

tral nervous system (CNS) progression in a cohort of patients undergoing surgical resection of melanoma brain metastases. **METHODS:** This retrospective, single-center study included patients undergoing first-time surgical resection of melanoma brain metastases. A multivariate Cox proportional model was used to estimate the association of patient and treatment factors with OS and CNS progression. **RESULTS:** 85 patients underwent first-time resection of 97 melanoma brain metastases with a median follow-up of 9.5 months. Checkpoint inhibitors (Pembrolizumab, Ipilimumab, and/or Nivolumab) were used in 55.1% of cases (19 pre-op; 47 post-op; median 9 cycles). Patients treated with checkpoint inhibitors had similar peri-op systemic disease status and KPS but had been treated with more systemic agents and had more instances of CNS progression prior to surgery. Median OS and time to CNS progression for the cohort were 1 year and 237 days, respectively. In a multivariate Cox regression model, age (HR 1.03 by decade;  $p=0.02$ ), treatment with a checkpoint inhibitor (HR 0.27;  $p<0.0001$ ), prior radiotherapy (HR 2.44;  $p=0.007$ ), and number of brain metastases at the time of surgery (HR 1.05 per metastasis;  $p=0.04$ ) were significant predictors of OS. Checkpoint inhibitor treatment was associated with longer OS from surgery (median 3 vs 0.5 yrs, log-rank  $p=0.004$ ). However, patients who underwent craniotomy after prior checkpoint inhibitor treatment had poor OS (median 0.56 yrs). Prior radiotherapy was also associated with poor OS (median 0.53 yrs). **CONCLUSIONS:** While checkpoint inhibitor treatment was associated with improved survival in this surgical cohort of melanoma brain metastases, patients who require surgical resection after checkpoint inhibitor treatment or radiotherapy are poor surgical candidates.

### 13. MANAGEMENT OF BRAIN METASTASES FROM SMALL CELL LUNG CANCER USING SRS

Daniel Koffler<sup>1</sup>, Sirisha Viswanatha<sup>1</sup>, Fatemeh Fekrmandi<sup>2</sup>, Zaker Rana<sup>1</sup>, Michael Schuller<sup>1</sup>, and Anuj Goenka<sup>1</sup>; <sup>1</sup>Northwell Health Cancer Institute, Lake Success, NY, USA, <sup>2</sup>Princess Margaret Cancer Center, Toronto, ON, Canada

**PURPOSE/OBJECTIVE(S):** The management of brain metastases in patients with SCLC has become controversial in the MRI era. We examine our institutional experience treating patients with SCLC with stereotactic radiosurgery. We hypothesize that an SRS strategy in well-selected patients with close MRI surveillance will result in acceptable tumor control, and without disproportionate future neurological symptoms associated with intracranial disease. **MATERIALS/METHODS:** Patients with a diagnosis of high grade neuroendocrine lung cancer who had undergone SRS between 2013 and 2019 were identified and divided into two groups: SRS-primary and SRS-salvage. SRS-primary was defined as patients who, at time of SRS, had not received previous PCI or WBRT. SRS-salvage was defined as patients who had received previous PCI or WBRT. Primary outcome was intracranial progression free survival. Secondary outcomes included overall survival and neurologic symptom free survival (N-SFS), defined as time to development of neurologic symptoms attributed disease. **RESULTS:** Twenty patients were identified with median follow-up of 14.1 months. 11 patients were identified as SRS-primary, 9 as SRS-salvage. Among SRS-primary, median PFS and OS were 6.1 months (range 0.9 – 14.5 months) and 15.6 months (4.1–43.5) respectively. N-SFS was 11.2 months (range 3.6–40.0). 3 of 11 patients developed neurological symptoms attributable to disease. 3 underwent salvage SRS and 2 salvage WBRT. None died from intracranial disease. Among SRS-salvage, median PFS following PCI/WBRT was 9.8 months (range 1.8 – 23.6 months) and OS following salvage SRS 5.5 months (range 1.1 – 27.8 months). 3 of 9 patients developed further brain metastases post-SRS. 1 patient died from intracranial disease. **CONCLUSION:** Among well-selected patients followed with MRI surveillance, our data suggest SRS as primary management of brain metastases from SCLC may be reasonable. Symptomatic intracranial disease was uncommon after SRS, and no patients undergoing upfront SRS died from intracranial disease. Prospective data are required to validate these results.

### 14. DELAYED MRI RESPONSE TO LITT IN PATIENTS UNDERGOING IMMUNOTHERAPY

Christopher Hong and Veronica Chiang; Yale University, New Haven, CT, USA

Laser interstitial thermal therapy (LITT) is an effective treatment for regrowing lesions after previous radiosurgery to brain metastases, typically resulting in decreased perilesional edema within weeks followed by delayed reduction in lesion size. We have anecdotally observed that patients on immunotherapy (IT) at time of LITT may exhibit a delayed edema resolution response to laser ablation. Post-operative imaging for cases of LITT, performed by the senior author from June 2012–July 2019, for regrowing lesions after prior radiosurgery for brain metastases were retrospectively reviewed. The IT group was defined as any patient receiving IT treatment within 3 months of LITT. Post-operative MRIs obtained at serial time points

