

Comparison of Dysplasia and Regenerative Atypia in Endoscopic Gastric Biopsies

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Abstract - Gastric cancer remains the second most common cause of cancer related deaths worldwide. In tumor progression model of intestinal type of gastric carcinoma, the cellular changes progress from initial inflammation and chronic gastritis to metaplasia, dysplasia and adenocarcinoma. Distinguishing regenerative atypia from dysplasia and carcinoma is the most daunting challenge for a pathologist. Focusing on cytological features can provide an opportunity for early diagnosis and may improve patient's survival.

Key Words: (Gastric Cancer), (Endoscopic Biopsies), (Dysplasia)

1. INTRODUCTION

Gastric epithelial dysplasia is a non-invasive neoplastic lesion, associated with increased risk of gastric adenocarcinoma. It can arise either in metaplastic mucosa (intestinal metaplasia) or in native mucosa.

The histologic diagnosis of gastric dysplasia can be diagnostic challenge to pathologists all over the world due to multiple causes including

- I. Interobserver variation
- II. Specimen orientation
- III. Sampling issues
- IV. Difficulty in distinguishing dysplasia from reactive atypia.

2. MATERIALS AND METHODS

2.1 Study settings

This study was conducted in the Department of Pathology, Kasturba Medical College and Hospital, Manipal. Gastric biopsy samples reported in the department in the period between JAN 2011 –DEC 2012 were included in the study as per the inclusion and exclusion criteria.

2.2 Study design

Cross sectional study

2.3 Study Procedure

A detailed case proforma was prepared before the start of the study. A total of 400 gastric biopsies were included as per the inclusion and exclusion criteria. The endoscopic findings of all these gastric biopsy samples were also recorded in the case proforma sheet.

The following are the histological parameters for,

2.4 Inclusion Criteria

Foveolar hyperplasia, Gastric polyp, Gastric ulcer, Chronic gastritis associated with or without regenerative atypia, Presence or absence of Helicobacter pylori, Gastric atrophy, Intestinal metaplasia, Low grade dysplasia, High grade dysplasia.

2.5 Exclusion Criteria

Normal biopsies and those with frank invasive malignancies.

All gastric biopsy specimens obtained from endoscopy were fixed in 10% formalin, embedded in paraffin and sectioned at 5µ thickness.

Table -1: ASSOCIATION BETWEEN SEX & CASES OF REGENERATIVE ATYPIA AND DYSPLASIA(N=66)

DIAGNOSIS	Sex		Total
	Female	Male	
DYSPLASIA	4	15	19
REGENERATIVE ATYPIA	11	36	47
Total	15	51	66

3. RESULTS AND ANALYSIS

In the present study a total of 400 gastric biopsies were studied after excluding normal biopsies and biopsies with malignancies. Out of 400 biopsies, 66 cases showed cellular atypia.

SITE OF BIOPSY:

All Endoscopic biopsies were taken from antrum and body of stomach.

AGE AND SEX DISTRIBUTION:

The age and sex distribution of the patients is shown (Table-1). In general, there was a male preponderance.

CELLULAR ATYPIA:

In cases with cellular atypia, regenerative atypia cases were 47(71.21 %) and dysplasia cases were 19(28.79%). The criteria given by Lewin & Goldstein was used to separate regenerative atypia from dysplasia.

Following are the types & percentile of cases of individual category which were encountered in cases with cellular atypia in this study (Chart -1)

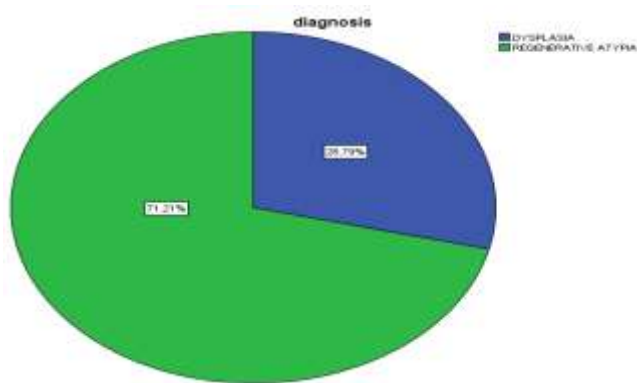


Chart -1: Cases with Cellular Atypia (N=66)

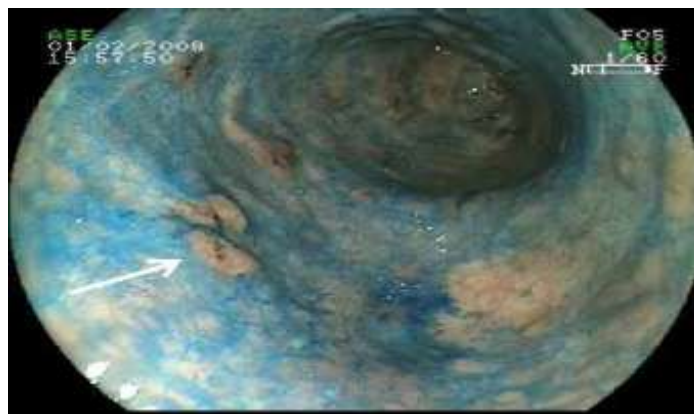


Fig -1:Endoscopic Picture Of Low Grade Dysplasia Case.

