CYRAMZA® (ramucirumab) is used with a chemotherapy combination called FOLFIRI (irinotecan, folinic acid, and 5-fluorouracil) to treat metastatic colorectal cancer (CRC) in patients whose cancer has progressed after being treated with other certain types of chemotherapy.

**WARNING: SEVERE BLEEDING, TEARS IN THE STOMACH OR BOWEL WALL, AND DIFFICULTY IN WOUND HEALING**

Severe bleeding has occurred with CYRAMZA. These events are sometimes fatal. Tell your doctor right away if you have bleeding or symptoms of bleeding, including lightheadedness. Your treatment will have to be permanently stopped if severe bleeding occurs.

Tears in the stomach or bowel wall may occur with CYRAMZA. These events are sometimes fatal. Tell your doctor if you have severe diarrhea, vomiting, or severe abdominal pain. Your treatment will have to be permanently stopped if this occurs.

Wounds may have difficulty healing with drugs like CYRAMZA. Tell your doctor if you have a wound that doesn’t heal properly. If you develop a wound that won’t heal during treatment, your doctor will stop CYRAMZA until the wound is fully healed. Also tell your doctor if you have planned surgery, as CYRAMZA treatment should be interrupted prior to surgery. Your doctor may restart CYRAMZA after your surgical wound has healed.

Please see Important Safety Information, including Boxed Warning for severe bleeding, tears in the stomach or bowel wall, and difficulty in wound healing, on pages 5–6. See full Prescribing Information for additional information about CYRAMZA.
QUESTIONS TO ASK MY DOCTOR

It is important that you ask your treatment team every question that comes to mind. No matter how minor they may seem to you, they want to answer all of your questions. Below is a list of questions to consider asking your treatment team, followed by an area where you can write in other questions you want to ask.

About CYRAMZA® (ramucirumab) combination therapy

• How is CYRAMZA in combination with FOLFIRI different from my past treatments? ____________________________________________
  _______________________________________________________________
  _______________________________________________________________

• What should I do to prepare for treatment? _____________________________________________________________
  _______________________________________________________________
  _______________________________________________________________
  _______________________________________________________________

• What side effects might occur with treatment? _________________________________________________________
  _______________________________________________________________
  _______________________________________________________________
  _______________________________________________________________

• How can I tell the difference between side effects of the treatment and symptoms of the cancer? _____________
  _______________________________________________________________
  _______________________________________________________________
  _______________________________________________________________

• Are there ways to help manage side effects? __________________________________________________________
  _______________________________________________________________
  _______________________________________________________________
  _______________________________________________________________

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QUESTIONS TO ASK MY DOCTOR  (continued)

• What can CYRAMZA® (ramucirumab) in combination with FOLFIRI offer as a treatment? _______________________

___________________________________________________________________________________________________

___________________________________________________________________________________________________

• What can you tell me about how CYRAMZA and FOLFIRI are given? ________________________________

___________________________________________________________________________________________________

___________________________________________________________________________________________________

• How long should I plan for my treatment visits to take? ________________________________

___________________________________________________________________________________________________

___________________________________________________________________________________________________

About eating right and keeping up your strength

• How can I keep myself as healthy as possible during the treatment? ________________________________

___________________________________________________________________________________________________

___________________________________________________________________________________________________

• What kind of foods should I eat and what should I avoid? ________________________________

___________________________________________________________________________________________________

___________________________________________________________________________________________________

• Will I need any extra food supplements? ________________________________

___________________________________________________________________________________________________

___________________________________________________________________________________________________

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QUESTIONS TO ASK MY DOCTOR (continued)

• Are there activities I can do to keep up my strength?

___________________________________________________________________________________________________
___________________________________________________________________________________________________
___________________________________________________________________________________________________

About support services

• What support services are available to me and my family and friends?

___________________________________________________________________________________________________
___________________________________________________________________________________________________
___________________________________________________________________________________________________

• Are there financial resources that may help with the cost of treatment?

