

## Histomorphological Evaluation of Colonoscopic Mucosal Biopsy with Chronic Gastrointestinal Disorders

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DOI: <https://doi.org/10.3329/jafmc.v13i2.41377>

### Abstract

**Introduction:** Colonoscopic biopsy is important in the diagnosis and treatment of suspected colonic diseases as it is a diagnostic procedure of choice for patients with gastrointestinal disorders lasting for several weeks to months. Histomorphological evaluation of colonic biopsy is important for specific diagnosis, for determining the extent of the disease and its response to therapy and for detecting complications.

**Objective:** To evaluate colorectal mucosal lesion by histopathological examination and to confirm the diagnosis of different gastrointestinal diseases including malignancies.

**Materials and Methods:** This descriptive cross-sectional study was conducted on 330 colonoscopic biopsies with different gastrointestinal disorders from January 2014 to December 2016 in the Department of Pathology, Armed Forces Hospital, Kuwait.

**Results:** About 75.2% of cases were male with average age was 45 years with ranging 10-80 years. Out of 330 biopsies specific diagnosis was made in 63% cases and among the specific diagnosis, tubular adenoma was the commonest (21.22%) followed by inflammatory bowel diseases 13.94% cases. Total malignant cases were 7.60% and majority of them in the age group of 41-60 years.

**Conclusion:** The importance of colonoscopic biopsies lies especially in some chronic diarrhoea, alteration of bowel habit and per rectal bleeding. Moreover with the early histopathological diagnosis by detecting the precancerous lesion, like Ulcerative colitis, adenomatous polyp, patient can get rid of developing cancer and thereby saved life.

**Key-words:** Colonoscopy, Inflammatory bowel diseases, Colitis, Neoplasia.

### Introduction

Since 1970, Colonoscopy is one of the most important tools to evaluate the large bowel and to screen colorectal diseases, especially neoplasia and polyps<sup>1</sup>. It also makes it possible to identify early lesions in risk groups, to investigate signs and

symptoms of abdominal pain, gastrointestinal bleeding, changes in bowel habits, chronic diarrhoea, unexplained refractory iron deficiency anemia and abdominal masses to follow up the patients treated for colorectal cancer or inflammatory bowel disease<sup>2</sup>. Through colonoscopy, it is possible to visualize the mucosa of the terminal ileum, colon and rectum and to check for macroscopic lesions<sup>3</sup>. In addition, several procedures can be performed, especially biopsies. The introduction of associated technologies, such as chromoscopy and image magnification has extended the use colonoscopy, making it easier to identify subtle lesions and benefiting a greater number of patients<sup>4</sup>. In fact, colonoscopy is currently one of the most complete tools for colorectal disease investigation<sup>5</sup>.

Examination of colorectal biopsy specimen is a reliable method for diagnosing inflammatory bowel disease<sup>6</sup> though many factors lead to variation in biopsy interpretation between reporting histopathologists<sup>7</sup>. Passing generations and the accompanying lifestyle changes is associated with an increased incidence of gastrointestinal diseases. Epithelial tumors of colon are a major cause of morbidity and mortality worldwide. Colorectal cancer is the fourth ranking cancer worldwide, accounting for approximately 9% of all cancers<sup>8</sup>. Adenocarcinomas are the commonest malignancies arising in the colorectal region, other being carcinoid and melanoma. Colonoscopy is currently considered to be gold standard for cancer surveillance<sup>9</sup>. The development of fibre-optic colonoscope has enabled the clinician to visualize the mucosa of the rectum, entire colon and terminal ileum, to screen the intestinal abnormalities and to get representative biopsy for the definitive diagnosis<sup>10</sup>. Histomorphological examinations of colorectal biopsies reveal a spectrum of lesions ranging from non-neoplastic lesion to neoplastic tumours including benign and malignant tumour. All these lesions often require colonoscopic biopsies for their conclusive diagnosis<sup>11</sup>. Both macroscopic and microscopic appearance along with clinical correlation helps in definitive diagnosis of the lesion which helps in early treatment and better outcome of the patient. The objective of the study is to evaluate colorectal mucosal lesions by histopathological examination and to confirm the diagnosis of different gastrointestinal diseases including malignancies.

