# Cognitive functioning during long-term tamoxifen treatment in postmenopausal women with breast cancer

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#### Abstract

Objective: Endocrine therapy is widely used—often for many years—in women with breast cancer. Yet little is known about cognitive functioning after long-term use of tamoxifen. We examine cognitive sequelae, approximately 3 years after diagnosis, in postmenopausal women with breast cancer who were treated with adjuvant tamoxifen.

Methods: Data from participants who underwent surgical operation with or without radiotherapy, participants who received adjuvant tamoxifen, and healthy controls were collected. Neuropsychological tests were administered, and participants completed questionnaires on health-related quality of life (Quality of Life Questionnaire Core 30 and Breast Cancer-Specific Quality-of-Life Questionnaire), menopausal symptoms (Functional Assessment of Cancer Therapy—Breast endocrine symptom subscale), and anxiety and depression (Hopkins Symptom Checklist).

Results: In total, 107 women participated (adjuvant tamoxifen group, n=20; surgical operation/radiotherapy group, n=43; healthy control group, n=44). Women in the adjuvant tamoxifen group had received tamoxifen for a mean (SD) of 31.5 (18.6) months (range, 15-79 mo) and performed worse on verbal memory than the surgical operation/radiotherapy group (P < 0.05) and the healthy control group (P < 0.05). Participants in the adjuvant tamoxifen group performed worse on measures of fluency than healthy controls (P < 0.05). Furthermore, women in the adjuvant tamoxifen group reported worse cognitive functioning (P < 0.05) than women in the surgical operation/radiotherapy group or the healthy control group.

Conclusions: Our results provide insights into cognitive functioning in women who receive long-term adjuvant tamoxifen treatment. By adding the surgical operation/radiotherapy group, we could control for the mental and physical influences of the diagnosis and treatment of breast cancer. Cognitive domains that rely on verbal abilities (verbal memory and fluency) seem to be at risk for deterioration after treatment with tamoxifen.

Key Words: Tamoxifen - Cognition - Postmenopausal - Breast cancer.

White increased survival, cognitive functioning after breast cancer therapy becomes of greater importance to the well-being of women. In postmenopausal women, approximately 75% of breast tumors are hormonesensitive. In this group of women, endocrine therapy is widely used and often prescribed for many years.

Endocrine therapy aims to inhibit the growth of estrogen receptor (ER)-positive tumors by inhibiting or blocking estrogen. This can be achieved by administering either selective ER modulators (SERMs), which allow selective inhibition or stimulation of estrogen-like action in various tissues, or aromatase inhibitors (Als), which block the synthesis of estrogen, thus lowering estrogen levels. These endocrine therapies may be provided in a switch schedule (initially SERMs for a number of years followed by an AI up to a total of 5-7 y of endocrine therapy).2.3 Common adverse effects of endocrine therapy, such as hot flashes and mood disturbances, are related to estrogen deprivation.4 Increasing evidence suggests that estrogens play a role in cognitive functioning.7 Although the mechanisms of action of estrogens on the brain are not entirely understood, neuroscientific studies indicate that estrogens exert neurotrophic and neuroprotective actions." Estropens seem to favor differentiation and survival by acting through ERs,7 but estrogens also act through membrane receptors to influence the functioning of the brain." These ERs are present in structures that are important for cognitive functioning, such as the frontal lobe and the hippocampus." Therefore, it is biologically plausible that endocrine therapy could affect cognitive functioning. Moreover, because estrogen is known to interact with the cholinergic system, 10 it seems probable that cognitive functions such as attention, verbal learning, and memory may be particularly vulnerable.

Although much is already known about the cognitive effects of chemotherapy,11 many women with breast cancer receive

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# **Long Term Tamoxifen Treatment For Breast Cancer**

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#### **Long Term Tamoxifen Treatment For Breast Cancer:**

**Long-term Tamoxifen Treatment for Breast Cancer** Virgil Craig Jordan, 1994 During the past twenty years tamoxifen has become the most widely prescribed and most successful drug used in the treatment of breast cancer In this volume editor V Craig Jordan provides articles that trace the development pharmacology and clinical research surrounding this drug which by the year 2000 could be used to treat as many as one million women annually Drawing from research conducted by specialists in the United States the United Kingdom and Italy the series of articles describes the clinical testing of tamoxifen highlighting the benefits Studies show that tamoxifen lowers cholesterol and can potentially protect women against osteoporosis and fatal coronary heart disease Equally important is a discussion of side effects and possible drug interactions and how these issues relate to patient concerns An investigation of the development of a new class of drugs for use after tamoxifen fails provides valuable insight into future treatments as the contributers consider possible resistance to tamoxifen This volume provides invaluable information for physicians and surgeons who care for patients with breast cancer and for women interested in exploring this therapeutic dimension Tamoxifen and Breast Cancer Michael W. DeGregorio, Valerie J. Wiebe, 1999-01-01 A discussion of the diagnosis of breast cancer and the risks benefits and limitations of treatment alternatives particularly tamoxifen This edition contains information on developments in the use of tamoxifen especially in the results of the Breast Cancer Prevention Trial Journal of the National Cancer Institute ,2005

