

Endoscopic biopsy yielding upper gastrointestinal malignancies

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Abstract

Objective: Aim of this study is to find the optimal number of endoscopic biopsies needed to diagnose the upper gastrointestinal malignancies in the patient who undergoes endoscopic evaluation. **Methods:** This is an observational study conducted in Government Staney Medical College. Patients with upper gastrointestinal symptoms underwent esophago gastroduodenoscopy using forward viewing scope after getting the proper consent from them. Procedure was done by well experienced endoscopist. In patients with suspected lesion of malignancy in the tract, biopsies are taken. Number of biopsies aimed are eight and serially taken biopsies are labelled in four separate vials. Each vial contains two consecutive samples in the 10% formal saline solution. Details of the site, extent, and type of the lesion were recorded. In case of haemorrhage or any complications the procedure is terminated with proper monitoring of patient until discharged. **Results:** The yield of endoscopic biopsy specimens from 50 patients after combining the results from successive vials. The yield in the first vial is 94% and the cumulative percentage of the second vial yielded 100%, which means the malignancy in the specimen is proved without doubt in the first two vials itself for all the patients. **Conclusion:** In conclusion, this study shows that four biopsy specimens are likely to yield a 100% diagnosis in advanced upper gastrointestinal malignancies. Whereas the endoscopic biopsy yield in the early stages of carcinomas should be evaluated in further studies.

Keywords: Endoscopic Biopsies, Gastrointestinal Malignancies, Endoscopic Evaluation.

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Introduction

Upper gastrointestinal carcinomas are becoming increasingly common in Indian population accounting to the change in food habits and environmental changes occurring rapidly in the country. Of patients whom we included in study from the total of fifty patients half of the patients had gastric carcinoma, Others comprising esophageal carcinoma and periampullary carcinoma. There is a gradual increase in esophageal carcinoma incidence compared to the past incidence. Periampullary carcinomas contribute to 12% of the total upper gastrointestinal malignancies. Gastric carcinomas account for the 52% of total malignancies in the upper gastrointestinal carcinomas. Esophageal carcinomas account for about 36% of carcinomas included in this study [1-3].

Materials and methods

Tissue-Sampling Techniques

Biopsy and brushing

By combining the various tissue biopsy techniques to the endoscopy visualisation, the utility of the endoscopy is enhanced one step ahead. In diagnosis of infections in the upper gastrointestinal tract is very difficult in past, but with the cytology of the tract by endoscopy is particularly useful for confirming the infections by bacterial, fungal and Helicobacter pylori infection. In upper gastrointestinal malignancy evaluation, it can add a yield of 10% to endoscopic biopsy alone. Particularly in the malignancy of upper gastrointestinal brush cytology is very useful as sensitivity is around 80% to 90% and specificity is around 100%. In case of touch

cytology the standard biopsy samples itself is processed by rolling it in the slide and fixing it for the study. In evaluation of malignancy and particularly the infections is the main advantage of this cytology. This technique can be used as an adjunct to the biopsy alone.

Standard biopsy techniques

In case of foregut malignancies the yield is maximum with standard biopsy as disease represents at the mucosal level. It yields a maximum results when targeted correctly. For H. Pylori disease evaluation it the diagnostic yields as been shown to be comparable in all parts of the stomach. A pediatric colonoscope can be used for the jejuna limbs, after gastrojejunostomy to enter into the jejunal limbs. Infected patients are being diagnosed by combining three biopsy samples, taken from the antrum pyloric region, near the incisura in lesser curvature and body of stomach greater curvature. With malignant lesions, when the biopsy are taken from the rim of the ulcers and also from the base with number of biopsies being 8 to 10, the yield is maximum in diagnosis. These specimens retained in the endoscopy channel can be used for the brush cytology and salvage cytology. With these procedures malignancy can be diagnosed upto 100%

1. Sheathed brush is passed through the endoscopic channel which is used for the brush cytology. Sheath is positioned near to the area to be examined and extend the brush [5]

A Cells in the brush can be dislodged by moving it vigorously to and fro.

