

CYTOTOXIC TREATMENT OF METASTATIC BREAST CANCER

Which drugs and drug combinations to use?

PER PYLIFTER, SØREN COLD and CARSTEN ROSE

Although the optimal efficacy of cytotoxic therapy plateaued in the 1970s, no consensus yet exists regarding which cytotoxic combination to offer as first-line therapy for metastatic breast cancer. Comparison of a combination of CAF/CEF with other cytotoxic combinations reveals that an anthracycline-containing regimen not only increases response rate, but also improves time to progression and survival. Regarding appropriate duration of cytotoxic therapy, randomized trials indicate that therapy in excess of 6 months is beneficial. Although the main objective of cytotoxic therapy in patients with metastatic disease is to palliate symptoms at the least toxic cost, the finding that adjuvant cytotoxic therapy improves survival provides a clinical and ethical rationale for continued research into new drugs and combinations in the hope that new strategies can be employed not only against metastatic breast cancer but may be applied with benefit also in the adjuvant situation.

Key words: Metastatic breast cancer, cytotoxic therapy, survival, duration of therapy.

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As the incidence of breast cancer continues to rise, it is encouraging that adjuvant therapy of the primary disease improves survival in both pre- and postmenopausal patients (1, 2). Among women less than 50 years of age, cytotoxic treatment improves the 10-year survival by about 10%.

Although efficacy of cytotoxic therapy has reached a plateau in the 1970s with response rates of 60% and median times to progression (TTP) of 8-12 months, no consensus yet exists regarding which cytotoxic combination to offer as first-line therapy for metastatic disease. This uncertainty arises from the fact that some investigators have found a survival prolongation with cytotoxic therapy (3, 4), while others have stated that the survival

advantage for metastatic breast cancer patients undergoing such therapy is at best minimal (5, 6, 7).

In the present review, we summarize the aims of cytotoxic therapy in patients with metastatic breast cancer and discuss briefly the most effective drugs and drug combinations. Finally, we address the clinical trials investigating optimal duration of cytotoxic therapy.

Aim of cytotoxic therapy in metastatic breast cancer

Although most patients experience temporary relief of symptoms, only a minority survive more than 10 years after diagnosis of metastatic disease. The main objective of cytotoxic therapy in these patients is thus to palliate symptoms at the least toxic cost. Furthermore, the finding that adjuvant cytotoxic therapy improves survival provides a clinical and ethical basis, for continued research on new active drugs and combinations, in the hope that new strategies can be employed not only against metastatic breast cancer, but possibly be applied with benefit also in the adjuvant situation.

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Correspondence to: Dr Carsten Rose, Department of Oncology B, Odense University Hospital DK-5000 Odense C, Denmark.
Address: As above (all authors).

Cytotoxic Drugs In The Treatment Of Cancer

JS Bruner



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Table of Contents Cytotoxic Drugs In The Treatment Of Cancer

1. Understanding the eBook Cytotoxic Drugs In The Treatment Of Cancer
 - The Rise of Digital Reading Cytotoxic Drugs In The Treatment Of Cancer
 - Advantages of eBooks Over Traditional Books
2. Identifying Cytotoxic Drugs In The Treatment Of Cancer
 - 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 Cytotoxic Drugs In The Treatment Of Cancer
 - User-Friendly Interface
4. Exploring eBook Recommendations from Cytotoxic Drugs In The Treatment Of Cancer
 - Personalized Recommendations
 - Cytotoxic Drugs In The Treatment Of Cancer User Reviews and Ratings
 - Cytotoxic Drugs In The Treatment Of Cancer and Bestseller Lists
5. Accessing Cytotoxic Drugs In The Treatment Of Cancer Free and Paid eBooks
 - Cytotoxic Drugs In The Treatment Of Cancer Public Domain eBooks
 - Cytotoxic Drugs In The Treatment Of Cancer eBook Subscription Services

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