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Influence of treatment-related lymphopenia on the efficacy of immune checkpoint inhibitors in lung cancer: a meta-analysis

Ye Zhang, Cheng Huang and Shangqing Li*

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Background: Treatment-related lymphopenia (TRL) is common in patients with lung cancer, particularly in those with radiotherapy. However, the influence of TRL on the efficacy of immune checkpoint inhibitors (ICIs) for patients with lung cancer remains poorly understood. We performed a systematic review and meta-analysis to investigate the influence of TRL on survival of lung cancer patients on ICIs.

Methods: In order to accomplish the aim of the meta-analysis, a comprehensive search was conducted on databases including PubMed, Embase, Cochrane Library, and the Web of Science to identify observational studies with longitudinal follow-up. The Cochrane Q test was employed to evaluate heterogeneity among the included studies, while the I^2 statistic was estimated. Random-effects models were utilized to merge the results, considering the potential impact of heterogeneity.

Results: Ten cohort studies with 1130 lung cancer patients who were treated with ICIs were included. Among them, 427 (37.8%) had TRL. Pooled results showed that compared to patients without TRL, patients with TRL were associated with poor progression-free survival (hazard ratio [HR]: 2.05, 95% confidence interval [CI]: 1.62 to 2.60, $p < 0.001$; $I^2 = 2230$) and overall survival (HR: 2.69, 95% CI: 2.10 to 3.43, $p < 0.001$; $I^2 = 0\%$). Sensitivity analysis limited to patients with non-small cell lung cancer showed similar results (HR: 2.66 and 2.62, both $p < 0.05$). Moreover, subgroup analyses according to the diagnostic criteria of TRL, regression analysis model (univariate or multivariate), and indications of ICIs (for locally advanced or advanced lung cancer) showed consistent results (p for subgroup difference all > 0.05).

Conclusion: TRL was associated with poor survival of lung cancer patients who were treated with ICIs.

KEYWORDS

lung cancer, lymphopenia, immunotherapy, survival, meta-analysis

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Table of Contents Lymphopenia Treatment Manual Guide

1. Understanding the eBook Lymphopenia Treatment Manual Guide
 - The Rise of Digital Reading Lymphopenia Treatment Manual Guide
 - Advantages of eBooks Over Traditional Books
2. Identifying Lymphopenia Treatment Manual Guide
 - Exploring Different Genres
 - Considering Fiction vs. Non-Fiction
 - Determining Your Reading Goals
3. Choosing the Right eBook Platform
 - Popular eBook Platforms
 - Features to Look for in an Lymphopenia Treatment Manual Guide
 - User-Friendly Interface
4. Exploring eBook Recommendations from Lymphopenia Treatment Manual Guide
 - Personalized Recommendations

- Lymphopenia Treatment Manual Guide User Reviews and Ratings
- Lymphopenia Treatment Manual Guide and Bestseller Lists
- 5. Accessing Lymphopenia Treatment Manual Guide Free and Paid eBooks
 - Lymphopenia Treatment Manual Guide Public Domain eBooks
 - Lymphopenia Treatment Manual Guide eBook Subscription Services
 - Lymphopenia Treatment Manual Guide Budget-Friendly Options
- 6. Navigating Lymphopenia Treatment Manual Guide eBook Formats
 - ePub, PDF, MOBI, and More
 - Lymphopenia Treatment Manual Guide Compatibility with Devices
 - Lymphopenia Treatment Manual Guide Enhanced eBook Features
- 7. Enhancing Your Reading Experience
 - Adjustable Fonts and Text Sizes of Lymphopenia Treatment Manual Guide
 - Highlighting and Note-Taking Lymphopenia Treatment Manual Guide
 - Interactive Elements Lymphopenia Treatment Manual Guide
- 8. Staying Engaged with Lymphopenia Treatment Manual Guide
 - Joining Online Reading Communities
 - Participating in Virtual Book Clubs
 - Following Authors and Publishers Lymphopenia Treatment Manual Guide
- 9. Balancing eBooks and Physical Books Lymphopenia Treatment Manual Guide
 - Benefits of a Digital Library
 - Creating a Diverse Reading Collection Lymphopenia Treatment Manual Guide
- 10. Overcoming Reading Challenges
 - Dealing with Digital Eye Strain
 - Minimizing Distractions
 - Managing Screen Time
- 11. Cultivating a Reading Routine Lymphopenia Treatment Manual Guide
 - Setting Reading Goals Lymphopenia Treatment Manual Guide
 - Carving Out Dedicated Reading Time
- 12. Sourcing Reliable Information of Lymphopenia Treatment Manual Guide
 - Fact-Checking eBook Content of Lymphopenia Treatment Manual Guide

- Distinguishing Credible Sources
- 13. Promoting Lifelong Learning
 - Utilizing eBooks for Skill Development
 - Exploring Educational eBooks
- 14. Embracing eBook Trends
 - Integration of Multimedia Elements
 - Interactive and Gamified eBooks

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