Zydelig®
(idelalisib) 150 mg 100 mg tablets

Zydelig REMS

FDA REQUIRED UPDATED REMS SAFETY INFORMATION

Boxed Warning Risk of:

- **Fatal and/or serious hepatotoxicity – *updated***
- **Fatal and/or serious and severe diarrhea or colitis – *updated***
- **Fatal and/or serious pneumonitis**
- **Fatal and/or serious infections – *updated***
- **Fatal and serious intestinal perforation**

January 2018

Dear Healthcare Provider:

The FDA has required this safety notice as part of the Zydelig REMS (**R**isk **E**valuation and **M**itigation **S**trategy) to inform you about the recent **update to the incidence rates of fatal and serious toxicities in the Zydelig Boxed Warning** as follows:

WARNING: FATAL AND SERIOUS TOXICITIES: HEPATIC, SEVERE DIARRHEA, COLITIS, PNEUMONITIS, INFECTIONS and INTESTINAL PERFORATION

- Fatal and/or serious hepatotoxicity occurred in 16% to 18% of ZYDELIG-treated patients. Monitor hepatic function prior to and during treatment. Interrupt and then reduce or discontinue ZYDELIG.
- Fatal and/or serious and severe diarrhea or colitis occurred in 14% to 20% of ZYDELIG-treated patients. Monitor for the development of severe diarrhea or colitis. Interrupt and then reduce or discontinue ZYDELIG.
- Fatal and/or serious pneumonitis occurred in 4% of ZYDELIG-treated patients. Monitor for pulmonary symptoms and interstitial infiltrates. Interrupt or discontinue ZYDELIG.
- Fatal and/or serious infections occurred in 21% to 48% of ZYDELIG-treated patients. Monitor for signs and symptoms of infection. Interrupt ZYDELIG if infection is suspected.
- Fatal and serious intestinal perforation can occur in ZYDELIG-treated patients across clinical trials. Discontinue ZYDELIG if intestinal perforation is suspected.

The updated incidence rates of fatal and serious toxicities in the Zydelig Boxed Warning reflect data from patients treated with Zydelig in combination with rituximab or with unapproved combination therapies.

Please see the enclosed **Zydelig REMS Fact Sheet**, a non-promotional fact sheet reviewed by the FDA, for more detailed safety information. Be sure to give the **Zydelig Patient Safety Information Card** to all patients. This card, additional copies of the fact sheet, and other important information are available at: www.ZydeligREMS.com.

Zydelig is a kinase inhibitor indicated for the treatment of patients with:

- Relapsed chronic lymphocytic leukemia (CLL) in combination with rituximab, in patients for whom rituximab alone would be considered appropriate therapy due to other co-morbidities.
- Relapsed follicular B-cell non-Hodgkin lymphoma (FL) in patients who have received at least two prior systemic therapies
- Relapsed small lymphocytic lymphoma (SLL) in patients who have received at least two prior systemic therapies

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Limitation of use:

Zydelig is not indicated and is not recommended for first-line treatment of any patient. Zydelig is not indicated and is not recommended in combination with bendamustine and/or rituximab for the treatment of FL.

This letter does not contain the complete safety profile for Zydelig. Please review the enclosed Prescribing Information.

Reporting Adverse Events

You are encouraged to report negative side effects of prescription drugs to the FDA. Visit www.fda.gov/medwatch or call 1-800-FDA-1088. Healthcare Providers should report all suspected adverse events associated with Zydelig to the FDA or to Gilead at 1-800-445-3235.

Sincerely,

William Guyer, Pharm.D.
Senior Vice President, Medical Affairs