B To avoid sample loss while the thing being withdrawn through the endoscope biopsy channel, just retract the brush from the sheath.

C The slides for the cytologic review is prepared with these samples. Another method of contributing to the sample yield is washing the brush in balanced salt solution. Forceps biopsy allows us to get adequate tissue (generally limited to the mucosa) for the diagnosis by histological examinations. Different kinds of biopsy for cepsare

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present, by choosing the proper forceps our intended sample can be obtained

1. Spiked forceps: In the single passage itself a chunk of multiple biopsies can be obtained by use of this forceps as it is provided with multiple tiny projections. The endoscopist's ability to get tissue that is oriented tangentially to the endoscope may be enhanced by helping the forceps to firmly engage the tissue to be sampled.

2. Large cupped forceps, or jumbo forceps: These forceps are very useful in taking large biopsies from the sites. It is useful when only a few attempts can be taken for a biopsy. For this large diameter endoscopes are needed for the passage of biopsy forceps.

3. Endoscopic mucosal resection: It is a very useful technique in case of early malignancies and also when large areas are to be examined. It helps in the complete removal of suspicious areas. It can be combined with the endoscopic ultrasound when it can also serve as a therapeutic tool too. In high incidence countries like Japan it is a very useful method. Hypertonic saline is used to elevate the lesion where the biopsy is targeted. It helps in raising the lesion and easy snaring of the lesion.

Disadvantages

Risk of bleeding

Perforation

Endoscopic Ultrasound: It has revolutionised the diagnostic technique in the gastro-intestinal malignancies particularly in the upper

digestive tract. This endoscopic ultrasound has a main role in staging of disease which can be difficult by any other investigations. Its main applications are in 1. diagnosis and staging of gastrointestinal cancers 2. submucosal pathology and biopsy. 3. Common bile duct stones and also in pancreatic malignancies. EUS-guided fine-needle aspiration cytology is gaining more importance in adjunct to the standard endoscopy. The yield in diagnosis is increased to 100% when it is combined with the aging is more accurate standard biopsy technique. It is more useful in:

1. Esophageal, pancreatic, gastric and even in pulmonary neoplasia.

2. Staging is more accurate as compared to the radiological investigations as CT and MRI [6]

3. The submucosal lesions are easily diagnosed by this endoscopic gas. The lesions are difficult to find in the radiologically in early stages. The stromal tumors, neuroendocrine tumors are easily identified and at the early stages with the use of this endo usg. Drawbacks are cost and long learning curve.

Consent: Written consent was obtained from the relatives of patients after explaining them the nature and purpose of the study. They were assured that confidentiality would be strictly maintained. The option to withdraw from the study was always open.

Observation chart

Table 1: Gastrointestinal Malignancies

Parts Of Upper GI	Number Of cases
Esophageal Carcinoma	18
Gastric Carcinoma	26
Periampullary Carcinoma	6

Table 2: Esophageal Carcinoma

Upper Esophagus	1
Middle Esophagus	8
Lower Esophagus	6

Table 3: Gastric Carcinoma

Site	Number of cases
Fundus	0
Body	3
Antropylorus	23

Gastric Carcinoma: Tumor location in the stomach has a greater significance for the diagnosis, management and prognosis of the patient. Proximal tumors of the stomach are more common in the western world due to increased incidence of gastroesophageal reflux disease and obesity in the western population. They account for more than half of the patients with gastric cancer. But the trend in the developing countries is totally different from the western world incidence of gastric cancers. It corresponds to the incidence of Helicobacter pylori

infection in the stomach. The type of tumor has also a greater impact on the outcome of patients. Distal gastric tumors are common in the developing countries like India where we have a higher incidence of Helicobacter pylori infection which correlates to the higher incidence of tumor in the distal stomach. The intestinal types are most common in the distal tumors of the stomach as compared to the diffuse type in the proximal tumors [7-10]

