

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

FACULTY INTERVIEWS

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Lung Cancer Update

A Continuing Medical Education Audio Series

OVERVIEW OF ACTIVITY

Traditional chemotherapy, surgery and radiation therapy have had a modest effect on long-term outcomes for patients with lung cancer. However, the advent of biologic and immunotherapeutic agents has led to recent improvements in disease-free and overall survival in select populations. In order to offer optimal patient care, including the option of clinical trial participation, clinicians must be well informed of these advances. Featuring information on the latest research developments, this program is designed to assist medical and radiation oncologists with the formulation of up-to-date strategies for the care of patients with lung cancer.

LEARNING OBJECTIVES

- Compare and contrast the mechanisms of action, efficacy and safety/toxicity of approved and investigational
 anti-PD-1/PD-L1 antibodies for the treatment of lung cancer to determine the current and/or potential utility of
 each in clinical practice.
- Formulate management strategies for small cell lung cancer, considering systemic therapy in addition to current research studies evaluating novel immunotherapeutic and targeted approaches.
- Appreciate the FDA approval of durvalumab and available Phase III data documenting the benefit of sequential
 anti-PD-L1 therapy after the completion of chemoradiation therapy for unresectable Stage III non-small cell lung
 cancer, and consider the role of this therapeutic approach for appropriate patients.
- Develop a genomic testing algorithm to assist in identifying appropriate patients eligible for protocol and clinical targeted treatment options.
- Consider published safety and efficacy data with available and emerging therapeutic strategies, and appropriately
 incorporate targeted therapies into the care of patients with identified tumor driver mutations or alterations.
- Educate patients about the side effects associated with recently approved novel agents and immunotherapeutic
 approaches, and provide preventive strategies to reduce or ameliorate these toxicities.

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Interview with David R Spigel, MD

Tracks 1-20

Track 1	Case: A 73-year-old man with recurrent small cell lung cancer (SCLC) receives ipilimumab/nivolumab on a clinical trial	Track 11	Clinical implications of the KEYNOTE-042 results; perspective the future clinical utility of TPS			
Track 2	Management of immune checkpoint inhibitor-associated rash	Track 12	Evolution of first-line checkpoint inhibitor-based treatment for metastatic nonsquamous NSCLC			
Track 3	Correlation between toxicity and treatment benefit for patients receiving immune checkpoint inhibitors		with and without targetable tumor mutations			
		Track 13	Selection of checkpoint inhibitor- based regimens for patients experi-			
Track 4	Clinical experience with dermatologic side effects of checkpoint inhibitors		encing disease progression on an EGFR tyrosine kinase inhibitor (TKI)			
Track 5	Second-line therapy options for metastatic SCLC	Track 14	Case: A 57-year-old man with Stage IIIA squamous NSCLC receives chemoradiation therapy followed by			
Track 6 Track 7	Perspective on ipilimumab/nivolumab dosing and therapy-associated toxicities Activity, side effects and ongoing investigation of the antibody-drug conjugate rovalpituzumab tesirine (Rova-T) in DLL3-positive SCLC		durvalumab			
		Track 15	Ongoing studies of checkpoint inhibitors in the (neo)adjuvant setting			
		Track 16	PACIFIC trial: Efficacy and tolerability of durvalumab after chemoradiation therapy for unresectable Stage III NSCLC			
Track 8	Clinical experience with Rova-T-associated edema	Track 17	Management of chemoradiation therapy-associated pneumonitis			
Track 9 Track 10	Results of the Phase III KEYNOTE-407 trial evaluating the addition of pembrolizumab to carboplatin with paclitaxel or nab paclitaxel as first-line therapy for metastatic squamous non-small cell lung cancer (NSCLC) KEYNOTE-042: Overall survival benefit with pembrolizumab versus platinum-based chemotherapy as first-line treatment for locally advanced or metastatic NSCLC with a PD-L1 tumor proportion score (TPS)	Track 18	Perspective on the synergy of durvalumab and chemoradiation therapy			
		Track 19	Use of chemoradiation therapy followed by durvalumab for patients with Stage III NSCLC and a targetable tumor mutation			
		Track 20	Case: A woman in her early fifties with advanced "pan-negative" nonsquamous NSCLC experiences a near complete response with 1 dose of ipilimumab/nivolumab			