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**Materials and Methods**

This descriptive cross-sectional study was conducted on 330 biopsies from patients attending the Gastroenterology OPD who presented with lower gastrointestinal tract symptoms. Information was retrieved from digital clinical records at Jaber Al Ahmed Armed Forces Hospital, Kuwait covering the period January 2014 to December 2016. Clinical details along with a detailed description of the colonoscopic findings were obtained. An attempt was made by the clinician to give a colonoscopic diagnosis in all the cases. The samples were collected in 10% neutral buffered formalin processed and embedded in paraffin with the mucosal surface being uppermost and 4µ thick serial sections were prepared. All tissues were stained with H&E and special stains like Periodic Acid Schiff (PAS), Alcian blue (PAS/Alb), Reticulin, Ziehl Nielsen (ZN) along with Immunohistochemistry (IHC) such as cytokeratin(CK), Mucicarmin (MUC) for colorectal adenocarcinoma, Chromogranin and synaptophysin for neuroendocrine tumour were done as and when required. The diagnosis of colorectal biopsies was made on the basis of clinical presentation, colonoscopic findings and light microscopic features of H&E and special stained sections. The lesions were classified as non-neoplastic lesions, benign neoplastic lesions and malignant tumors. The tumors were classified as per WHO classification and observations were compared with other studies. Statistical analysis was done by software SPSS version 17.0. Categorical variables are expressed as frequencies and percentages. Chi-square test was done and p<0.05 was considered statistically significant.

**Results**

Biopsies were performed on patients of all age groups with complains of gastrointestinal disorders, the youngest was a 10 year old child and oldest was an 85 year old male. Average age was 45 irrespective of age and sex; the overall common symptoms of presentation were altered bowel habits, per rectal bleeding, anaemia, diarrhoea, pain abdomen and colonic ulcer, mass or growth. However, most of the patients had more than one symptom during presentation. In Table-I shows diarrhoea (43.33%), altered bowel habit (33.64%) and per rectal bleeding (11.52%) were common predominant symptoms.

**Table-I:** Distribution of patients according to predominant symptoms (n=330)

Symptoms	No of patients	(%)
Diarrhoea	143	43.33%
Altered bowel habit	111	33.64%
Pain abdomen	28	8.48%
Per rectal bleeding	38	11.52%
Anaemia	08	2.42%
Intestinal obstruction	02	0.61%
Total	330	100%

Among 330 cases, 248(75.15%) cases were male and 82 (24.85%) were female. Male to female ratio 3:1. The type of lesions broadly diagnosed as specific and non specific diagnosis. Specific diagnosis based on definitive changes like tubular adenoma, adenocarcinoma, carcinoid tumour etc. Non specific diagnosis included non specific colitis and no significant histomorphological changes. Specific diagnosis was made in 63% and nonspecific diagnosis was 37% cases (Table-II).

**Table-II:** Age and sex distribution of histopathological diagnosis of colonic mucosal lesion (n=330)

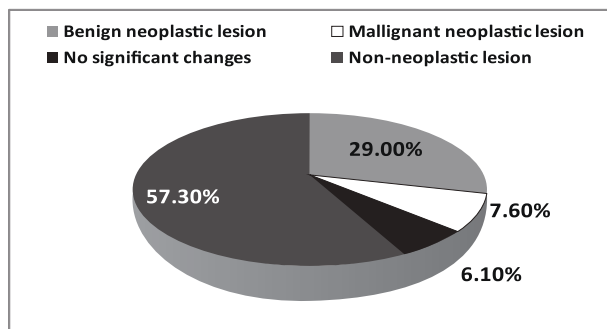
Type of lesions		SEX		Distribution of ages (yrs)				n=330	%
		Male	Female	10-20	21-40	41-60	>60 yrs		
Non specific changes	No significant changes	14	6	10	5	4	1	20	6.06
	Nonspecific colitis	82	20	18	30	26	28	102	30.92
Specific diagnosis	Microscopic colitis	04	02	02	01	02	01	06	1.82
	Ulcerative colitis	25	07	02	17	09	04	32	9.70
	Corhn disease	9	5	3	7	3	1	14	4.24
	Lymphocytic colitis	02	0	0	02	0	0	02	0.60
	Lymphoid hyperplasia	03	01	03	01	0	0	04	1.22
	Hyperplastic polyp	18	04	15	05	02	0	22	6.66
	Tubular adenoma	54	16	15	16	30	09	70	21.22
	Villous adenoma	06	04	02	03	02	03	10	3.03
	Tubulovillous adenoma	10	06	02	04	06	04	16	4.85
	Radiation colitis	01	0	0	0	0	01	01	0.30
	Collagen colitis	01	0	0	01	0	0	01	0.30
	Solitary rectal ulcer	04	01	0	02	02	01	05	1.51
	Adenocarcinoma	12	08	01	02	10	07	20	6.06
Carcinoid	03	02	01	02	01	01	05	1.51	
<b>Total</b>		<b>248</b>	<b>82</b>	<b>74</b>	<b>98</b>	<b>97</b>	<b>61</b>	<b>330</b>	<b>100.00</b>

