# CHROMOENDOSCOPY OF THE GASTEROINTESTINAL TRACT

#### **THESIS**

# SUBMITTED IN PARTIAL FULFILLMENT FOR M.D DEGREE OF INTERNAL MEDICINE

BY
Dr. AMEL EL-SAYED AHMED SALEM
(M.B., B.CH.M.Sc.)
Assistant lecturer of Internal Medicine

Faculty of Medicine Cairo University

**Supervised BY** 

### Dr. MOHAMMED REDA AWADEIN

Professor of Internal Medicine Faculty of Medicine Cairo University

### Dr.ALI HASSAN FARAG

Lecturer of Internal Medicine Faculty of Medicine Cairo University

### Dr. SOLIMMAN SABA SOLIMMAN

Professor of PATHOLOGY Faculty of Medicine Cairo University

Faculty of Medicine Cairo University

### **Acknowledgement**

In the name of GOD the most merciful and the mostgraceful I would like to express my deepest gratitude and profound appreciation to **Professor Mohammed Reda Awadeen**, Internal Medicine Department, Cairo University. Working under the supervision of such an eminent professor was a privilege that gave me great honor and much pride. His care, perfectionism and invaluable experience were of much guidance to me. He sincerely supported me throughout this work and offered any possible help to overcome every difficulty I faced through the rough times.

It was an impressive opportunity to work and learn under the supervision of **Professor Ali Farag**, Internal Medicine Department, Cairo University, who taught me how to own a tireless will in discovering and improving my work style. His inexhaustible patience and calm enthusiasm were more than admirable.

**Professor, Solimman Saba**, Pathology Department., Cairo University owes me a lot for the faithful care and kindness he showed me. He sincerely and faithfully guided me in every step and facilitated lots of difficulties with his comprehensive co-operation and humanistic experience.

**To all my Professors** at the Internal Medicine Department Cairo University, which I have the honor to belong, Iam truly grateful for your care, support, encouragement and concern that started the moment I stepped into the department and throughout my progress.

This work would have never been complete without the co-operation of the nursing and working staff of the Endoscopy Unit, Cairo University.

My family has been offering me as always endless and continuing help and support. Their understanding and patience were much appreciable.

This work was done by and for the sake of patients. May God alleviate their sufferings and accept our honest intentions to dedicate this work for the sake of their own benefit.

Amel Salem

# **CONTENTS**

No	Content	Page
	List of abbreviation	
	List of tables	
	List of figures	
I	Introduction and aim of the work	1
	II Review of literature	
	Barrett's esophagus	4
		2.1
	Colonic polyp	21
	Chromoendoscopy	40
	magnification endoscopy	66
	Narrow band imaging (NBI)	73
III	Patients and methods	85
IV	Results	95
V	Discussion	144
VI	Summary and conclusion	153
VII	References	157
VIII	Arabic summary	

## **LIST OF ABBREVIATIONS**

ADR	Adenoma Detection Rate
AGA	American Gastroenterology Association
AMSP	Absent microstructural pattern
BE	Barrett's esophagus
CCD	Charged coupled device
CIMP	CpG Island Methylator Phenotype
COX	Cyclooxygenase
CRC	Colorectal carcinoma
CTC	Computed tomographic colonography
CUC	Chronic ulcerative colitis
DM	Diabetes mellitus
EME	Enhanced magnification endoscopy
EMR	Endoscopic mucosal resection
FICE	Fujinon Intelligent Color Enhancement
FJP	Familial juvenile Polyposis
FOBT	Fecal occult blood testing
GEJ	Gastroesophageal junction
GERD	Gastroesophageal reflux disease
GI	Gastrointestinal
GIT	Gastrointestinal tracte
H and E	Hematoxylin –Eosin
H2RAs	H2 Receptor Antagonists
HGIN	High -grade intraepithelial neoplasia
HNPCC	Hereditary non-polyposis colorectal cancer
HPS	Hyperplastic Polyposis syndrome
IBD	Inflammatory bowel disease
IMSP	Irregular microstructural pattern
IMVP	Irregular microvascular patterns
JPC	Juvenile Polyposis coli
k	kappa
MB	Methylene blue
NBI	Narrow band imaging
NSAIDs	Nonsteroidal anti-inflammatory drugs
PIVI	Preservation and Incorporation of Valuable
	Endoscopic Innovation
PJS	Peutz-Jeghers syndrome
PPI	Proton pump inhibitor

RGB	Red, green, blue
RMVP	Regular microvascular patterns
SAs	Serrated adenomas
SD-WL	Standard-definition white-light
SIM	Specialized intestinal metaplasia
SSPs	Sessile serrated polyps
UC	Ulcerative colitis
WLE	White –light endoscopy
WT	Withdrwal time

