

Lymphoma and Chronic Lymphocytic Leukemia™

U P D A T E

Conversations with Oncology Investigators
Bridging the Gap between Research and Patient Care

FACULTY INTERVIEWS

Laurie H Sehn, MD, MPH

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EDITOR

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Lymphoma and Chronic Lymphocytic Leukemia™

U P D A T E

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Lymphoma and Chronic Lymphocytic Leukemia Update

A Continuing Medical Education Audio Series

OVERVIEW OF ACTIVITY

The treatment of hematologic cancer remains a challenge for many healthcare professionals and patients despite recent gains in the management of this group of diseases. Determining which treatment approach is most appropriate requires careful consideration of patient characteristics, physician expertise and available health-system resources. To bridge the gap between research and patient care, this program features one-on-one discussions with leading hematology-oncology investigators. By providing information on the latest clinical developments in the context of expert perspectives, this activity assists medical oncologists, hematologists and hematology-oncology fellows with the formulation of evidence-based and current therapeutic strategies, which in turn facilitates optimal patient care.

LEARNING OBJECTIVES

- Evaluate recent data on therapeutic advances and changing practice standards in Hodgkin and non-Hodgkin lymphoma, including chronic lymphocytic leukemia (CLL), and integrate this information, as appropriate, into current clinical practice.
- Individualize the selection and sequence of systemic therapy for patients with newly diagnosed and relapsed/refractory CLL, considering the clinical presentation and disease characteristics.
- Consider current and emerging clinical research data in the formulation of therapeutic recommendations for patients with newly diagnosed and relapsed/refractory follicular lymphoma and diffuse large B-cell lymphoma.
- Integrate new and existing therapeutic strategies into the best-practice management of Hodgkin lymphoma.
- Review emerging clinical trial data on the efficacy and safety of brentuximab vedotin for Hodgkin lymphoma and other CD30-positive lymphomas, and use this information to prioritize protocol and nonresearch options for patients.

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Interview with Laurie H Sehn, MD, MPH

Tracks 1-19

Track 1	Case: A 66-year-old man who presents with nausea, vomiting and weight loss is diagnosed with germinal center B-cell diffuse large B-cell lymphoma (DLBCL)	Track 11	AETHERA: Efficacy and safety results from a Phase III trial of brentuximab vedotin for patients with HL at high risk of relapse or disease progression after autologous stem cell transplant (ASCT)
Track 2	Management of “double-hit” DLBCL	Track 12	Activity of immune checkpoint inhibitors in R/R HL
Track 3	CNS International Prognostic Index: A risk model for CNS relapse in patients with DLBCL treated with R-CHOP	Track 13	Treatment options for elderly patients with newly diagnosed HL
Track 4	CNS prophylaxis for patients with DLBCL	Track 14	Immune-related adverse events associated with checkpoint inhibitors
Track 5	Assessing the risk of CNS involvement and relapse for patients with primary testicular DLBCL	Track 15	Case: A 72-year-old woman with Grade I/II follicular lymphoma (FL) receives up-front bendamustine/rituximab (BR) followed by maintenance rituximab
Track 6	Incidence and risk of bowel perforation in patients with DLBCL and gastrointestinal involvement	Track 16	Primary results of the Phase III GALLIUM study: Obinutuzumab-based induction and maintenance therapy prolongs progression-free survival in comparison to rituximab-based therapy for previously untreated FL
Track 7	Case: A 62-year-old man with relapsed/refractory (R/R) Stage IVB activated B-cell-like DLBCL	Track 17	Side effects associated with bendamustine in combination with obinutuzumab or rituximab in the GALLIUM trial
Track 8	Integration of chimeric antigen receptor (CAR) T-cell therapy into the treatment algorithm for patients with R/R DLBCL	Track 18	StiL NHL7-2008 MAINTAIN: A Phase III trial evaluating 4 versus 2 years of maintenance rituximab after BR for previously untreated FL
Track 9	Case: A 31-year-old man with Stage IIB classical Hodgkin lymphoma (HL) experiences disease relapse 18 months after receiving ABVD (doxorubicin, bleomycin, vinblastine and dacarbazine)	Track 19	Perspective on the use of rituximab or obinutuzumab monotherapy for patients with FL
Track 10	ECHELON-1: Results of a Phase III trial comparing brentuximab vedotin with AVD to ABVD as front-line therapy for Stage III or IV classical HL		

