Assessment. Pharmacists must also complete **Module B** and the Module B Knowledge Assessment.

Dear Sodium Oxybate REMS Program Certified Pharmacy Staff,

Welcome to the Sodium Oxybate REMS Program, which has been approved by the Food and Drug Administration (FDA) as a Risk Evaluation and Mitigation Strategy (REMS).

The Sodium Oxybate REMS Program

The FDA has determined that a REMS is necessary to ensure that the benefits of sodium oxybate oral solution outweigh the risks of serious adverse outcomes resulting from inappropriate prescribing, misuse, abuse, and diversion of sodium oxybate by:

1. Informing prescribers, pharmacists, and patients of:
 - The risk of significant central nervous system (CNS) and respiratory depression associated with sodium oxybate
 - The contraindication of use of sodium oxybate with sedative hypnotics and alcohol
 - The potential for abuse, misuse, and overdose associated with sodium oxybate
 - The safe use, handling, and storage of sodium oxybate

2. Ensuring that pharmacy controls exist prior to filling prescriptions for sodium oxybate that:
 - Screen for concomitant use of sedative hypnotics and other potential interacting agents
 - Monitor for inappropriate prescribing, misuse, abuse, and diversion of sodium oxybate
 - Notify prescribers when patients are receiving concomitant contraindicated medications or when there are signs of potential abuse, misuse, or diversion.

This training provides information about the Sodium Oxybate REMS Program that includes important information about sodium oxybate and the responsibilities of certified pharmacy staff involved in the dispensing of sodium oxybate.

Sodium oxybate is approved for:

- Treatment of cataplexy in narcolepsy
- Treatment of excessive daytime sleepiness (EDS) in narcolepsy

Sodium oxybate may be prescribed only by prescribers certified in the Sodium Oxybate REMS Program and dispensed only to patients enrolled in the Sodium Oxybate REMS Program and dispensed by pharmacies certified in the Sodium Oxybate REMS Program.

Sincerely,

Sodium Oxybate sponsors

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Sodium Oxybate REMS Program

Certified Pharmacy Training Module A

Training for Pharmacy Staff Involved in the Sodium Oxybate REMS Program

All pharmacy staff within a Sodium Oxybate REMS Program certified pharmacy must complete training on **Module A** and successfully complete the Module A Knowledge Assessment. Training must be completed annually.

MODULE A: SODIUM OXYBATE REMS PROGRAM

Important Safety Information

Indications and Usage

Sodium oxybate oral solution is a central nervous system (CNS) depressant that is indicated for the following:

- Treatment of cataplexy in narcolepsy
- Treatment of excessive daytime sleepiness (EDS) in narcolepsy

Sodium oxybate may be prescribed only by prescribers certified in the Sodium Oxybate REMS Program and dispensed only to patients enrolled in the Sodium Oxybate REMS Program.

How Supplied

Sodium oxybate is shipped from a Sodium Oxybate REMS Program certified pharmacy directly to patients. Each shipment to a patient will contain:

- The prescribed amount of medication, contained in one or more bottles of sodium oxybate
- A press-in-bottle adaptor (PIBA) inserted into the bottle at the certified pharmacies
- A sodium oxybate-specific grams-based oral measuring device (plastic syringe) to measure out each nightly dose
- Two empty pharmacy vials with child-resistant caps for preparation of both nightly doses (sodium oxybate dose mixed with water)
- A sodium oxybate Medication Guide

Controlled Substance Scheduling

The active ingredient in sodium oxybate is sodium oxybate or gamma-hydroxybutyrate (GHB, a known drug of abuse). GHB has been used to facilitate sexual assaults. Because of its rapid sedative effects (particularly when mixed with alcohol) and its colorless and odorless appearance, GHB has been used to "spike" the drinks of unsuspecting victims. Because of its abuse potential, GHB is designated a controlled substance by the Drug Enforcement Administration (DEA) and has been placed in a bifurcated federal schedule:

- GHB products approved by the FDA, such as sodium oxybate, and used as prescribed for therapeutic purposes are Schedule III drugs

The active ingredient of sodium oxybate is classified as a Schedule I controlled substance when used for any other reason or by anyone other than for whom it was prescribed. Federal law prohibits the transfer of sodium oxybate to any persons other than the patient for whom it was prescribed.

Boxed Warning

WARNING: CENTRAL NERVOUS SYSTEM DEPRESSION and MISUSE AND ABUSE

Sodium oxybate is a CNS depressant. In clinical trials at recommended doses obtundation and clinically significant respiratory depression occurred in sodium oxybate-treated patients. Almost all of the patients who received sodium oxybate during clinical trials in narcolepsy were receiving central nervous system stimulants.

Sodium oxybate is the sodium salt of gamma hydroxybutyrate (GHB). Abuse of GHB, either alone or in combination with other CNS depressants, is associated with CNS adverse reactions, including seizure, respiratory depression, decreases in the level of consciousness, coma, and death.

Because of the risks of CNS depression, abuse and misuse, sodium oxybate is available only through a restricted distribution program called the Sodium Oxybate REMS Program, using certified pharmacies. Prescribers must certify and patients must enroll in the Sodium Oxybate REMS Program. For further information go to www.SodiumOxybateREMSProgram.com or call 855-705-2424.

Contraindications

- Sodium oxybate is contraindicated in:
 - Patients who take sedative hypnotic agents.
 - Patients who drink alcohol while using sodium oxybate.
 - Patients with succinic semi aldehyde dehydrogenase deficiency, a rare disorder of inborn error of metabolism variably characterized by mental retardation, hypotonia, and ataxia.

Warnings and Precautions

CNS Depression

- Sodium oxybate is a CNS depressant.
- Concurrent use of sodium oxybate with other CNS depressants, including but not limited to opioid analgesics, benzodiazepines, sedating antidepressants or antipsychotics, sedating anti-epileptic drugs, general anesthetics, muscle relaxants, and/or illicit CNS depressants, may increase the risk of respiratory depression, hypotension, profound sedation, syncope, and death.
 - If use of these CNS depressants in combination with sodium oxybate is required, dose reduction or discontinuation of one or more CNS depressants (including sodium oxybate) should be considered.
 - If short-term use of an opioid (e.g., post- or perioperative) is required, interruption of treatment with sodium oxybate should be considered.
- Patients who have sleep apnea or compromised respiratory function may be at a higher risk of developing respiratory depression, loss of consciousness, coma, and death with sodium oxybate use.
- Healthcare providers should caution patients about operating hazardous machinery for the first 6 hours after taking a dose of sodium oxybate.

Abuse, Misuse, and Diversion

- Sodium oxybate or GHB, is a Schedule I controlled substance. Abuse of illicit GHB, either alone or in combination with other CNS depressants, is associated with CNS adverse events, including seizure, respiratory depression, decreases in the level of consciousness, coma, and death.
- The rapid onset of sedation, coupled with the amnesic features of sodium oxybate, particularly when combined with alcohol, has proven to be dangerous for the voluntary and involuntary user (e.g. assault victim).
- Patients should be carefully evaluated for a history of substance abuse. Patients with a history of drug abuse should be closely monitored for signs of misuse or abuse of GHB (e.g. increase in dose or frequency of dosing, drug-seeking behavior, feigned cataplexy).

For complete safety information, please see the Prescribing Information for sodium oxybate.

Sodium Oxybate REMS Program Requirements

Sodium oxybate may be prescribed only by prescribers certified in the Sodium Oxybate REMS Program and dispensed only to patients enrolled in the Sodium Oxybate REMS Program. Because of the risks of central nervous system depression, abuse, misuse, and diversion, sodium oxybate is available only through a restricted distribution program called the Sodium Oxybate REMS Program.

Required components of this program include:

- Use of a certified pharmacy.
- Healthcare Providers who prescribe sodium oxybate must have completed the *Sodium Oxybate REMS Program Prescriber Enrollment Form* and must comply with the requirements of the Sodium Oxybate REMS Program.
- To receive sodium oxybate, patients must be enrolled in the Sodium Oxybate REMS Program and be counseled on the serious risks and safe use of sodium oxybate treatment. Patients are enrolled by certified prescribers who must fill out and submit the *Sodium Oxybate REMS Program Patient Enrollment Form*. Prescribers must also complete and submit the *Sodium Oxybate REMS Program Prescription Form* to one of the certified pharmacies for all new sodium oxybate prescriptions and for sodium oxybate prescriptions for patients restarting sodium oxybate treatment after not receiving sodium oxybate for 6 months or more.
- Further information is available at www.SodiumOxybateREMSProgram.com.