Table 4: Sex distribution in gastric carcinoma

Male	16
Female	10
Total	26

Table 5: Periampullary Growth

Male	3
Female	3
Total	6

Table 6: Esophageal Carcinoma

Squamous Cell Carcinoma	11
Adenocarcinoma	7

In the past squamous cell carcinoma accounted for more than 95% of cases but in recent years due to increased incidence of Barrett's esophagus the incidence of adenocarcinoma arising from it. In the United States squamous cell carcinoma is around 1.5 to 7 cases per 100,000 people. In India it is around 100 to 500 per 100,000 people. Squamous cell carcinoma is five times more common among Africans and Americans than in whites, whereas adenocarcinoma occurs

approximately three to four times more common in whites, particularly in men. In this study the incidence of squamous cell carcinoma is 69% and adenocarcinoma is 39%, which correlates with the Indian population. Squamous cell carcinoma is also common in the upper and middle esophagus and adenocarcinoma more common in the lower segment.

Table 7:Squamous Cell Carcinoma

Well Differentiated	0
Moderately Differentiated	11
Poorly Differentiated	0
Total	11

Table 8:Adenocarcinoma of esophagus

Well Differentiated	1
Moderately Differentiated	4
Poorly Differentiated	3

Table 9:Esophageal Carcinoma

	Squamous Cell Carcinoma	Adenocarcinoma
Upper	1	0
Middle	7	1
Lower	3	6

Table 10:Age wise distribution of upper GI malignancies

Age	Esophagus	Stomach	Periampullary
20-30	1	1	0
30-40	1	1	0
40-50	3	10	2
50-60	6	4	1
60-70	5	9	3
70-80	2	1	0
Total	18	26	6

Table 11:Yield in Specimens

Vial Number	No .of Positive Patients
First	47 OUT OF 50 (94%)
Second	44 OUT OF 50 (88%)
Third	27 OUT OF 35(77.14%)
Fourth	5 OUT OF 12 (41.66%)

Table 12:Positive numbers and Percentage

Vial Number	No.of Positive Patients	% of Positivity
I	47	94%
I+ II	50	100%
I+ II+III	50	100%
I+ II+ III+IV	50	100%

Result

Esophageal carcinomas has increased rapidly in the western countries in few decades. In the esophagus most common site of the malignancy is middle third of esophagus, as it continues in this study also but there is a alarming rise in the incidence of lower esophageal carcinomas. Its increase is greater than any other malignancy. Upper esophageal growth accounts for 6% of esophageal tumors which is the lowest and middle esophageal tumors accounts for maximum number with 44% of cases in this study. Lower esophageal growth consists of 33% of tumors in this study. OG junction tumors are include in these parate entity and account for 17% of tumors in this study. Proximal 5cm tumors of gastric is also included in the esophageal tumors only based on the new classification. Although the esophageal carcinomas are uncommon compared to other malignancies it is gaining importance due to advents in the endoscopic diagnosis and early interventional modalities. Sex distribution in the esophageal carcinomas is very important compared to other malignancies. Males are show a striking predominance in esophageal carcinomas for a long time. In white males esophageal adenocarcinomas was double that of the Hispanics and four fold higher than that of blacks. Asians, and native Americans. In all races the incidence of esophageal carcinoma is very less compared to the male incidence of tumor. Males have more incidence of squamous cell carcinomas compared to females. In this study also incidence is more in the males compared to females. Males account for 61% of esophageal tumors and females consist of 39% of tumors. Adenocarcinomas of cardia incidence is low among the API males compared with white males, but it was higher compared with black

