

tases (BrM) or surgical cavities larger than >4 cc: 127 and 237 treated with surgery plus stereotactic radiosurgery (S+SRS) and SRS alone, respectively. We compared the 2 treatment arms using propensity score-matched (PSMA) and multivariate analyses (MVA). P values <0.05 were considered statistically significant. RESULTS: Median target volume was 6.6cc (4-36.9cc) for intact BrM and 15cc (4-54) for cavities. Median OS was 19 and 12 months for the S+SRS and SRS groups, respectively [HR 1.73 (1.35-2.22) (P<0.001)]. On UVA, number of BrM [HR 1.13 (1.06-1.22) (P<0.001)], ECOG 3-4 [HR 2.78 (1.73-4.46) (P<0.001)], and extracranial disease (ECD) at BrM treatment [HR 1.82 (1.37-2.40) (P<0.001)], correlated inversely with OS. GPA [HR 0.61 (0.52,0.70) (P<0.001)] and receipt of systemic therapy after BrM treatment [HR 0.58 (0.45-0.75) (P<0.001)] correlated to improved OS. On MVA, S+SRS [HR 1.81 (1.19,2.74) (P<0.0054)], reduced target volume [HR 1.03 (1.01,1.06) (P<0.0042)], and receipt of immune/targeted therapy [HR 0.68 (0.50,0.93) (P<0.015)] correlated with OS. PSMA comparing the treatment arms matched by ECD, number of BrM, ECOG, and SRS target volume, demonstrated that treatment arm remained correlated to OS [HR 1.62 (1.20-2.19) (P=0.0015)]. The cumulative incidence (CI) of LF requiring surgical resection at 12 months was 3% versus 7% for S+SRS and SRS groups, respectively [(HR 2.04 (0.89-4.69) (P=0.091)]. CI of PMD at 12 months was 16% versus 0% for S+SRS and SRS groups, respectively. CONCLUSION: Reduced SRS target volume, treatment with systemic therapy following BrM treatment, and surgical resection prior to SRS correlate with survival in patients with large BrM. PSMA supports the hypothesis that surgery prior to SRS improves survival in patients with large BrM.

MMAP-10

ADVERSE RADIATION EFFECT AFTER STEREOTACTIC RADIOSURGERY AND IMMUNOTHERAPY/TARGETED THERAPY FOR MELANOMA BRAIN METASTASES

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BACKGROUND: Safety of immunotherapy (IO) and targeted therapy (TT) with stereotactic radiosurgery (SRS) in melanoma brain metastases (MBM) treatment remains incompletely understood. We aim to identify whether timing of IO/TT in relation to SRS impacts rates of adverse radiation effect (ARE) in MBM. METHODS: Retrospective review of patients with MBM treated with SRS and IO/TT within three months prior and one year after SRS, from 2011-2021 at a single institution with at least two months MRI follow-up, identified 108 patients with 939 unique MBM meeting criteria. ARE was confirmed on independent imaging review. Concurrent IO/TT was defined as receiving IO/TT within 4 weeks before or after SRS. Data analysis was performed with the univariate cox proportional hazard model and Kaplan-Meier method. RESULTS: Median radiographic follow-up from time of SRS was 16months. IO/TT was initiated prior to SRS for 681 (72.5%) metastases and after SRS for 258 (27.5%) metastases. 837 (89.1%) metastases received concurrent IO/TT. Most common IO agents were ipilimumab (n=416), nivolumab (n=448), and pembrolizumab (n=203). Most common TT agents were dabrafenib (n=548), trametinib (n=540), and vemurafenib (n=81). 2-year local progression-free survival (PFS), distant intracranial PFS, and overall survival were 94.1%, 33.3%, and 55.2%, respectively. 55 (5.9%) metastases in 33 (30.6%) patients experienced ARE. Median time to ARE was 5mo (IQR 4-9mo). Of those who experienced ARE, 22 (66.7%) patients were symptomatic and treated with steroids; 12 (36.4%) patients underwent surgical intervention. ARE rates were not impacted by concurrent vs nonconcurrent IO/TT (5.5% vs 4.9%, p=0.34) nor IO/TT initiation pre vs post SRS (6.0% vs 5.4%, p=0.61). CONCLUSION: IO/TT in conjunction with SRS resulted in low ARE rates as compared to historical controls in the pre-IO/TT era. Timing of IO/TT in relation to SRS may not significantly impact ARE rates in MBM treatment.