Treatment of the Postmenopausal Woman Rogerio A. Lobo, 2007-06-05 For anyone who treats postmenopausal women this latest edition of Rogerio Lobo s classic work combines the best from two well known references Menopause and the second edition of Treatment of the Postmenopausal Woman By adding significant discussions of the basic science behind menopause it is possible to objectively assess the clinical value and limitations of current approaches to treatment and provide a basis and rationale for strategies that will result in better individualized and specialized care Not only does the third edition discuss diagnosis and treatment of menopause but it covers biological anatomical physiological pathobiological and pharmacological aspects as well bringing together in one source all of the information needed to understand and treat postmenopausal conditions Over 50% new material representing the vast amount of information available since the Women s Health Initiative WHI clinical trials were completed making this the most up to date reference on postmenopausal women Includes several new sections on comparisons between clinical trials and observational data urology and pelvic support Each section is preceded by a preface to put the area into context with many chapters having suggested treatment regimens

**Tamoxifen Tales** V. Craig Jordan,2022-01-12 Tamoxifen Tales Suggestions for Scientific Survival presents a case study describing the academic journey of teams behind major advances in medical sciences highlighting lessons learned that are applicable to the next generation of scientists This book provides a manual on the successful mentoring of young scientists including stories describing how training experience shaped careers to become leaders in academia and the pharmaceutical

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Holland-Frei Cancer Medicine 8 James F. Holland,2010 Holland Frei Cancer Medicine serves as a quick reference to current information on an extensive list of cancers including breast lung thyroid colorectal ovarian prostate and gastric cancer to name but a few Presented as an accessible pocket sized handbook the chapters are organized in an outline format offering only the most essential information on the etiology staging including TNM staging and treatment for each cancer type Individual chapters are devoted to the molecular biology of cancer cancer prevention cancer screening the mechanisms of chemotherapy and diagnostic imaging in cancer Additionally each chapter lists all the major phase III clinical trials and therefore serves as an excellent reference of the major randomized controlled trials for each cancer reported to date Specific chapters are also dedicated to the discussion of oncologic emergencies pain and palliation and prescription complications At the conclusion of the book a glossary of oncologic terms and chemotherapeutic drug programs a table of common cancer incidences and an overview of the mechanisms common uses and related toxicities of various anti cancer agents are featured In addition performance status tables mathematical formulas and a listing of common biomedical cancer web sites are highlighted

Clinical Ocular Toxicology E-Book Frederick T. Fraunfelder, Frederick W. Fraunfelder, Wiley A. Chambers, 2008-06-12 Written by international authorities in ocular toxicology including the Founder of The National Registry of Drug Induced Side Effects and its current Director this essential resource provides the clinically relevant information you need to effectively diagnose and manage herbal chemical and drug related ocular problems Comprehensive coverage of all drugs generic and trade drug names primary uses ocular and systemic side effects and clinical significance make this book like its best selling predecessor Drug Induced Ocular Side Effects the ideal reference for quick on the spot consultation Leaders in the field provide need to know information on all aspects of ocular toxicology all in one concise reference Data from the National Registry of Drug Induced Ocular Side Effects Casey Eye Institute Portland OR and the World Health Organization Uppsala Sweden help you recognize and avoid drug induced ocular side effects A highly templated format makes retrieval of essential knowledge quick and easy A wealth of full color photographs provide vivid visual diagnostic guidance The latest information on approved medications helps you stay up to date and provide state of the art care Extensive coverage of principles of therapy ocular drug delivery methods to evaluate drug induced visual side effects and the role of electrophysiology and psychophysics gives you the knowledge you need to manage any challenge in ocular toxicology Authoritative guidance on ocular drugs and their use in pregnancy helps you safely manage the unique needs of these patients The inclusion of the WHO classification system helps you determine whether a particular side effect is certain probable or likely to occur

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