Interview with Andrew D Zelenetz, MD, PhD

Tracks 1-16

Track 1	Management of chronic lymphocytic leukemia (CLL)	Track 5	Therapeutic options for patients with CLL in the front-line setting
Track 2	Approach to R/R CLL after disease progression on ibrutinib	Track 6	Mitigation of the infusion-related reactions associated with obinutuzumab
Track 3	Efficacy and tolerability of ibrutinib and venetoclax for CLL	Track 7	Efficacy and side effects associated with idelalisib for CLL
Track 4	Activity of venetoclax with rituximab for R/R CLL		

Interview with Dr Zelenetz (continued)

- | | | | |
|----------|--|----------|---|
| Track 8 | Case: A frail 84-year-old man with del(17p) CLL receives venetoclax and rituximab after disease progression on multiple lines of therapy, including ibrutinib | Track 13 | Role of the lenalidomide/rituximab (R ²) regimen in FL |
| Track 9 | Clinical implications of the GALLIUM trial | Track 14 | Case: A 57-year-old woman with ALK-negative anaplastic large cell lymphoma |
| Track 10 | Perspective on the toxicity associated with bendamustine on the GALLIUM study | Track 15 | Ongoing Phase II study of high-dose chemotherapy with ASCT followed by maintenance romidepsin for patients with T-cell non-Hodgkin lymphoma |
| Track 11 | Selection of therapy for patients with R/R FL | Track 16 | Approach to CD30 testing and the use of brentuximab vedotin for patients with T-cell lymphomas |
| Track 12 | Case: A 57-year-old man with FL experiences early relapse on BR and enrolls on a clinical trial of an investigational PI3K delta inhibitor | | |

Video Program

View the corresponding video interviews with (from left) Drs Sehn and Zelenetz by Dr Love at www.ResearchToPractice.com/LymphomaCLLUpdate118/Video



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- Marcus R et al. **Obinutuzumab for the first-line treatment of follicular lymphoma.** *N Engl J Med* 2017;377(14):1331-44.
- Moskowitz CH et al; AETHERA Study Group. **Brentuximab vedotin as consolidation therapy after autologous stem-cell transplantation in patients with Hodgkin's lymphoma at risk of relapse or progression (AETHERA): A randomised, double-blind, placebo-controlled, phase 3 trial.** *Lancet* 2015;385(9980):1853-62.
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- Schmitz N et al. **CNS International Prognostic Index: A risk model for CNS relapse in patients with diffuse large B-cell lymphoma treated with R-CHOP.** *J Clin Oncol* 2016;34(26):3150-6.
- Seymour JF et al. **Venetoclax plus rituximab in relapsed or refractory chronic lymphocytic leukaemia: A phase 1b study.** *Lancet Oncol* 2017;18(2):230-40.
- Thompson PA, Wierda WG. **Eliminating minimal residual disease as a therapeutic end point: Working toward cure for patients with CLL.** *Blood* 2016;127(3):279-86.
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- Woyach JA et al. **BTKC481S-mediated resistance to ibrutinib in chronic lymphocytic leukemia.** *J Clin Oncol* 2017;35(13):1437-43.

QUESTIONS (PLEASE CIRCLE ANSWER):

1. The pivotal study evaluating the combination of idelalisib and rituximab versus rituximab alone for patients with relapsed CLL who are not eligible for standard chemotherapy demonstrated that the combination resulted in a significant improvement in _____.
 - a. Progression-free survival (PFS)
 - b. Overall survival
 - c. Both a and b

2. According to the CNS International Prognostic Index risk model, approximately _____ of patients with DLBCL are in the category of high risk for CNS involvement.
 - a. 10%
 - b. 25%
 - c. 40%

3. Which of the following statements is true regarding the Phase III ECHELON-1 trial comparing brentuximab vedotin in combination with AVD to ABVD for advanced classical HL?
 - a. Patients with R/R disease were evaluated
 - b. Brentuximab vedotin with AVD was superior to ABVD in terms of PFS
 - c. Both a and b

