

Follicular Lymphoma™

U P D A T E

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

FACULTY INTERVIEWS

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Follicular Lymphoma™

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Follicular Lymphoma Update

A Continuing Medical Education Audio Series

OVERVIEW OF ACTIVITY

Follicular lymphoma (FL) is an indolent form of non-Hodgkin lymphoma that can vary significantly in its clinical presentation. Because of this variety, no single standard approach to the initial management of the disease has been established and available options range from watchful waiting, radiation therapy and rituximab monotherapy to various combinations of chemoimmunotherapy. Recent data sets and corresponding FDA actions have led to the emergence of new therapeutic targets and regimens, thereby altering management algorithms, and in order to offer optimal patient care, including the option of clinical trial participation, the practicing medical oncologist must be well informed of these advances. 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 emerging research data and recent FDA approvals when designing an optimal therapeutic approach for patients with newly diagnosed FL requiring active therapy.
- Recall published research data and other clinical factors in the best-practice selection, sequencing or combining of available therapeutic agents in the nonresearch care of patients with relapsed/refractory (R/R) FL.
- Compare and contrast the efficacy and safety of the PI3-kinase inhibitors approved for the treatment of R/R FL to determine the current role of each in clinical practice.
- Develop practical strategies to prevent, recognize and ameliorate the toxicities associated with therapies routinely used in the management of FL.
- Identify ongoing clinical trials evaluating innovative investigational approaches for FL, and obtain consent from appropriate patients for study participation.

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Please note, this program has been specifically designed for the following ABIM specialty: **medical oncology**.

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Interview with Nathan H Fowler, MD

Tracks 1-26

- Track 1** Overview of current treatment approaches in follicular lymphoma (FL)
- Track 2** Mechanism of action of the anti-CD20 antibodies rituximab and obinutuzumab
- Track 3** Activity of lenalidomide and possible correlation of cereblon expression with response in FL
- Track 4** Inhibition of the PI3-kinase pathway in low-grade lymphomas
- Track 5** Activity of BTK inhibitors in FL
- Track 6** Rationale for the Phase III RELEVANCE trial: Lenalidomide/rituximab (R²) versus rituximab/chemotherapy, each followed by maintenance rituximab, as first-line therapy for FL
- Track 7** Design of the RELEVANCE study for patients with previously untreated FL
- Track 8** Duration of up-front R² and maintenance rituximab
- Track 9** Adherence to long-term lenalidomide therapy
- Track 10** Perspective on the 3-year progression-free survival results from the RELEVANCE trial
- Track 11** Tolerability of R² versus rituximab/chemotherapy
- Track 12** Clinical experience with lenalidomide-associated neutropenia, rash and diarrhea
- Track 13** Second-line therapy options for patients who receive first-line R²
- Track 14** **Case:** A 52-year-old woman with symptomatic Grade I/II FL attains a complete remission (CR) with first-line R² therapy
- Track 15** **Case:** A 53-year-old woman who previously received several lines of therapy, including R-CHOP, BR, R² and idelalisib, receives chimeric antigen receptor (CAR) T-cell therapy on a clinical trial
- Track 16** Activity of CAR T-cell therapies in FL
- Track 17** Potential synergy of lenalidomide and CAR T-cell therapy
- Track 18** Activity of idelalisib for patients with FL
- Track 19** Side effects and quality of life with idelalisib
- Track 20** **Case:** A 73-year-old woman with relapsed/refractory FL receives copanlisib
- Track 21** Selection of PI3-kinase inhibitor for patients with FL
- Track 22** Risks and benefits with idelalisib versus copanlisib
- Track 23** Treatment options for patients with FL and early disease relapse
- Track 24** **Case:** A 43-year-old man presents with Grade IIIA FL, receives R-CHOP and experiences a CR lasting for 5 years
- Track 25** Response to second-line lenalidomide/obinutuzumab
- Track 26** Perspective on the use of CAR T-cell therapy or immune checkpoint inhibition for patients with transformed FL

Interview with Gilles A Salles, MD, PhD

Tracks 1-20

- Track 1** **Case:** A 56-year-old man with Stage IV FL achieves a CR with the R² regimen as first-line therapy on a clinical trial
- Track 2** Therapeutic approach to Grade III FL
- Track 3** Deciding between treatment and observation for patients with newly diagnosed FL
- Track 4** RELEVANCE: A Phase III trial evaluating R² versus rituximab/chemotherapy, each followed by maintenance rituximab, for previously untreated FL
- Track 5** Dosing and administration schedule of the R² regimen
- Track 6** Side effects and quality of life with R² versus rituximab/chemotherapy
- Track 7** Activity and tolerability of R² in patients with untreated FL

Interview with Dr Salles (continued)