___________________________________________________________________________________________________
___________________________________________________________________________________________________
___________________________________________________________________________________________________

• What if I can’t find a ride to an appointment?

___________________________________________________________________________________________________
___________________________________________________________________________________________________
___________________________________________________________________________________________________

• Are there other people affected by cancer I can talk to?

___________________________________________________________________________________________________
___________________________________________________________________________________________________
___________________________________________________________________________________________________

My other questions to ask

___________________________________________________________________________________________________
___________________________________________________________________________________________________

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What is the most important information I should know about CYRAMZA® (ramucirumab)?

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- Infusion reactions related to injecting CYRAMZA have occurred. Most of these reactions happened during or after the first or second CYRAMZA infusion. Signs and symptoms of infusion reactions include shaking or stiffness of the body, back pain or spasms, chest pain or tightness, chills, flushing, difficulty breathing, wheezing, becoming blue due to lack of oxygen, and tingling or numbness of the skin. In severe cases, rapid heartbeat, low blood pressure, and severe trouble breathing may occur. Your healthcare team will monitor you for these side effects. In the case of a severe infusion reaction, your CYRAMZA treatment will have to be immediately and permanently stopped.

- CYRAMZA may worsen certain types of liver disease. Tell your doctor if you have or have had liver disease or other liver problems.

- RPLS (reversible posterior leukoencephalopathy syndrome), a very rare but serious brain disorder, has been reported in clinical trials with CYRAMZA. Signs and symptoms of RPLS may include headache, seizures, visual changes, and changes in mental function. These symptoms usually stop or improve within days, but some patients can experience continuing changes in mental function or death.

- Too much protein in the urine (proteinuria) has been reported in clinical trials with CYRAMZA. This may be a sign of kidney damage. Your doctor will monitor your urine protein levels during treatment. If you develop protein in your urine, your doctor may interrupt your treatment and decrease your dose of CYRAMZA. In severe cases your treatment will have to be permanently stopped.

- Thyroid gland problems have been reported in clinical trials with CYRAMZA. Your doctor will do blood tests to monitor your thyroid gland function during treatment.

- CYRAMZA can harm your unborn baby. You should avoid getting pregnant, and use adequate contraception while receiving CYRAMZA and for at least 3 months after stopping CYRAMZA.

Please see full Prescribing Information for additional information about CYRAMZA, including Boxed Warning for severe bleeding, tears in the stomach or bowel wall, and difficulty in wound healing.
What are the most common side effects of CYRAMZA® (ramucirumab)?

- The most common side effects reported in patients treated with CYRAMZA when given in combination with FOLFIRI (irinotecan, folinic acid, and 5-fluorouracil) were diarrhea; low white blood cell count; decreased appetite; nosebleeds; mouth sores; low platelet count; high blood pressure; swelling in the arms, legs, hands, or feet; too much protein in the urine; hand-foot syndrome; bleeding in the gastrointestinal tract; and low albumin (a protein in the blood). The most common serious adverse events with CYRAMZA plus FOLFIRI were diarrhea, blockage of the intestine, and low white blood cell count with fever. Treatment to increase white blood cell count called granulocyte colony-stimulating factors was given to 20% of patients treated with CYRAMZA plus FOLFIRI.

You are encouraged to report negative side effects of prescription drugs to the FDA. Visit www.fda.gov/safety/medwatch or call 1-800-FDA-1088.

What should I tell my doctor before receiving treatment with CYRAMZA?

Before you receive CYRAMZA, tell your doctor if you:

- Have had or are at high risk for strokes or heart attack.
- Have high blood pressure or have blood pressure problems.
- Are planning to have surgery of any kind.
- Have or have had liver problems, including liver cirrhosis or other diseases of the liver.
- Are pregnant or may be pregnant: CYRAMZA can harm your unborn baby. You should avoid getting pregnant, and use adequate contraception while receiving CYRAMZA and for at least 3 months after stopping CYRAMZA.

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