# **LIST OF TABLES**

No	TABLE	Page
1	Show staining agents for Chromoendoscopy.	41
2	Mucosal morphology according to the 3main classification system described for Barrett's esophagus	76
3	Group 1 patient demographic characteristics	95
4	Patients with suspected Barrett's esophagus using chromoendoscopy with Methylene blue 0.5%	100
5	Patients with suspected Barrett's esophagus in standard endoscopy using Narrow Band Imaging (N.B.I)	103
6	Group 2 patient demographic characteristics	106
7	Patients with Colonic Polyp and Chromoendoscopy using Indigocarmine 0.4% and Narrow Band Imaging	108
8	Age of patients in the first group	114
9	Gender of patients	114
10	Associated medical conditions	114
11	History of cigarette smoking	115
12	Reason for Endoscopy	115
13	Categories of patients based on the length of Barrett's mucosa with SWLE	116
14	Suspected dysplasia in Barrett's mucosa detected with SWLE	116
15	Confirmed intestinal metaplasia (Barrett's mucosa) by histopathology	117
16	Histopathology confirmed dysplasia in Barrett's mucosa	117
17	Barrett's mucosa detected with methylene bleu chromoendoscopy	117
18	Dysplasia in Barrett's esophagus detected after methylene bleu chromoendoscopy	118
19	Correlation between standard endoscopy (Barrett's length) and histopathological confirmation of Barrett's (Crosstabulation)	119
20	Correlation between standard endoscopy suspected and histopathological confirmation of Barrett's (Crosstabulation)	120
21	Correlation between standard endoscopy suspected and histopathological confirmation of dysplasia (Crosstabulation)	121
22	Correlation between chromoendoscopy suspected and histopathological confirmation of Barrett's (Crosstabulation)	122
23	Correlation between chromoendoscopy suspected and histopathological confirmation of dysplasia (Crosstabulation)	123
24	Barrett's mucosa detected with N.B.I	124
25	Dysplasia in Barrett's esophagus detected after N.B.I	124

26	Correlation between N.B.I suspected and histopathological confirmation of Barrett's (Crosstabulation)	125
27	Correlation between N.B.I suspected and histopathological confirmation of dysplasia (Crosstabulation)	126
28	Age of patients in the second group	130
29	Gender of patients of second group	130
30	History of cigarette smoking in the second group	130
31	Associated medical conditions in the second group	131
32	Reason for colonoscopy	131
33	Size of polyps	132
34	Shape of polps	132
35	Site of polyps	133
36	chromoendoscopy of polyps	133
37	N.B.I. of polyps	133
38	Histopathological confirmation of polyps	134
39	Correlation between chromoendoscopy suspected and histological confirmation of adenomatous polyps (crosstabulation)	135
40	Correlation between chromoendoscopy suspected and histological confirmation of hyperplastic polyps (crosstabulation)	136
41	Correlation between NBI suspected and histological confirmation of adenomatous polyps (crosstabulation)	137
42	Correlation between NBI suspected and histological confirmation of hyperplastic polyps (crosstabulation)	138
43	Correlation between standard endoscopy suspected and histological confirmation of hyperplastic& adenomatous polyps (crosstabulation)	139
44	Correlation between chromoendoscopy suspected and histological confirmation of hyperplastic& adenomatous polyps (crosstabulation)	140
45	Correlation between NBI suspected and histological confirmation of hyperplastic& adenomatous polyps (crosstabulation)	141

### **LIST OF FIGURES**

No	Figures	Page
1	Diagnosis of Barrett's esophagus	9
2	Landmarks for the Diagnosis of Barrett's esophagus	10
3	Gastroesophageal junction	11
4	Esophagectomy specimen with Barrett's esophagus	17
5	Histopathology of Barrett's esophagus	20
6	colonic polyps	22
7	Low (left) and high (right) power views of a biopsy of a normal colon	25
8	Medium power view of a hyperplastic colonic polyp	25
9	Chromoendoscopy with Lugol's solution	44
10	Endoscopic views of short-segment Barrett's esophagus (left panel) and long-segment Barrett's esophagus (right panel)	46
11	Kudo Pit Pattern Classification	48
12	A, Endoscopic image of long-segment Barrett's esophagus (BE) with no apparent cancer B, Endoscopic image of long-segment BE from the same patient at a separate procedure after methylene blue (MB) staining	54
13	A, Colonoscopic view of hyperplastic polyp stained with 0.9% indigo carmine dye. B, Colonoscopic view of adenomatous polyp stained with 0.9% indigo carmine dye.	56
14	Pit pattern classification of colonic crypts following dye spray application	62
15	Hyperplastic polyp revealing a stellar-type pit pattern (type II) as seen with magnification following spraying with 0.4% Indigocarmine	62
16	Magnification endoscopy following instillation of acetic acid	68
17	An area with nondysplastic Barrett's mucosa at the squamo-columnar junction imaged with high resolution endoscopy (left) and with narrow band imaging (right).	77
18	A detailed view of an early cancer area in a Barrett's esophagus	78
19	Diagnostic criterion for superficial gastric neoplasia by magnifying endoscopy combined with narrow-band imaging	80
20	Polyp surface mucosal and vascular patterns with narrow-band imaging	82
21	Representative screen shot: hyperplastic polyp. NBI, narrow-band imaging	84
22	Representative screen shot: adenoma. NBI, narrow-band imaging	84
24	Barrett's mucosa by standard endoscopy	127
25	Chromoendoscopy of the same Barrett's mucosa	128
26	Barrett's mucosa before N.B.I.	128
27	Barrett's mucosa after N.B.I.	129
28	adenomatous polyp by WLE	142
29	adenomatous polyp by N.B.I	142
30	adenomatous polyp by N.B.I	143