Among 46 cases of IBD 32 of them were Ulcerative colitis, 14 cases were Crohn's disease and 25 cases were found malignant tumor with variable gastrointestinal symptoms. Significant ( $p < 0.001$ ) correlation of gastrointestinal symptoms with crohn's disease, ulcerative colitis and malignant tumor was found (Table-III).

**Table-III:** Types of lesion related to the predominant symptoms of patient (n=330)

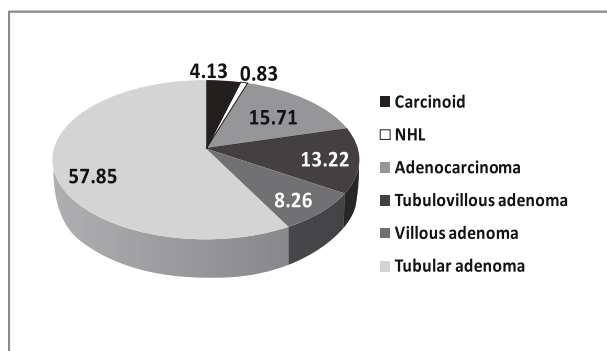
	Chronic unexplained diarrhoea (n=143)	Altered bowel habit (n=111)	Bleeding per rectum (n=38)	Others symptoms (38)	Total	Statistics
Non Specific colitis	67	20	03	12	102	$\chi^2 = 112.6$ df = 12 P < 0.001
Inflammatory bowel diseases (UC, CD)	16	21	06	03	46	
Adenomatous polyp	34	31	10	21	96	
Malignant tumor	04	05	15	01	25	
Others lesions	22	34	04	01	61	
<b>Total</b>	<b>143</b>	<b>111</b>	<b>38</b>	<b>38</b>	<b>330</b>	

Out of 330 cases, 20(6.10%) cases did not show any significant histomorphological changes, 189(57.30%) were non-neoplastic and 121(36.70%) were neoplastic lesion where 96(29%) benign neoplastic polyps and 25(7.60%) cases were malignant lesion. It was observed that non-neoplastic lesions (57.30%) were predominant over neoplastic lesions (Fig-1).



**Fig-1:** Histopathological changes of colonic mucosal biopsies lesion

Colonic polyps are broadly categorized in non- neoplastic and neoplastic polyps. Out of 118 polyps, 22 cases were non neoplastic (hyperplastic polyps) and 96 were neoplastic polyps. Neoplastic polyps included Tubular adenoma, Tubulovillous adenoma and Villous adenoma. In Fig-2 shows among the neoplastic lesions, most common is the tubular adenoma 57.85%, followed by tubulovillous adenoma 13.22%, villous adenoma 8.26%. Among the malignant lesion, adenocarcinoma was 15.71% and carcinoid 4.13%.



**Fig-2:** Distribution of colorectal neoplastic lesion (n=121)

Among the malignant lesion adenocarcinoma were more commonly present in sigmoid colon (08 cases) followed by rectum (05 cases). Thus adenocarcinoma showed a left sided predilection. Most of the carcinoid is in the appendix (Table-IV).

**Table-IV:** Site distribution of malignant neoplastic lesions (n=25)

Site	Adenocarcinoma	Carcinoid	Others
Caecum	02		01
Appendix	01	03	
Ascending colon	02	-	
Transverse colon	01	-	
Sigmoid colon	08	-	
Rectum	05	02	
<b>Total</b>	<b>19</b>	<b>05</b>	<b>01</b>

Out of 25 malignant lesions, most common malignant tumour was moderately differentiated adenocarcinomas followed by well differentiated adenocarcinoma and carcinoid (Table-V).