males. However ethnic differences are important in the esophageal carcinoma because of increasing trend of adenocarcinoma of lower esophagus. In the esophagus the site of tumor mostly correlates with type of malignancy. Squamous cell carcinoma occurs most commonly in the upper and middle third of the esophagus and adenocarcinomas in the lower third of esophagus. In this study in upper third carcinomas all are squamous cell carcinoma and in middle third of esophagus most of the tumors are squamous cell carcinoma. In lower third of esophagus the incidence is changing as adenocarcinomas are more common than squamous cell malignancies. Pathology of the esophageal carcinoma is very important on which treatment is decided. Squamous cell carcinoma is more common and the histology of this is very important in prognosis and treatment. Mostly the tumors are well to moderately differentiated in squamous cell carcinoma. In this study all the patients had moderately differentiated carcinoma. Adenocarcinomas are increasing at a very high rate compared to any other malignancy. In this study also the adenocarcinomas are comparatively more in number. About 50% of adenocarcinomas are moderately differentiated and next to it 38% are poorly differentiated. Only 12% of tumors are well differentiated adenocarcinomas in this study. These differentiation affects the outcome and treatment. The incidence of esophageal adenocarcinoma increases with age, with a median age at diagnosis of 55 to 60 years and a striking male preponderance (7:1). The incidence stomach cancer also increases with age starting in the fourth decade of life and generally peaks in the seventh decade. In the United States, demographic risk factors for periampullary cancer include age, with the majority of patients in or beyond their sixth decade of life; sex, with a

slight male preponderance. In this study esophageal and stomach cancers are seen in age groups from 20 years to 80 years. Esophageal cancer occurring mostly in the fifth decade and stomach cancer in the fourth decade. Periapillary cancers are occurring from fourth decade to sixth decade, with peak during the sixth decade. In all the fifty patients four samples are obtained with few patients had the difficulty in getting after that. Most of the patients presented to us are with advanced lesions and there was bleeding while taking biopsies from the advanced lesions. So third vial specimen was taken for only 35 patients and fourth vial taken for 12 patients only. The patients who had bleeding are monitored continuously with fluids and vitals monitoring. Afterwards patients are discharged safely with instructions. The yield of endoscopic biopsy specimens from 50 patients after combining the results from successive vials. The yield in the first vial is 94% and the cumulative percentage of these cond vial yielded 100%, which means the malignancy in the specimen is proved without doubt in the first two vials itself for all the patients.

Statistical Analysis

Data was compiled using MS excel 2007 and analysis was done with the help of Epi-Info 7 software. Frequency and percentage were calculated & statistical test (Chi Square) was applied wherever applicable; $p < 0.05$ was taken as statistically significant.

Discussion

Gastric carcinomas account for the 52% of total malignancies in the upper gastrointestinal carcinomas. Esophageal carcinomas account for about 36% of carcinomas included in this study. Most of the carcinomas are moderately differentiated carcinomas. In esophagus most of the tumors are squamous cell carcinomas though there is increase in adenocarcinomas. In stomach most of the tumors are distal tumors in antro pyloric region and all are adenocarcinomas with predominantly moderately differentiated cancers. The endoscopy findings of most of the patients is advanced nodulo proliferative or ulcer o proliferative lesions.

Sancho Poch et al found that only one of 66 cases of gastric cancer in which eight specimens had been obtained was negative. However, these authors also did not take into account the order in which biopsy specimens had been taken. Misumi et al [13] found that the diagnostic accuracy was 100% in gastric cancers when six or more biopsy specimens were obtained. Most of the other studies are also in gastric cancers and the number of biopsy specimens varied from 4-10. There are many studies of the diagnostic accuracy of endoscopically performed cytological techniques in the diagnosis of carcinoma of the oesophagus. In these cytological studies, the various authors have either not mentioned the number of biopsy specimens or have taken varying numbers, leaving this to the judgement of the endoscopist rather than evaluating the optimal number of specimens needed to obtain the maximum yield. Graham et al conducted a study in which biopsy and cytology specimens were obtained from 202 consecutive patients, 27 of whom had carcinoma of the oesophagus. In each instance the authors obtained seven biopsy specimens in three groups. Group A contained the first biopsy specimen; group B biopsy specimens 2, 3, and 4; and group C biopsy specimens 5, 6, and 7. In Graham's study, the first biopsy yielded a correct diagnosis in 92-6% of patients with oesophageal cancer; with four specimens accuracy went up to 96% and with seven biopsy specimens it reached 96-3%. Seven biopsy and cytology specimens yielded a diagnosis in all the cases.