Interview with Justin F Gainor, MD

of 1% or higher

Tracks 1-17

Track 1	Case: A 76-year-old man and never smoker presents with metastatic NSCLC with an EGFR L858R tumor mutation, a low PD-L1 TPS and brain metastases and receives first-line osimertinib	Track 3	Stereotactic radiosurgery, whole-brain radiation therapy (WBRT) and EGFR TKIs for patients with EGFR tumor mutations and brain metastases
		Track 4	Incidence and pathophysiology of neurocognitive effects of WBRT
Track 2	Activity and tolerability of first-line osimertinib	Track 5	Optimal sequencing of EGFR TKIs

Interview with Dr Gainor (continued)

- Track 6 Mechanism of action, benefits and limitations of the second-generation EGFR inhibitor decomitinib
- Track 7 Investigational strategies for patients experiencing disease progression on osimertinib
- Track 8 Bevacizumab with erlotinib as first-line therapy for patients with metastatic NSCLC and an EGFR tumor mutation
- Track 9 Rationale for combining first- and third-generation EGFR TKIs to potentially treat tumors with resistance mutations
- Track 10 Case: A 57-year-old woman and never smoker with crizotinib-refractory NSCLC with an ALK rearrangement receives alectinib
- Track 11 Sequencing of ALK inhibitors for patients with metastatic NSCLC with an ALK rearrangement
- Track 12 Second-line therapy options for patients with metastatic NSCLC with an ALK rearrangement

- Track 13 Case: A 64-year-old woman and never smoker with NSCLC with brain and bone metastases initially treated with carboplatin/pemetrexed is found to harbor a RET rearrangement
- Track 14 Case: A 48-year-old man with heavily pretreated nonsquamous NSCLC whose tumor is positive for an NTRK gene fusion receives entrectinib on a clinical trial
- Track 15 Efficacy of the TRK inhibitors entrectinib and larotrectinib
- Track 16 Case: A 71-year-old man and 35 pack-year smoker with metastatic nonsquamous NSCLC and a BRAF V600E tumor mutation, renal insufficiency and a high TPS receives dabrafenib/trametinib
- Track 17 Consideration of immune checkpoint inhibitor-based regimens as second-line therapy for patients with metastatic disease and renal insufficiency

Video Program

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SELECT PUBLICATIONS

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Lung Cancer Update — Volume 15, Issue 2

QUESTIONS (PLEASE CIRCLE ANSWER):

first-line osimertinib to either erlotinib or gefitinib for advanced NSCLC with an EGFR

1. The Phase III FLAURA study comparing

a. 1% or higher

b. 20% or higher

c. 50% or higher

d. All of the above

gefitinib for advanced NSCLC with an E tumor mutation demonstrated a signific improvement in progression-free surviva (PFS) for patients who received osimert a. True b. False	ant ALK rearrangement. a. Alectinib
2. Which of the following categories reflec the mechanism of action of Rova-T? a. Antibody-drug conjugate b. Anti-PD-1 antibody c. RET inhibitor	ts 7 is a promising investigational agent that targets TRK kinases in adult and pediatric patients with cancers harboring an NTRK gene fusion. a. Entrectinib b. Larotrectinib
3. Results of a Phase III trial evaluating	c. Both a and b
dacomitinib versus gefitinib as first-line therapy for patients with locally advance or metastatic NSCLC and an EGFR turn mutation demonstrated a significant impent in with dacomitinib. a. Overall survival b. PFS c. Both a and b 4. The results of the Phase III IMpower150 trial of atezolizumab and/or bevacizuma added to carboplatin and paclitaxel as first-line therapy for patients with metas	8. The Phase III KEYNOTE-407 trial evaluating the addition of pembrolizumab to carbopatin with paclitaxel or <i>nab</i> paclitaxel as first-line therapy for metastatic squamous NSCLC demonstrated prolonged median overall survival and PFS with the addition of pembrolizumab to conventional chemotherapy across all PD-L1 expression subgroups. a. True b. False
nonsquamous NSCLC failed to demonst any statistically significant improvement in overall survival or PFS with the addit of atezolizumab and bevacizumab to carboplatin/paclitaxel. a. True b. False	trate 9. The Phase III PACIFIC trial of durvalumab versus placebo for patients with locally
Results of the Phase III KEYNOTE-042 demonstrated a significant improvemen overall survival with single-agent pembr zumab compared to platinum-based che	trial b. PFS t in c. Objective response rate oli- emo-
therapy as first-line treatment for locally advanced or metastatic NSCLC in patie with a PD-L1 TPS of	