4. CONCLUSIONS

The present study attempted to evaluate the histopathological parameters to define dysplasia and regenerative hyperplasia and their association with other conditions in gastric biopsies.

2) A total of 400 gastric endoscopic biopsies reported in the department of Pathology, Kasturba Medical College and Hospital, Manipal in the period between Jan 2011 – Dec 2012 were included in the study as per the inclusion and exclusion criteria.

3) All endoscopic biopsies studied were taken from antrum and body of stomach.

4) Out of 400 biopsies, 66 cases showed cellular atypia. In cases with cellular atypia, regenerative atypia were 47(71.21 %) and dysplasia were 19(28.79%).

5) Most of them (83%) presented with abdominal symptoms like nausea, vomiting, abdominal pain, diarrhoea, and melaena.

6) Among 19 case of dysplasia, 15 were males and 4 were females. Dysplasia was more common in the age group from 40 to 70 years (mean:55).

7) On endoscopy of the 47 cases of regenerative atypia, 24 cases were normal study, 8 cases showed gastric ulcer, and 6 cases showed duodenal ulcer; and out of 19 cases of dysplasia, 9 showed gastric ulcer, and 5 showed normal study.

8) Among the type of associated inflammation, chronic inflammation was the commonest (47/66 cases).

9) Foveolar hyperplasia was seen in 19/47 cases of regenerative atypia and 10/19 cases of dysplasia.

10) Helicobacter pylori was present in 28/47 cases of the regenerative atypia and 7/19 cases of dysplasia.

11) Intestinal metaplasia was present in 11/47 cases of regenerative atypia and 15/19 cases of dysplasia. Association of intestinal metaplasia with dysplasia had statistical significance (p<0.05).

12) Gastric atrophy was present in 4/47 cases of regenerative atypia and 2/19 cases of dysplasia.

13) Gastric polyp was present in 4/47 cases of regenerative atypia and 2/19 cases of dysplasia. Among the two cases of polyp with dysplasia, one case was associated with high grade dysplasia.

14) Most of the cases with regenerative atypia showed maturation at mucosal surface, mitosis restricted to base of glands / pits, uniform nuclear size and chromatin, basal location of nucleus, regular nucleoli and non-uniform accentuation of histological change near area of inflammation (statistical significance (p<0.05).

15) Most of the cases with dysplasia showed absence of maturation at mucosal surface, mitosis at surface of glands / pits, irregular nuclear size, irregular clumping of nuclear

chromatin, nuclear stratification, irregular macronucleoli and regionally uniform histological change (statistical significance ($p < 0.05$).

