HIGHLIGHT ON GASTROINTESTINAL TUMORS

Essay

Submitted for Partial Fulfillment of Master Degree in Tropical Medicine

By

Mary Michel Zekry Iskandarous (M.B., B.CH.)

Under Supervision of **Professor Dr. Wahid Abd El Monsef**

Professor of Tropical Medicine Faculty of Medicine, Ain Shams University

Dr. Iman Fawzy Montasser

Lecturer of Tropical Medicine Faculty of Medicine, Ain Shams University

Faculty of Medicine
Ain Shams University
2016





First and foremost, thanks to "ALLAH" for granting me the power accomplish this work;

I would like to express my endless gratitude and appreciation to **Professor Dr. Wahid Abd El Monsef** Professor of Tropical Medicine Faculty of Medicine, Ain Shams University for giving me the honour of working under his supervision and providing me with a lot of encouragement and support.

My deepest gratitude and appreciation are to **Dr. Iman Fawzy Montasser,** Lecturer of Tropical Medicine Faculty of Medicine, Ain Shams University for her acceptance to supervise my work for her continuous support, her valuable advice and encouragement.

Special thanks and gratitude to all my family for their kind guidance and great help during the preparation of the work.

Mary Michel Zekry Iskandarous

CONTENTS

Subject	Page No.
Contents	I
List of Tables	ii
List of Figures	iii
List of Abbreviations	iv
Introduction	1
Aim of the essay	4
Upper Gastrointestinal Malignancy	5
Esophageal tumors	7
Gastric Tumors	31
Lower Gastrointestinal Malignancy	60
Small Bowel Tumors	60
Polyps of the Colon	83
Colorectal cancer	102
Anal Cancer	125
Hepato-biliary pancreatic Malignancy	138
Liver and intrahepatic bile_ducts tumors	138
Biliary Tract Cancer	167
Pancreatic cancer	192
Summary and Conclusion	210
References	211
اللخص العربي	1
	

LIST OF TABLES

Table No.	Title	Page No.
Table (1): TN	NM staging for Esophageal Cancer	17
	FNM staging of gastric cancer (7th Editi	
` '	ummary of methods for evaluation of small- nors	
` '	TNM classification of small into	
Table (5): Tu	mor stages of small intestine adenocarcinoma	ı79
Table (6): Feat	tures of colorectal polyps.	84
Table (7): Pat	thologic staging systems for colorectal cancer	115
Table (8): TN	NM classification of Colorectal cancer	116
	TNM Classification of Anal Carcinoma (Exclanoma, Carcinoid, and Sarcoma)	
	Staging and TNM Classification of Anal Carc	
Table (11): T	The BCLC staging system and treatment strate	gy152
Table (12): A	AJCC 7th Edition Staging for ICC	154
	AJCC 7th Edition Staging System for Gallb enocarcinoma	
Edi	Jarnagin–Blumgart Classification and AJC tion Staging System for Pe plangiocarcinoma	erihilar
	AJCC 7th Edition Staging System for blangiocarcinoma	
Table (16): A	AJCC cancer staging for ampulla cancers	186
Table (17): A	AJCC TNM staging of Pancreatic cancer	200
Table (18): U	JICC staging and survival of PC	201

LIST OF FIGURES

Figu	re No.	Title	Page No.
Fig(1)	Layers of esophagus an	d stages of esophageal	cancer18
Fig(2)	Algorithm for Therapy	of EC	19
Fig(3)	Algorithm for the manag	gement of gastric cancer.	43
Fig(4)	Diagnostic approach to	small bowel tumors	70
Fig(5)	Strategy and treatment cancer	C	
Fig(6):	The BCLC staging and trea	tment strategy updated in	2011155
Fig(7):	Flow chart showing of GBA	•	· ·
Fig(8)	Flowchart showing c extrahepatic cholangioc	•	•

5-FU : 5-fluorouracil 90Y : yttrium-90

ACs : adenocarcinomas

AFAP : Attenuated Familial adenomatous polyposis

AFI : Autofluorescence imaging

AFP : alpha-fetoprotein

AJCC : American Joint Committee on Cancer

ALT : alanine aminotransferase **AP** : alkaline phosphatase

APC : adenomatous polyposis coli
 APE : abdominoperineal excision
 AST : aspartate aminotransferase
 BCLC : Barcelona Clinic Liver Cancer