after surgery (2 weeks, 6 weeks, 3 months, 6 months, and 12 months) were reviewed for treatment response to LITT, defined as change in surrounding edema on T2 FLAIR and change of lesion size on T1-weighted post-contrast images. Out of 60 ablated lesions, 22 were in the IT and 38 were in the non-IT groups. There were no differences in distribution of original cancer pathology (IT: 9 melanoma, 8 lung, 5 other, non-IT: 6 melanoma, 20 lung, 12 other;  $p>0.05$ ). Time to lesion size response on T1-weighted post-contrast MRI neared but did not reach statistical significance between the IT and non-IT groups: median 3.0 versus 2.25 months (HR 1.5, 0.8–2.5, 95% CI,  $p=0.08$ ), respectively. However, time to reduction of perilesional edema on T2-weighted MRI was significantly longer in the IT group, compared to the non-IT group: median 2.25 versus 1.5 months (HR 1.5, 0.9–2.5, 95% CI,  $p=0.04$ ), respectively. These data suggest that IT around the time of LITT may lead to delayed edema reduction on MRI after LITT. We hypothesize IT may enhance normal immune-mediated mechanisms thus increasing perilesional inflammation after LITT. Further studies are needed to corroborate our observations and explore the underlying pathophysiology.

### 16. GAMMA KNIFE CLINICAL DOSE PROFILE FOR EXTENSIVE BRAIN METASTASES

Greg Bowden<sup>1</sup>, Jong Kim<sup>2</sup>, Andrew Faramand<sup>2</sup>, Kevin Fallon<sup>3</sup>, John Flickinger<sup>2</sup>, and L. Dade Lunsford<sup>2</sup>; <sup>1</sup>University of Alberta, Edmonton, AB, Canada, <sup>2</sup>UPMC, Pittsburgh, PA, USA, <sup>3</sup>MUSC, Charleston, SC, USA

**BACKGROUND:** The use of Gamma Knife stereotactic radiosurgery (GKSRS) for the treatment of extensive intracranial metastases has been expanding due to its superior dosimetry and efficacy. However, there remains a dearth of data regarding the dose parameters in actual clinical scenarios. We endeavored to calculate the radiation dose to the brain when treating >15 brain metastases with GKSRS. **METHODS:** This retrospective analysis reviewed dosage characteristics for patients requiring single session GKSRS for the treatment of <sup>3</sup>15 brain metastases. Forty-two patients met the inclusion criteria between 2008 and 2017. The median number of tumors at the initial GKSRS procedure was 20 (15–39) which accounted for 865 tumors in this study. The median aggregate tumor volume was 3.1cm<sup>3</sup>(0.13–13.26) and the median marginal dose was 16Gy (14–19Gy). **RESULTS:** The median of the mean brain dose was 2.58Gy (range 0.95–3.67Gy) and 79% of patients had a dose <3Gy. The 12Gy dose volume was a median of 12.45cm<sup>3</sup>, which was equivalent to 0.9% of the brain volume. The median percentage of brain receiving 5Gy and 3Gy was 6.7% and 20.4%, respectively. There was no correlation between the number of metastases and the mean dose to the brain ( $p=0.8$ ). A higher tumor volume was significantly associated with an increased mean brain dose ( $p<0.001$ ). The median of the mean dose to the bilateral hippocampi was 2.3Gy. Sixteen patients had supplementary GKSRS, resulting in an additional mean dose of 1.4Gy (0.2–3.8Gy) to the brain. **CONCLUSION:** GKSRS is a viable means of managing extensive brain metastases. This procedure provides a relatively low dose of radiation to the brain, especially when compared to traditional whole brain radiation protocols.

### 17. MELANOMA BRAIN METASTASIS: PRESENTATION, TREATMENT AND OUTCOMES IN THE AGE OF TARGETED- AND IMMUNO-THERAPIES

Evan D. Bander<sup>1,2</sup>, Melissa Yuan<sup>1</sup>, Joseph A. Carnevale<sup>1,2</sup>, Anne S. Reiner<sup>3</sup>, Katherine S. Panageas<sup>3</sup>, Michael A. Postow<sup>4,5</sup>, Viviane Tabar<sup>1</sup>, and Nelson S. Moss<sup>1</sup>; <sup>1</sup>Memorial Sloan Kettering Cancer Center, Department of Neurological Surgery, New York, NY, USA, <sup>2</sup>NewYork Presbyterian Hospital/Weill Cornell Medical College, Department of Neurological Surgery, New York, NY, USA, <sup>3</sup>Memorial Sloan Kettering Cancer Center, Department of Epidemiology & Biostatistics, New York, NY, USA, <sup>4</sup>Memorial Sloan Kettering Cancer Center, Department of Medicine, New York, USA, <sup>5</sup>Weill Cornell Medical College, Department of Medicine, New York, USA

**BACKGROUND:** Melanoma brain metastasis (MBM) prognosis has historically been dismal. However, breakthroughs in targeted and immunotherapies have improved long-term survival in advanced melanoma. As such, MBM presentation, prognosis and multimodality CNS-directed treatment use were reassessed in this contemporary age of treatment. **METHODS:** This retrospective study evaluated patients treated at Memorial Sloan Kettering Cancer Center between 2010–2019 with a diagnosis of melanoma brain metastases (MBM). Kaplan-Meier methodology was used to describe overall survival (OS). Recursive partitioning analysis (RPA) and time-dependent multivariable Cox modeling were used to assess prognostic variables and associate CNS-directed treatments with OS. **RESULTS:** Four hundred and twenty-five patients with 2,488 MBM were included. Median OS from MBM diagnosis was 8.9 months (95%CI: 7.9–11.3). RPA demonstrated significantly longer survival in patients diagnosed with MBM between 2015–2019 versus 2010–2014 (13.0 months [95%CI: 10.47–17.06]