

*Original Article*

## Histopathological Grades of Colorectal Cancer: A One-Year Single-Center Study

Sumaya Taher Draa<sup>\*1</sup>, Fadia T.M Oshkondale<sup>2</sup>, Adel M Abdallaa<sup>3</sup>, Mohamed Khalifa Salem Al-Kawash<sup>4</sup>, Mohamed Mohamed Ali Altoumi<sup>4</sup>, Kholoud Abd Al-Razzaq Kardmain<sup>5</sup>, Khitam Mohamed Al-Shein<sup>5</sup>

1. Academic surgical department, Faculty of Medicine, University OF Zawia, Libya

2. Pharmacological Department, Faculty of Medicine, University OF Zawia, Libya

3. Advanced Center for Plant Research and Complementary Medicine, Zawia University, Zawia, Libya

4. General practitioner, Subrata National Cancer Institute (SNCI), Subrata, Libya

5. General practitioner, Zawia Medical Center, Zawia, Libya

Corresponding Author Sumaya Taher Draa : E-mail: [S.draa@zu.edu.ly](mailto:S.draa@zu.edu.ly)

Received: 15/05/2025 Accepted: 27/06/2025, Published: 07/07/2025, DOI: <https://doi.org/10.54361/LJMR.19.2.08>

### ABSTRACT

**Background:** Colorectal cancer (CRC) poses a major global health burden. It necessitates careful assessment for prognosis and treatment planning. Histopathological grading plays a key role in prognosis and treatment planning. Local studies are essential to understand CRC characteristics in specific populations and guide national health strategies. This study aimed to assess the histopathological grading patterns among individuals diagnosed with CRC during a one-year interval at the National Cancer Institute in Subrata, Libya, in 2024. **Material and Methods:** A cross-sectional retrospective analytic study was conducted involving 202 patients consecutively selected from the medical records of Subrata National Cancer Institute in 2024 using descriptive statistics and the chi-square test for association through SPSS software. **Results:** This study reveals that the majority of patients (78.7%) had Grade II tumors, with the highest frequency (35.1%) in patients over 60 years of age. Grade I and Grade III accounted for 17.8% and 3.5% of cases, respectively. A statistically significant association was found between histopathological grade and age ( $\chi^2 = 14.9$ ,  $p = 0.021$ ). **Conclusion:** These findings underscore the urgent need for implementing targeted regional strategies, including enhanced public awareness campaigns, robust screening programs for early detection, and strengthening of the diagnostic infrastructure to improve CRC outcomes in Libya.

**Keywords:** histopathology grade, age, colorectal cancer, Libya.

**How to cite this article:** Draa .S.T, Oshkondale .F.T, Abdallaa .A.M, Al- Kawash .M.K, Altoumi .M.A, Kardmain .K.A, Al-Shein .K.M. Histopathological Grades of Colorectal Cancer: A One-Year Single-Center Study