BMI : body mass index

BRPC : "borderline resectable pancreatic cancer"BRRS : Bannayan–Riley–Ruvalcaba syndrome

BTC : Biliary tract cancer

Ca19-9 : carbohydrate antigen 19-9

CCA : cholangiocarcinomaCE : Capsule endoscopy

CEA : carcinoembryonic antigen
CEUS : Contrast-enhanced ultrasound

CIN : chromosomal instability

CLE : Confocal laser endomicroscopyCMT : combined modality therapy

CRT : ChemoradiotherapyCTC : CT-colonographyCTx : chemotherapy

dCC : distal cholangiocarcinoma

DEB-TACE : drug-eluding bead transcatheter arterial

chemoembolization

dMMR : defective mismatch repairDRE : digital rectal examination

EASL : European Association for the Study of the Liver

EATL : Enteropathy associated T cell intestinal

lymphoma

EC : Esophageal cancer

ECF : epirubicin, cisplatin, and fluorouracil

ECOG-PS: performance status according to Eastern

Cooperative Group

ECX : epirubicin, cisplatin, and capecitabine

EEC : Early esophageal cancer
 EGD : Esophagogastroduodenoscopy
 EGFR : epidermal growth factor receptor
 EMR : Endoscopic mucosal resection
 EMR : endoscopic mucosal resection
 EOX : epirubicin/oxaliplatin/capecitabine

ERCP : endoscopic retrograde cholangiopancreatography

ESCC : esophageal squamous cell carcinoma ESD : endoscopic submucosal dissection

ESMO : the European Society for Medical Oncology

EUA : examination under anaesthesia
 EUS-FNA : EUS guided fine needle aspiration
 FAP : Familial adenomatous polyposis
 FDA : food and drug administration

FIT : fecal immunohistochemical testing

FOBT : fecal occult blood testing s **FS** : flexible sigmoidoscopy

GC : Gastric cancer

GEJ : Gastroesophageal junction

GGT : gamma-glutamyl-transpeptidase

GI : Gastrointestinal

GISTs : gastrointestinal stromal tumors

GIT : gastrointestinal tract
H. pylori : Helicobacter pylori
HBV : Hepatitis B Virus

HCC : Hepatocellular carcinoma

HCT : haematopoietic cell transplantation

HCV : Hepatitis C Virus

HER-2 : Human epidermal growth factor receptor 2 **HIPEC** : hyperthermic intraperitoneal chemotherapy

HIV : human immunodeficiency virus

HL : Hepaticolithiasis

HMPS : Hereditary mixed polyposis syndromeHNPCC : hereditary nonpolyposis colorectal cancer

HPV : human papillomavirus

IBD : Inflammatory Bowel diseaseICC : Intrahepatic cholangiocarcinoma

IHC : immunohistochemistry

IMRT : intensity modulated radiation therapy

IPSID : Immunoproliferative small intestinal disease

IVC : inferior vena cava

JCCRC : Japanese Classification of Colorectal

Carcinoma

JGCA : The Japanese Gastric Cancer Association

JPS : Juvenile polyposis syndrome

LCSGJ : the Liver Cancer Study Group of Japan

LVLs : Lugol-voiding lesions

MAGIC : Medical Research Council Adjuvant Gastric

Infusional Chemotherapy

MALT : mucosa-associated lymphoid tissue,

MAP : MUTYH-associated polyposis

MDCT : multidetector CT

MEN-1 : Multiple endocrine neoplasia type 1MIE : Minimally invasive esophagectomy

MMC : mitomycin C

MOSAIC : The Multicenter International Study of

Oxaliplatin/5-Fluorouracil/Leucovorin in the

adjuvant Treatment of Colon Cancer

MRCP : magnetic resonance cholangiopancreatography

MRI : magnetic resonance imaging
 MSI : microsatellite instability
 MSM : men who have sex with men

NAFLD : nonalcoholic fatty liver disease

NASH : nonalcoholic steatohepatitis

NBI : Narrow band imaging

NCCN : The National Comprehensive Cancer

Network

NET : Neuroendocrine Tumor

NICE : National Institute for Health and Clinical

Excellence

OLT : orthotopic liver transplantation

OS : overall survival

PanIN: pancreatic intraepithelial neoplasia

PBD : Preoperative biliary drainage
 PBM : pancreaticobiliary maljunction
 pCC : perihilar cholangiocarcinoma

PE : Push Endoscopy

PEG : Percutaneous endoscopic gastrostomy

PEI : percutaneous ethanol injectionPET : Positron emission tomography

PPAP : Polymerase proofreading associated polyposis

PS : Performance status

PSC: primary sclerosing cholangitis

PTBD : percutaneous transhepatic biliary drainage
PTC : percutaneous transhepatic cholangiography
PTCS : percutaneous transhepatic cholangioscopy

PV : portal vein PV : portal vein.