Overview of Certified Pharmacy Responsibilities

Enrollment Verification

- The *Sodium Oxybate REMS Program Prescriber Enrollment Form* and the *Sodium Oxybate REMS Program Patient Enrollment Form* are sent to the Sodium Oxybate REMS Program by the prescriber.
- Information from the enrollment forms is maintained in the appropriate Sodium Oxybate REMS Program database by the Sodium Oxybate REMS Program.
- No duplicate patients may be enrolled.
- Patients must confirm that they have been counseled on the serious risks and safe use of sodium oxybate; their certified pharmacy will provide counseling with every sodium oxybate prescription dispensed.
- The Sodium Oxybate REMS Program will notify the prescriber of successful certification in the Sodium Oxybate REMS Program, and that he or she is eligible to prescribe sodium oxybate.
 - If there is a delay in shipping while a question about the prescriber's credentials is being resolved, the patient will be notified by their certified pharmacy.
 - If the prescription cannot be filled because a question about the prescriber's credentials could not be resolved, the patient will be notified by their certified pharmacy.
 - The prescriber will be notified by the Sodium Oxybate REMS Program that he/she cannot be certified due to credential verification failure.
- The Sodium Oxybate REMS Program will notify the prescriber of successful patient enrollment in the Sodium Oxybate REMS Program.
- Enrollment status is maintained in the Sodium Oxybate REMS Program.
 - The Sodium Oxybate REMS Program will confirm that the prescriber's DEA and state license numbers are active and that the prescriber has provided all REMS-required attestations.
 - A prescriber may be disenrolled from the program for expired DEA or state licensures or for non-compliance with the Sodium Oxybate REMS Program.
 - Following enrollment, the patient remains in the Sodium Oxybate REMS Program unless their certified pharmacy, and/or certified prescriber determine that the patient should be disenrolled.
- A certified prescriber and/or a certified pharmacy can direct that a patient be disenrolled from the Sodium Oxybate REMS Program.
 - A patient may be disenrolled from the program for non-compliance with the Sodium Oxybate REMS Program, including for multiple suspicious early refill requests, or other information that indicates abuse, misuse, or diversion.
 - The Sodium Oxybate REMS Program will contact a prescriber if an enrollment form is received for a patient previously disenrolled from the program, or for suspicions of abuse, misuse, or diversion, and will provide the prescriber with all relevant patient history.

Prescription Processing

- A certified pharmacy must validate all prescriptions prior to dispensing sodium oxybate. This includes obtaining a Pre-Dispense Authorization (PDA) from the Sodium Oxybate REMS Program for each prescription upon receipt of a *Sodium Oxybate REMS Program Prescription Form*. The issuance of PDA informs the pharmacy that the prescriber is certified and patient is enrolled in the Sodium Oxybate REMS Program and the patient has no other active sodium oxybate prescriptions.
 - The certified pharmacy will process all sodium oxybate prescriptions, including cash payments, through the pharmacy management system (PMS) and obtain a PDA via electronic telecommunication verification to verify the prescriber is certified, the patient is enrolled in the Sodium Oxybate REMS Program and the patient has no other active sodium oxybate prescriptions.
 - To verify the safe use conditions electronically through the PMS, the following prescription information, at a minimum, is required to be submitted upon processing every sodium oxybate prescriptions:
 - Patient First Name
 - Patient Last Name
 - Patient Date of Birth
 - Patient Zip Code
 - Prescriber Identifier on prescription (NPI or DEA)
 - Date of Fill
 - Days' Supply
 - Quantity
 - Product/NDC
 - If all safe use conditions are met, a PDA will be generated by the Sodium Oxybate REMS Program. The PDA will be maintained in the Sodium Oxybate REMS Program patient database, and does not need to be recorded by the pharmacy. The pharmacy is authorized to dispense sodium oxybate upon receiving a PDA.
 - If the safe use conditions are not met, a PDA will not be issued and the pharmacy will be notified of the reason why:
 - Pharmacy is not certified
 - Prescriber is not certified
 - Patient is not enrolled
 - Patient has a known active, overlapping prescription for sodium oxybate
- Before ordering sodium oxybate from your distributor/wholesaler, you must obtain a PDA from the Sodium Oxybate REMS Program. Distributors/wholesalers are required to verify with the Sodium Oxybate REMS Program that you have a PDA before sending sodium oxybate to your pharmacy. Each bottle of Sodium Oxybate is to be ordered on a per-patient basis only.
- Before a prescription for sodium oxybate can be shipped to a patient, the pharmacy must:
 - Verify that the *Sodium Oxybate REMS Program Prescription Form* is complete and signed by the prescriber.
 - Verify the *Sodium Oxybate REMS Program Prescription Form* was received from the prescriber's office.
 - Verify the prescription is dated according to state controlled prescription regulations.
 - Verify the prescription is for only a one-month supply on a patient's first sodium oxybate fill and no more than a 3-month supply on subsequent fills.
 - Verify there are no discrepancies or concerns with the dosing and titration.
 - If there are discrepancies or concerns, the certified pharmacy must contact the prescriber to revise and resubmit the *Sodium Oxybate REMS Program Prescription Form*.
 - Review the patient information contained in the Sodium Oxybate REMS Program patient database using the secure web viewing portal and the *Sodium Oxybate REMS Program Prescription Form* including:
 - Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents that either are unknown to the prescriber or pose a high risk of serious interaction with sodium oxybate.
 - If patient use of a contraindicated medication or other potentially interacting agent is confirmed and if the prescriber has not indicated prior knowledge, then the pharmacist will notify and consult the prescriber about the risks of concomitant medication use and document the call and the prescriber's treatment rationale on the *Sodium Oxybate Patient Counseling Checklist*.
 - Alerts and Sodium Oxybate REMS Program RMRs regarding potential abuse, misuse, or diversion.
 - Contact the Xyrem REMS Program by phone to:
 - Verify that the patient has no other active prescriptions for sodium oxybate that overlap with the current prescription.
 - Verify that the patient/prescriber has not been disenrolled in the Xyrem REMS Program for suspected abuse, misuse, or diversion.
 - Report all prescriptions filled for sodium oxybate.
 - Document that the call to the Xyrem REMS Program was completed using the *Sodium Oxybate REMS Program Prescription Form*.
- If a certified pharmacy receives overlapping prescriptions for sodium oxybate for a patient, the certified pharmacy responsible for dispensing the current prescription will notify and consult each prescriber.

- Prescriptions are considered overlapping when more than one prescription for sodium oxybate is received for a patient from multiple prescribers within an overlapping timeframe.
 - If a certified pharmacy suspects abuse, misuse, or diversion, the prescription should not be filled, the certified pharmacy must complete and submit a *Sodium Oxybate REMS Program RMR* to the Sodium Oxybate REMS Program, and the prescriber will be notified.
 - There are valid reasons why a patient may have overlapping prescriptions, including if the patient moves or changes prescribers, or if the prescriber sends in a new prescription prior to the completion of all refills.
 - A certified pharmacy responsible for dispensing sodium oxybate to a patient must ensure that under these situations a patient does not receive multiple overlapping shipments of sodium oxybate.
- Once a PDA is obtained from the Sodium Oxybate REMS Program, review of the patient information in the patient database using the secure web viewing portal has been performed, and the Xyrem REMS Program has been contacted, the certified pharmacy will contact the patient to schedule shipment and complete the required counseling.
 - For a new patient, the certified pharmacy provides the *Sodium Oxybate REMS Program Patient Quick Start Guide*.
 - A pharmacist must counsel the patient by completing the *Sodium Oxybate REMS Program Patient Counseling Checklist* prior to every dispense of sodium oxybate.
 - The certified pharmacy must submit the *Sodium Oxybate REMS Program Patient Counseling Checklist* to the Sodium Oxybate REMS Program online at www.SodiumOxybateREMSProgram.com or complete a print version and fax to the Sodium Oxybate REMS Program at 800-353-0987.

Shipping

All sodium oxybate is shipped to patients (or their adult designee) by an overnight service with receipt signature required. Certified pharmacies must provide confirmation of receipt of each prescription of sodium oxybate to the Sodium Oxybate REMS Program by accessing the Sodium Oxybate REMS Program website (www.SodiumOxybateREMSProgram.com), or calling the Sodium Oxybate REMS Program (855-705-2424).

- The patient may request an alternate shipping address, which is subject to approval by a pharmacist.
- See [How Supplied](#) for details of the contents of each sodium oxybate shipment.
- Daily tracking reports are generated to confirm the receipt of each order shipped.
- Lost shipments are investigated.