### Libyan 19.2

## INTRODUCTION:

Colorectal cancer (CRC), an aggressive malignancy originating in the large bowel (colon and rectum), is a leading cause of cancer-related sickness and death across the globe [1,2]. The global burden is growing, especially in developing nations where lifestyle changes and insufficient screening programs contribute to rising incidence rates. [3] The histological subtype (grade) of colorectal cancer (CRC) is clinically essential for prognostic categorization of affected patients, highlighting the necessity of early detection methods in high-risk populations. Colorectal adenocarcinoma is the most common histological subtype in more than 90% of colorectal cancer (CRC) patients. These adenocarcinomas are commonly classified into four categories (I-IV) using the Broder classification system: highly differentiated (Grade I), moderately differentiated (Grade II), poorly differentiated (Grade III), and undifferentiated (Grade IV). This rating is based on the amount of glandular tissue impacted [4]. While most CRCs are differentiated adenocarcinomas, about 10% are categorized as poorly differentiated adenocarcinoma (Por), 10% as mucinous carcinoma (Muc), and 1% as signet-ring cell carcinoma [5,6]. The Por/Muc/Sig histological subtypes of CRC demonstrate a strong association with deficient mismatch repair (dMMR), resultant microsatellite instability (MSI), and B-rapid accelerated fibrosarcoma (BRAF) gene mutations. Given that these genetic alterations correlate with poor prognosis in Stage III, the histological classification itself could function as a prognostic factor, possibly obviating the need for routine dMMR/MSI/BRAF analysis in this context [7-10]. Histopathological grading of CRC is a major component in staging and prognosis, influencing therapeutic methods and patient outcomes [4]. Regional studies are essential to identify local disease characteristics and inform policy. In Libya, epidemiological data on CRC are scarce, with few researches describing the histological patterns and grades of tumors observed in clinical practice. Given histopathology's critical role in directing treatment, there is an urgent need for localized investigations that describe tumor features in varied populations. This study aims to fill that gap by presenting a one-year retrospective analysis of CRC histopathological grading from the National Cancer Institute in Subrata, Libya a leading center for cancer care in the region. We aim to describe the grading of the tumor according to age group,

contributing to a better understanding of CRC pathology in the Libyan context.

## MATERIALS AND METHODS:

This retrospective, analytical cross-sectional study was conducted at the Subrata National Cancer Institute. The study population consisted of 202 patients who were consecutively selected from the institute's database. The primary inclusion criterion was a histologically confirmed diagnosis of primary colorectal cancer (CRC). Patients were excluded if their medical records contained incomplete information or if their histopathological findings were inadequate for analysis. Data were collected from medical records for the year 2024. The variables analyzed included patient age and the histological type of adenocarcinoma. The histopathological grade of CRC was categorized as Grade I (Well-differentiated), Grade II (Moderately differentiated), and Grade III (Poorly differentiated). Statistical analysis was performed using SPSS software. Descriptive statistics were employed to summarize the data, and the Chi-Square test was used to assess the statistical distribution and association among the histopathological grades. Ethical approval for the study was granted by the Director General of the National Cancer Institute in Subrata City. To maintain patient privacy, all data were fully anonymized by removing personal identifiers, ensuring strict confidentiality. The use of all collected data was restricted solely to the purposes of this research.

## RESULTS:

Based on data collected from the medical records of the histopathology laboratory of the Subrata Nation Cancer Institute (SNCI) in 2024, the total number of colorectal malignancies was 202 patients. From table 1, 36 (17.8%) patients were diagnosed with Grade I adenocarcinoma, 159 (78.7%) of patients with Grade II adenocarcinoma, and 7 (3.5%) of the total cases were Grade III adenocarcinoma. This study showed that the majority of cases presented with Grade II adenocarcinoma in the age group > 60 (35.1% of total cases). The chi-square test value (14.9) and significance level (0.021) suggest a statistically significant association between age and histopathological classification, indicating that cancer grading may be influenced by patient age.

**Table 1:** Distribution of CRC Grades by Age Group

Age Group	Grade I	Grade II	Grade III	Total
≤40	1 (0.5%)	14 (6.9%)	3 (1.5%)	18 (8.9%)
41-50	7 (3.5%)	21 (10.4%)	1 (0.5%)	29 (14.4%)
51-60	15 (7.4%)	53 (26.2%)	0 (0%)	68 (33.7%)
>60	13 (6.4%)	71 (35.1%)	3 (1.5%)	87 (43.1%)
Total	36 (17.8%)	159 (78.7%)	7 (3.5%)	202 (100.0%)
Chi-Square Test	14.9			
Significance Level	0.021			

