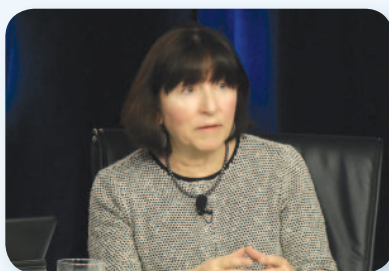


# New Biologic Insights and Recent Therapeutic Advances in the Management of Acute and Chronic Leukemias and Myelodysplastic Syndromes

## *Proceedings from a Clinical Investigator Think Tank*



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# *New Biologic Insights and Recent Therapeutic Advances in the Management of Acute and Chronic Leukemias and Myelodysplastic Syndromes*

## A Continuing Medical Education Audio Program

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### OVERVIEW OF ACTIVITY

Hematologic cancers include the lymphomas, the leukemias, multiple myeloma and other related disorders (eg, myelodysplastic syndromes [MDS] and myeloproliferative diseases) stemming from lymphoid and myeloid progenitor cell lines. Taken together, it is estimated that approximately 162,020 new lymphoid, myeloid and leukemic cancer cases will be identified in the United States in the year 2015 and 56,630 individuals will die from these diseases. Although an extensive list of treatment options is available for these patients, this poses a challenge to the practicing clinician who must maintain up-to-date knowledge of appropriate clinical management strategies across a vast spectrum of diseases. To address this issue, this CME program brings together leading clinical investigators to provide biologic insights into the recent therapeutic advances in the management of acute and chronic leukemias and MDS. By reviewing the available clinical trial data and relevant case scenarios, this initiative will provide perspectives on gaps in medical knowledge and illuminate treatment ambiguities pertinent to the treatment of these diseases.

### LEARNING OBJECTIVES

- Appraise recent data on therapeutic advances and changing practice standards in the management of select acute and chronic leukemias and MDS, and refine or validate existing treatment algorithms based on discussion of this information.
- Recognize evidence-based therapeutic options for patients with progressive chronic myeloid leukemia.
- Appreciate the FDA approvals of novel targeted agents indicated for the treatment of newly diagnosed and relapsed or refractory chronic lymphocytic leukemia, and discern how these treatments can be appropriately integrated into clinical practice.
- Review existing and evolving clinical trial data to recommend safe therapeutic alternatives for patients with acute myeloid leukemia, including acute promyelocytic leukemia, and increase knowledge regarding investigational options designed for patients who are not candidates for intensive therapy.
- Apply the results of emerging clinical research to optimize treatment for young adult and adult patients with newly diagnosed and recurrent acute lymphoblastic leukemia.
- Counsel patients with MDS about supportive and systemic treatment options to manage disease-related cytopenias and minimize leukemic progression.

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*New Biologic Insights and Recent Therapeutic Advances in the Management of Acute and Chronic Leukemias and Myelodysplastic Syndromes*

**QUESTIONS (PLEASE CIRCLE ANSWER):**

1. Which of the following agents approved for the treatment of CML is classified as a protein synthesis inhibitor?
  - a. Bosutinib
  - b. Dasatinib
  - c. Omacetaxine mepesuccinate
  - d. Ponatinib
  - e. All of the above
2. Which of the following is an adverse event associated with idelalisib treatment in patients with CLL?
  - a. Transaminitis
  - b. Pneumonitis
  - c. Colitis
  - d. Both a and b
  - e. All of the above
3. The achievement of partial responses with lymphocytosis in patients with CLL may be observed as part of the clinical pattern of response associated with \_\_\_\_\_ treatment.
  - a. Single-agent idelalisib
  - b. Single-agent ibrutinib
  - c. Both a and b
4. Lenalidomide therapy did not result in improved rates of transfusion independence when compared to placebo in patients with low/intermediate-risk MDS without del(5q) who were unresponsive or refractory to erythropoiesis-stimulating agents.
  - a. True
  - b. False
5. Initial management of high-risk APL (high white blood cell count, bleeding complications) should include \_\_\_\_\_.
  - a. All-trans retinoic acid
  - b. Steroids
  - c. Anthracycline
  - d. All of the above
6. Data from the Phase II SORAML study indicate that clinical benefit with the addition of sorafenib to standard therapy in younger patients with newly diagnosed AML was restricted to only those with FLT3-ITD mutation-positive disease.
  - a. True
  - b. False
7. In elderly patients with AML who were unfit for intensive induction therapy, the addition of volasertib, a polo-like kinase inhibitor, to low-dose cytarabine improved efficacy in terms of \_\_\_\_\_ when compared to low-dose cytarabine alone.
  - a. Response rate
  - b. Median event-free survival
  - c. Median overall survival
  - d. Both b and c
  - e. All of the above
8. Which of the following is the mechanism of action of venetoclax (ABT-199)?
  - a. Bcl-2 inhibitor
  - b. FLT3 inhibitor
  - c. Protein synthesis inhibitor
9. Intensive treatment regimens developed for pediatric patients with ALL have also been shown to be efficacious for the treatment of older adolescents and young adults with this disease.
  - a. True
  - b. False
10. In adults with relapsed/refractory ALL, the bispecific T-cell engaging monoclonal antibody \_\_\_\_\_ resulted in high complete remission rates.
  - a. Obinutuzumab
  - b. Blinatumomab
  - c. Ofatumumab



**EDUCATIONAL ASSESSMENT AND CREDIT FORM**

*New Biologic Insights and Recent Therapeutic Advances in the Management of Acute and Chronic Leukemias and Myelodysplastic Syndromes*

Research To Practice is committed to providing valuable continuing education for oncology clinicians, and your input is critical to helping us achieve this important goal. Please take the time to assess the activity you just completed, with the assurance that your answers and suggestions are strictly confidential.

