Agreement between different pathologists in histopathologic diagnosis of 128 gastric polyps

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ABSTRACT – Background – Gastric polyps are elevated mucosal lesions. Most of them are less than 1 cm and when larger than 2 cm, has a high malignancy probability. The histopathological types are mainly fundic gland polyps, hyperplastic polyps and adenomatous polyps. Objective – To evaluate the agreement between three different pathologists in the histopathological diagnosis of 128 biopsied gastric polyps in Digestive Endoscopy Unit from Walter Cantídeo University Hospital, between May 2010 to May 2012. Methods – To describe the intensity of agreement between observers, we use kappa index that is based on the number of concordant measures between them. Results – There was substantial agreement in the diagnosis of adenoma (kappa=0.799, CI: 0.899-0.698) and fundic glands (kappa=0.655, CI: 0.755-0.555). Regarding to hyperplastic polyps (kappa=0.415, CI: 0.515-0.315) and inflammatory (kappa=0.401, CI: 0.501-0.301), we obtained a moderate agreement. Regarding the presence of Helicobacter pylori in biopsy of the polyp, there was a low agreement (kappa=0.219, CI: 0.319-0.119). Conclusion – It is clear that the agreement between pathologists depends on the histological type of the biopsied polyp and this agreement is more substantial in adenoma, or fundic gland polyps.

INTRODUCTION

The term polyp is strictly used to describe lesions that originate from the epithelial layer of the stomach. Retrospective studies based on necropsies have estimated a prevalence of 0.12% to 0.8% of stomach polyps in the general population⁹. Studies performed in digestive endoscopy services find that stomach polyps have a prevalence of 3% to 8% in exams³⁸. Most patients carrying stomach polyps are asymptomatic, and they are generally found incidentally during an endoscopy exam. Nevertheless, some cases show severe clinical symptoms, such as abdominal pain, digestive hemorrhage, and obstructive symptoms⁴⁰. Endoscopically, polyps are usually all similar; thus, histopathological studies are fundamental to assess the malignancy risk⁹. The majority of stomach polyps are smaller than 1 cm (60%-82%) and those larger than 2 cm have a high probability of malignancy⁵⁰. The main histopathological types are fundic gland polyps (16%-41%), hyperplastic polyps (17%-71%), and adenomatous polyps (0.7%-23%)⁴⁰. In one Brazilian university hospital, Morais et al. showed that in 26,000 consecutive upper digestive endoscopies, there were 153 patients with gastric polyps. In this series, hyperplastic polyps were the most frequent and accounted for 71.3% of the cases, whereas fundic gland polyps accounted for 16.33% and adenomatous polyps for 12.4%⁴⁰. The fundic gland polyps are not aggressive, with dysplasia rarely being observed (1%)⁸. The hyperplastic polyps rarely affect the normal mucosa, are strongly related to the use of proton pump inhibitors (PPIs), and they rarely undergo neoplastic transformation⁹. On the other hand, adenomas, by definition, contain epithelial proliferative dysplasia and thus have a malignancy potential of around 30%⁹. The malignancy risk is associated with the size and degree of cellular atypia. Taking into account the malignancy potential, a detailed study of the mucosa in adenoma cases through complete resection of the lesion is necessary⁸.

The agreement among pathologists in sample analysis may be affected by observation bias, random errors of the observer, and sampling mistakes². There is a lack of studies assessing the correlation between pathologists in regard to endoscopic diagnoses. Therefore, we assessed the agreement between three experienced pathologists in the histopathological diagnosis of a series of 128 stomach polyps.