## DISCUSSION:

Colorectal carcinoma is a type of colon cancer that results from uncontrolled cell proliferation [11-13]. Distant metastasis is the leading cause of death in CRC, whereas the liver is the most common location, followed by the lungs and the peritoneum. According to many studies, several factors influence the prognosis of CRC patients, including sex, age, initial tumor resection, number of metastases, and tumor differentiation degree [14, 15]. The risk of CRC began to rise after the age of 40 and increased rapidly between the ages of 50 and 55, increasing every decade thereafter [16]. In our study, the majority of CRC patients treated at SNCI in 2024 (35.1%) were above the age of 60 and the study found that all 202 individuals with histopathologic examinations had adenocarcinoma (100%). The results of this study largely match global trends in CRC pathology, with moderately differentiated adenocarcinoma being the most common grade. This is consistent with findings from other Middle Eastern and North African nations (MENA). However, the comparatively high frequency of poorly differentiated tumors (24.2%) is significant and may indicate later-stage diagnosis or aggressive tumor characteristics in the local community [17]. Many previous researches are agreement with this study in that the degree of differentiation of the vast majority (78.7%) were grade II; moderately differentiation adenocarcinoma (35.1% of age group >60), followed by (17.8%) grade I; well differentiation

and (3.5%) grade III; poorly differentiation, [18-21]. These findings suggested that histological assessment is very significant for the evaluation of clinical and management of CRC patient. Besides that, this study showed a significant correlation between the adenocarcinoma differentiation and age ( $p < 0.001$ ) indicating that cancer grading may be influenced by patient age. Whereas a study done in 2023 demonstrated that a strong association between age groups (EAO versus LAO) and histological features ( $p < 0.001$ ) [22]. This aligns with broader research trends showing younger CRC patients often present with more aggressive tumor biology, including higher-grade malignancies [23, 24]. However, some studies, like the 2022 analysis from Cengkareng Hospital, found no significant age-differentiation correlation ( $p = 0.476$ ), highlighting the need for population-specific investigations [25].

## CONCLUSION:

In conclusion, while the precise relationship between advanced age and higher colorectal cancer grade necessitates further research controlling for confounders, this single-center Libyan study identifies a notable prevalence of aggressive tumor grades. This underscores the critical importance of robust early detection programs, improved diagnostic resources, and dedicated public health initiatives to decrease the CRC burden within this population and potentially inform similar contexts.

## REFERENCE:

- 1- Sung H, Ferlay J, Siegel RL, Laversanne M, Soerjomataram I, Jemal A, Bray F. Global Cancer Statistics 2020: GLOBOCAN Estimates of Incidence and Mortality Worldwide for 36 Cancers in 185 Countries. CA Cancer J Clin. 2021 May;71(3):209-249. doi: 10.3322/caac.21660. Epub 2021 Feb 4. PMID: 33538338.
- 2- Mattiuzzi C, Sanchis-Gomar F, Lippi G. Concise update on colorectal cancer epidemiology. Ann Transl Med. 2019

