# World Journal of Gastrointestinal Endoscopy

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CASE REPORT

# Giant Brunner's gland hyperplasia of the duodenum successfully resected en bloc by endoscopic mucosal resection: A case report

Makomo Makazu, Akiko Sasaki, Chikamasa Ichita, Chihiro Sumida, Takashi Nishino, Miki Nagayama, Shinichi Teshima

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# **Abstract**

#### **BACKGROUND**

Duodenal Brunner's gland hyperplasia (BGH) is a therapeutic target when complications such as bleeding or gastrointestinal obstruction occur or when malignancy cannot be ruled out. Herein, we present a case of large BGH treated with endoscopic mucosal resection (EMR).

# CASE SUMMARY

An 83-year-old woman presented at our hospital with dizziness. Blood tests revealed severe anemia, esophagogastroduodenoscopy showed a 6.5 cm lesion protruding from the anterior wall of the duodenal bulb, and biopsy revealed the presence of glandular epithelium. Endoscopic ultrasonography (EUS) demonstrated relatively high echogenicity with a cystic component. The muscularis propria was slightly elevated at the base of the lesion. EMR was performed without complications. The formalin-fixed lesion size was 6 cm × 3.5 cm × 3 cm, showing nodular proliferation of non-dysplastic Brunner's glands compartmentalized by fibrous septa, confirming the diagnosis of BGH. Reports of EMR or hot snare polypectomy are rare for duodenal BGH > 6 cm. In this case, the choice of EMR was made by obtaining information on the base of the lesion as well as on the internal characteristics through EUS.

# CONCLUSION

Large duodenal lesions with good endoscopic maneuverability and no evident muscular layer involvement on EUS may be resectable via EMR.

Key Words: Duodenum; Brunner's gland hyperplasia; Brunner's gland hamartoma; Brunner's gland adenoma; Endoscopic mucosal resection; Case report

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Core Tip: This is a report of a large Brunner's gland hyperplasia in an older female patient with anemia that was successfully resected en bloc using endoscopic mucosal resection (EMR). After obtaining information on the base of the lesion as well as on the internal characteristics using endoscopic ultrasonography, we chose EMR as a minimally invasive treatment.

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# INTRODUCTION

Brunner's glands are exocrine glands located in the duodenum, extending from the mucosal to the submucosal layer, which secrete alkaline mucus. Brunner's gland hyperplasia (BGH) is characterized by the proliferation of Brunner's gland without histological atypia. There are proposals to classify relatively large proliferative disorders of Brunner's glands as Brunner's gland hamartomas and smaller ones as BGH; however, there is no established consensus[1,2]. Since the term "hamartoma" is traditionally used for congenital disorders, we have used the term "Brunner's gland hyperplasia" in our study.

BGH is targeted for treatment when complications such as bleeding or gastrointestinal obstruction occur or when the existence of a malignant component cannot be ruled out. Although surgery is often the preferred treatment for large lesions, several recent reports have described endoscopic submucosal dissection (ESD)[3]. However, ESD for duodenal lesions involves longer treatment time and a higher complication rate[4].

To the best of our knowledge, few reports on the criteria for large BGHs suitable for endoscopic mucosal resection (EMR) have been published. This report aims to present a case of large BGH treated with EMR, as well as to illustrate the criteria for determining the appropriateness of EMR.

# CASE PRESENTATION

#### Chief complaints

An 83-year-old woman presented at our hospital with dizziness and anemia.

#### History of present illness

She had been complaining of dizziness for approximately two weeks.

# History of past illness

Her medical history included dementia, diabetes mellitus, dyslipidemia, hypertension, and an appendectomy. The patient was not taking any antithrombotic drugs.

#### Personal and family history

The personal and family history of the patient was unremarkable.

# Physical examination

The patient's vital signs were stable.

# Laboratory examinations

Laboratory tests revealed normocytic normochromic anemia with a hemoglobin level of 5.6 g/dL.

# Imaging examinations

Esophagogastroduodenoscopy (EGD) identified a 6.5 cm lesion protruding from the anterior wall of the duodenal bulb (Figure 1A and B). The lesion surface was mostly covered with normal mucosa; however, a depression with a mucosal defect was observed in a portion of the lesion (Figure 2A). Magnified narrowband imaging of this depressed area revealed structures resembling small round glandular openings and band-like septa suggestive of fibrous components



