Adenocarcinoma of the third and fourth portions of the duodenum: The capsule endoscopy value

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Abstract

Primary adenocarcinoma of the small intestine occurs in over 50% of cases in the duodenum. However, its location in the third and fourth duodenal portions occurs rarely and is a diagnostic challenge. The aim of this work is to report an adenocarcinoma of the third and fourth duodenal portions, emphasizing its diagnostic difficulty and the value of video capsule endoscopy. A man, 40 years old, with no medical history, with abdominal discomfort and progressive fatigue, presented four months ago with one episode of moderate melena. The physical examination was normal, except for mucosal pallor. Blood tests were consistent with microcytic, hypochromic iron deficiency anemia with 7.8 g/dL hemoglobin. Upper and lower endoscopy were normal. Additional work-up with video capsule endoscopy showed a polypoid lesion involving the third and fourth portions of the duodenum. Biopsy showed a moderately differentiated adenocarcinoma. Abdominal computed tomography showed a wall thickening from the third duodenal portion to the proximal jejunum, without distant metastasis. The patient underwent segmental resection (distal duodenum and proximal jejunum) with duodenojejunostomy. The surgical specimen histology confirmed the biopsy diagnosis, with transmural infiltration, without nodal involvement. Conclusion: Adenocarcinoma of the third and fourth portions of the duodenum is difficult to diagnose and capsule endoscopy is of great value.

Key words: Duodenum; Duodenal cancer; Adenocarcinoma; Endoscopy; Video capsule endoscopy

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Core tip: Third and/or fourth duodenal portion adenocarcinoma is a rare disease, associated with a vague clinical picture and a diagnostic challenge. Capsule endoscopy has shown a higher accuracy compared to conventional endoscopic methods. This case reports the occurrence of adenocarcinoma of the third and fourth duodenal portions and the value of capsule endoscopy to minimize the diagnostic difficulty.

Paquissi FC, Lima AHFBP, Lopes MFNV, Diaz FV. Adenocarcinoma of the third and fourth portions of the duodenum: The capsule endoscopy value. World J Gastroenterol 2015; 21(31): 9437-9441 Available from: URL: http://www.wjgnet.com/1007-9327/full/v21/i31/9437.htm DOI: http://dx.doi.org/10.3748/wjg.v21.i31.9437

INTRODUCTION

The small intestine is approximately 75% of the length and 90% of the mucosal surface of the gastrointestinal tract but represents only 2% to 5% of all primary malignant gastrointestinal tumors. Tumors in the small intestine are about 13 to 18 times less common than colon cancer; despite its exposure to a variety of endogenous and exogenous harmful substances. Histologically, there are four subtypes of malignant tumors of the small intestine: adenocarcinomas (around 40%); neuroendocrine tumors (35% to 40%); lymphomas (15%) and sarcomas (11% to 13%). When distributed by segments, adenocarcinomas are more common in the duodenum and proximal jejunum, neuroendocrine tumors and lymphomas are more common in the distal portions, while sarcomas have diffuse distribution.

Duodenal adenocarcinoma represents approximately 0.5% of all malignant gastrointestinal tumors and the most studied of them are those located in the first and second (most periampullary) portions. The location of this tumor in the third and/or fourth duodenal portion is rare, presents with non-specific symptoms and is of difficult diagnosis using conventional endoscopic methods. In this paper is presented an adenocarcinoma of the third and fourth portions of the duodenum, with review of the literature, emphasizing the difficulties and the value of video capsule endoscopy for diagnosis.