**PART 1 — Please tell us about your experience with this educational activity**

**How would you characterize your level of knowledge on the following topics?**

4 = Excellent    3 = Good    2 = Adequate    1 = Suboptimal

	BEFORE	AFTER
Efficacy of sorafenib in patients with AML based on FLT3 mutation status	4 3 2 1	4 3 2 1
Selection of patients with CML for whom discontinuation of TKI therapy is appropriate	4 3 2 1	4 3 2 1
Response rates in a Phase II trial of volasertib combined with low-dose cytarabine in elderly patients with AML	4 3 2 1	4 3 2 1
Response and survival outcomes for patients with untreated CLL and comorbidities on the Phase III CLL11 trial evaluating obinutuzumab/chlorambucil or rituximab/chlorambucil versus chlorambucil alone	4 3 2 1	4 3 2 1
Incidence and management of pneumonitis, colitis, transaminitis and other toxicities associated with idelalisib in CLL	4 3 2 1	4 3 2 1
Magnitude of clinical responses observed with chimeric antigen receptor T-cell therapy in ALL	4 3 2 1	4 3 2 1
Results of the Phase III MDS-005 trial evaluating lenalidomide in patients with transfusion-dependent, lower-risk MDS without del(5q)	4 3 2 1	4 3 2 1
Benefits of the Erwinia-based preparation of asparaginase for patients with ALL	4 3 2 1	4 3 2 1

**Practice Setting:**

- Academic center/medical school     Community cancer center/hospital     Group practice  
 Solo practice     Government (eg, VA)     Other (please specify).....

**Was the activity evidence based, fair, balanced and free from commercial bias?**

- Yes     No    If no, please explain: .....

**Please identify how you will change your practice as a result of completing this activity (select all that apply).**

- This activity validated my current practice  
 Create/revise protocols, policies and/or procedures  
 Change the management and/or treatment of my patients  
 Other (please explain): .....

**If you intend to implement any changes in your practice, please provide 1 or more examples:**

.....

**The content of this activity matched my current (or potential) scope of practice.**

- Yes     No    If no, please explain: .....

**Please respond to the following learning objectives (LOs) by circling the appropriate selection:**

4 = Yes    3 = Will consider    2 = No    1 = Already doing    N/M = LO not met    N/A = Not applicable

**As a result of this activity, I will be able to:**

- Appraise recent data on therapeutic advances and changing practice standards in the management of select acute and chronic leukemias and MDS, and refine or validate existing treatment algorithms based on discussion of this information . . . . . 4 3 2 1 N/M N/A
- Recognize evidence-based therapeutic options for patients with progressive chronic myeloid leukemia. . . . . 4 3 2 1 N/M N/A
- Appreciate the FDA approvals of novel targeted agents indicated for the treatment of newly diagnosed and relapsed or refractory chronic lymphocytic leukemia, and discern how these treatments can be appropriately integrated into clinical practice. . . . . 4 3 2 1 N/M N/A
- Review existing and evolving clinical trial data to recommend safe therapeutic alternatives for patients with acute myeloid leukemia, including acute promyelocytic leukemia, and increase knowledge regarding investigational options designed for patients who are not candidates for intensive therapy . . . . . 4 3 2 1 N/M N/A

**EDUCATIONAL ASSESSMENT AND CREDIT FORM (continued)**

**As a result of this activity, I will be able to:**

- Apply the results of emerging clinical research to optimize treatment for young adult and adult patients with newly diagnosed and recurrent acute lymphoblastic leukemia . . . . . 4 3 2 1 N/M N/A
- Counsel patients with MDS about supportive and systemic treatment options to manage disease-related cytopenias and minimize leukemic progression . . . . . 4 3 2 1 N/M N/A

**Please describe any clinical situations that you find difficult to manage or resolve that you would like to see addressed in future educational activities:**

**Would you recommend this activity to a colleague?**

Yes       No      If no, please explain: .....

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- Yes, I am willing to participate in a follow-up survey.
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**PART 2 — Please tell us about the faculty and moderator for this educational activity**

	4 = Excellent	3 = Good	2 = Adequate	1 = Suboptimal		
<b>Faculty</b>					<b>Knowledge of subject matter</b>	<b>Effectiveness as an educator</b>
Jennifer R Brown, MD, PhD	4	3	2	1	4	3 2 1
Hagop M Kantarjian, MD	4	3	2	1	4	3 2 1
Charles A Schiffer, MD	4	3	2	1	4	3 2 1
B Douglas Smith, MD	4	3	2	1	4	3 2 1
David P Steensma, MD	4	3	2	1	4	3 2 1
Wendy Stock, MD, MA	4	3	2	1	4	3 2 1
<b>Moderator</b>					<b>Knowledge of subject matter</b>	<b>Effectiveness as an educator</b>
Neil Love, MD	4	3	2	1	4	3 2 1

**Please recommend additional faculty for future activities:**

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