PVE : Portal vein embolization

QoL : quality of life RCTx : chemo-radiation

RFA : Radiofrequency ablation

RTx : radiation

SCC : squamous cell carcinoma

SE : Spiral enteroscopy

SEER : The Surveillance, Epidemiology and End

Results

SEMS : Self-expanding metal stents

SPS : Serrated/hyperplastic polyp syndrome

SSA/P : sessile serrated adenoma/polyp
TACE : transarterial chemoembolization

TAE : transarterial embolization

TNM : tumor, lymph node, metastasis **TSA** : traditional serrated adenoma

UGI/SBFT : Upper GI series with small bowel follow-

through

UICC : the International Union Against CancerVEGF : vascular endothelial growth factor

VEGFR : Vascular endothelial growth factor receptor

VIN : vulvar intraepithelial neoplasia

WCRF/AICR: The World Cancer Research Fund/American

Institute for Cancer Research

WHO : World Health Organization

Introduction

Gastrointestinal tumors, from the commonly presenting neoplasms of the stomach and large intestine to the rarer lesions within the small intestine, are an area of keen professional interest. Indeed, research in this area generates a considerable body of scientific literature covering the entire spectrum from bench to bedside. Yet gastrointestinal tumors still pose a number of major conceptual and practical challenges. Lesions in the small intestine, for example, often elude early detection and their general symptoms frequently perplex clinicians. (Fang & Malfertheiner, 2014)

The epidemiology of digestive cancers is under dynamic changes owing to the changing prevalence and distribution of etiological factors. Colorectal cancer (CRC) incidence in many developing countries, including Asian countries, has increased 2- to 4-fold over the last two decades and has now reached a rate comparable to that of developed countries, with Westernization of diet playing a key role in this trend. Obesity, an increasingly prevalent global health issue, is also expected to become an important risk factor of CRC. (Wu & Sung, 2013)

The worldwide incidence and mortality rates of Gastric Cancer (GC) are decreasing, probably caused by increased availability of fresh fruits and vegetables and decreased consumption of salted and preserved foods, reduction in pylori chronic Helicobacter infection, and increased screening activities in high-risk countries. some Nevertheless, GC remains the fourth most common cancer and the second leading cause of cancer death in the world. (Wu & Sung, 2013)

Although small bowel polyps and tumors are rare, their incidence has increased significantly over the past 30 years. The relative 'rarity' of these tumors has led to stagnation in the development of effective curative or adjuvant therapies. Thus, the prognosis of most of these tumors is still dismal. Nevertheless, hope is now on the horizon as new methods such as capsule endoscopy and balloon-assisted enteroscopy have contributed to a rise in the diagnosis of these lesions and a diagnosis at earlier stages. (Fry et al., 2014)

The majority of GI tumors are epithelial in origin, and most patients present with advanced (regional or distant) disease (~60% patients for colorectal and esophageal cancer) with poor prognosis and low survival rates. Despite

and chemotherapy, surgery, radiotherapy, advances in treatment for most patients is palliative. Indeed, the life expectancy for patients with advanced gastric cancer (with or without chemotherapy) is only 6 to 9 months. (Yamaguchi et al., 2014)

Gastrointestinal cancers remain one of the main causes of death in developed countries. The main obstacles to combating these diseases are the limitations of current diagnostic techniques and the low stability, availability, and/or_specificity of pharmacological treatment. (Prados et al., 2014)

Our deeper knowledge of the genetics and pathobiology of these tumors has yielded breakthroughs in diagnostic and regimens, extending patient therapeutic survival improving quality of life. However, there is still much work that needs to be done to improve the prognosis of patients diagnosed with these malignancies. (Fang & Malfertheiner, 2014)

AIM OF THE ESSAY

The aim of this essay is to review the recent advances in gastrointestinal tumors and to highlight the current concepts for their prevention, screening, early diagnosis, staging and therapeutic regimens.