CASE REPORT

A man, 40 years old, with no medical history, presented with epigastric and mesogastric discomfort and progressive fatigue, with four months of evolution; and reported an episode of melena in moderate quantity. Physical examination was normal except for mucosal pallor. The laboratory findings were consistent with microcytic, hypochromic iron deficiency anemia with $3.49 \times 10^{12}$/mm$^3$ RBC, 7.8 g/dL hemoglobin, 27.5% hematocrit, 79 fl MCV, 22.5 pg MCH, 28.4 g/dL MCHC, 22.6% RDW, 915000/mm$^3$ platelets, and 13 ng/mL serum ferritin. Urine analysis, serum liver function test (LFT), hemolysis markers, and serum electrolytes were unremarkable. Upper gastrointestinal endoscopy (until the 2nd duodenal portion) and colonoscopy were normal. Further investigation, using video capsule endoscopy, in the outpatient setting, showed polypoid lesions involving the third and fourth portions of the duodenum (Figure 1). The biopsy showed a moderately differentiated adenocarcinoma. Abdominal CT showed a wall thickening involving the third and fourth portions of the duodenum and proximal jejunum, with no clear cleavage lines with adjacent structures without evidence of nodal and distant metastasis (Figure 2). The patient underwent a segmental resection of the duodenum (third and fourth portions) and proximal jejunum, with duodenoejejunostomy. The pathological examination of the surgical specimen confirmed a moderately differentiated adenocarcinoma infiltrating the wall of the organ without lymph node metastasis (Figure 3). The patient underwent a follow-up by oncology.

DISCUSSION

This case represents a rare location of primary duodenal adenocarcinoma in a younger patient compared to the average peak incidence of duodenal adenocarcinoma shown in literature (seventh decade of life), with a slight predominance for males. The patient was younger, and there was no known condition associated with early occurrence, such as inflammatory bowel disease, familial adenomatous polyposis, or hereditary nonpolyposis colorectal cancer, in which cancer presents earlier (median 39 years) in his personal and/or family medical history.

The clinical picture of adenocarcinoma in the third and fourth portions occurs with rather non-specific symptoms. Unlike periampullary tumors, whose main clinical picture is jaundice and other clinical aspects from the obstruction of the hepatobiliary-pancreatic system, in third and fourth portion tumors there are non-specific symptoms such as vague abdominal pain, weight loss, anemia symptoms, but no frank bleeding, and more rarely, bowel obstruction dominates the clinical picture. In this case, the duration of symptoms before diagnosis was 4 mo, that is within the average literature range (from 1.4 to 8 mo). One study showed worse 2-year survival rate associated with 4 mo or longer duration.

In routine work-up, both upper and lower endoscopy were normal. This situation is a substratum for missing tumors in the third and/or fourth portions, and is often worsened by the low index of clinical suspicion, which usually delays the diagnosis, resulting in advanced disease at diagnosis and decreasing
the rate of potentially curative resections\textsuperscript{[7,15]}. After nondiagnostic conventional endoscopic tests, in the setting of iron deficiency anemia, it is worth having a high index of suspicion for tumors beyond the second portion and to carry on the work-up using a method of greater accuracy for these tumors, the endoscopic capsule\textsuperscript{[8,16]}.

Capsule endoscopy is a non-invasive procedure that uses a wireless endoscopic device that allows imaging of the gastrointestinal tract. In this case, it was a valuable tool that allowed complete small bowel exploration in the ambulatory setting. The main indications for its use are the evaluation of obscure gastrointestinal bleeding, and Crohn's disease\textsuperscript{[17,18]}. Its sensitivity and specificity for diagnosing a small bowel tumor is 88.9\% to 95\% and 75\% to 95\% respectively, in the setting of obscure gastrointestinal bleeding\textsuperscript{[16,19]}. Tumors are found in about 3\% to 9\% of patients undergoing this procedure for evaluation of obscure gastrointestinal bleeding, and 50\% to 60\% were malignant\textsuperscript{[18]}. Video capsule endoscopy has also been used for the evaluation of patients with certain disorders that increase the risk of tumors of the small intestine, such as familial adenomatous polyposis\textsuperscript{[18]}.

Treatment of primary duodenal adenocarcinoma depends on the location and staging. In this case, a segmental resection of the duodenum (3\textsuperscript{rd} and 4\textsuperscript{th} portions) and the proximal jejunum (20 cm from duodenojejunal flexure), with primary duodenojejunostomy was performed. This approach was preferred to more extensive resection, because it provides equivalent survival rates to extensive resections (since it is possible to achieve negative margins), with the benefit of lower morbidity than that associated with pancreaticoduodenectomy\textsuperscript{[20]} and even better survival, as was shown in one study\textsuperscript{[21]}. Currently, extensive pancreaticoduodenectomy applies more to tumors of the proximal duodenum (1\textsuperscript{st} and 2\textsuperscript{nd} portions)\textsuperscript{[22]}.

