

Age-related differences in inflammation may shed light on the limit of immunotherapies

December 6 2021

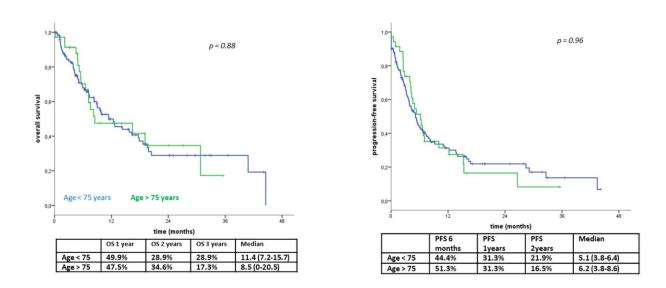


Figure 1. Kaplan–Meyer curves. (a) Overall survival (OS) and (b) progression-free survival (PFS) of metastatic non-small-cell lung cancer (mNSCLC) patients under treatment with nivolumab or atezolizumab divided into two age-related groups: age 75 years (green). In our series, we did not find any correlation with age for both overall survival (OS, patients \leq 74 years median OS 11.4 months, 95% CI 7.2–15.7 months vs. patients \leq 74 years median OS 8.5 months, 95% CI 0–20.5 months, p-value: 0.88) and progression-free survival (PFS, patients \leq 74 years median PFS 5.1 months, 95% CI 3.8–6.4 months vs. patients \geq 75 years median OS 6.2 months, 95% CI 3.8–8.6 months, p-value: 0.88). Credit: DOI: 10.3390/life11111235

Scientists at the Sbarro Institute for Cancer Research and Molecular



Medicine, Center of Biotechnology, located at Temple University's College of Science and Technology, working with collaborators in Italy, have described the age-related differences in inflammatory response to an emerging immunotherapy treatment for non-small-cell lung cancer (NSCLC) called PD-1 blockade. These age-related changes to immune system response, known as immunosenescence, may have implications for the efficacy or the safety of immunotherapies for a variety of conditions and disease.

The paper, titled "Distinctive Role of the Systemic Inflammatory Profile in Non-Small-Cell Lung Cancer: Younger and Elderly Patients Treated with a PD-1 Immune Checkpoint Blockade: A Real-World Retrospective Multi-Institutional Analysis," was published recently in *Life*.

The authors write, "It has been speculated that a chronic status of systemic inflammation as well as the immunosenescence physiologically occurring in <u>elderly patients</u> may affect the efficacy of the treatment and the occurrence of irAEs. We performed a multi-institutional retrospective study aimed at evaluating the effects of these mAbs in 117 mNSCLC patients younger (90 cases) and older (27 cases) than 75 years in correlation with multiple inflammatory parameters (NLR, CRP, ESR, LDH and PCT). No differences were observed when the cohorts were compared in terms of the frequency of PFS, OS, inflammatory markers and immune-related adverse events (irAEs). Similarly, the occurrence of irAEs was strictly correlated with a prolonged OS survival in both groups.

"On the contrary," the authors continue, "a negative correlation between the high baseline levels of inflammatory markers and OS could be demonstrated in the younger cohort only. Overall, PD-1/PD-L1-blocking mAbs were equally effective in young and elderly mNSCLC patients; however, the detrimental influence of a systemic inflammation at the



baseline was only observed in young patients, suggesting different agingrelated inflammation immunoregulative effects."

More information: Valerio Nardone et al, Distinctive Role of the Systemic Inflammatory Profile in Non-Small-Cell Lung Cancer Younger and Elderly Patients Treated with a PD-1 Immune Checkpoint Blockade: A Real-World Retrospective Multi-Institutional Analysis, *Life* (2021). DOI: 10.3390/life11111235

Provided by Sbarro Health Research Organization

Citation: Age-related differences in inflammation may shed light on the limit of immunotherapies (2021, December 6) retrieved 4 May 2024 from https://medicalxpress.com/news/2021-12-age-related-differences-inflammation-limit-immunotherapies.html

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