The concordance of endoscopic and histologic findings of 530 pediatric patients suspicious for celiac disease in a referral center, Isfahan, Iran

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Introduction:
The purpose of this study is evaluation of concordance of histologic and endoscopic findings in pediatrics referred to Imam Hossein children's hospital, in Isfahan for evaluation of celiac disease. An accurate prediction of the likelihood of a histologic finding in the setting of a visually normal endoscopy can help prepare families for unanticipated results.

Patients and Methods:
This is a retrospective study conducted from 2015-2017. The sampling method is census and all eligible patients are included in the study. At the beginning of the work, obtaining consent for entering the project has been done through interviews. After that, patients information such as file number, age and gender according to the file were entered into the data gathering sheet. Data were analyzed using SPSS software version 22. P-value < 0.05 was considered significant.

Results:
Around 530 patients suspected of having celiac disease participated in the present study. The mean age of the subjects was 6.88 ± 4.31 years. Among them, 57.6% (305 patients) were girls and 42.4% (225 patients) were boys. According to the results, the kappa agreement coefficient is equal to 0.42. There was moderate agreement between endoscopy and pathology findings (kappa = 0.42).

Conclusion:
The results of the study and the kappa Cohen coefficient indicate the appropriateness of using endoscopy in patients with suspected celiac disease.

Key point
In a retrospective investigation on 530 patients suspected of having celiac disease, we found a moderate agreement between endoscopy and pathology findings (kappa = 0.42).

Abstract

Introduction: The purpose of this study is evaluation of concordance of histologic and endoscopic findings in pediatrics referred to Imam Hossein children's hospital, in Isfahan for evaluation of celiac disease. An accurate prediction of the likelihood of a histologic finding in the setting of a visually normal endoscopy can help prepare families for unanticipated results.

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disease can also be neglect if the disorder is not supposed and biopsy sampling not handled. Therefore, in these conditions, the role of the endoscopist is important, due to the significant impact of the macroscopic feature of the duodenum (14-16).

**Objectives**

Due to the above issues and the increasing number of patients with celiac disease, a comprehensive study with a higher sample size and a more standard method is necessary. Therefore, the purpose of this study is evaluation of concordance of histologic and endoscopic features in pediatrics suspicious for celiac disease who referred to an educational hospital in Isfahan.

**Patients and Methods**

**Study design**

This was a retrospective study conducted at Imam Hossein hospital from 2015-2017.

**Inclusion and exclusion criteria**

*Inclusion criteria:* Pediatric patients clinically suspicious for celiac disease, age between one month and 18 years.

*Exclusion criteria:* Having a defect in the file information, existence of other gastrointestinal diseases, and history of intervention that has changed the intestinal tissue.

**Sampling method and sample size**

The sampling method is census and all eligible patients were included in the study.

**Data collection**

At the beginning of the work, obtaining consent to enter the project has been done through interviews by the project executors. After that, for all patients suspicious of celiac disease, patients' information such as file number, age and gender according to the file were entered into the questionnaire. In celiac disease, if the results of blood tests show celiac disease, the patient undergoes an endoscopy to evaluate the small intestine and remove small tissue samples (biopsies). The sample is examined in a laboratory to determine the histologic status.

After collecting the necessary information, the results of histology and endoscopy in patients have been evaluated and compared. Positive histologic findings were classified according to the Marsh classification; Marsh I; increased number of intraepithelial lymphocytes (IELs) with normal mucosa architecture; Marsh II; proliferation of the crypts and increased number of IELs and Marsh III; destructive lesion including partial or complete villous atrophy and crypt hypertrophy with increased number of IELs. In cases who the pathology report is in favor of celiac disease but the diagnosis is not definitive, the status of their disease was determined by contacting patients.

**Ethical issues**

The research followed the tenets of the Declaration of Helsinki. The institutional ethical committee at Isfahan University of Medical Sciences approved all study protocols (IR.MUI.MED.REC.7397.231). Accordingly, written informed consent was taken from all participants before any intervention.

**Statistical analysis**

The collected data were imported into SPSS software version 22 and analyzed with appropriate statistical tests. The significance level in the present study was less than 0.05.