- Nov;7(21):609. doi: 10.21037/atm.2019.07.91. PMID: 32047770; PMCID: PMC7011596.
- 3- Xi Y, Xu P. Global colorectal cancer burden in 2020 and projections to 2040. *Transl Oncol.* 2021 Oct;14(10):101174. doi: 10.1016/j.tranon.2021.101174. Epub 2021 Jul 6. PMID: 34243011; PMCID: PMC8273208.
  - 4- Chen S, Zhang M, Wang J, Xu M, Hu W, Wee L, Dekker A, Sheng W, Zhang Z. Automatic Tumor Grading on Colorectal Cancer Whole-Slide Images: Semi-Quantitative Gland Formation Percentage and New Indicator Exploration. *Front Oncol.* 2022 May 11;12:833978. doi: 10.3389/fonc.2022.833978. PMID: 35646672; PMCID: PMC9130480.
  - 5- Dubansky B, Lewis S, Telgenhoff D. Classification and Histological Characteristics of Colorectal Cancer. *Am Soc Clin Lab Sci.* 2024 Sep 18. doi:10.29074/ascls.2022003206. Epub ahead of print.
  - 6- Miyakawa T, Kawamura H, Honda M, Takano Y, Kinuta S, Kamiga T, Yamazaki S, Muto A, Shiraso S, Yamashita N, Iwao T, Kono K, Konno S. Impact of histological subtype on prognosis in stage IV colorectal cancer: A population-based cohort study. *PLoS One.* 2022 Mar 3;17(3):e0264652. doi: 10.1371/journal.pone.0264652. PMID: 35239725; PMCID: PMC8893698.
  - 7- Xiao H, Yoon YS, Hong SM, Roh SA, Cho DH, Yu CS, Kim JC. Poorly differentiated colorectal cancers: correlation of microsatellite instability with clinicopathologic features and survival. *Am J Clin Pathol.* 2013 Sep;140(3):341-7. doi: 10.1309/AJCP8P2DYNKGRBVI. PMID: 23955452.
  - 8- Verhulst J, Ferdinande L, Demetter P, Ceelen W. Mucinous subtype as prognostic factor in colorectal cancer: a systematic review and meta-analysis. *J Clin Pathol.* 2012 May;65(5):381-8. doi: 10.1136/jclinpath-2011-200340. Epub 2012 Jan 18. PMID: 22259177.
  - 9- Kazama Y, Watanabe T, Kanazawa T, Tanaka J, Tanaka T, Nagawa H. Microsatellite instability in poorly differentiated adenocarcinomas of the colon and rectum: relationship to clinicopathological features. *J Clin Pathol.* 2007 Jun;60(6):701-4. doi: 10.1136/jcp.2006.039081. PMID: 17557871; PMCID: PMC1955052.
  - 10- Venderbosch S, Nagtegaal ID, Maughan TS, Smith CG, Cheadle JP, Fisher D, Kaplan R, Quirke P, Seymour MT, Richman SD, Meijer GA, Ylstra B, Heideman DA, de Haan AF, Punt CJ, Koopman M. Mismatch repair status and BRAF mutation status in metastatic colorectal cancer patients: a pooled analysis of the CAIRO, CAIRO2, COIN, and FOCUS studies. *Clin Cancer Res.* 2014 Oct 15;20(20):5322-30. doi: 10.1158/1078-0432.CCR-14-0332. Epub 2014 Aug 19. PMID: 25139339; PMCID: PMC4201568.
  - 11- Alzahrani SM, Al Doghaither HA, Al-Ghafari AB. General insight into cancer: An overview of colorectal cancer (Review). *Mol Clin Oncol.* 2021 Dec;15(6):271. doi: 10.3892/mco.2021.2433. Epub 2021 Nov 1. PMID: 34790355; PMCID: PMC8591689.
  - 12- Siegel RL, Miller KD, Wagle NS, Jemal A. Cancer statistics, 2023. *CA Cancer J Clin.* 2023 Jan;73(1):17-48. doi: 10.3322/caac.21763. PMID: 36633525.
  - 13- Duan B, Zhao Y, Bai J, Wang J, Duan X, Luo X, Zhang R, Pu Y, Kou M, Lei J, Yang S. Colorectal Cancer: An