METHODS

This is a cross-sectional retrospective study based on information collected from databases at the endoscopy and pathology units of the Federal University of Ceará from May 26, 2010 to May 8, 2012. A total of 128 stomach polyps from 121 patients were analyzed. The study was approved by the ethics committee of the University Hospital under protocol number 45868215.7.0000.5045. Patients with stomach polyps who underwent upper digestive endoscopy and polypectomy with a biopsy clamp or diathermic loop, depending on the size of the lesion, were selected. By using a standardized questionnaire, the demographic, endoscopic, and...
histopathological aspects were analyzed. The quantity, location, and size of each polypoid lesion were measured. In addition, experienced pathologists meet to delineate the study, then evaluated the same same routine slide to make the histopathological subclassification of these polypoid lesions, according the classification described before(16). To evaluate the presence H pylori infection it was used the coloration of HE, and the giemsa. The pathologist looked independently the same routine slide. The gold-standard was consensus diagnosis.

We used the kappa coefficient to describe the agreement among the three experienced pathologists. It is based on the number of concordant measurements between the three observers. The kappa coefficient is a measure of interobserver agreement, and measures the degree of agreement beyond what would be expected by chance alone. This measure of agreement has a maximum value of 1, where a 1 represents total agreement, 0.80–1 corresponds to an excellent agreement, 0.60–0.79 to a substantial agreement, 0.40–0.59 to a moderate agreement, 0.2–0.39 a considerable agreement, 0–0.19 a slight agreement, and the values near or below 0, indicate no agreement, or the agreement was exactly as expected by chance(17).

RESULTS

In the present work, we analyzed 128 polyps, which were collected from 121 patients who underwent upper digestive endoscopy at Walter Cantídeo University Hospital from Federal University of Ceará. There were more women (72%) compared to men (28%). The majority had a single lesion (57%), but some had multiple lesions (43%). The polyps were found predominantly in the gastric body (38%) and antrum (23.5%), and more rarely found in the fundus (11%), fundus and body (8%), and antrum and body (9.5%). The majority of the polyps were smaller than 1 cm (90%).

Regarding the histological subtypes (consensus), we can see in Figure 1 that there is a higher prevalence of hyperplastic polyps (63.8%), followed by inflammatory polyps (18.0%), fundic glands (9.9%), and adenomas (8.3%).

In the evaluation by the pathologists, there was a substantial agreement in the diagnosis of adenomas (kappa=0.799, 95% CI: 0.899–0.698) and fundic glands (kappa=0.655, 95% CI: 0.755–0.555). Regarding the diagnosis of hyperplastic polyps (kappa=0.415, 95% CI: 0.515–0.315) and inflammatory polyps (kappa=0.401, 95% CI: 0.501–0.301), we observed a moderate agreement. Regarding the presence of Helicobacter pylori in in the biopsy of the polyp, we observed a low agreement (kappa=0.219, 95% CI: 0.319–0.119) (Table 1).

TABLE 1. Agreement among the three pathologists in the histopathological diagnosis of 128 polyps at the Endoscopy Unit of Walter Cantídeo University Hospital, Federal University of Ceará

<table>
<thead>
<tr>
<th></th>
<th>Kappa coefficient</th>
<th>CI</th>
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<tr>
<td>Adenoma</td>
<td>0.799</td>
<td>0.899–0.698</td>
</tr>
<tr>
<td>Fundic glands</td>
<td>0.655</td>
<td>0.755–0.555</td>
</tr>
<tr>
<td>Hyperplastic</td>
<td>0.401</td>
<td>0.515–0.315</td>
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<tr>
<td>Inflammatory</td>
<td>0.401</td>
<td>0.501–0.301</td>
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<tr>
<td>H. pylori</td>
<td>0.219</td>
<td>0.319–0.119</td>
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DISCUSSION

In our case series, there was a predominance of polyps involving the body (38%) followed by the antrum (23.5%), and they were less common in the fundus (11%). This data is congruent with the literature that shows a predominance of stomach polyps in the body (38% to 40%) and antrum (35% to 40%), and less commonly in the fundus (21% to 24%)18. Similarly, it has been established in the literature that the majority of stomach polyps are smaller than 1 cm (60% to 82%), with larger lesions being more uncommon (18% to 40%)19. This result is also congruent with our study where 90% of polyps were smaller than 1 cm. The small disparities found between our results and those in the literature may be due to the different types of studies performed, as well as due to the different populations studied, with distinct environmental characteristics, such as a prevalence of H. pylori10.