Conclusion

Our study differs from that of others in that we evaluated more patients and the diagnostic yield of four biopsy specimens was

100%; two specimens were placed in each of the four vials rather than one, three and three pieces in three vials. In conclusion, this study shows that four biopsy specimens are likely to yield a 100% diagnosis in advanced upper gastrointestinal malignancies. Whereas the endoscopic biopsy yield in the early stages of carcinomas should be evaluated in further studies.

References

- Graham DY, Schwartz JT, Cain GD, Gyorkey F. Prospective evaluation of biopsy number in the diagnosis of esophageal and gastric carcinoma. *Gastroenterology* 1982;82:228-31.
- Reilly RW, Kirner JB, Cockerhans L. Direct vision endoscopic cytology and biopsy in the diagnosis of oesophageal and gastric tumours. Current experience. *Acta Cytol* 1977; 21:399-403.
- Young JA, Hughes HE, Lee FD. Evaluation of endoscopic brush and biopsy touch smear cytology and biopsy histology in the diagnosis of carcinoma of the lower esophagus and cardia. *JT Clin Pathol* 1980;33:811-4.
- Kasugai T, Kobayashi, Nobuyoshi K. Endoscopic cytology of oesophagus stomach and pancreas. *Acta Cytol* 1978; 22: 327-31.
- Bhansali SK. Geographical distribution of gastrointestinal cancer in India. *Ind J Surg* 1968;30:33-7.
- Inawer SJ, Posner G, Lightdale CJ, Sherlock P, Melamed M, Fostner JG. Endoscopic diagnosis of advanced gastric cancer. Factors influencing yield. *Gastroenterology* 1975;69: 1183-7.
- Chambers LA, Clark WE. The endoscopic diagnosis of gastroesophageal malignancy: A cytological review. *Acta Cytol* 1986; 30:110-4.
- Bhasin DK, Kochhar R, Rajwansi A, Gupta SK, Mehta SK. Endoscopic Suction Cytology in Upper Gastrointestinal Tract Malignancy. *Acta Cytol* 1988;32:452-4.
- Bhasin DK, Raiwansi A, Kochhar R, Mehta SK. Brush cytology for colorectal cancer. *Lancet* 1989; i:1133-4.
- Kochhar R, Bhasin DK, Rajwansi A, Gupta SK, Malik AK, Mehta SK. Crush smears preparations of gastroesophageal malignancies. *Acta Cytol* 1990; 34: 214-6.
- Dekker W, Tytgat GN. Diagnostic accuracy of fiberendoscopy in the detection of upper intestinal malignancy. A followup analysis. *Gastroenterology* 1977;73:710-4.
- Sancho-Poch FJ, Balanzo J, Ocana J. An evaluation of gastric biopsy in the diagnosis of gastric cancer. *Gastrointest Endosc* 1978; 24: 281-2.
- Misumi A, Mori K, Ikeda T, et al. Evaluation of fiber gastroscopic biopsy in the diagnosis of gastric cancer. A study of 339 cases. *Gastroenterol Jpn* 1978;13:255-63.
- Hatfield ARW, Slavin G, Segal AW. Importance of the site of endoscopic gastric biopsy in ulcerating lesions of the stomach. *Gut* 1975; 16:884-6.
- Johansen A, Sikjaer B. Gastroscopic biopsy. Reliability of histological diagnosis with special reference to the single biopsy. *Sc and J Gastroenterol* 1975;10:453-8.
- Witzel L, Halter F, Grelillat PA, Scheurer U, Keller M. Evaluation of specific value of endoscopic biopsies and brush cytology for malignancies of the oesophagus and stomach. *Gut* 1976; 17:375-7.
- Keighley MRB, Thompson H, Moore J, et al. Comparison of brush cytology before or after biopsy of diagnosis of gastric carcinoma. *Br J Surg* 1979;66:246-7.

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