- Track 8** **Case:** A 78-year-old woman who receives idelalisib for relapsed FL experiences gastrointestinal and hepatic toxicity
- Track 9** Efficacy and tolerability of idelalisib
- Track 10** Monitoring and management of hepatic toxicity and neutropenia associated with idelalisib
- Track 11** Risk of opportunistic infections and colitis with idelalisib
- Track 12** Comparison of the PI3-kinase isoforms targeted and the tolerability of idelalisib, duvelisib and copanlisib
- Track 13** Perspective on the differences in efficacy among PI3-kinase inhibitors
- Track 14** Hyperglycemia associated with copanlisib
- Track 15** Therapeutic options for patients with relapsed FL
- Track 16** Approach to patients with FL who experience relapse early in their treatment course
- Track 17** Perspective on the efficacy of obinutuzumab-based chemotherapy versus rituximab-based chemotherapy for FL
- Track 18** **Case:** A 44-year-old man receives rituximab for Stage II FL and experiences disease transformation to diffuse large B-cell lymphoma
- Track 19** Efficacy of CAR T-cell therapy for patients with FL
- Track 20** **Case:** A 55-year-old man experiences a prolonged CR after receiving rituximab for relapsed FL

Video Program

View the corresponding video interviews with (from left) Drs Fowler and Salles by Dr Love at www.ResearchToPractice.com/FLUpdate119/Video



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QUESTIONS (PLEASE CIRCLE ANSWER):

- 1. Idelalisib may be associated with an increased risk for _____.**
 - a. Neutropenia
 - b. Colitis
 - c. Hepatotoxicity
 - d. Opportunistic infections
 - e. All of the above
- 2. Which of the following statements is true regarding the Phase III RELEVANCE study comparing R² to rituximab/chemotherapy, each followed by maintenance rituximab, for previously untreated FL?**
 - a. R² significantly improved progression-free survival
 - b. R² significantly improved the CR rate
 - c. R² and rituximab/chemotherapy demonstrated similar efficacy
- 3. The phase III GALLIUM study comparing first-line obinutuzumab-based chemotherapy to rituximab-based chemotherapy, followed by obinutuzumab or rituximab maintenance, demonstrated no difference in progression-free survival.**
 - a. True
 - b. False
- 4. A recent study by Schuster and colleagues published in *The New England Journal of Medicine* evaluating the efficacy of CAR T-cell therapy in patients with refractory diffuse large B-cell lymphoma or FL reported a CR rate of approximately _____ in the FL cohort.**
 - a. 30%
 - b. 50%
 - c. 70%
- 5. The ongoing AUGMENT trial is evaluating the R² regimen versus rituximab alone for patients with indolent non-Hodgkin lymphoma in which setting?**
 - a. First line
 - b. Relapsed/refractory disease
- 6. Side effects associated with lenalidomide include _____.**
 - a. Neutropenia
 - b. Constipation
 - c. Diarrhea
 - d. All of the above
 - e. Both a and c
- 7. What is the mechanism of action of tazemetostat, an agent that has shown promising response rates in the treatment of relapsed/refractory FL?**
 - a. Immune checkpoint inhibitor
 - b. Antibody-drug conjugate
 - c. EZH2 inhibitor
- 8. In the RELEVANCE trial, the toxicity profile of the R² regimen was observed to be similar to that of rituximab/chemotherapy, with similar rates of neutropenia, diarrhea and alopecia.**
 - a. True
 - b. False
- 9. Which statement is true regarding copanlisib in the treatment of FL?**
 - a. It is administered orally
 - b. It inhibits the alpha and delta isoforms of PI3 kinase
 - c. It may be associated with hyperglycemia and hypertension
 - d. All of the above
 - e. Both a and b
 - f. Both b and c
- 10. What was the duration of maintenance therapy with rituximab in both the PRIMA and RELEVANCE trials?**
 - a. 12 months
 - b. 18 months
 - c. 24 months

EDUCATIONAL ASSESSMENT AND CREDIT FORM

Follicular Lymphoma Update — Volume 1, Issue 1

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						
Results and clinical implications of the RELEVANCE study of R ² for patients with previously untreated FL	4	3	2	1			4	3	2	1			
Efficacy and side effects of PI3-kinase inhibitors for patients with FL	4	3	2	1			4	3	2	1			
Activity and tolerability of the R ² regimen for patients with FL	4	3	2	1			4	3	2	1			
Emerging data with CAR T-cell therapies in FL	4	3	2	1			4	3	2	1			
Efficacy of obinutuzumab-based chemotherapy versus rituximab-based chemotherapy in FL	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).....

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

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:

- Evaluate emerging research data and recent FDA approvals when designing an optimal therapeutic approach for patients with newly diagnosed FL requiring active therapy. 4 3 2 1 N/M N/A
- Recall published research data and other clinical factors in the best-practice selection, sequencing or combining of available therapeutic agents in the nonresearch care of patients with relapsed/refractory (R/R) FL. 4 3 2 1 N/M N/A
- Compare and contrast the efficacy and safety of the PI3-kinase inhibitors approved for the treatment of R/R FL to determine the current role of each in clinical practice. 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:

- Develop practical strategies to prevent, recognize and ameliorate the toxicities associated with therapies routinely used in the management of FL. 4 3 2 1 N/M N/A
- Identify ongoing clinical trials evaluating innovative investigational approaches for FL, and obtain consent from appropriate patients for study participation. 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:

PART 2 — Please tell us about the faculty and editor for this educational activity									
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Faculty		Knowledge of subject matter				Effectiveness as an educator			
Nathan H Fowler, MD		4	3	2	1	4	3	2	1
Gilles A Salles, 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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Follicular Lymphoma™

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