REFERENCES

- [1] Antonioli DA. Precursors of gastric carcinoma: a critical review with a brief description of early (curable) gastric cancer. *Hum Pathol* 1994; 25: 994-1005.
- [2] Butts KP, Barwick KW. The Esophagus and Stomach. In: Weidner N, editor. *The Difficult Diagnosis in Surgical Pathology*. 1st edition.
- [3] Cheng You W, Blot WJ, Li J, et al. Precancerous gastric lesions in a population at high risk of stomach cancer. *Cancer Res* 1993; 53: 1317-21.
- [4] Corral JM, Mindan JP, Razquin S, Ojeda C. Risk of cancer in patients with gastric dysplasia; follow-up study of 67 patients. *Cancer* 1990; 65: 2078-85.
- [5] Correa P, Tahara E. Stomach. In: Henson DE, Albores - Saavedra J, editors. *Pathology of incipient Neoplasia*. 2nd edition. Philadelphia: WB Saunders Company; 1993. p. 85-103.
- [6] Correa P. Helicobacter pylori and gastric cancer: state of the art. *Cancer epidemiol Biomarkers Prev*. 1996; 5(6):477-81.
- [7] Cuello C, Lopez J, Correa P, Murray J, Zarama G, Gordillo G. Histopathology of gastric dysplasias: correlations with gastric juice chemistry. *Am J SurgPathol* 1979; 3: 491-500.
- [8] Dombal de FT, Price AB, Thompson H, Williams GT, Morgan AG, Softle A et al. The British Society of Gastroenterology early gastric cancer / dysplasia survey: an interim report. *Gut* 1990; 31: 115-120.
- [9] Enchev VG, Raichev RD. Cytomorphometric studies on normal stomach mucosa, precancerous diseases and stomach cancer. *Arch Geschwulstforsch*. 1982; 52(8):641-7.
- [10] Farrands P, Blake J, Ansell I, Coton R. Endoscopic review of patients who have had gastric surgery. *Br Med J* 1983; 286; 755-8.
- [11] Fillipe ML. Gastrointestinal carcinoma and precursor lesions. Filipe MI, Lake BD, editors. *Histochemistry in pathology*. 2nd edition London: Churchill Livingstone; 1990. p. 175-198.
- [12] Ghandur ML, Paz J, Roldan E, Cassady J. Dysplasia of non metaplastic gastric mucosa. A proposal for its classification and its possible relationship to diffuse - type gastric carcinoma. *Am J SurgPathol* 1988; 12; 96-114.
- [13] Goldstein NS, Lewin KJ. Gastric epithelial dysplasia and adenoma: historical review and histological criteria for grading. *Hum Pathol* 1997; 28: 127-133.
- [14] Jarvis LR, Whitehead R. Morphometric analysis of gastric dysplasia. *JPathol*. 1985 Oct; 147(2):133-8.
- [15] Jass JR. Role of intestinal metaplasia in the histogenesis of gastric carcinoma. *J ClinPathol* 1980; 33: 801-810.
- [16] Koktysz R, Zieliński KW, Kulig A. Grading of gastric epithelial dysplasia. An interobserver study and analysis of diagnostic criteria. *Pol J Pathol*. 1997; 48(1):5-14.
- [17] Lansdown M, Quirke P, Dixon MF, Axon ATR, Johnston D. High grade dysplasia of the gastric mucosa: a marker for gastric carcinoma. *Gut* 1990; 31: 977-983.
- [18] Lansdown M, Quirke P, Dixon MF, Axon ATR. High grade dysplasia of the gastric mucosa: A marker for gastric carcinoma. *Gut* 1990; 31: 977-83.
- [19] Lauwers GY, Riddell RH. Gastric epithelial dysplasia. *Gut*. 1999 Nov; 45(5):784-90.
- [20] Lewin KJ. Nomenclature problems of gastrointestinal epithelial neoplasia. *Am J SurgPathol* 1998; 22: 1043-1047.
- [21] M Hiroshi, K Masahiro, E Munetomo, M Norikazu, H Yukiaki. Changes in gastric mucosa that antedate gastric carcinoma. *Cancer*, 2003 ; 66(9):2017 – 2026.
- [22] Ming SC, Bajtai A, Correa P, Elster K, Jarvi OH, Munoz N et al. Gastric dysplasia: Significance and Pathologic criteria. *Cancer* 1984; 54: 1794-1801.
- [23] Ming SC. Cancer of the gastrointestinal tract. Histogenesis and premalignant lesions. *JAMA* 1974; 228: 886-888.
- [24] Ming SC. The importance of follow up studies in gastric dysplasia, the pathologist's view. *Endoscopy* 1993; 4: 294-5.
- [25] Monroe LS, Boughton GA, Sonimers SC. The association of gastric epithelial hyperplasia and cancer. *Gastroenterology* 1964; 46: 267-272.
- [26] Morson BC, Sobin LH, Grunclmann E, Johansen A, Nagayo T, Serck -Hanssen A. Precancerous conditions and epithelial dysplasia in the stomach. *J ClinPathol* 1980; 33: 711-721.

- [27] Owen DA. The stomach. In: Steinberg SS, editor. Diagnostic Surgical Pathology. 3rd edition. Philadelphia: Lippincott Williams and Wilkins; 1999. P. 1311-1347.
- [28] Rosai J. Gastrointestinal tract. (Stomach). In-Rosai J, editor. Ackerman's Surgical Pathology. 8th edition. Singapore Harcourt Brace and company Asia Pte Ltd; 1996. p. 616-666.
- [29] Rugge M, Correa P, Dixon MF, Hatton T, Leandro G, Lewin K et al. Gastric dysplasia: The Padova international Classification. Am J SurgPathol 2000; 24: 167-176.
- [30] Rugge M, Farinati F, Mario FD, et al. Gastric epithelial dysplasia: A prospective multicenter follow-up study from the interdisciplinary group on gastric epithelial dysplasia, Hum Pathol 1991; 22; 1002-8.
- [31] Saraga EP, Gordiol D, Costa J. Gastric dysplasia; a histological follow-up study. AmSurg Pathol 1987; 11; 788-96.
- [32] Shao L. Morphometric analysis of gastric dysplasia and malignancy. Zhonghua Zhong Liu ZaZhi. 1992;14(4):264-6.
- [33] Srivastava A, Lauwers GY. Gastric epithelial dysplasia: the Western perspective. Dig Liver Dis. 2008 Aug;40(8):641-9.
- [34] Stolte M. The new Vienna classification of epithelial neoplasia of the gastrointestinal tract: advantages and disadvantages. Virchows Arch 2003;442:99-106.
- [35] Weinstein WM, Goldstein NS. Gastric dysplasia and its management. Gastroenterology 1994; 107: 1543-1545.