4. Which of the following observations was made in the Phase III GALLIUM study evaluating obinutuzumab- versus rituximab-based induction and maintenance therapy for previously untreated FL?
 - a. No difference in PFS
 - b. PFS favoring rituximab
 - c. PFS favoring obinutuzumab

5. Side effects associated with ibrutinib include _____.
 - a. Hypertension
 - b. Myalgias/arthralgias
 - c. Diarrhea
 - d. All of the above

6. A recent study by Seymour and colleagues investigating the combination of venetoclax and rituximab for R/R CLL demonstrated that patients who achieved a complete response and a minimal residual disease-negative status had durable remissions after discontinuing therapy.
 - a. True
 - b. False

7. Which of the following categories reflects the mechanism of action of copanlisib?
 - a. Anti-PD-1/PD-L1 antibody
 - b. Bruton tyrosine kinase inhibitor
 - c. Bcl-2 inhibitor
 - d. PI3 kinase inhibitor

8. Results of the Phase III AETHERA trial evaluating brentuximab vedotin versus placebo as consolidation therapy after ASCT for patients with HL at high risk of relapse or progression demonstrated a statistically significant advantage in _____ with brentuximab vedotin.
 - a. Overall survival
 - b. PFS
 - c. Both a and b
 - d. Neither a nor b

9. Which of the following statements is true regarding immune checkpoint inhibitors for the treatment of HL?
 - a. They elicit high overall response rates
 - b. Remissions are usually not durable
 - c. Both a and b

10. The transaminitis associated with idelalisib _____.
 - a. Is often observed in patients with heavily pretreated disease
 - b. Is immune mediated
 - c. Requires permanent discontinuation of the drug in all cases
 - d. All of the above
 - e. Both a and b

EDUCATIONAL ASSESSMENT AND CREDIT FORM

Lymphoma and Chronic Lymphocytic Leukemia Update — Volume 1, Issue 3

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 and safety results from the Phase III GALLIUM trial and FDA approval of obinutuzumab for untreated FL	4	3	2	1
Available research data with and clinical role of CAR T-cell therapy for aggressive lymphomas	4	3	2	1
Top-line efficacy and safety results from the Phase III ECHELON-1 trial evaluating brentuximab vedotin with AVD versus ABVD as front-line therapy for advanced classical HL	4	3	2	1
The CNS International Prognostic Index risk model for CNS relapse in patients with DLBCL	4	3	2	1
Activity of venetoclax with rituximab for R/R CLL	4	3	2	1
Efficacy and tolerability of immune checkpoint inhibitors for R/R HL	4	3	2	1

Practice Setting:

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

Approximately how many new patients with the following do you see per year?

CLL..... HL..... FL.....
 Mantle cell lymphoma..... DLBCL..... T-cell lymphoma.....

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).

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

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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:

- Evaluate recent data on therapeutic advances and changing practice standards in Hodgkin and non-Hodgkin lymphoma, including chronic lymphocytic leukemia (CLL), and integrate this information, as appropriate, into current clinical practice. 4 3 2 1 N/M N/A
- Individualize the selection and sequence of systemic therapy for patients with newly diagnosed and relapsed/refractory CLL, considering the clinical presentation and disease characteristics. 4 3 2 1 N/M N/A
- Consider current and emerging clinical research data in the formulation of therapeutic recommendations for patients with newly diagnosed and relapsed/refractory follicular lymphoma and diffuse large B-cell lymphoma. 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:

- Integrate new and existing therapeutic strategies into the best-practice management of Hodgkin lymphoma. 4 3 2 1 N/M N/A
- Review emerging clinical trial data on the efficacy and safety of brentuximab vedotin for Hodgkin lymphoma and other CD30-positive lymphomas, and use this information to prioritize protocol and nonresearch options for patients. . . . 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?

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.....

.....

PART 2 — Please tell us about the faculty and editor for this educational activity									
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Laurie H Sehn, MD, MPH	4	3	2	1	4	3	2	1	
Andrew D Zelenetz, MD, PhD	4	3	2	1	4	3	2	1	
Editor	Knowledge of subject matter				Effectiveness as an educator				
Neil Love, MD	4	3	2	1	4	3	2	1	

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U P D A T E

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