Monitoring for Inappropriate Prescribing, Abuse, Misuse, and Diversion

Certified pharmacies must conduct detailed monitoring on an ongoing basis of patients and prescribers for signs of inappropriate prescribing, abuse, misuse and diversion. Each certified pharmacy will:

- Document early refill requests and instances of patient and prescriber behavior that suggest potential abuse, misuse, or diversion by completing and submitting a *Sodium Oxybate REMS Program RMR* to the Sodium Oxybate REMS Program online at www.SodiumOxybateREMSProgram.com or complete a print version and fax to the Sodium Oxybate REMS Program at 800-353-0987. This information is maintained in the prescriber and/or patient databases in the Sodium Oxybate REMS Program.
 - Direct the Sodium Oxybate REMS Program to disenroll a patient that has demonstrated behavior that suggests potential abuse, misuse, or diversion by completing and submitting a *Sodium Oxybate REMS Program RMR* to the Sodium Oxybate REMS Program. The Sodium Oxybate REMS Program will notify the Xyrem REMS Program that the patient has been disenrolled.
 - Recommend that a prescriber who has demonstrated behavior that suggests potential abuse, misuse, or diversion be disenrolled by submitting a *Sodium Oxybate REMS Program RMR* to the Sodium Oxybate REMS Program. The Sodium Oxybate REMS Program will notify the Xyrem REMS Program that the prescriber has been disenrolled if disenrollment is determined to be the appropriate corrective action.
- Review the patient's Sodium Oxybate REMS Program RMR history and alerts in the Sodium Oxybate REMS Program using the secure pharmacy web viewing portal for the patient database prior to granting an early refill request or if abuse, misuse, or diversion is suspected.
- Discuss early refill requests or other patient incidents with the prescriber so that the prescriber can make a decision to allow or deny the early refill, or to take some other action based on the patient's behavior and history.
- Report all Sodium Oxybate REMS Program RMRs to the Sodium Oxybate REMS Program by completing and submitting the *Sodium Oxybate REMS Program RMR*.
- Determine whether an alert should be placed in the patient's profile in the patient database within the Sodium Oxybate REMS Program for repeated reports of lost, stolen, destroyed, or spilled drug for review prior to shipping sodium oxybate.
- Inform a pharmacist immediately if certified pharmacy staff suspects patients or prescribers of abuse, misuse, or diversion.

Adverse Event Reporting

- Everyone on staff in each certified pharmacy has an essential role to play in the process of collecting information on potential adverse events for reporting to the Sodium Oxybate REMS Program.

- Report all potential adverse events reported by all sources, including any CNS depression, respiratory depression, loss of consciousness, coma, and death, and any cases of suspected abuse, misuse, or diversion by calling the Sodium Oxybate REMS Program at 855-705-2424.
- Report all potential adverse events related to suspected abuse, misuse, or diversion, by completing and submitting the *Sodium Oxybate REMS Program RMR* to the Sodium Oxybate REMS Program online at www.SodiumOxybateREMSProgram.com or by fax to 800-353-0987.

Ongoing Patient Education

Patients in the Sodium Oxybate REMS Program have access to ongoing education while taking sodium oxybate through:

- 24-hour toll-free telephone help line staffed by a pharmacist trained in the Sodium Oxybate REMS Program.
- Continued contact with the certified pharmacy for every refill.
- Sodium Oxybate REMS Program website (www.SodiumOxybateREMSProgram.com).

Sodium Oxybate REMS Program

Certified Pharmacy Training Module B

Sodium Oxybate REMS Program Training for Pharmacists Involved in the Dispensing of Sodium Oxybate

All Sodium Oxybate REMS Program certified pharmacy pharmacists must complete training on Module B (in addition to Module A) and successfully complete the associated Knowledge Assessment. For all pharmacists who dispense sodium oxybate, training must be completed annually.

MODULE B: SODIUM OXYBATE REMS PROGRAM TRAINING FOR PHARMACISTS

All pharmacists involved in dispensing sodium oxybate must complete the following additional training at least annually. The Sodium Oxybate REMS Program requires that pharmacists within a certified pharmacy are thoroughly trained on the requirements of the Sodium Oxybate REMS Program. Training will be conducted by reviewing the Sodium Oxybate REMS Program materials and successfully completing a Knowledge Assessment with 100% accuracy on the requirements of certified pharmacies and pharmacists working within a certified pharmacy. These duties will include:

- Review of PI
- Review of certified pharmacy's internal processes and procedures established to support the Sodium Oxybate REMS Program with an experienced pharmacist
- Execution of the Sodium Oxybate REMS Program Patient Counseling Checklist
- Detailed monitoring including completion of a *Sodium Oxybate REMS Program RMR*, as needed
- Follow-up interactions with patients and prescribers
- Sodium Oxybate REMS Program documentation and processes

Sodium Oxybate REMS Program Requirements

Sodium oxybate may be prescribed and dispensed only to patients enrolled in the Sodium Oxybate REMS Program. Because of the risks of CNS depression, abuse, misuse, and diversion, sodium oxybate is available only through a restricted distribution program called the Sodium Oxybate REMS Program.

Required components of this program include:

- Use of a certified pharmacy.
- Healthcare providers who prescribe sodium oxybate must complete and submit the following to the Sodium Oxybate REMS Program:
 - *The Sodium Oxybate REMS Program Prescriber Enrollment Form*
 - *The Sodium Oxybate REMS Program Patient Enrollment Form*
- Healthcare providers who prescribe sodium oxybate must complete prescriptions for sodium oxybate on the *Sodium Oxybate REMS Program Prescription Form* and submit the completed form to one of the certified pharmacies.
 - After completion of prescription processing, the pharmacy will fax the prescription form to the Sodium Oxybate REMS Program and retain the original.
 - Prescription refills and renewals may be conveyed by phone or fax and must be documented in the Sodium Oxybate REMS Program.
- To receive sodium oxybate, patients must be:
 - Enrolled in the Sodium Oxybate REMS Program.
 - Prescribed sodium oxybate by a prescriber certified in the Sodium Oxybate REMS Program.
 - Counseled on the serious risks and safe use of sodium oxybate.
 - Have only one active sodium oxybate prescription.

Certified Pharmacy Responsibilities

Certified pharmacies will:

- Limit the first prescription fill to a one-month supply of sodium oxybate and no more than a 3-month supply for subsequent prescription fills.
- Report potential adverse events to the Sodium Oxybate REMS Program.
- Notify prescribers when there are signs of potential abuse or misuse or when patients are taking sedative hypnotics, other CNS depressants, or other potentially interacting agents of which the prescriber is not already aware.
- Certified pharmacies must complete and submit a *Sodium Oxybate REMS Program RMR* to the Sodium Oxybate REMS Program for all instances of potential abuse, misuse, or diversion.
- Utilize the Sodium Oxybate REMS Program, which has access to the secure, validated, separate and distinct Sodium Oxybate REMS Program databases (patient database, certified prescriber database, certified pharmacy database, and disenrolled prescriber database) that will only be queried independently through electronic verification, to verify the following:
 - Complete patient enrollment information
 - Complete prescriber certification information
 - Patient information including:
 - Name and two additional identifiers (date of birth, phone number, address, gender)
 - Current and previous prescribers
 - Comorbid conditions and concomitant medications reported by the patient

- Prescription history
- Prescription information including:
 - Date
 - Dose
 - Titration instructions
 - Number of refills
 - Directions
 - Total quantity (volume and number of days' supply)
 - Concomitant medications
- Sodium Oxybate REMS Program Risk Management Reports (RMRs)
- Shipment information, including:
 - Dates of shipments
 - Dates of shipment receipts
 - Patient addresses
 - Designee information
 - Number of shipments sent daily
 - Quantity of sodium oxybate dispensed daily
- Documentation of interactions with prescribers, patients, and other parties.

These data must be available to the Sodium Oxybate REMS Program for review on an ongoing basis to ensure that sodium oxybate is dispensed to enrolled patients only after completion and documentation of safe use conditions. In certain cases, a pharmacist must access a patient's or prescriber's historical data in the Sodium Oxybate REMS Program using the certified pharmacy secure web viewing portal for the patient database and review it prior to dispensing sodium oxybate.

