Evaluation of the management of uterine sarcoma

Thesis

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Dedication

To my Mother and Father

for encouragement, support, understanding and love.

To My Sister and my Brothers

for supporting, understanding and giving me the help to finish this work

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LIST OF ABBREVIATIONS

CA 125 Cancer Antigens 125

CD 10 Cluster Designation 10

CD 44 ver 3 Cluster Designation 44 version 3

CS Carcinosarcoma

CT Computed Tomography

DFS Disease Free Survival

DNA Deoxyribonucleic acid

D&C Dilatation and Curettage

EBRT External Beam Radiotherapy

ER Estrogen Receptor

ESS Endometrial Stromal Sarcoma

FBC Full Blood Count

FIGO Federation of International Gynecologic Oncology

GOG Gynecologic Oncology Group

HDR High Dose Radiotherapy

HGSS High Grade Stromal Sarcoma

HPF High Power Field

IMA Inferior Mesenteric Artery

IVC Inferior Vena Cava

LDR Low Dose Radiotherapy

LGSS Low Grade Stromal Sarcoma

LMS Leiomyosarcoma

LND Lymph Node Dissection

MMMT Malignant Mixed Müllerian Tumour

MRI Magnetic Resonance Imaging

NCI National Cancer Institute

OS Overall Survival

PLND Paraaortic Lymph Node Dissection

PR Progesterone Receptor

PTEN Piedmont Triad Entrepreneurial Network

RR Response Rate

SD Standard Deviation

TAH & BSO Total Abdominal Hysterectomy and Bilateral Salpingioophorectomy

VAC Vincristine, Actinomycin D, Cyclophosphamid

WHO World Health Organization

Abstract

Uterine sarcomas include a heterogeneous group of rare tumours that usually have an aggressive clinical behavior and poor prognosis. Clinical staging, histologic cell types, grades, mitotic index and DNA ploidy are important prognostic factors predicting the overall and disease free survival. This retrospective study highlights the experience of management and surgical approaches offered uterine sarcoma in National Cancer Institute, Cairo University during the period January 2000- December 2007. The study included fifty seven patients with pathologically proven uterine sarcoma and subjected to different types of management at our institute. In our study FIGO staging is the most significant prognostic affecting both overall and disease free survival. In our study, external beam radiotherapy with or without brachytherapy seemed improved local disease control with better results in early stage. Despite using adjuvant treatment survival rates were very poor with 2- year survival 37.6% and 4-year survival 22.2%.

Key words: uterine sarcoma, surgical approaches, FIGO staging, external beam radiotherapy, brachytherapy, overall survival, disease free survival.



INTRODUCTION

Introduction

Uterine sarcoma represents a small percentage of all uterine malignancies and about 1% of all female genital tract neoplasms (*Tavassoli and Devilee*, *2003*). It represents (4–9%) of all invasive uterine cancers, and has an annual incidence

rate of less than 2 per 100,000 females (Brooks et al., 2004).

Survival rates for uterine sarcoma patients have been uniformly poor with most series have reported a 5-year survival of 30-48% (*Livi et al.*, 2004)

In general, two distinct tissue components give rise to malignant mesenchymal tumors; myometrial smooth muscle is the tissue origin for leiomyosarcoma (LMS) and endometrial stroma for stromal sarcoma (ESS), whilst both muscle and stromal tissue types give rise to malignant mixed müllerian (MMMT) sarcoma. In addition, uterine sarcoma is sub classified into homologous (consisting of cells native to the uterus) or heterogonous (cells usually not found in the uterus). The mixed Müllerian tumour consists of malignant elements of both epithelial and stromal elements, hence it is also known as carcinosarcoma. (*Papadopoulos and Kenney, 2001*).

There are important differences in the ways these tumours grow and metastasize. Carcinosarcoma tends to have a higher incidence of lymphatic spread and lymph node metastases, whereas leiomyosarcoma is more likely to have early hematogenous spread usually in the lungs or the liver (*Gadducci et al.*, 2002).

However, the metastatic potential is very wide and distant lesions can be found anywhere (*Falconi et al.*, 2006).

Clinical staging, histologic cell types, grades, mitotic index and DNA ploidy are important prognostic factors predicting the overall and disease free survival (*Giuntoli et al.*, 2003).

The most significant prognostic factor is the stage at diagnosis .The FIGO staging for uterine sarcoma is identical to that for the corpus uteri, with two-thirds being stage I and one-fifth stage IV at diagnosis. (*Bodner et al.*, 2003)

In leiomyosarcoma, the size of the tumour affects prognosis, where lesions greater than 5 cm have a poorer outcome. One study demonstrated that leiomyosarcoma has a poorer prognosis than mixed müllerian tumours when adjusted for other known prognostic factors. (*Olah et al.*, 1992)

In early stage disease of mixed müllerian tumours some investigators have demonstrated that the following features influence the prognosis; tumour size, cervical spread, depth of myometrial invasion and lymph nodal disease. (*Major et al.*, 1993)

Proper Surgery should involve a minimum of total abdominal hysterectomy, bilateral salpingo-oophorectomy, careful and thorough inspection of the abdominal and pelvic organs, sampling of fluid for cytological assessment of the washings and perhaps -little more contentiously- pelvic and/or para-aortic lymph node dissection (PLND) (*Amant*, 2005).

The poor disease control is in itself an indication for adjuvant therapy. In most cases, radiotherapy improves local control but not survival. (*Livi et al.*, 2004)

On the basis of the results of a recent clinical trial, there is no difference in terms of overall survival or disease free survival for sarcomas, but radiotherapy seems to provide a superior local disease control for mixed müllerian tumours and no benefit with respect to LMS (*Reed et al.*, 2003).

In recurrent uterine sarcomas the response rates in different chemotherapeutic regimens have been between 0–57 percent (*Kanjeekal et al.*, 2005).



REVIEW OF LITERATURE