

**Y**ou are about to begin your treatment with ELOXATIN®, a chemotherapy drug. ELOXATIN® is given together with 2 other drugs: 5-fluorouracil (5-FU) and leucovorin. This combination is given to treat adults with stage III colon cancer after surgery to remove the tumor, or to treat adults with advanced colorectal cancer.

- Your treatment will be given through one of your veins. This is called an intravenous, or **IV**, infusion
- Before your first treatment, your doctor may insert a soft, flexible tube into a large vein and attach a **port**
  - A port is a tiny disk with an opening, completely hidden under your skin, that will remain in place during the course of your treatment
- Every time you receive a treatment, the IV will be inserted into that port
  - This means fewer needle sticks
  - It also helps protect your skin from direct exposure to the chemotherapy



In most cases, chemotherapy with ELOXATIN® is given every 2 weeks. This is called a **cycle** of treatment. Your doctor will determine the best approach to your treatment.

## What happens during your 14-day treatment cycle?

### Day 1

You will be given ELOXATIN® and leucovorin by IV infusion over a 2-hour period. After that, you will receive 2 doses of 5-FU. The first dose is given by IV infusion as soon as the leucovorin infusion is over. The second dose is given as a steady, slow drip over the next 22 hours, using a pump device.

### Day 2

You will not be given ELOXATIN® on Day 2. That day you will receive an IV infusion of leucovorin for 2 hours. After that, you will receive 2 doses of 5-FU. The first dose is given by IV infusion as soon as the leucovorin infusion is over. The second dose is given as a steady, slow drip over the next 22 hours, using a pump device.

### Day 3

Your pump device will be disconnected.

### Days 4 to 14

No treatments or procedures are scheduled.

***If you have questions about how your treatment is given, speak to your doctor or nurse.***

Please see full prescribing information, including Boxed WARNING, enclosed in kit.

**Eloxatin®**  
(OXALIPLATIN injection)

## Indications and Usage

Eloxatin® (oxaliplatin injection), used in combination with infusional 5-FU/LV, is indicated for

- Adjuvant treatment of stage III colon cancer patients who have undergone complete resection of the primary tumor. The indication is based on an improvement in disease-free survival, with no demonstrated benefit in overall survival after a median follow-up of 4 years
- Treatment of advanced carcinoma of the colon or rectum

## Clinical Safety Considerations

**ELOXATIN should be administered under the supervision of a physician experienced in the use of cancer chemotherapeutic agents.**

**Appropriate management of therapy and complications is possible only when adequate diagnostic and treatment facilities are readily available.**

**Anaphylactic-like reactions to ELOXATIN have been reported and may occur within minutes of ELOXATIN administration. Epinephrine, corticosteroids, and antihistamines have been employed to alleviate symptoms, and discontinuation of ELOXATIN therapy may be required.**

- ELOXATIN should not be administered to patients with a history of known allergy to ELOXATIN or other platinum compounds. Hypersensitivity and anaphylactic/anaphylactoid reactions to ELOXATIN have been reported and were similar in nature and severity to those reported with other platinum compounds (ie, rash, urticaria, erythema, pruritus, and, rarely, bronchospasm and hypotension). These reactions occur within minutes of administration and should be managed with appropriate supportive therapy. Drug-related deaths from this reaction have been reported
- ELOXATIN may cause fetal harm when administered to a pregnant woman. Women of childbearing potential should be advised not to become pregnant while receiving ELOXATIN. It is not known whether ELOXATIN or its derivatives are excreted in human milk
- ELOXATIN has been associated with pulmonary fibrosis (<1% of study patients), which may be fatal. The combined incidence of cough and dyspnea was 7.4% (<1% grade 3, no grade 4) in the ELOXATIN plus 5-FU/LV arm compared to 4.5% (no grade 3, 0.1% grade 4) in the 5-FU/LV alone arm in the adjuvant colon cancer study. In this study, one patient died from eosinophilic pneumonia in the ELOXATIN combination arm. The combined incidence of cough, dyspnea, and hypoxia was 43% (7% grade 3 and 4) in the ELOXATIN plus 5-FU/LV arm compared to 32% (5% grade 3 and 4) in the irinotecan plus 5-FU/LV arm in patients with previously untreated colorectal cancer. In case of unexplained respiratory symptoms, ELOXATIN should be discontinued until pulmonary investigation excludes interstitial lung disease or pulmonary fibrosis
- ELOXATIN is associated with two types of primarily peripheral sensory neuropathy: an

acute, reversible type of early onset and a persistent type (>14 days). In patients with advanced colorectal cancer, paresthesias occurred in 77% (all grades) and 18% (grade 3/4) of previously untreated patients. In previously treated patients, acute neuropathy occurred in 56% (all grades) and 2% (grade 3/4) of patients; persistent neuropathy occurred in 48% (all grades) and 6% (grade 3/4) of patients. In patients with stage II and III colon cancer, paresthesia was seen in 92% (all grades) and 13% (grade 3/4) of patients; 21% (all grades) and 0.5% (grade 3/4) of patients had residual paresthesia at 18-month follow-up

- Hepatotoxicity, as evidenced in the adjuvant study by increase in transaminases and alkaline phosphatase, was observed more commonly in the ELOXATIN combination arm. The incidence of increased bilirubin was similar on both arms. Changes noted on liver biopsies include: peliosis, nodular regenerative hyperplasia or sinusoidal alterations, perisinusoidal fibrosis, and veno-occlusive lesions. Hepatic vascular disorders should be considered and, if appropriate, investigated in case of abnormal liver function test results or portal hypertension not explained by liver metastases
- Monitoring of white blood cell count with differential, hemoglobin, platelet count, and blood chemistries (including ALT, AST, bilirubin, and creatinine) is recommended before each ELOXATIN cycle
- The safety and effectiveness of ELOXATIN plus 5-FU/LV in patients with renal impairment have not been evaluated. Since the primary route of platinum elimination is renal, this combination should be used with caution in patients with preexisting renal impairment. Clearance of these products may be decreased by coadministration of potentially nephrotoxic compounds, although this has not been specifically studied
- The incidence of diarrhea, dehydration, hypokalemia, leukopenia, fatigue, and syncope was higher in patients  $\geq 65$  years old
- Extravasation may result in local pain and inflammation that may be severe and lead to complications, including necrosis. Injection site reaction, including redness, swelling, and pain, has been reported
- There have been reports of prolonged prothrombin time and INR occasionally associated with hemorrhage in patients receiving ELOXATIN plus 5-FU/LV while on anticoagulants. Patients receiving ELOXATIN plus 5-FU/LV and requiring oral anticoagulants may require closer monitoring
- The most common adverse reactions in patients with stage II or III colon cancer receiving adjuvant therapy were peripheral sensory neuropathy, neutropenia, thrombocytopenia, anemia, nausea, increase in transaminases and alkaline phosphatase, diarrhea, emesis, fatigue, and stomatitis. The most common adverse reactions in patients with advanced colorectal cancer were peripheral sensory neuropathy, fatigue, neutropenia, nausea, emesis, and diarrhea



sanofi aventis

© 2006 sanofi-aventis U.S. LLC  
US.OXA.06.04.071 June 2006 Printed in USA

www.ELOXATIN.com

**Eloxatin**<sup>®</sup>  
(OXALIPLATIN injection)