- Overview. In: Morgado-Diaz JA, editor. *Gastrointestinal Cancers* [Internet]. Brisbane (AU): Exon Publications; 2022 Sep 30. Chapter 1. PMID: 36343150.
- 14- Page AJ, Cosgrove DC, Herman JM, Pawlik TM. Advances in understanding of colorectal liver metastasis and implications for the clinic. *Expert Rev Gastroenterol Hepatol*. 2015 Feb;9(2):245-59. doi: 10.1586/17474124.2014.940897. Epub 2014 Jul 18. PMID: 25033964.
  - 15- Jin LJ, Chen WB, Zhang XY, Bai J, Zhao HC, Wang ZY. Analysis of factors potentially predicting prognosis of colorectal cancer. *World J Gastrointest Oncol*. 2019 Dec 15;11(12):1206-1217. doi: 10.4251/wjgo.v11.i12.1206. PMID: 31908725; PMCID: PMC6937433.
  - 16- Pilleron S, Sarfati D, Janssen-Heijnen M, Vignat J, Ferlay J, Bray F, Soerjomataram I. Global cancer incidence in older adults, 2012 and 2035: A population-based study. *Int J Cancer*. 2019 Jan 1;144(1):49-58. doi: 10.1002/ijc.31664. Epub 2018 Oct 30. PMID: 29978474.
  - 17- Jiang Y, Yuan H, Li Z, Ji X, Shen Q, Tuo J, Bi J, Li H, Xiang Y. Global pattern and trends of colorectal cancer survival: a systematic review of population-based registration data. *Cancer Biol Med*. 2021 Sep 6;19(2):175–86. doi: 10.20892/j.issn.2095-3941.2020.0634. Epub ahead of print. PMID: 34486877; PMCID: PMC8832952.
  - 18- Shiraj-Um-Mahmuda S, Begum F, Rahman MM, Rahman P, Islam T, Shabnam US, et al. Demographic and clinicopathological evaluation of colorectal adenocarcinoma in Bangladesh at a tertiary level hospital. *Cancer Stud Ther J*. 2023;8:1-8. DOI: 10.31038/CST.2023811.
  - 19- Barresi V, Reggiani Bonetti L, Ieni A, Caruso RA, Tuccari G. Histological grading in colorectal cancer: new insights and perspectives. *Histol Histopathol*. 2015 Sep;30(9):1059-67. doi: 10.14670/HH-11-633. Epub 2015 May 25. PMID: 26004398.
  - 20- Kobayashi T, Ishida M, Miki H, Yamamoto N, Harino T, Yagyu T, Hori S, Hatta M, Hashimoto Y, Kotsuka M, Yamasaki M, Inoue K, Hirose Y, Sekimoto M. Prognostic scoring system based on indicators reflecting the tumor glandular differentiation and microenvironment for patients with colorectal cancer. *Sci Rep*. 2024 Jun 20;14(1):14188. doi: 10.1038/s41598-024-65015-2. PMID: 38902294; PMCID: PMC11189912.
  - 21- Ji X, Kang M, Zhao X, Li X, Guo Y, Xie P, Yu Y, Tian Z. Poorly differentiated cluster grade-a vital predictor for lymph node metastasis and oncological outcomes in patients with T1 colorectal cancer: a retrospective study. *BMC Gastroenterol*. 2022 Sep 5;22(1):409. doi: 10.1186/s12876-022-02492-7. PMID: 36064316; PMCID: PMC9442993.
  - 22- Lukman K, Gunawan GG, Rudiman R, Sribudiani Y, Hasibuan LY, Dewayani BM, Nugraha P, Primastari E. Relationship between age and the histopathological features to chemotherapy response in colorectal cancer patients: a prospective observational study. *J Coloproctol (Rio J)*. 2023;43(4):e300-e309. doi:10.1055/s-0043-1776890.
  - 23- Yao H, Li C, Tan X. An age stratified analysis of the biomarkers in patients with colorectal cancer. *Sci Rep*. 2021 Nov 17;11(1):22464. doi:

10.1038/s41598-021-01850-x. PMID: 34789836; PMCID: PMC8599678.

- 24- Carbajal-López B, Coronel-Hernández J, Herrera M, Ruiz-García E, Miyagui-Adame SM, Díaz-Romero C, et al. Age as a predictor of overall survival in colorectal cancer. *Diagnostics (Basel)* [Internet]. 2024 Nov 14 [cited 2025 Jul 2];14(22):2550.

Doi: 10.3390/diagnostics14222550

- 25- Sugiarto S, Nurfitriana A, Marindawati M. Relationship between age and sex with histopathological differentiation of colorectal adenocarcinoma. *Muhammadiyah Med J*. 2022;3(2):67-73. doi:10.24853/mmj.3.2.67-73