Patient Counseling and Screening

- Certified pharmacies must complete the *Sodium Oxybate REMS Program Patient Counseling Checklist* and submit to the Sodium Oxybate REMS Program for each patient prior to every shipment of sodium oxybate.
- Each time a pharmacist completes the *Sodium Oxybate REMS Program Patient Counseling Checklist*, the pharmacist must:
 - Verify that early refill requests have been thoroughly questioned and approved through the Sodium Oxybate REMS Program RMR procedure (see below).
 - Screen the patient for concomitant use of contraindicated medications (sedative hypnotics), alcohol, other CNS depressants, and other potentially interacting agents.
 - The pharmacist asks the patient if he or she is taking any other medications and can consult external pharmacy databases to identify drug interactions or prescriptions for other drug products that might have been filled at different pharmacies before filling the prescription.
 - If patient use of a contraindicated medication or other potentially interacting agent is confirmed, and if the prescriber has not indicated prior knowledge, then the pharmacist will notify and consult the prescriber about the risks of concomitant medication use prior to shipping sodium oxybate.
 - Screen the patient for other medical conditions.
 - The pharmacist asks the patient what other medical conditions he or she has.
 - If the patient indicates that he or she has a certain medical condition listed on the *Sodium Oxybate REMS Program Patient Counseling Checklist*, the pharmacist counsels the patient, and notifies the prescriber about the medical condition prior to shipping sodium oxybate.
 - Document the results of the patient screening, all reported concomitant medications and comorbid medical conditions, the action(s) taken, and the date the *Sodium Oxybate REMS Program Patient Counseling Checklist* is completed in the Sodium Oxybate REMS Program.
 - Submit the *Sodium Oxybate REMS Program Patient Counseling Checklist* to the Sodium Oxybate REMS Program online at www.SodiumOxybateREMSProgram.com or complete a print version and fax to the Sodium Oxybate REMS Program at 800-353-0987.
- Certified pharmacies must provide patients with 24/7 access to a pharmacist.

Clinical Usage Clarifications

The pharmacist must:

- Review the information on each *Sodium Oxybate REMS Program Prescription Form*.
- Notify and consult the prescriber if there are any clinical usage clarifications required, such as:
 - Dose over maximum recommended dose (9 g/night)
 - Non-standard doses or instructions
 - Possible errors in dosing or titration amounts or directions

If the issue is not resolved with the prescriber, the pharmacist may consult with the Pharmacist in Charge at their certified pharmacy and with the Sodium Oxybate REMS Program.

Prescription Refills

- Up to 5 refills are allowed on a sodium oxybate prescription (per DEA regulations for CIII controlled substances).
- Refills may be conveyed by phone or fax from the prescriber or patient and communicated to the Sodium Oxybate REMS Program by obtaining a PDA.
- For information on the prescription processing requirements see Module A – [Prescription Processing](#)
- Changes in dose require a new prescription.
- Refill orders should be opened at a patient's certified pharmacy when the patient has approximately 10 days of therapy remaining from the previous shipment.
 - A certified pharmacy technician will contact the patient and schedule a shipment. The technician will ask the patient if there has been any change in his or her medications or medical history.
 - The technician will transfer him or her to a pharmacist who must complete the *Sodium Oxybate REMS Program Patient Counseling Checklist*. The patient should be counseled on the use or diagnosis of:
 - Sedative hypnotics (for example, diazepam, phenobarbital, or zolpidem)
 - CNS depressants: including but not limited to opioid analgesics, benzodiazepines, sedating antidepressants or antipsychotics, sedating anti-epileptic drugs, general anesthetics, and muscle relaxants
 - Alcohol
 - Sleep apnea
 - Asthma, COPD, or other conditions affecting his or her breathing
 - Other current medical conditions
 - The pharmacist must complete refill counseling and confirmation of prescriber consultation or notification by completing and submitting the *Sodium Oxybate REMS Program Patient Counseling Checklist* to the Sodium Oxybate REMS Program online (www.SodiumOxybateREMSProgram.com) or by fax (800-353-0987).
- All patient requests for early refills are to be questioned and documented by the pharmacist.
 - An early refill request is a request for sodium oxybate shipment prior to the date of the next shipment.
 - Requests to accommodate shipment logistics (scheduled delivery date falls on a Sunday, holidays, and vacations) are not considered early refills.
 - If the early refill is required due to a dosage increase, a pharmacist must:
 - Confirm the new dosage with the prescriber prior to processing the prescription.
 - If an early refill is requested for any other reason, a pharmacist must:
 - Discuss the request with the patient to evaluate his/her compliance with therapy, assessing for misuse, abuse, and diversion.
 - Evaluate the patient's record in the Sodium Oxybate REMS Program using the certified pharmacy secure web viewing portal for the patient database and review the patient's prior Sodium Oxybate REMS Program RMR history to identify previous reports of early refills or other incidents suggestive of abuse, misuse, and diversion.
 - Contact the prescriber to discuss the request and any prior early refill requests or incidents suggestive of abuse, misuse, and diversion.
 - Send new shipments of sodium oxybate to the patient only if approved by the prescriber.
 - Send new shipments to replace sodium oxybate reported stolen by a patient only after obtaining a copy of the police report filed by the patient.
 - Document the discussion and outcome by completing and submitting the *Sodium Oxybate REMS Program RMR* to the Sodium Oxybate REMS Program online (www.SodiumOxybateREMSProgram.com) or by fax (800-353-0987).

Monitoring and Assessing for Signs of Abuse, Misuse, and Diversion

- Risk management events must be documented in the Sodium Oxybate REMS Program.
 - Risk management events are reported or discovered events outside the norm that give rise to a reasonable suspicion of abuse, misuse, or diversion
 - Examples of events that should generate a Sodium Oxybate REMS Program RMR include but are not limited to:
 - Requests for early refills
 - Patient's misuse or abuse of product
 - Lost, stolen, destroyed, or spilled drug
 - Delivery to incorrect address and not returned
 - Patient claims that product was not delivered while carrier shows receipt of delivery
 - Product tampering
 - Counterfeit product
 - Contaminated product
 - Inquiries and/or arrests by law or regulatory enforcement agencies associated with the misuse, abuse, or diversion of the product
 - Crimes related to the product
 - Sodium Oxybate REMS Program RMRs must document:
 - Patient and/or prescriber identifying information (patient names to be concealed)
 - Reason for report
 - Certified Pharmacy actions
 - Prescriber contact
 - Supporting documentation (if applicable, such as a police report, fire report, DEA Form 106, or shipper investigation report)
 - Pharmacies can request that a patient is monitored by the Sodium Oxybate REMS Program if serious or repeated events give rise to reasonable suspicion of misuse, abuse or diversion.
 - If abuse, misuse, or diversion is suspected, the pharmacist must review the patient's Sodium Oxybate REMS Program RMR history and discuss the incident with the prescriber prior to shipping sodium oxybate.
 - Repeated reports of lost, stolen, destroyed, or spilled drug will be documented as an alert to the patient record stored in the patient database of the Sodium Oxybate REMS Program and will be accessible to the dispensing pharmacist using the secure web viewing portal for the patient database for review prior to shipping drug.
 - Certified pharmacies and/or prescribers may direct the Sodium Oxybate REMS Program to disenroll a patient after review and discussion of incidents suggestive of abuse and misuse, or diversion by completing and submitting a *Sodium Oxybate REMS Program RMR Form* to the Sodium Oxybate REMS Program.. All requests from prescribers to disenroll a patient will be submitted to a certified pharmacy. The certified pharmacy is required to intake the request, then complete and submit the *Sodium Oxybate REMS Program RMR* to the Sodium Oxybate REMS Program to notify of disenrollment.
 - Pharmacies may recommend that a prescriber be disenrolled by submitting a Sodium Oxybate REMS Program RMR Form to the Sodium Oxybate REMS Program. Sodium Oxybate sponsors will review the information and determine if the prescriber should be disenrolled.
 - All *Sodium Oxybate REMS Program RMRs* must be reported to the Sodium Oxybate REMS Program online (www.SodiumOxybateREMSProgram.com) or by fax (800-353-0987).

Shipping Procedures

- Sodium oxybate must be shipped via an overnight service with receipt signature required.
 - Sodium oxybate is shipped directly to the patient or adult designee (18 years, or 21 years if required by carrier) if the patient is not available to receive the order.
- The patient may request an alternate shipping address, which is then subject to approval by a pharmacist.
- If the patient requests Saturday delivery, his or her certified pharmacy will verify with the overnight shipping service that it is available for the shipping address.
- Each sodium oxybate shipment must include:
 - The prescribed amount of medication, contained in one or more bottles of sodium oxybate
 - A press-in-bottle adaptor (PIBA) inserted into the bottle at the certified pharmacies
 - A sodium oxybate-specific grams-based oral measuring device (plastic syringe) to measure out each nightly dose
 - Two empty pharmacy vials with child-resistant caps for preparation of both nightly doses (sodium oxybate dose mixed with water)
 - A sodium oxybate Medication Guide
- Daily tracking reports must be generated by each certified pharmacy to confirm the receipt of each order shipped during the previous 48 hours. Saturday deliveries are confirmed the following Monday.
 - A patient will be contacted if there is no proof of patient or designee signature, if the patient or designee on file did not

sign for the shipment, or if there is a potential incomplete delivery.

