## Highlights of This Issue 2017

### SPECIAL FEATURES

#### CCR Translations

- **2019** New Approaches but the Same Flaws in the Search for Prognostic Signatures  
  Ramon Salazar and Josep Tabernero  
  See related article, p. 2159

- **2023** Emerging Immunologic Biomarkers: Setting the (TNM-Immune) Stage  
  Janis M. Taube  
  See related article, p. 2147

- **2026** SETting OP449 into the PP2A-Activating Drug Family  
  Paolo Neviani and Danilo Perrotti  
  See related article, p. 2092

#### CCR Perspectives in Drug Approval

- **2029** U.S. Food and Drug Administration Approval: Crizotinib for Treatment of Advanced or Metastatic Non–Small Cell Lung Cancer That Is Anaplastic Lymphoma Kinase Positive  

#### CCR Drug Updates

- **2035** Dabrafenib and Trametinib, Alone and in Combination for BRAF-Mutant Metastatic Melanoma  
  Alexander M. Menzies and Georgina V. Long

#### Molecular Pathways

- **2044** Molecular Pathways: Interleukin-15 Signaling in Health and in Cancer  
  Anjali Mishra, Laura Sullivan, and Michael A. Caligiuri

- **2051** Molecular Pathways: Molecular Basis for Sensitivity and Resistance to JAK Kinase Inhibitors  
  Sara C. Meyer and Ross L. Levine

### Reviews

- **2060** Molecular Biomarkers in Advanced Renal Cell Carcinoma  
  Pablo Maroto and Brian Rini

- **2072** Hepatocellular Carcinoma: Reasons for Phase III Failure and Novel Perspectives on Trial Design  
  Josep M. Llovet and Virginia Hernandez-Gea

### HUMAN CANCER BIOLOGY

- **2080** High-Throughput Detection of Clinically Relevant Mutations in Archived Tumor Samples by Multiplexed PCR and Next-Generation Sequencing  
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### CANCER THERAPY: PRECLINICAL

- **2092** Antagonism of SET Using OP449 Enhances the Efficacy of Tyrosine Kinase Inhibitors and Overcomes Drug Resistance in Myeloid Leukemia  
  Anupriya Agarwal, Ryan J. MacKenzie, Raffaella Fippa, Christopher A. Eide, Jessica Oddo, Jeffrey W. Tyner, Rosalie Sears, Michael P. Vitk, María D. Odero, Dale J. Christensen, and Brian J. Druker  
  See related article, p. 2026

- **2104** Rational Combination Therapy of Vintafolide (EC145) with Commonly Used Chemotherapeutic Drugs  
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- **2115** Inhibition of RET Increases the Efficacy of Antiestrogen and Is a Novel Treatment Strategy for Luminal Breast Cancer  
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2237 Proteomic Markers of DNA Repair and PI3K Pathway Activation Predict Response to the PARP Inhibitor BMN 673 in Small Cell Lung Cancer—Response
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ABOUT THE COVER

The cover shows a class of recurrent hotspot mutations in PIK3R1 and PIK3CA from endometrial cancer patients that are clustered at the interface between the ISH2 domain of PIK3R1 and the C2 domain of PIK3CA. Alteration of some of these crucial amino acids has been shown to be sufficient to disrupt the inhibitory contact by PIK3R1 and may represent a novel mechanism of oncogenic activation of PIK3CA. For details, see the article by Bourgon and colleagues on page 2080 of this issue.