MMAP-11

VOLUMETRIC STUDY OF BRAIN METASTASES IN EGFR-POSITIVE NSCLC TREATED WITH OSMERTINIB WITH OR WITHOUT CNS-DIRECTED RADIATION THERAPY

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BACKGROUND: In patients with brain metastases (BM) from EGFR-positive non-small cell lung cancer (NSCLC), recent data indicated that treating with CNS-penetrant tyrosine kinase inhibitors such as osimertinib

may enable deferring radiotherapy (RT) in select patients. The purpose of this study was to describe the radiographic response of newly diagnosed BM to osimertinib with or without stereotactic radiosurgery or whole brain radiotherapy, to identify parameters that may guide early versus delayed salvage RT. METHODS: In this single-institution retrospective study, 35 patients with 186 newly diagnosed BM started on osimertinib between 2014 and 2020 were reviewed. BM with tumor volume ≥ 0.1 cm³ were included in the volumetric analyses (N=106 BM). Survival was estimated with the Kaplan-Meier method, and univariable analysis was performed using log-rank tests. Cox proportional hazards was used for multivariable analyses for local control (LC), distant brain failure (DBF), and overall survival (OS). RESULTS: Of the 35 patients, 8 (23%) received osimertinib alone. Median follow-up was 29 months. The 1- and 2-year LC rates were 94% and 86%. The 1- and 2-year OS rates were 89% and 66%. Median time to DBF was 24 months. Patients treated with osimertinib and RT were more likely to have a significant radiographic volumetric response at early follow-up (4-12 weeks after treatment initiation) compared to osimertinib alone (median volumetric response of -80% vs. -41%, p=0.05). On per lesion analysis, early volumetric response of $\geq 80\%$ was associated with improved LC (3-year LC 98% vs 72%, p=0.04). CONCLUSIONS: The combination of osimertinib and CNS RT is associated with greater early volumetric response in patients with BM from EGFR-positive NSCLC compared to osimertinib alone. BM with significant initial radiographic response remain well-controlled in the long term. Patients whose BM demonstrate limited initial volumetric response may benefit from targeted RT to provide long term control.

FINAL CATEGORY: NEUROIMAGING

NEIM-01

INCIDENCE AND DIAGNOSTIC TECHNIQUES FOR LEPTOMENINGEAL DISEASE IN PATIENTS WITH BRAIN METASTASIS

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BACKGROUND: Leptomeningeal disease (LMD) is malignant infiltration of the pia mater and cerebrospinal fluid (CSF) space. LMD carries a poor prognosis with median survival of a few months. Annually, 110,000 patients in the United States are diagnosed with LMD. The incidence is rising because of improvements in control of primary cancers and recognizing that LMD is a late sequelae of some malignancies. Definitive diagnosis of LMD is made by CSF cytology and/or spine MRI, although neither tool shows robust sensitivity. The diagnostic challenges for LMD have led to a lack of uniformity in the diagnostic approach. METHODS: A systematic chart review of brain metastasis patients was conducted at Froedtert Hospital between 2019-2021. Information on primary cancer, LMD suspicion, work up, confirmation, treatment, and survival were collected and analyzed. RESULTS: Among 151 patients with brain metastasis, 86 were suspected and 29 were confirmed to have LMD. Of the confirmed patients, the most common primary cancers were lung (n=8, 27.6%) and breast (n=8, 27.6%). Most patients (n=24, 82.8%) underwent both LP and MRI. LMD was confirmed by positive cytology in a minority of cases (n=9, 31%), with most patients being confirmed by positive MRI or clinical findings alone (n=20, 69%). All LPs had over 10 mL of CSF sent to analysis. A median of 2 LPs were required before a positive cytology confirmed the diagnosis. Due to small sample size, no statistical analysis was made to correlate positive LP with primary cancer sites. CONCLUSION: Less than one third of cancer patients with confirmed LMD have positive cytology, despite the majority (>80%) of them undergoing LP. The dissonance between diagnostic strategies and confirmatory results is expected considering the low sensitivity of LPs; however, it highlights the need for more precise diagnostic tools, and development of a data-based strategy for LMD confirmation.

NEIM-02

DEVELOPMENT OF A DEEP LEARNING MODEL FOR DISCRIMINATING TRUE PROGRESSION FROM PSEUDOPROGRESSION IN GLIOBLASTOMA PATIENTS

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INTRODUCTION: Glioblastomas (GBMs) are highly aggressive tumors. Despite multimodal treatment, its median overall survival ranges between 16 and 20 months. The standard treatment regimen consists of surgical resection followed by concurrent chemoradiotherapy and adjuvant temozolomide. Despite temozolomide's effectiveness, it may cause the clinical challenge of