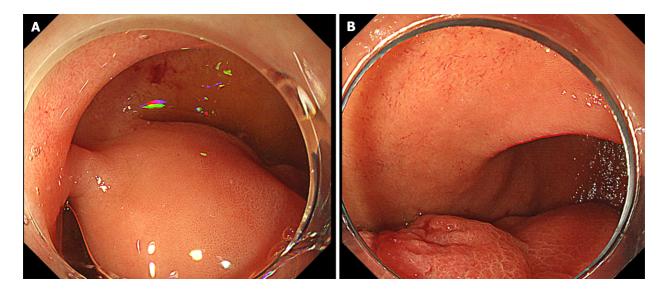


Figure 1 Endoscopic images of the duodenal lesion. A: Endoscopic image of the lesion; B: Endoscopic image of the middle to the tip of the

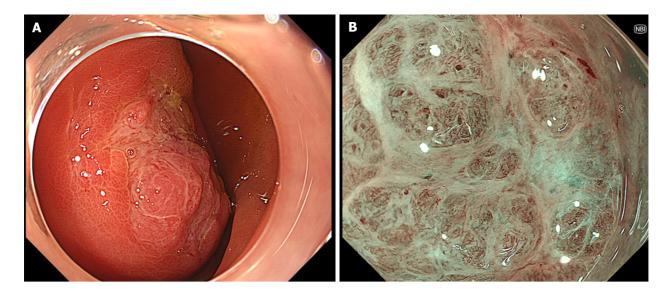


Figure 2 A depression with a mucosal defect in the lesion. A: In a portion of the lesion, a depression with a mucosal defect was seen; B: Magnified narrow-band imaging observation of this depressed area revealed structures resembling small round glandular openings and band-like septum-like structures suggestive of fibrous components.

(Figure 2B). A boring biopsy of the depressed area revealed the presence of glandular epithelium. Endoscopic ultrasonography (EUS) revealed that the lesion had a relatively high echogenicity with an internal cystic component (Figure 3A). At the base of the lesion, a slight elevation of a low-echogenicity layer, suggestive of the muscular layer, was observed (Figure 3B).

Abdominal contrast-enhanced computed tomography (CT) revealed a 7 cm protruding lesion extending from the bulb to the descending portion of the duodenum (Figure 4). The lesion showed minimal enhancement in the arterial phase and a mosaic-like enhancement pattern in the venous phase.

# **FINAL DIAGNOSIS**

Differential diagnoses included BGH, gastrointestinal stromal tumor (GIST), and cancer. However, considering endoscopic morphology, biopsy results, and EUS findings, the likelihood of malignancy was low.

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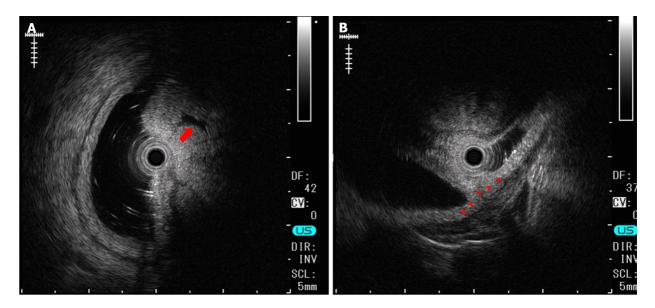


Figure 3 Endoscopic ultrasonography. A: Endoscopic ultrasonography revealed that the lesion had relatively high echogenicity with a cystic component (arrow) inside; B: At the base of the lesion, a slight elevation of a low echogenicity layer that was suggestive of the muscular layer was seen (arrowhead).

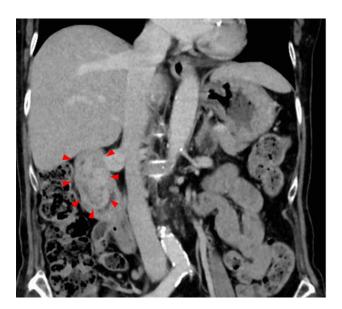


Figure 4 Abdominal contrast-enhanced computed tomography. It revealed a 7 cm protruding lesion (arrowhead) hanging from the bulb to the descending portion of the duodenum. The lesion showed minimal enhancement in the arterial phase and a mosaic-like enhancement pattern in the venous phase.

# **TREATMENT**

Upon considering the following conditions associated with the present case, we chose EMR as the treatment method: preference for a non-invasive approach due to the patient's advanced age; EUS and biopsy findings suggesting a low probability of malignancy; the relatively favorable maneuverability of the endoscope; the lesion having a wide base but being thin, with visibility across its entire circumference upon endoscopic examination; and minimal retraction of the muscle layer into the lesion as observed on EUS.

We injected hypertonic saline epinephrine solution at the base of the lesion and grasped it with a snare measuring 33 mm in diameter (Captivator II; Boston Scientific, Tokyo, Japan). We attempted to excise the lesion using the ENDO CUT Q mode (effect, 3; cut duration, 2; cut interval, 4) of the high-frequency generator VIO300D (Erbe Elektromedizin, Tübingen, Germany); however, there was no electrical activation. We relocated the constriction site with the snare slightly away from the base. We attempted to resect the lesion again in the same ENDO CUT Q mode as before (Figure 5A). At this time, satisfactory electrical conduction was achieved, and the lesion was excised entirely. There were no macroscopic residual tumors on the ulcerated surface after lesion excision and no muscle layer injuries or perforations (Figure 5B). The excised lesion was grasped using a snare and extracted orally. The ulceration after excision was completely closed using endoclips (Figure 5C).