**Table-V:** Categorization of malignant tumour (n=25)

Categories of malignant tumour	n
Well differentiated adenocarcinoma	05
Moderate differentiated adenocarcinoma	09
Poorly differentiated adenocarcinoma	02
Mucinous adenocarcinoma	02
Signet ring adenocarcinoma	01
Non Hodgkin Lymphoma	01
Carcinoid tumour	05
<b>Total</b>	<b>25</b>

## Discussion

In present study (Table-II), Out of 330 biopsies, 63.33% were non-neoplastic and 36.66% biopsies were neoplastic lesion. These finding are similar to other studies where non-neoplastic lesions were detected more than the neoplastic lesions (66.3% vs. 28.9%, 61.3% vs. 38.7%)<sup>12,13</sup>. Among all non-neoplastic lesions, 53.97% biopsies comprised of chronic non specific colitis, 24.34% were IBD and others made up of 21.69% cases. But colitis was found in other previous study series where it was 47.3% and 38.3% biopsies<sup>12,14</sup>. From this study, it was observed

an incidence of 13.94% cases of IBD among all lesions (n=330), 9.7% being Ulcerative colitis (UC) and 4.24% Crohn's disease (CD). Wool rich and colleagues<sup>15</sup> showed that low grade dysplasia (LGD), like HGD, is also predictive factor of future carcinoma. Their study showed that 18% of those with LGD progressed to carcinoma within an average of 6.3 years. Therefore, it is important to document presence of dysplasia in UC and CD, which will determine the course of treatment in these patients. Rangaswamy R et al<sup>16</sup> observed that IBD comprised 24.4% of non-neoplastic lesions in their study. Karve SH et al<sup>17</sup> in their study observed 11% cases of Ulcerative colitis and 4% cases of Crohn's disease. They also identified dysplasia in 62% cases of UC, 72% showing low grade dysplasia(LGD) and 28% was high grade dysplasia (HGD). In the present study, 65% cases of UC showed variable degree of dysplasia, among them 70% cases of LGD and the other 30% cases of HGD. These patients require follow up and different modalities of treatment. Thus our observations were comparable to these studies.

In the present study, colonic biopsies enabled to identify histopathological causes of unexplained chronic diarrhoea in nearly 43.33% of a sample of patients whom colonoscopic findings were normal, approximately 20.60% of them could be treated by an appropriate therapy or surgical removal based on the findings of UC, CD, hyperplastic polyps, adenomatous polyp, adenocarcinoma and different types of colitis like collagenous colitis, lymphocytic colitis etc. Adenomas are the precursor conditions of colorectal carcinoma Most common site for adenomatous polyps was sigmoid colon followed by rectum. Konishi F et al<sup>18</sup> in their study of colorectal adenomas found that 81 % were tubular adenomas while only 19% were villous and tubulo-villous type. Tony J and Harish K et al<sup>19</sup> found that in Southern India adenomatous polyps were the most common polyps (79.8%) in the age group of 23-82 years with M:F ratio of 2.5:1. They also found severe dysplasia in 12% of tubular adenomas and 43% of villous adenomas which is consistent with this study. In high grade dysplasia patients requires more attention as colectomy may required. Other adenomas require only polypectomy and further follow up in future.

In this study malignancies were seen only in 7.5% of all biopsies. Majority of the malignant cases were males (60%). The most common site of occurrence of the malignant tumors was sigmoid colon followed by rectum. The number of malignant lesion are less than other different study series of Sudarshan et al<sup>20</sup> and Laishram RS et al<sup>21</sup>. It was evident that due to health awareness and frequent medical checkup and early biopsy taken from suspected lesion that reduced the number of malignant lesion in this region.

## Conclusion

The histomorphological evaluation of colorectal biopsies has a wide spectrum, ranging from infectious conditions, inflammatory disorders and precancerous lesions to colorectal malignancies. This study emphasizes the need for early diagnosis of these diseases through histopathology which when correlated clinically will help the surgeon/clinician to implement the appropriate treatment and improve the survival of the patient. Patients with normal colonoscopy as in present study that contributed the more specific diagnosis like lymphocytic colitis, eosinophilic colitis, ulcerative colitis and pseudo-membranous colitis. Moreover with the early histomorphological diagnosis by detecting the precancerous lesion, like Ulcerative colitis, adenomatous polyp, patient can get rid of developing cancer and thereby saved life.