**Results**

Around 530 patients suspected of having celiac disease participated in the present study. The mean age of the subjects was 6.88 ± 4.31 years. Among them, 57.5% (305 patients) are girls and 42.4% (225 patients) are boys.

Based on endoscopic results, 24.1% (128 patients) of patients with suspected celiac disease were positive (reveal atrophy or scalloping) and 75.8% (402 patients) of patients with suspected celiac disease were negative. Additionally, based on the pathological results, 11.9% (63 patients) of patients with suspected celiac disease were positive (intraepithelial lymphocytosis and/or crypt hyperplasia and villi atrophy) and 88.1% (467 patients) of patients with suspected celiac disease were negative.

In group of positive histologic findings, 50 patients had Marsh 3 (10.5%), 6 patients Marsh 2 (1.2%) and 7 patients Marsh 1 (1.4%) of celiac disease.

**Table 1.** p Kappa agreement coefficient and sensitivity and specificity

<table>
<thead>
<tr>
<th>Pathology</th>
<th>Kappa agreement coefficient</th>
<th>Sensitivity</th>
<th>Specificity</th>
<th>PPV</th>
<th>NPV</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Endoscopy</strong></td>
<td></td>
<td></td>
<td></td>
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</tr>
<tr>
<td>Positive</td>
<td>0.42</td>
<td>0.73</td>
<td>0.83</td>
<td>0.38</td>
<td>0.96</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Negative</td>
<td>0.79</td>
<td>0.61</td>
<td>0.77</td>
<td>0.59</td>
<td>0.87</td>
<td>&lt;0.001</td>
</tr>
</tbody>
</table>

PPV, positive predictive value; NPV, Negative predictive value.
**Discussion**

Celiac disease is an autoimmune disease with significant prevalence in the general population that can manifest at any age and shows a rather varied clinical presentation. Due to the large number of patients with nonspecific gastrointestinal and extra-intestinal symptoms or even asymptomatic, diagnosis is often challenging (17). Currently, an increase in the prevalence of the disease in its atypical manifestations is well known and less than one third of adult patients present chronic diarrhea (18,19). Therefore, the aim of this study was to determine the agreement between endoscopic and pathological findings in patients with suspected celiac disease and to predict the likelihood of a positive histologic finding in the setting of a visually normal endoscopy.

The final result of the study showed that the agreement between the endoscopic and pathological findings was equal to 0.42. Besides, the sensitivity is equal to 0.68 and the specificity is equal to 0.76.

In follow up all the patients with positive histologic findings, had elevated serum transglutaminase antibody (tTG) and the diagnosis of Celiac, confirmed for them.

In a study by Kar et al, the correlation between endoscopic and histological findings was 66% in 400 patients.

Other studies in adults have suggested that routine duodenal biopsies be performed (20). Previous pediatric studies have supported routine biopsy in endoscopy. Studies in children have reported a mismatch between endoscopic and histological results.

In another study of 94 patients, the sensitivity and specificity of duodenal endoscopy were 82% and 27%, respectively, and gastric trunk endoscopy was 57% and 47% sensitive and specific (21).

In another study in Italy, endoscopy often provided inaccurate results with less deterioration than biopsy (22). Oderda et al found a 13.8% correlation between 32 biopsy responses and their endoscopic responses (23).

**Conclusion**

Based on the results of the study the amount of agreement between the endoscopic and pathological findings was equal to 0.44. The kappa Cohen coefficient indicates the appropriateness of using endoscopy in individuals with supposed celiac disease.

**Limitations of the study**

One of the limitations of the present study was the small sample size. It is suggested to use more sample size in other studies and also to use clinical trial and or follow-up studies.

**Authors’ contribution**

MA and AN and AB were the principal investigators of the study. MA and AN were included in preparing the concept and design. EP and AN revisited the manuscript and critically evaluated the intellectual contents. All authors participated in preparing the final draft of the manuscript, revised the manuscript and critically evaluated the intellectual contents. All authors have read and approved the content of the manuscript and confirmed the accuracy or integrity of any part of the work.

**Ethical considerations**

Ethical issues (including plagiarism, data fabrication, double publication) have been completely observed by the authors.

**Conflicts of interest**

The authors declare that they have no competing interests.

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None.

**References**