- If a shipment is reported lost, an investigation will be launched to find it.
- Receipt of each shipment of sodium oxybate by a patient must be reported to the Sodium Oxybate REMS Program through the website (www.SodiumOxybateREMSProgram.com) or by calling (855-705-2424) by the patient's certified pharmacy.

Inventory Control

The sodium oxybate inventory must be reconciled at the start and end of each business day and recorded in the pharmacy management system. A physical count must match the count in the pharmacy management system). If the sodium oxybate inventory cannot be reconciled for any reason, no other patient orders can be processed until an investigation is completed by the Pharmacist in Charge and approved by the Sodium Oxybate REMS Program. Internal procedures for reconciling the sodium oxybate inventory are subject to audit.

(To be completed by the pharmacist online at www.SodiumOxybateREMSProgram.com or complete a print version and fax to the Sodium Oxybate REMS Program at 800-353-0987 prior to dispensing each sodium oxybate shipment. Include additional requirements (if any) per federal or state requirements that need to be collected as part of the patient counseling process.)

Step 1: Patient Information

- New/restart
- Scheduled refill
- Early refill approved through Risk Management Report (RMR) process

Patient Name: _____ Patient ID Number: _____

Include Pharmacist Name and Date Time Stamp for each section completed

Step 2: Counseling

- Verify that the patient will receive the *Sodium Oxybate REMS Program Patient Quick Start Guide* (if not already received) and that the drug shipment to the patient will include the sodium oxybate Medication Guide.

_____ (Pharmacist Name) ____/____/____ (Date Time)

- Verify that patient has been counseled on **Therapy Expectations** below:
 - During clinical trials with sodium oxybate, many patients with narcolepsy saw some improvement with excessive daytime sleepiness and/or cataplexy in the first weeks after beginning sodium oxybate therapy. However, the response to sodium oxybate varies from patient to patient. It may also take time to find the right dose that works for you. Your doctor will determine the dose that is appropriate for you.
 - Be sure to talk to your doctor about any troubling side effects or if you don't feel any benefits while taking sodium oxybate.
 - For any changes to your prescription, have your doctor call or fax the new prescription change to the pharmacy and NEVER attempt to change the dose yourself.

_____ (Pharmacist Name) ____/____/____ (Date Time)

□ Verify that patient has been counseled on **Preparation and Administration** information below:

- Sodium oxybate should be taken as directed by your doctor (review prescriber's instructions with patient).
 - Prepare each of your doses by placing _____ grams of sodium oxybate in one of the provided pharmacy containers and place ____ grams in the second container. Add 1/4 cup of water to each pharmacy container. The water does not come with sodium oxybate. You can use either tap or bottled water. The solution should remain clear and it will taste salty. Place the child-resistant cap onto the containers and put them in a safe place, out of the reach of children or pets, by your bed.
- Feel free to call your certified pharmacy if you have any questions regarding preparation or how to take your sodium oxybate doses. The Sodium Oxybate REMS Program is also available Monday through Friday, from 8 am to 8 pm Eastern Time, at 855-705-2424, and a pharmacist is always available 24 hours a day, 7 days a week at your certified pharmacy, if needed.

Refer to the Medication Guide for additional information on preparation of your sodium oxybate doses.

- Set alarm to go off 2.5 to 4 hours after you take your first dose.
- When you are ready to go to sleep, sit at your bedside and drink one dose of sodium oxybate and then lie down.
 - Your first dose of sodium oxybate should be taken at least 2 hours after eating as food will decrease the amount of sodium oxybate that your body absorbs.
 - Patients usually fall asleep in about 5 to 15 minutes, although some patients have reported falling asleep more quickly (without first feeling drowsy) and others may take longer to fall asleep. The time that it takes to fall asleep might be different from night to night.
 - Upon waking up, take the second dose of medication as prescribed by your physician.
 - A minimum of 2.5 hours must separate each dose.
 - If you happen to miss a dose, NEVER take two doses of sodium oxybate at once.
- The diluted medication MUST be used within 24 hours of preparation. Discard any unused medication down the sink drain or toilet.
- When you can no longer draw medication out of the bottle with the dispensing device, dispose of your bottle. Use a marker or pen to deface the bottle to protect your confidentiality.
- Be sure to store sodium oxybate in the original bottle in a safe and secure place out of the reach of children and pets. Get emergency help (call 911) right away if a child drinks your sodium oxybate.
- Sodium oxybate should be stored at room temperature.

_____ (Pharmacist Name) ____ / ____ / _____ (Date Time)

- Verify that patient has been counseled on **Precautions needed for sodium oxybate use:**
 - Sodium oxybate is classified as a controlled substance medication. Sodium oxybate must be used only by the person for whom it is prescribed and as directed by the physician. All lost or stolen medication must be reported.
 - Federal law prohibits the transfer of this drug to any person other than the patient for whom it was prescribed.
 - The active ingredient is sodium oxybate. Sodium oxybate is converted to gamma-hydroxybutyrate (GHB) in the body. GHB has been used as a substance of abuse and has been associated with drug-facilitated sexual assault (date rape).
 - Abuse of GHB can lead to dependence (a physical need to take the drug), craving for the medicine, and severe withdrawal symptoms (symptoms that start when the drug is stopped, especially when it is stopped suddenly). Abuse of GHB, with or without other CNS depressants (for example, nortriptyline, oxycodone, or heroin) including alcohol can lead to seizure, trouble breathing, decreases in the level of consciousness, coma, and death.
 - Tell your doctor if you:
 - Are pregnant or plan to become pregnant. It is not known if sodium oxybate can affect your unborn baby.
 - Are breastfeeding. It is not known whether sodium oxybate can pass through the breast milk. Talk to your doctor about the best way to feed your baby if you take sodium oxybate.
 - Have or had depression or tried to harm yourself. You should be watched for new signs of depression.
 - Have liver problems. Your dose may need to be adjusted.
 - Have sleep apnea (short periods of not breathing while you sleep), snoring, or breathing or lung problems. You may have a higher chance of serious breathing problems with sodium oxybate.
 - Have mental health problems.
 - Walk in your sleep.
 - Are on a salt-restricted diet, have high blood pressure, heart failure, or kidney problems. Sodium oxybate contains sodium (salt) and may not be right for you.

_____ (Pharmacist Name) ___/___/____ (Date Time)

□ Verify that patient has been counseled on **Side Effects**:

- In clinical trials, the most commonly observed side effects associated with the use of sodium oxybate included: headache, nausea, dizziness, sleepiness, vomiting, urinary incontinence, and inflammation of the area around the nostrils and the back of the mouth. Some side effects may be more likely to be observed with higher doses of sodium oxybate.
- Sodium oxybate can cause serious side effects, including trouble breathing while asleep, confusion, unusual or disturbing thoughts, depression, and passing out, even at recommended doses. Tell your doctor if you have any of these problems while taking sodium oxybate.
- Remember that you must not drive a car, operate heavy machinery, or perform any activity that is dangerous or that requires mental alertness or motor coordination for the first 6 hours after taking a dose of sodium oxybate.
- When taking sodium oxybate, do not drink alcohol or take medicines that make you sleepy, including antidepressants, antipsychotics, anti-epileptics, opioids, general anesthetics, muscle relaxants and/or illicit CNS depressants (for example, heroin or GHB).
- These are not all of the side effects that you might experience. Contact your doctor if you are concerned about any possible side effects. Refer to the Medication Guide for additional information on possible side effects.

_____ (Pharmacist Name) ____/____/____ (Date Time)

Step 3: Screening

1. Is the patient taking sedative hypnotics (for example, diazepam, phenobarbital, or zolpidem)?

- Yes Counseled Patient
 No

Please list the drug(s) and dose of each:

2. Is the patient taking sedating antidepressants, antipsychotics, or anti-epileptics such as divalproex sodium (Depakote); general anesthetics; muscle relaxants; opioid analgesics; or illicit CNS depressants (for example, heroin or GHB)?

