# 17P: Treatment Sequence for Non-small-cell Lung Cancer with Brain Oligometastases does not Impact Overall Survival



Icahn School of Medicine at Mount Sinai

Arvind Kumar<sup>1</sup>, Sangkavi Kuhan<sup>2</sup>, Alexandra L. Potter<sup>2</sup>, Camille Mathey-Andrews<sup>2</sup>, Hugh G. Auchincloss<sup>2</sup>, David E. Kozono<sup>3</sup>, Chi-Fu Jeffrey Yang<sup>2</sup> <sup>1</sup>Icahn School of Medicine at Mount Sinai, USA; <sup>2</sup>Massachusetts General Hospital, USA; <sup>3</sup>Brigham and Women's Hospital, USA

## INTRODUCTION

- Current treatment guidelines for oligometastatic Stage IV NSCLC recommend aggressive local treatment to the primary lung and metastatic site
- However, the optimal sequence of primary thoracic and metastatic treatment is not well-established

## **OBJECTIVES**

To evaluate long-term survival of patients with NSCLC with brain oligometastases who underwent initial brain metastatic treatment followed by primary thoracic treatment versus initial primary thoracic treatment followed by brain metastatic treatment

## **METHODS**

- **Data Source:** National Cancer Database (2010-2019)
- Inclusion Criteria: Patients with cT1-4, N0-3, M1b-c NSCLC with synchronous limited metastatic disease involving only the brain who underwent systemic chemotherapy and definitive (surgery or radiation) primary and metastatic treatment
- **Exclusion Criteria:** Previous unrelated malignancy, other distant metastasis, whole brain radiation therapy, palliative therapy
- **Definition of Comparison:** Upfront treatment of brain metastatic disease followed by local treatment to the lung primary site versus upfront local treatment to the lung primary site followed by treatment of brain metastatic disease
- Statistical Analysis: Kaplan-Meier analysis; Multivariable Cox proportional hazards modeling; Propensity score matching on 10 common prognostic variables (2:1 variable ratio with replacement, greedy nearest neighbor matching algorithm, caliper of 0.01)



# RESULTS

### **Baseline Characteristics:** Stratified by Treatment Sequence

|                    | Optront        | Upfront          |        |
|--------------------|----------------|------------------|--------|
|                    | Treatment      | Treatment        |        |
|                    | of Brain       | of Primary       |        |
|                    | Metastatic     | Thoracic         |        |
|                    | Disease        | Disease          |        |
|                    | (N=893)        | (N=237)          | Ρ      |
| Treatment of Brain |                |                  | <0.001 |
| Metastatic Disease |                |                  |        |
| Neurosurgery       | 71.7%          | 21.9%            |        |
| Brain SRS          | 28.3%          | 78.1%            |        |
| Thoracic Treatment |                |                  | <0.001 |
| Surgery            | 29.7%          | 46.8%            |        |
| Radiation          | 70.3%          | 50.2%            |        |
| Age, median (IQR)  | 61 (54, 67)    | 63 (57, 69)      | <0.001 |
| Female Sex         | 50.6%          | 51.9%            | 0.73   |
| Race               |                |                  | 0.70   |
| White              | 85.8%          | 87.8%            |        |
| Black              | 10.3%          | 8.4%             |        |
| Other              | 3.9%           | 3.8%             |        |
| Charlson-Deyo      |                |                  | 0.06   |
| Comorbidity Score  |                |                  |        |
| 0                  | 65.7%          | 69.2%            |        |
|                    | 21.3%          | 22.8%            |        |
| 2                  | 8.1%           | 3.0%             |        |
| 5+<br>Histology    | 4.9%           | 5.1%             | <0.001 |
| Adonasarsinama     |                | 67 10/           | <0.001 |
| Squamous           | /0.U%<br>1/20/ | 02.1%<br>22.70/  |        |
| Other              | 14.5%<br>0 00/ | 23.170<br>11 70/ |        |
| Tumor Location     | 9.0%           | 14.270           | 0.20   |
|                    | 17 10/         | 26 1%            | 0.20   |
| RMI                | 42.470<br>1 6% | Δ 2%             |        |
| RII                | 12 8%          | 18 7%            |        |
|                    | 29 5%          | 28.7%            |        |
|                    | 10.7%          | 12 0%            |        |
| Clinical T Status  | 10.770         | 12.070           | 0.02   |
| T1                 | 29.3%          | 21.5%            | 0.02   |
| T2                 | 37.1%          | 35.4%            |        |
| T3                 | 19.5%          | 23.2%            |        |
| T4                 | 14.1%          | 19.8%            |        |
| Clinical N Status  |                |                  | 0.06   |
| NO                 | 42.7%          | 39.7%            |        |
| N1                 | 16.1%          | 11.0%            |        |
| N2                 | 32.8%          | 37.6%            |        |
| N3                 | 18.4%          | 11.8%            |        |
|                    | -              |                  |        |

# **RESULTS (cont.)**

### Tumor and Treatment Characteristics: Stratified by Primary Thoracic Treatment Modality

|                           | Thoracic            | Thoracic |        |
|---------------------------|---------------------|----------|--------|
|                           | Radiation           | Surgery  |        |
|                           | (N=754)             | (N=376)  | Р      |
| Clinical T Status         |                     |          | <0.001 |
| T1                        | 22.9%               | 37.2%    |        |
| T2                        | 34.4%               | 41.5%    |        |
| Т3                        | 22.7%               | 15.4%    |        |
| Т4                        | 20.0%               | 5.9%     |        |
| Clinical N Status         |                     |          | <0.001 |
| NO                        | 27.3%               | 71.5%    |        |
| N1                        | 15.9%               | 13.3%    |        |
| N2                        | 43.4%               | 14.6%    |        |
| N3                        | 13.4%               | 0.5%     |        |
| Treatment of Brain        |                     |          | <0.001 |
| Metastatic Disease        |                     |          |        |
| Neurosurgery              | 65.4%               | 52.9%    |        |
| Brain SRS                 | 34.6%               | 47.1%    |        |
| Surgery Type              |                     |          | N/A    |
| Segmentectomy             |                     | 2.9%     |        |
| Lobectomy                 | N/A                 | 75.8%    |        |
| Pneumonectomy             |                     | 4.3%     |        |
| Other                     |                     | 9.8%     |        |
| <b>Radiation Modality</b> |                     |          |        |
| SBRT                      | 10.5%               | N/A      | N/A    |
| Dose (Gy), median (IQR)   | 60 (45 <i>,</i> 60) | N/A      | N/A    |
| Fractions, median (IQR)   | 30 (10, 32)         | N/A      | N/A    |

### **Overall Survival after Upfront Treatment of Brain Metastatic versus Primary Thoracic Disease:**





not impact overall survival.

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# **RESULTS (cont.)**

**Overall Survival after Upfront Treatment of Brain Metastatic versus Primary Thoracic Disease:** 

MASSACHUSETTS

GENERAL HOSPITAL

**THORACIC SURGERY** 

# **SUMMARY**

79.0% of patients with Stage IV M1b-c NSCLC with limited metastases involving only the brain underwent upfront treatment of brain metastatic disease followed by local treatment to the lung primary site

Upfront treatment of brain metastatic disease was more common in patients who underwent neurosurgical resection or thoracic radiation

In unadjusted and propensity score-matched analyses, upfront treatment of brain metastatic versus primary thoracic disease did not impact overall survival

# CONCLUSIONS

The findings of this study suggest that for patients presenting with NSCLC with synchronous limited metastatic disease isolated to the brain who can tolerate aggressive treatment of both the primary lung and brain metastatic sites, treatment sequence does