CNS Cancer U P D A T E

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

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CNS Cancer Update

A Continuing Medical Education Audio Series

OVERVIEW OF ACTIVITY

Brain tumors are a diverse group of neoplasms arising from different cells within the central nervous system (CNS) or from systemic tumors that have metastasized to the CNS. Primary brain tumors include a number of histologic types with markedly different tumor growth rates and are divided into anaplastic gliomas (anaplastic astrocytoma, anaplastic oligodendroglioma and anaplastic oligoastrocytoma) and glioblastoma multiforme (GBM) based on their histopathologic features. Despite treatment, the median survival for anaplastic oligodendroglioma is 2 to 3 years, and patients with GBM can succumb to their disease within a year of its onset. Thus, clinical education regarding standard and evolving best-practice therapeutic management of these neoplasms is essential to improving patient outcomes. To bridge the gap between research and patient care, this issue of CNS Cancer Update features one-on-one discussions with leading neuro-oncologists and neurosurgeons. By providing information on the latest research developments in the context of expert perspectives, this activity assists medical oncologists with the formulation of state-of-the-art clinical management strategies, which in turn facilitates optimal patient care.

LEARNING OBJECTIVES

- Utilize case-based learning to facilitate the development of evidence-based and individualized management strategies for newly diagnosed and recurrent primary CNS cancers.
- Identify strategies to distinguish between true disease progression and radiographic pseudoprogression in patients with malignant glioma who have undergone chemoradiation therapy.
- Evaluate the prognostic and/or predictive role of molecular profiling (eg, MGMT methylation status) in primary CNS cancers.
- Recall the rationale for and activity of novel targeted agents under investigation for the treatment of CNS cancers.
- Adhere to guideline-based approaches for the use of prophylactic anticonvulsant therapy for patients with primary CNS cancer or brain metastases who have no history of seizures.
- Incorporate key recent clinical trial data into treatment planning for patients with primary CNS lymphomas.
- Counsel appropriately selected patients with CNS cancer about participation in ongoing clinical trials.

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QUESTIONS (PLEASE CIRCLE ANSWER):

- Up to 40% of patients with MGMTmethylated GBM have been observed to develop recurrence outside of the original tumor site.
 - a. True
 - b. False
- 2. Which of the following is generally true about pseudoprogression in GBM?
 - a. Occurs more frequently in MGMTmethylated tumors
 - b. Tends to occur within 3 months of finishing temozolomide (TMZ) and radiation therapy (RT)
 - c. Tends to occur within the radiation field
 - d. All of the above
- 3. Which of the following was demonstrated in a French randomized trial of 1 course of RT with best supportive care (BSC) versus BSC alone for elderly patients with GBM?
 - a. RT improved overall survival
 - b. RT resulted in a clinically significant decline in cognitive functioning
 - c. RT resulted in decrements in quality of life
- 4. A randomized study with patients with primary CNS lymphoma demonstrated an improvement in radiographic response with the addition of cytarabine to methotrexate compared to methotrexate alone.
 - a. True
 - b. False
- In an NCI-sponsored pilot trial, no patients with relapsed primary CNS lymphoma experienced objective radiographic response to rituximab monotherapy.
 - a. True
 - b. False

- 6. Which of the following was observed in the Phase III RTOG-0525 study of standard adjuvant TMZ versus dosedense TMZ for patients with newly diagnosed GBM?
 - Dose-dense TMZ resulted in an improvement in median overall survival and progression-free survival compared to standard TMZ
 - Dose-dense and standard TMZ were equivalent with regard to median overall survival
 - c. MGMT methylation status was strongly prognostic
 - d. Both a and c
 - e. Both b and c
- In patients with GBM whose disease progresses on bevacizumab, the use of fractionated stereotactic radiation therapy may result in increased responsiveness to subsequent bevacizumab/ chemotherapy.
 - a. True
 - b. False
- 8. A highly infiltrative pattern of disease recurrence will emerge among a proportion of patients with GBM who receive bevacizumab.
 - a. True
 - b. False
- Studies have demonstrated that VEGF tyrosine kinase inhibitors result in similar efficacy to bevacizumab in recurrent GBM.
 - a. True
 - b. False
- In an uncontrolled, retrospective study, the development of hypertension and proteinuria in patients with recurrent GBM treated with bevacizumab was associated with
 - a. Increased overall survival
 - b. Decreased overall survival
 - c. No effect on overall survival