- Yes Counseled Patient
 No

Please list the drug(s) and dose of each:

3. What other prescription and non-prescription medications is the patient taking?

Please list the drug(s) and dose of each:

4. Does the patient drink alcohol?

- Yes Counseled Patient
 No

5. Has the patient been diagnosed with sleep apnea (short periods of not breathing while asleep)?

- Yes Counseled Patient
 No

6. Does the patient have a diagnosis of or suffer from asthma, COPD, or other conditions affecting his/her breathing (slower breathing, trouble breathing)?
- Yes
 - No
 - Counseled Patient

Please list the drug(s) used to treat and dose of each, if known:

7. Does the patient have any other current medical conditions for which the patient is under a healthcare provider's care?
- Yes
 - No
 - Counseled Patient

Please list the condition(s), if known:

8. Does the patient have any clinical questions about sodium oxybate?
- Yes
 - No
 - Counseled Patient
 - Referred Patient to Prescriber

Please list the question(s):

_____ (Pharmacist Name) ___ / ___ / _____ (Date Time)

Step 4: Concomitant Medication & Comorbidity Summary

Medication Type:

- Sedative hypnotics
- Alcohol
- Other potentially interacting agents:
 - Sedating antidepressants, antipsychotics, or anti-epileptics
 - General anesthetics
 - Muscle relaxants
 - Opioid analgesics
 - Divalproex sodium or other valproate drug (e.g., valproic acid)
 - Illicit CNS depressants (e.g., heroin or GHB)

Medical Conditions:

- Sleep apnea
- Asthma
- COPD
- Other conditions affecting their breathing
- History of depression or suicidality
- History of drug or alcohol abuse
- Seizure disorders
- Hepatic impairment
- High blood pressure, heart problems, kidney problems, or are on a salt-restricted diet

If any of the medication types or medical conditions listed above are checked, or any of the questions in Section 3 were answered yes and there is no confirmation of prior prescriber knowledge, call the prescriber to consult:

Is a prescriber consult required? Yes No

If no, please provide reason: _____

If yes, action(s) taken (check all that apply and document details in Prescriber consult outcome section below):

Called prescriber: ____/____/____

Other: ____/____/____

Name of prescriber consulted: _____

Prescriber NPI or DEA: _____

Prescriber consult outcome: _____

_____(Pharmacist Name) ____/____/____(Date Time)

Step 5: Completion Summary

Checklist Completed: Yes No (Sodium oxybate is not shipped until checklist is completed.)

If yes, date checklist completed: _____/_____/_____ (Date Time)

If no, document the reason for non-completion:

_____ (Pharmacist Name) ____/____/_____ (Date Time)

SODIUM OXYBATE REMS PROGRAM RISK MANAGEMENT REPORT

Sodium oxybate oral solution 500 mg/mL

Sodium Oxybate
REMS Program

Instructions

Risk Management Reports (RMRs) are filled out by pharmacies that are certified in the Sodium Oxybate REMS Program to document and report events that give rise to a reasonable suspicion of abuse, misuse, diversion, or any behavior or information that may indicate the drug is not being used according to the prescriber's instructions. For immediate reporting, RMRs can be completed by the pharmacist online at www.SodiumOxybateREMSProgram.com. Alternatively, a pharmacist can complete a print version and fax to the Sodium Oxybate REMS Program at 800-353-0987.

The RMR history of a patient allows for the review of prior events of suspected abuse, misuse, or diversion and gives the pharmacist a more complete picture of the patient's history. The availability of individual patient RMRs enables the pharmacist to track and monitor for trends suggesting abuse, misuse, or diversion in individual patients. A trend or pattern of behavior in a patient's RMR history can be an indicator of abuse, misuse, or diversion and identifies patients who may require additional scrutiny when another event, such as an early refill request, occurs. In these cases, the RMR history informs actions of the pharmacist.

Examples of events that would require completion of an RMR under the Sodium Oxybate REMS Program include, but are not limited to, the following:

- Patient requests for early refills.
- Patient's loss/misuse of the product.
- Patient claim that he or she did not receive the product but the delivery service shows receipt of delivery, or that the shipment was lost, stolen, or delivered to an incorrect address and was not returned.
- Tampering with or counterfeiting or contamination of the product.
- Inquiries and/or arrests by law and regulatory enforcement agencies associated with the misuse or diversion of the product, or crimes related to the product.
- Prescribers whose DEA and/or state license numbers cannot be validated and the prescriber is submitting a *Sodium Oxybate REMS Program Prescriber Enrollment Form, Prescription Form, or Patient Enrollment Form*.

To complete a RMR:

- Contact the Sodium Oxybate REMS Program to assign a unique Control Number to each report in the Sodium Oxybate REMS Program.
- Complete investigation of the event, which may include contacting the patient, prescriber, law enforcement agency, or other parties.
- Attach any additional documentation required to support the investigation, including but not limited to the following: DEA 106 Form, police or fire report, or report from the shipping service.
- Complete review, follow-up, and sign-off of the RMR.
 - When the event involves suspected abuse, misuse, or diversion, the prescriber will be contacted and an alert may be placed in the prescriber database or patient database of the Sodium Oxybate REMS Program to ensure prescriber and pharmacist awareness.
 - The Sodium Oxybate REMS Program will monitor any associated patient or prescriber activity during the course of the investigation and for a period after the investigation, where appropriate.
 - The certified pharmacies will complete and submit the RMR to the Sodium Oxybate REMS Program. The Sodium Oxybate REMS Program will work with the sponsors to determine the need to notify local, state, or federal authorities.
- Send the RMR to the Sodium Oxybate REMS Program within one business day.
- If the RMR includes a potential adverse event, the potential adverse event is reported to the FDA through the Sodium Oxybate sponsors. If the RMR includes a product complaint, the event is also reported to the FDA through the Sodium Oxybate sponsors.

Sodium Oxybate REMS Program Risk Management Report

Date:	Control Number:	Type of Reporter:	<input type="checkbox"/> Patient	<input type="checkbox"/> Prescribing Physician	<input type="checkbox"/> Pharmacist	<input type="checkbox"/> Other
Name of Reporter (if not a patient):			Name and Address of Pharmacy:			
Nature of Report (e.g., early refill request, lost or stolen bottle, package not received, other):						
Identification Number (patient and/or prescriber ID associated with RMR):					Date Enrolled in Program:	
Have the alerts and RMR history been reviewed with the patient?			<input type="checkbox"/> Yes	<input type="checkbox"/> No	Date(s) of RMR Event (Start/End):	
RMR Event (please provide detail):						
Early Refill Requested? <input type="checkbox"/> Yes <input type="checkbox"/> No						
If yes, reason for early refill request (e.g., dose increase, spilled medication, lost/stolen product):						
Prescriber Contacted?		<input type="checkbox"/> Yes	If yes, what was the outcome of the conversation?			
		<input type="checkbox"/> No	If no, what is the reason?			
Was early refill approved?			<input type="checkbox"/> Yes	<input type="checkbox"/> No	Early refill status reason:	
Potential adverse event associated with report?			<input type="checkbox"/> Yes	<input type="checkbox"/> No	If yes, AE number:	
Summary of investigation: <input type="checkbox"/> Yes <input type="checkbox"/> No						
Attachments (check all that apply): <input type="checkbox"/> DEA 106 Form <input type="checkbox"/> Police/Fire Report <input type="checkbox"/> Shipping Service Report <input type="checkbox"/> Other (specify):						
Should patient be monitored (alert placed)? <input type="checkbox"/> Yes <input type="checkbox"/> No <input type="checkbox"/> N/A						
Are you requesting disenrollment for suspected abuse, misuse, or diversion?		For the patient? <input type="checkbox"/> Yes <input type="checkbox"/> No		For the prescriber? <input type="checkbox"/> Yes <input type="checkbox"/> No		
<input type="checkbox"/> Yes <input type="checkbox"/> No		Patient Name: _____		Prescriber Name: _____		
Pharmacist in Charge Name:		Signature:		Date:		

Sodium Oxybate
REMS Program

Username Password [Sign in](#)

[Forgot Username?](#) [Forgot Password?](#) [Need an Account?](#)

[Safety Information](#) [Prescribing Info](#) [Medication Guide](#)

[Prescriber](#) [Pharmacy](#) [Patient](#) [Resources](#)

Risk Evaluation and Mitigation Strategy (REMS)

The goal of the Sodium Oxybate REMS Program is to mitigate the risk of serious adverse outcomes resulting from inappropriate prescribing, misuse, abuse, and diversion of sodium oxybate by:

1. Informing prescribers, pharmacists, and patients of:
 - i. The risk of significant Central Nervous System (CNS) and respiratory depression associated with sodium oxybate
 - ii. The contraindication of use of sodium oxybate with sedative hypnotics and alcohol
 - iii. The potential for abuse, misuse, and overdose associated with sodium oxybate
 - iv. The safe use, handling, and storage of sodium oxybate
2. Ensuring that pharmacy controls exist prior to filling prescriptions for sodium oxybate that:
 - i. Screen for concomitant use of sedative hypnotics, and other potentially interacting agents
 - ii. Monitor for inappropriate prescribing, misuse, abuse, and diversion of sodium oxybate
 - iii. Notify prescribers when patients are receiving concomitant contraindicated medications or there are signs of potential abuse, misuse, or diversion

Sodium Oxybate REMS Program Overview

- All prescribers and pharmacies must enroll in the Sodium Oxybate REMS Program and comply with requirements for prescribing and dispensing sodium oxybate
- All patients must be enrolled in the Sodium Oxybate REMS Program to receive sodium oxybate
- All patients are required to be counseled on the serious risks and safe use of sodium oxybate
- Sodium oxybate will be dispensed only by pharmacies that are specially certified through the Sodium Oxybate REMS Program

Sodium oxybate is approved for:

- Treatment of cataplexy in patients with narcolepsy
- Treatment of excessive daytime sleepiness (EDS) in patients with narcolepsy

If you require any additional assistance or information, please call the Sodium Oxybate REMS Program at 855-705-2424

[FAQs](#) | [Contact Us](#) | [Privacy Policy](#) | © 2016

Contact Us

Sodium Oxybate REMS Program

Phone

U.S. Phone 855 705 2424

Outside the U.S. Phone +1 855 705 2424

Fax

800 353 0987

Mailing Address

Sodium Oxybate REMS Program

P.O. Box XXXXX

City, ST XXXXX XXXX

If you require any additional assistance or information, please call the Sodium Oxybate REMS Program at 855-705-2424

Create an Account

Pharmacists and prescribers can create a web account in the Sodium Oxybate REMS Program by completing the fields below. The Username you specify must be unique within this website. All fields below are required unless otherwise indicated.

First Name	<input type="text"/>
Last Name	<input type="text"/>
Email Address	<input type="text"/>
Certification ID (opt)	<input type="text"/>
	<i>(If you certified via fax, please enter your Certification ID)</i>
Username	<input type="text"/>
	Suggest Username Check Username Availability
	<input type="checkbox"/> Use Email Address as Username
Password	<input type="password"/>
Confirm Password	<input type="password"/>
	<input type="button" value="Cancel"/> <input type="button" value="Submit"/>

If you require any additional assistance or information, please call the Sodium Oxybate REMS Program at 855-705-2424

Frequently Asked Questions

– Is it still possible to enroll offline?

Yes, you can still enroll offline. Enrollment forms are located under the [Resources](#) tab.

– Do I need to complete the online version of the Prescriber Enrollment Form if I am already enrolled in the Sodium Oxybate REMS Program?

No. If you are already enrolled in the Sodium Oxybate REMS Program or have completed the prescriber enrollment process offline and faxed it, you do not need to complete the online version of the prescriber enrollment form.

You can still use the online enrollment form to enroll your patients.

– How long does it take to process my enrollment so I can enroll patients?

Generally, the Sodium Oxybate REMS Program will process your enrollment within 2 to 3 business days.

– How will I be notified that my patient is enrolled?

Once you complete the patient enrollment form, both you and the patient have signed the form, and the form has been submitted, the Sodium Oxybate REMS Program will process the enrollment form within 2 to 3 business days.

Once the enrollment has been processed you will receive a notification from the Sodium Oxybate REMS Program.

– Will my or my patient's information be shared with any third parties?

Your or your patient's information will only be shared with other sodium oxybate REMS programs, its agents, contractors, and affiliates.

If you require any additional assistance or information, please call the Sodium Oxybate REMS Program at 855-705-2424

Prescriber Roles & Responsibilities

To become certified, each prescriber must complete a one-time enrollment by completing the Sodium Oxybate REMS Program Prescriber Enrollment Form and submitting it to the Sodium Oxybate REMS Program online or offline via fax or mail.

Enroll Now

Prescribers enrolled in the Sodium Oxybate REMS Program agree to:

1. Review the Prescribing Information (PI) and the Sodium Oxybate REMS Program Prescriber Brochure.
2. Screen each patient for:
 - History of alcohol or substance abuse
 - History of sleep-related breathing disorders
 - History of compromised respiratory function
 - Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
 - History of depression or suicidality
3. Counsel each patient prior to initiating therapy with sodium oxybate on the serious risks and safe use and handling of sodium oxybate using the Sodium Oxybate REMS Program Quick Start Guide.
4. Enroll each patient in the Sodium Oxybate REMS Program by completing the Sodium Oxybate REMS Program Patient Enrollment Form and submitting the form to the Sodium Oxybate REMS Program.
5. Evaluate each patient within the first three (3) months of starting sodium oxybate therapy, including an evaluation of the following. It is recommended that patients be re-evaluated every three (3) months thereafter while taking sodium oxybate.
 - a. Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
 - b. Serious adverse events
 - c. Signs of abuse and misuse, including:
 - i. An increase in dose or frequency of dosing
 - ii. Reports of lost, stolen, or spilled medication
 - iii. Drug-seeking behavior
6. Report all potential serious adverse events, including CNS depression, respiratory depression, loss of consciousness, coma, death, and any cases of suspected abuse, misuse, or diversion to the Sodium Oxybate REMS Program.

The prescriber will complete the *Sodium Oxybate REMS Program Prescription Form* for each new prescription and submit the form to one of the certified pharmacies, based on the patient's zip code, as indicated by the online certified pharmacy lookup on www.SodiumOxybateREMSProgram.com. By completing and signing this form, the prescriber acknowledges:

1. Having an understanding of:
 - a. The approved indications for sodium oxybate:
 - i. Treatment of cataplexy in narcolepsy
 - ii. Treatment of excessive daytime sleepiness in narcolepsy
 - b. The serious risks associated with sodium oxybate
 - c. The Prescribing Information and Sodium Oxybate REMS Program Prescriber Brochure
2. Having screened the patient for the following:
 - a. History of alcohol or substance abuse
 - b. History of sleep-related breathing disorders
 - c. History of compromised respiratory function
 - d. Concomitant use of sedative hypnotics, other CNS depressants, or other potentially interacting agents
 - e. History of depression or suicidality
3. Having counseled the patient on:
 - a. The serious risks associated with sodium oxybate
 - b. Contraindications (alcohol and sedative hypnotics) and implications of concomitant use of sodium oxybate with other potentially interacting agents
 - c. Preparation and dosing instructions for sodium oxybate
 - d. Risk of abuse and misuse associated with sodium oxybate
 - e. Risk of operating hazardous machinery including automobiles or airplanes for the first six (6) hours after taking a dose of sodium oxybate
 - f. Safe use, handling, and storage of sodium oxybate
4. That sodium oxybate is medically appropriate for the patient
5. Having listed all known prescription and non-prescription medications and doses on the Sodium Oxybate REMS Program Prescription Form

Materials for Prescriber

- [IMPORTANT UPDATE FOR PHARMACISTS ON Sodium Oxybate \(XOXYBATE\) ONLINE PHARMACY LOOKUP](#)
- [Sodium Oxybate Prescribing Information](#)
- [Sodium Oxybate REMS Program Prescriber Enrollment Form](#)
- [Sodium Oxybate REMS Program Patient Enrollment Form](#)
- [Sodium Oxybate REMS Program Prescriber Brochure](#)
- [Sodium Oxybate REMS Program Patient Enrollment Form](#)
- [Sodium Oxybate REMS Program Prescriber Brochure](#)

Pharmacy

Certification in the Sodium Oxybate REMS Program requires pharmacies to agree to:

1. Designate an authorized representative to complete and submit the Sodium Oxybate REMS Pharmacy Enrollment Form on behalf of the pharmacy
2. Ensure that the authorized representative oversees implementation and compliance with the Sodium Oxybate REMS Program by the following:
 - i. Ensure that all pharmacy staff involved in the Sodium Oxybate REMS Program complete the *Sodium Oxybate REMS Program Certified Pharmacy Training Program* Module A
 - ii. Ensure that all pharmacists who dispense sodium oxybate complete the *Sodium Oxybate REMS Program Certified Pharmacy Training Program* Modules A and B
3. Dispense sodium oxybate only to patients enrolled in the Sodium Oxybate REMS Program pursuant to a valid prescription written by a prescriber specially certified in the Sodium Oxybate REMS Program
4. Dispense only after obtaining a Pre Dispense Authorization (PDA) for each sodium oxybate prescription
5. Recertify in the Sodium Oxybate REMS Program if the pharmacy designates a new authorized representative
6. Provide 24/7 toll free access to a pharmacist at a Sodium Oxybate REMS Program specially certified pharmacy
7. Ship sodium oxybate directly to each patient or a patient authorized adult designee, and track and verify receipt of each shipment of sodium oxybate
8. Limit the first shipment for each patient to a one month supply of sodium oxybate, and subsequent shipments to no more than a three month supply of sodium oxybate
9. Report all potential adverse events reported by all sources, including any CNS depression, respiratory depression, loss of consciousness, coma, and death, and any cases of suspected abuse, misuse, or diversion to the Sodium Oxybate REMS Program for documentation

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If you require any additional assistance or information, please call the Sodium Oxybate REMS Program at 855-705-2424

Patient

Patients prescribed sodium oxybate must enroll in the Sodium Oxybate REMS Program. If you were previously registered in the XYREM REMS, your certified prescriber will need to enroll you in the Sodium Oxybate REMS Program.