The pathological examination of the surgical specimen confirmed a moderately differentiated adenocarcinoma, which is the most common histological grade\textsuperscript{[3,23]}, that infiltrates the three layers of the wall, without invasion of adjacent organs or metastasis.

\begin{figure}
\centering
\includegraphics[width=\textwidth]{figure1.png}
\caption{Capsule endoscopy findings in the 3\textsuperscript{rd} and 4\textsuperscript{th} portions of the duodenum. A: Polypoid and multilobular lesion; B, C: Partially obstructing the lumen; C, D: Ulcerated with low-flow bleeding.}
\end{figure}

\begin{figure}
\centering
\includegraphics[width=\textwidth]{figure2.png}
\caption{Abdominal computed tomography scan demonstrating a thickening of the wall involving the 3\textsuperscript{rd} and 4\textsuperscript{th} portions of duodenum, narrowing its lumen (arrows), without clear lines of cleavage with adjacent structures and with no evidence of nodal and distant metastasis.}
\end{figure}
to the lymph nodes, and surgical margins were negative for tumor cells. Therefore, it was a stadium II tumor (T3 N0 M0), that is the most frequent stage for adenocarcinoma in this site\(^{12,24}\). Despite negative margins and no lymph node involvement, the combination of 3 variables present in this case - tumor extension, histological grade and transmural invasion - are associated with poor prognosis\(^{6}\).

With regard to the adjuvant treatment, there is no established protocol for small bowel adenocarcinomas, due to the lack of randomized trials\(^{25}\); and the few available data from retrospective studies have shown no statistically significant overall survival benefit\(^{26}\). As with treatment, there is no established follow-up protocol for patients with resected adenocarcinoma of the small intestine. In this case, the patient continued follow-up by oncologist.

Malignant tumors of the small intestine, although rare, should be part of the differential diagnosis in the investigation of obscure gastrointestinal bleeding and the high index of suspicion and appropriate use of endoscopic capsule are of great value.

**COMMENTS**

**Case characteristics**

A 40-year-old male presenting with abdominal discomfort and progressive fatigue due to severe anemia by continuous bleeding from third and fourth portions duodenal cancer.

**Clinical diagnosis**

Small intestine examination with video capsule endoscopy revealed a multilobular tumor, ulcerated with low-flow bleeding lesion in the third and fourth duodenal portions.

**Differential diagnosis**

Upper gastrointestinal endoscopy (until the 2\(^{nd}\) duodenal portion) and colonoscopy were performed to rule out stomach and colon bleeding respectively.

**Laboratory diagnosis**

Blood tests demonstrated RBC 3.49 × 10\(^6\)/mm\(^3\); hemoglobin 7.8 g/dL; MCH 79 fl and serum ferritin 13 ng/mL. Metabolic panel and liver function tests were within normal limits.

**Imaging diagnosis**

Abdominal computed tomography demonstrated a wall thickening involving third and fourth duodenal portions and proximal jejunum, without evidence of nodal and distant metastasis.

**Pathological diagnosis**

The histopathological examination of the surgical specimen demonstrated a three layer infiltrating adenocarcinoma, without lymph node invasion, with free surgical margins.

**Treatment**

Segmental resection of the duodenum (3\(^{rd}\) and 4\(^{th}\) portions) and proximal jejunum (20 cm from duodenojejunal flexure) was performed, with primary duodenojejunostomy.
Experiences and lessons
Malignant tumors of the third and fourth duodenal portions are a diagnostic challenge using conventional endoscopic tests; a high index of suspicion and appropriate use of the endoscopic capsule is of great value for early diagnosis.

Peer-review
The authors reported primary adenocarcinoma of the 3rd/4th portions of the duodenum in a 40-year-old man. Blood tests revealed microcytic, hypochromic iron deficiency anemia. Upper gastrointestinal endoscopy and colonoscopy were normal, but the polypoid lesions with low-flow bleeding were observed by video capsule endoscopy.

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P-Reviewer: Charoenphandhu N  S-Editor: Ma YJ  L-Editor: O’Neill M  E-Editor: Liu XM