EDUCATIONAL ASSESSMENT AND CREDIT FORM

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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 ONE — Please tell us about your experience with this educational activity

| How would you characterize your level of knowledge on the following topics? | | | | | | | | |
|--|----------------------|-------------|--|--|--|--|--|--|
| 4 = Excellent $3 = Good$ $2 =$ | | | | | | | | |
| | BEFORE | AFTER | | | | | | |
| Results of RTOG-0525: A Phase III trial of standard adjuvant TMZ versus a dose-dense schedule in newly diagnosed GBM | 4 3 2 1 | 4 3 2 1 | | | | | | |
| Timing and characteristics of pseudoprogression after RT and TMZ | 4 3 2 1 | 4 3 2 1 | | | | | | |
| Effect of bevacizumab on edema and mass effect in second- and later-line treatment of recurrent GBM | 4 3 2 1 | 4 3 2 1 | | | | | | |
| Salvage fractionated stereotactic RT for GBM that progresses on bevacizumab | 4 3 2 1 | 4 3 2 1 | | | | | | |
| Prognosis and treatment advances for primary and secondary CNS lymphomas | 4 3 2 1 | 4 3 2 1 | | | | | | |
| Was the activity evidence based, fair, balanced and free from comm Yes No If no, please explain: | | | | | | | | |
| that apply). This activity validated my current practice; no changes will be made 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 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: Utilize case-based learning to facilitate the development of evidence-and individualized management strategies for newly diagnosed and reprimary CNS cancers. Identify strategies to distinguish between true disease progression an endiagraphic secundary progression in patients with malignant gligmants. | ecurrent 4 3 d | 2 1 N/M N/A | | | | | | |
| radiographic pseudoprogression in patients with malignant glioma wh have undergone chemoradiation therapy • Evaluate the prognostic and/or predictive role of molecular profiling | 4 3 | 2 1 N/M N/A | | | | | | |
| (eg, MGMT methylation status) in primary CNS cancers | | 2 1 N/M N/A | | | | | | |
| investigation for the treatment of CNS cancers | | 2 1 N/M N/A | | | | | | |
| anticonvulsant therapy for patients with primary CNS cancer or brain metastases who have no history of seizures | 4 3 | 2 1 N/M N/A | | | | | | |
| Incorporate key recent clinical trial data into treatment planning for pa with primary CNS lymphomas. Coursel prographical policy of the control o | 4 3 | 2 1 N/M N/A | | | | | | |
| Counsel appropriately selected patients with CNS cancer about particle in ongoing clinical trials. | | 2 1 N/M N/A | | | | | | |

EDUCATIONAL ASSESSMENT AND CREDIT FORM (continued)

| 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 | | | | | | | | | |
| ☐ Yes ☐ No If no, please explain: Additional comments about this activity: | | | | | | | | | |
| As part of our ongoing, continuous up surveys to assess the impact of indicate your willingness to particip Yes, I am willing to participate in No, I am not willing to participate | quality-imp our educati ate in such a a follow-u te in a follor | orovem ional i a sur p surv w-up s | nterve vey. vey. | fort, we contions on | profession | tactiv al pra | ity fo ctice. | llow- | |
| PART TWO — Please tell us ab | out the fac | culty a | nd ed | tor for thi | s education | nal ac | tivity | | |
| 4 = Excellent 3 | = Good | = Good 2 = Adequate | | | 1 = Suboptimal | | | | |
| Faculty | Knowled | ge of | subjec | t matter | Effectiveness as an educator | | | | |
| Tracy Batchelor, MD, MPH | 4 | 3 | 2 | 1 | 4 | 3 | 2 | 1 | |
| Tom Mikkelsen, MD | 4 | 3 | _ | 1 | 4 | 3 | 2 | 1 | |
| Editor | | nowledge of subject matter | | | | educator | | | |
| Neil Love, MD | 4 | 3 | 2 | 1 | 4 | 3 | 2 | 1 | |
| Other comments about the faculty a | | | | | | | | | |
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