Materials for Patients

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If you require any additional assistance or information, please call the Sodium Oxybate REMS Program at 855-705-2424

Medication Guide - Sodium Oxybate (soe' dee um ox' i bate) Oral Solution, CIII

Read this Medication Guide carefully before you start taking sodium oxybate and each time you get a refill. There may be new information. This information does not take the place of talking to your doctor about your medical condition or your treatment.

What is the most important information I should know about sodium oxybate?

Sodium oxybate can cause serious side effects including slow breathing or changes in your alertness. Do not drink alcohol or take medicines intended to make you fall asleep while you are taking sodium oxybate because they can make these side effects worse. Call your doctor right away if you have any of these serious side effects.

- The active ingredient of Sodium Oxybate Oral Solution is a form of gamma-hydroxybutyrate (GHB). GHB is a chemical that has been abused and misused. Abuse and misuse of sodium oxybate can cause serious medical problems, including:
 - seizures
 - trouble breathing
 - changes in alertness
 - coma
 - death
- Do not drive a car, use heavy machinery, fly an airplane, or do anything that is dangerous or that requires you to be fully awake for at least 6 hours after you take sodium oxybate. You should not do those activities until you know how sodium oxybate affects you.
- Sodium oxybate is available only by prescription and filled through the Sodium Oxybate REMS Program. Before you receive sodium oxybate, your doctor or pharmacist will make sure that you understand how to use sodium oxybate safely and effectively. If you have any questions about sodium oxybate, ask your doctor or call the Sodium Oxybate REMS Program at 1-800-XXX-XXXX.

What is sodium oxybate?

Sodium oxybate is a prescription medicine used to treat the following symptoms in people who fall asleep frequently during the day, often at unexpected times (narcolepsy):

- suddenly weak or paralyzed muscles when they feel strong emotions (cataplexy)
- excessive daytime sleepiness (EDS) in people who have narcolepsy

It is not known if sodium oxybate is safe and effective in children.

Sodium Oxybate Oral Solution is a controlled substance (CIII) because it contains sodium oxybate that can be a target for people who abuse prescription medicines or street drugs. Keep your sodium oxybate in a safe place to protect it from theft. Never give your sodium oxybate to anyone else because it may cause death or harm them. Selling or giving away this medicine is against the law.

Who should not take sodium oxybate?

Do not take sodium oxybate if you:

- take other sleep medicines or sedatives (medicines that cause sleepiness)
- drink alcohol
- have a rare problem called succinic semialdehyde dehydrogenase deficiency

Before you take sodium oxybate, tell your doctor if you:

- have short periods of not breathing while you sleep (sleep apnea).
- snore, have trouble breathing, or have lung problems. You may have a higher chance of having serious breathing problems when you take sodium oxybate.
- have or had depression or have tried to harm yourself. You should be watched carefully for new symptoms of depression.
- have liver problems.
- are on a salt-restricted diet. Sodium oxybate contains a lot of sodium (salt) and may not be right for you.
- have high blood pressure.
- have heart failure.
- have kidney problems.
- are pregnant or plan to become pregnant. It is not known if sodium oxybate can harm your unborn baby.
- are breastfeeding or plan to breastfeed. It is not known if sodium oxybate passes into your breast milk. You and your doctor should decide if you will take sodium oxybate or breastfeed.

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

Especially, tell your doctor if you take other medicines to help you sleep (sedatives). Do not take medicines that make you sleepy with sodium oxybate.

Know the medicines you take. Keep a list of them to show your doctor and pharmacist when you get a new medicine.

How should I take sodium oxybate?

- Read the **Instructions for Use** at the end of this Medication Guide for detailed instructions on how to take sodium oxybate.
- Take sodium oxybate exactly as your doctor tells you to take it.
- Never change your sodium oxybate dose without talking to your doctor.
- Sodium oxybate can cause sleep very quickly. You should fall asleep soon. Some patients fall asleep within 5 minutes and most fall asleep within 15 minutes. Some patients take less time to fall asleep and some take more time. The time it takes you to fall asleep might be different from night to night.
- Take your first sodium oxybate dose at bedtime while you are in bed. Take your second sodium oxybate dose 2.5 to 4 hours after you take your first sodium oxybate dose. You may want to set an alarm clock to make sure you wake up to take your second sodium oxybate dose. You should remain in bed after taking the first and second doses of sodium oxybate.
- If you miss your second sodium oxybate dose, skip that dose and do not take sodium oxybate again until the next night. Never take 2 sodium oxybate doses at 1 time.
- Wait at least 2 hours after eating before you take sodium oxybate.
- You should see your doctor every 3 months for a check-up while taking sodium oxybate. Your doctor should check to see if sodium oxybate is helping to lessen your symptoms and if you feel any side effects while you take sodium oxybate.
- If you take too much sodium oxybate, call your doctor or go to the nearest hospital emergency room right away.

What are the possible side effects of sodium oxybate?

Sodium oxybate can cause serious side effects, including:

- See **“What is the most important information I should know about sodium oxybate?”**
- **Breathing problems, including:**
 - slower breathing

- trouble breathing
- short periods of not breathing while sleeping (sleep apnea). People who already have breathing or lung problems have a higher chance of having breathing problems when they use sodium oxybate.
- **Mental health problems, including:**
 - confusion
 - seeing or hearing things that are not real (hallucinations)
 - unusual or disturbing thoughts (abnormal thinking)
 - feeling anxious or upset
 - depression
 - thoughts of killing yourself or trying to kill yourself

Call your doctor right away if you have symptoms of mental health problems.

- **Sleepwalking.** Sleepwalking can cause injuries. Call your doctor if you start sleepwalking. Your doctor should check you.

The most common side effects of sodium oxybate include:

- nausea
- dizziness
- vomiting
- bedwetting
- diarrhea

Your side effects may increase when you take higher doses of sodium oxybate. Sodium oxybate can cause physical dependence and craving for the medicine when it is not taken as directed.

These are not all the possible side effects of sodium oxybate. For more information, ask your doctor or pharmacist.

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 sodium oxybate?

- **Always store sodium oxybate in the original bottle or in the dosing cups with child-resistant caps provided with your prescription.**
- **Keep sodium oxybate in a safe place out of the reach of children and pets.**
- **Get emergency medical help right away if a child drinks your sodium oxybate.**
- Store sodium oxybate between 68°F to 77°F (20°C to 24°C). When you have finished using a sodium oxybate bottle:
 - empty any unused sodium oxybate down the sink drain
 - cross out the label on the sodium oxybate bottle with a marker
 - place the empty sodium oxybate bottle in the trash

General information about the safe and effective use of sodium oxybate

Medicines are sometimes prescribed for purposes other than those listed in a Medication Guide. Do not use sodium oxybate for a condition for which it was not prescribed. Do not give sodium oxybate to other people, even if they have the same symptoms that you have. It may harm them.

This Medication Guide summarizes the most important information about sodium oxybate. If you would like more information, talk with your doctor. You can ask your pharmacist or doctor for information about sodium oxybate that is written for health professionals.

For more information, call the Sodium Oxybate REMS Program at 1-800-XXX-XXXX.

What are the ingredients in Sodium Oxybate Oral Solution?

Active Ingredients: sodium oxybate

Inactive Ingredients: purified water, USP

This Medication Guide has been approved by the U.S. Food and Drug Administration.

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