The hyperplastic polyps were the main histological type of our series and the main category of polyp with disagreement between the evaluators. Hyperplastic polyp rarely affects the normal gastric mucosa, being commonly related to chronic or atrophic gastritis, infection by H. pylori, or pernicious anemia13. Histologically, these are represented by elongated and grossly distorted hyperplastic foveolar dilatation, over the richly vascularized stroma, with different levels of inflammation, and it is difficult to distinguish gastritis and reactive foveolar proliferation from a hyperplastic polyp20. The hyperplastic polyps rarely undergo neoplastic transformation. A relationship between H. pylori and hyperplastic polyps was observed in more than 25% of cases. Randomized and controlled studies demonstrated that hyperplastic polyps resolve after eradication of infection13. Regarding the diagnostic agreement in the present study, there was a moderate agreement between the three pathologists in the histological type.
and a low agreement about the presence of infection by *H. pylori* in the biopsy sample of the polyp.

The fundic gland polyps represented 12% of our sample, and comprised, histologically, one or more oxyntic glands with cystic dilatation, covered by parietal cells or mucosa. These are rarely aggressive, with reports of dysplasia occurring in less than 1% of cases. Several authors suggested a relationship between the use of PPIs and the appearance of fundic gland polyps. The substantial agreement between pathologists regarding this histological type in our sample is probably due to the specific characteristics of this subtype of gastric polyp.

Despite being infrequent in our case series, the adenomatous polyps are very important since they are proliferous lesions; thus, they promote an increase in the number of tubular glandular structures and a proportional decrease in the quantity of interglandular stroma. We also observed a nuclear pseudostratification with a variable loss of polarization. The nuclei have an increased size, with a pleomorphism that varies from soft to severe, and more prominent nucleolus. The adenomas may be lesions promoting the formation of gastric adenocarcinomas and in 30% of cases, there are synchronous lesions. In polyps larger than 2 cm, there was a 50% rate of adenocarcinoma foci. Due to the high potential of malignancy, these lesions should be submitted for complete excision and follow up to ensure that there is no recurrence. In our work, there was a substantial agreement between pathologists for this histological type, with all lesions being removed completely without presenting dysplastic foci. The gastric adenomas have a high rate of malignant transformation, and a precise diagnosis of these lesions is fundamental, in order to perform a complete resection of these polyps. Thus, this fact demonstrates clearly the relevance of our study.

The low agreement in regard to *H. pylori* was mainly due to the use of PPIs by patients, since the use of this class of drugs is associated with a fluctuation in the number and distribution of *H. pylori*. Another factor to take into consideration is the drop in sensitivity of biopsies in patients with gastritis and high atrophy, regardless of the site subjected to biopsy. In addition, further histopathological analyses of the material through tissue stained with hematoxylin and eosin or Giemsa only, not subjected to silver staining, should be performed because it facilitates identification of the microorganisms on the slide.

Our study demonstrated that the agreement between observers depended on the histological type of the polyp subjected to biopsy, with high levels of agreement in the diagnosis of adenoma and fundic gland polyps.

**Authors’ contributions**

Barbosa SHB, Lázaro Filho GC and Souza MHLP: conception and design of research; Barbosa SHB, Lázaro Filho GC, Franco LM and Valença Junior JT: performed experiments; Barbosa SHB, Lázaro Filho GC, Nobre e Souza MA, and Souza MHLP: analyzed data; Barbosa SHB, Lázaro Filho GC, Nobre e Souza MA and Souza MHLP: interpreted results of experiments; Barbosa SHB and Souza MHLP: prepared figures; Nobre e Souza MA, Souza MHLP, Franco LM and Valença Junior JT, drafted manuscript; Barbosa SHB, Lázaro Filho GC, Nobre e Souza MA and Souza MHLP: edited and revised manuscript; Souza MHLP: approved final version of manuscript.
REFERENCES