_____ is a second-generation ALK inhibitor that is currently FDA approved for

alectinib for patients with _____ advanced

NSCLC with an ALK rearrangement.

a. Treatment-naïve

b. Previously treated

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	•	
How would you characterize your level of knowledge on the following topics:		Culpantin
4 = Excellent $3 = Good$ $2 = A$		
	BEFORE	AFTER
Results of the Phase III FLAURA trial and use of osimertinib as first-line therapy for advanced NSCLC with an EGFR tumor mutation	4 3 2 1	4 3 2 1
Clinical implications of the KEYNOTE-042 trial results: Overall survival benefit with pembrolizumab versus platinum-based chemotherapy as first-line treatment for metastatic NSCLC with a PD-L1 TPS of 1% or higher	4 3 2 1	4 3 2 1
Sequencing of FDA-approved ALK inhibitors for NSCLC with an ALK rearrangement	4 3 2 1	4 3 2 1
$\label{eq:pacific} \textbf{PACIFIC: Results of a Phase III trial of durvalumab as sequential treatment for locally advanced, unresectable Stage III NSCLC} $	4 3 2 1	4 3 2 1
Clinical implications of the Phase III KEYNOTE-407 trial evaluating the addition of pembrolizumab to carboplatin with paclitaxel or <i>nab</i> paclitaxel as first-line therapy for metastatic squamous NSCLC	4 3 2 1	4 3 2 1
Practice Setting:		
□ Academic center/medical school □ Community cancer center/host	spital \Box (Group practice
Solo practice ☐ Government (eg, VA) ☐ Other (please specification)	cify)	
Approximately how many new patients with lung cancer do you see per year?		patien
Nas the activity evidence based, fair, balanced and free from commercial b		
→ Yes → No If no, please explain:		
Please identify how you will change your practice as a result of completing apply).	this activity (se	lect all that
☐ 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):		
f you intend to implement any changes in your practice, please provide 1 o		
	-	
The content of this activity matched my current (or potential) scope of practive Yes No If no, please explain:		
Please respond to the following learning objectives (LOs) by circling the app		
4 = Yes 3 = Will consider 2 = No 1 = Already doing N/M = LO not r	riet IN/A = Not	applicable
As a result of this activity, I will be able to:		
 Compare and contrast the mechanisms of action, efficacy and safety/toxicity of approved and investigational anti-PD-1/PD-L1 antibodies for the treatment of cancer to determine the current and/or potential utility of each in clinical pract 	lung	2 1 N/M N
 Formulate management strategies for small cell lung cancer, considering syste therapy in addition to current research studies evaluating novel immunotherap and targeted approaches. 	peutic	2 1 N/M N
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therapy for unresectable Stage III non-small cell lung cancer, and consider the of this therapeutic approach for appropriate patients.	e roie 4 3	2 1 N/M N

EDUCATIONAL ASSESSMENT	AND CRE	DIT F	ORN	/I (continue	ed)				
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 Develop a genomic testing algorithm eligible for protocol and clinical targe 	to assist in eted treatmen	identifyi nt optio	ing ap	ppropriate p	oatients 	4	3 2 1	L N/M	N/A
Consider published safety and efficacy data with available and emerging therapeutic strategies, and appropriately incorporate targeted therapies into the care of patients									
with identified tumor driver mutation						4	3 2 1	l N/M	N/A
 Educate patients about the side effects associated with recently approved novel agents and immunotherapeutic approaches, and provide preventive strategies to reduce or ameliorate these toxicities						N/A			
Please describe any clinical situatio to see addressed in future education	ns that you	find di							
Would you recommend this activity t	o a colleagu	ıe?							
If no, please explain:									
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					1 = Suboptimal Effectiveness as an edu				
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Justin F Gainor, MD	4	3	2	1	4	3	2	1	
			_	-	'		_	-	
Editor		•	-	ct matter	Effective				or
Neil Love, MD	4	3	2	1	4	3	2	1	
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Lung Cancer

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