## References

1. Finlay AM, Prithi B. Colonoscopy and biopsy. *Gastroenterology* 1997; 11(1):65-82.
2. Badary DM, Hafez MZ. Role of Mucosal Colonic Biopsy in Patients with Chronic Unexplained Diarrhoea who their Colonoscopy is Normal. *Molecular biology* 2017; 2(6):1-4.
3. Bowles CJ, Leicester R, Romaya C et al. A prospective study of colonoscopy practice in the UK today: Are we adequately prepared for national colorectal cancer screening tomorrow? *Gut* 2004; 53:277-83.
4. Nossa FLC, Paula NBLBB. Diagnostic and therapeutic evaluation of colonoscopic biopsy. *Revista Brasileira de Coloproctologia* 1999; 19:168-71.
5. Kagueyama FM et al, Importance of biopsies and histological evaluation in patients with chronic diarrhea and normal colonoscopies. *ABCD Arq Bras Cir Dig* 2014; 27(3):184-7.
6. Dube AK, Cross SS, Lobo AJ. Audit of the histopathological diagnosis of non-neoplastic colorectal biopsies: Achievable standards for the diagnosis of inflammatory bowel disease. *J Clin Pathol* 1999; 51(5):378-81.
7. Berre NL, Heresbach D, Kerbaol M et al. Histological discrimination of Idiopathic Inflammatory Bowel Disease from other types of colitis. *J Clin Pathol* 1995; 48(8):749-53.
8. Fenoglio-Preiser CM, Noffsinger AE, Stemmerman GN et al. *Gastrointestinal pathology An Atlas and Text*. 3rd ed. Philadelphia: Lippincot Williams and Wilkins 2008; 14:899-1036.
9. Park DI, Kang MS, Oh SJ et al. Her-2/neu over expression is an independent prognostic factor in colorectal cancer. *Int J Colorectal Dis* 2007; 22:491-7.
10. Chercek and Berger. In: *Laboratory test and diagnostic procedure*. 5th ed; 2008:65-6.
11. Rajbhandari M, Karmacharya A, Khamal K et al. Histomorphological profile of colonoscopic biopsies and pattern of colorectal carcinomas in Kavre district. *Kathmandu Univ Med J* 2013; 43(3):196-200.
12. Qayyum A, Sawan AS. Profile of colonic biopsies in King Abdul Aziz University Hospital, Jeddah. *J Park Med Assoc* 2009; 59(9):608-11.
13. Winawer SJ, Leidner SD, Haidu SI. Colonoscopic biopsy and cytology in the diagnosis of colon cancer. *Cancer* 1978; 42(6):2849-53.

14. Dickinson RJ, Gilmour HM, McClelland DB. Rectal biopsy in patients presenting to an infectious unit with diarrhoeal disease. *Gut* 1979; 20(2):141-8.
15. Woolrich AJ, DaSilva MD, Korelitz BI. Surveillance in the routine management of ulcerative colitis: The predictive value of low-grade dysplasia. *Gastroenterology* 1992; 103:431-8.
16. Rangaswamy R, Sahadev R, Suguna BV et al. Clinico-colonoscopy and Histomorphological Spectrum of Colonic Diseases in an Academic Tertiary Care Centre. *Journal of Evolution of Medical and Dental Sciences* 2014; 6(3):1-9.
17. Karve SH, Vidya K, Shivarudrappa AS et al. The Spectrum of colonic lesions: A Clinico-pathological study of colonic biopsies. *Indian Journal of Pathology and Oncology* 2015; 2(4):189-209.
18. Konishi F, Morson BC. Pathology of colorectal adenomas. A colonoscopic survey. *J Clin Pathol* 1982; 35(8):830-41.
19. Tony J, Harish K, Ramachandran TM et al. Profile of colonic polyps in a Southern Indian population. *Indian J Gastroenterol* 2007; 26(3):127-29.
20. Sudarshan V, Hussain N, Gahine R et al. Colorectal cancer in young adults in a tertiary care hospital in Chattisgarh, Raipur. *Indian Journal of Cancer* 2013; 50(4):337-40.
21. Laishram RS, Kaiho N, Shimray R et al. Histopathological Evaluation of Colorectal Carcinomas status in Manipur, India. *International Journal of Pathology* 2010; 8(1):5-8.