### **Abstract**

**Background:** Chromoendoscopy, or Chromoscopy, refers to the topical application of stains or dye at the time of endoscopy in an effort to enhance tissue characterization, differentiation, or diagnosis. The continued rise in the incidence of adenocarcinoma of the esophagus has fueled resurgent interest in the use of a variety of endoscopic and nonendoscopic techniques to improve the diagnosis of Barrett's esophagus and associated dysplasia/cancer. Objective: To use chromoendoscopy & NBI for the characterization and the classification of the mucosal morphology in nondysplastic BE and in BE with HGIN& to discriminate neoplastic from non-neoplastic polyps at screening sigmoidoscopy will obviate the need for histologic diagnosis and could have the potential for great cost saving. Descriptive study. Patients & methods: We used chromoendoscopy & NBI to image and biopsy randomly selected areas in 80 patients with BE & in patients with colonic polyps. Results: Our study have shown a high significant benefit of MB chromoendoscopy over SD-WLE in identification of both intestinal metaplasia and dysplasia in Barrett's esophagus (P-value < 0.0001). The study have shown also a significant benefit of N.B.I over SD-WLE in identification of dysplasia in Barrett's esophagus (P-value 0.002) & There is a statistically significant difference in polyp histology prediction by using NBI or indigocarmine chromoendoscopy over SD-WL colonoscopy for both adenomatous & hyperplastic polyps (P-value < 0.001). Conclusions: Indigocarmine chromoendoscopy had the highest sensitivity, specificity& accuracy in predicting adenomas in real time during colonoscopy using a Kudo pit pattern classification & the use of MB chromoendoscopy& NBI resulted in the diagnosis of more neoplastic areas than with SD-WLE, with NBI had the highest sensitivity& MB chromoendoscopy had the highest specificity& accuracy in predicting dysplasia.

#### **Keywords:**

Chromoendoscopy
Methylene blue
Barrett's esophagus
Narrow band imaging (NBI)
Colonic polyps



### Introduction

Chromoendoscopy, or Chromoscopy, refers to the topical application of stains or dye at the time of endoscopy in an effort to enhance tissue characterization, differentiation, or diagnosis. Chromoendoscopy is distinguished from endoscopic tattooing, which involves the injection of a long-lasting pigment (e.g., India ink) into tissue for future localization (*Ginsberg GG*, *et al.*, 2002).

It provides additional diagnostic information regarding the epithelial morphology and pathophysiology. Based on experience gathered mainly in Japan, chromoendoscopy is now in more widespread use, particularly to identify preneoplastic and neoplastic lesions. The most promising technique is the depiction of squamous epithelium neoplasia of the esophagus with Lugol's solution, staining of Barrett's mucosa by methylene blue, including the potential to identify neoplasia, and the demarcation of neoplasia with indigocarmine in the stomach and colon for local endoscopic resection. Innovative application and refinement of the existing ones are soon to be expected (*Peitz U., 2002*).

Studies on chromoendoscopy have reported different classifications for the mucosal patterns in nondysplastic BE. Little, however, is known about the surface-pattern characteristics of high-grade intraepithelial neoplasia (HGIN). In addition, chromo endoscopy is operator dependant and labor intensive, requiring the use of staining solutions, spraying catheters, and multiple water rinses (*Connor MJ*, et al., 2004).

The mucosal morphology consists of both mucosal and vascular patterns. The detection of the superficial vascular patterns, however, is difficult with chromoendoscopy, because the staining solution may obscure visualization of blood vessels.

In Narrow Band Imaging NBI, RGB (red, green, blue) filters with narrow Band pass ranges and a higher relative intensity of blue light are used.

These filters enable better visualization of the mucosal patterns, because blue light allows for optimal superficial imaging. In addition, NBI reveals the superficial vasculature, because of absorption of the blue light by Hb (*Gono K*, et al., 2004).