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Immune-Related Adverse Events (irAE) in Cancer Immune Checkpoint Inhibitors (ICI) and Survival Outcomes Correlation: To Rechallenge or Not?

Heidar J. Albandar, Jacob Fuqua, Jasim M. Albandar, Salahuddin Safi, Samuel A. Merrill and Patrick C. Ma

Table S1. Type of irAEs after Initial ICI Reinitiation.

| IrAE (n = 41) | n (%) |
|--------------------------------|--------|
| Hypothyroidism/hyperthyroidism | 9 (22) |
| Diarrhea/colitis | 8 (20) |
| Rash | 6 (15) |
| Pneumonitis | 5 (12) |
| Adrenal Insufficiency | 4 (10) |
| Hepatitis | 4 (10) |
| Nephritis | 1 (2) |
| Myalgia | 1 (2) |
| Other ** | 3 (7) |

^{**} Other irAE included arthralgia, MAHA, and SIADH.

Table S2. Multivariable Cox Proportion Analysis of Overall Survival Comparing irAEs with or without Rechallenge vs Non-irAE Groups.

| Effect | Categories | Hazard Ratio * | Lower CI | Upper CI | p |
|-------------------------|--|----------------|----------|----------|---------|
| Group | No immune-related adverse events | Ref | | | |
| | IrAE and not rechallenged (non-reinitiated + interruption) | 0.40 | 0.22 | 0.52 | 0.0024 |
| | IrAE and rechallenged (reinitiated +/- interruption) | 0.34 | 0.22 | 0.52 | <0.0001 |
| Age | ≤60Y | Ref | | | |
| | >60Y | 1.48 | 1.01 | 2.17 | 0.04 |
| Gender | Female | Ref | | | |
| | Male | 0.77 | 0.53 | 1.11 | 0.16 |
| Immunotherapy type | Pembro/Nivo | Ref | | | |
| | Ipi & Ipi/Nivo | 1.89 | 0.88 | 3.47 | 0.1 |
| | Atezo/Durva/Avelu | 0.98 | 0.47 | 2.04 | 0.95 |
| Antibiotic use | No | Ref | | | |
| | Yes | 0.76 | 0.49 | 1.20 | 0.24 |
| ECOG Performance status | 0–1 | Ref | | | |
| | >1 | 2.00 | 1.35 | 2.98 | 0.0006 |
| Tumor Location | Lung | Ref | | | |
| | Melanoma | 0.31 | 0.16 | 0.62 | 0.0008 |
| | Other | 0.84 | 0.55 | 1.28 | 0.41 |

^{*} Ref: Reference for hazard ratio analysis and comparison.

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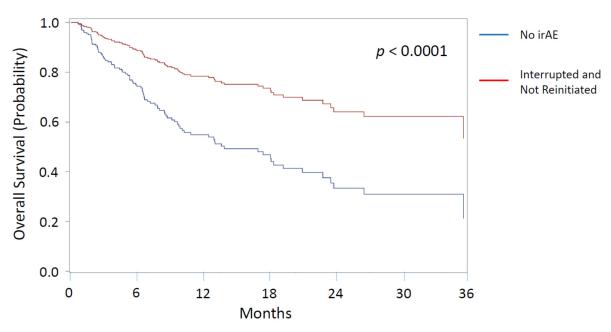


Figure S1. Kaplan-Meier Survival Analysis of Study Patients with No irAEs (blue) compared with those ICI "Interrupted and Not Reinitiated (i.e., Discontinued)" (red).

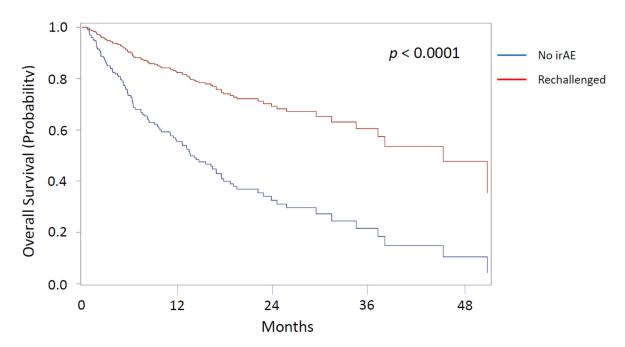


Figure S2. Kaplan-Meier Survival Analysis of the Study Patients with No irAEs (blue) compared with the "ICI Rechallenged" Cohort (ICI Reinitiated +/- Interruption) after irAEs.