Figure 5 Endoscopic mucosal resection of the lesion. A: After injection of hypertonic saline epinephrine solution at the base of the lesion, we grasped it with a snare. We attempted to excise the lesion using the cutting mode of the high-frequency generator. There was no electrical activation the first time. After relocating the constriction site with the snare slightly away from the base, we achieved satisfactory electrical conduction, and the lesion was excised entirely; B: There was no macroscopic residual tumor on the ulcerated surface after excising the lesion, and there were no muscle layer injuries or perforations; C: The ulceration after excision was completely closed using endoclips.

# **OUTCOME AND FOLLOW-UP**

EGD on the day following treatment showed that the resection site was maintained with clip suturing, with no bleeding or perforation. The patient resumed diet on the second day after treatment and was discharged without any problems on the third day. Thereafter, the patient experienced no recurrence of anemia.

The size of the lesion after formalin fixation was 6 cm  $\times$  3.5 cm  $\times$  3 cm (Figure 6A). Histopathological examination revealed nodular proliferation of the Brunner's glands without atypia, partitioned by fibrous septa (Figure 6B). This site was positive for MUC6 (Figure 6C). The pathological diagnosis was BGH. The area observed as a depressed region during EGD showed superficial erosion and a regenerating epithelium (Figure 6D). At the resection margin, thickened muscularis mucosa, fibrosis in the submucosal layer, and normal Brunner's glands were observed, whereas the Brunner's gland hyperplastic area was completely excised (Figure 6E).

#### DISCUSSION

We encountered a large BGH in an older female patient with anemia and successfully resected it en bloc using EMR. After obtaining information about the internal characteristics of the lesion and its base using EUS, we chose EMR as a minimally invasive treatment.

Several differential diagnoses should be considered when a subepithelial lesion (SEL) is identified in the duodenum. Malignant conditions include cancer and GIST, whereas benign conditions include lipomas and BGH. The diagnostic rate of boring biopsy for gastrointestinal SEL has been reported to be approximately 25%-65% [5,6]. EUS-guided sampling has been reported to account for approximately 60%-70% of EUS-guided fine-needle aspiration (FNA) and 80%-90% of EUSguided fine-needle biopsy (FNB)[7,8]. In the present case, we did not perform EUS-FNA/FNB after obtaining the pathological result of "glandular epithelium" on a boring biopsy. This is because the EUS findings inside the lesion did not strongly suggest the presence of a malignant disease.

EUS can delineate the origin layer and the internal characteristics of SELs. This information helps hypothesize the nature of such SEL. BGH originating from the third layer is often depicted on EUS. In the present case, the lesion originated in the third layer. In terms of the EUS image of BGH, Hizawa et al[9] described, in a summary of six cases of BGH, that the EUS features of Brunner's gland hamartoma are of a heterogeneous solid or cystic mass within the submucosa. Additionally, in their study of EUS findings in 10 cases of BGH, Lee et al[10] classified them into three types: (1) Heterogeneous appearance with large cysts; (2) heterogeneous echogenicity without cystic lesions; and (3) heterogeneous mixed echogenicity with very small cysts. They stated that these findings reflect diverse histological images of the BGH. In the present case, the lesion exhibited relatively high echogenicity on EUS with internal cystic components. This finding is consistent with previously reported EUS findings of BGH.

Methods of BGH resection include surgery and endoscopic treatment. Although numerous cases of BGH resection have been reported, there are no established guidelines for selecting the appropriate resection method. When the possibility of concomitant malignancy cannot be ruled out or when endoscopic maneuverability is poor, surgery is considered the preferred approach to BGH treatment. Factors such as the patient's overall condition, comorbidities, and preferences also influenced the treatment choice. Historically, large BGHs have primarily been resected surgically[11]. In older adult patients, as in the present case, surgery may be excessively invasive. Compared to surgery, endoscopic treatment of duodenal SELs offers advantages such as less invasiveness, shorter hospital stays, and lower complication rates[12]. In recent years, there have been several reports of ESD for BGH[13]. ESD for non-ampullary duodenal epithelial tumors has a higher en bloc resection rate than EMR but is associated with longer treatment times and higher complication rates[14]. In the present case, we chose EMR for a large BGH. One of the bases for this decision was EUS finding of the lesion. Based on the EUS findings, it was possible to assess that the involvement of the muscle layer in the lesion was

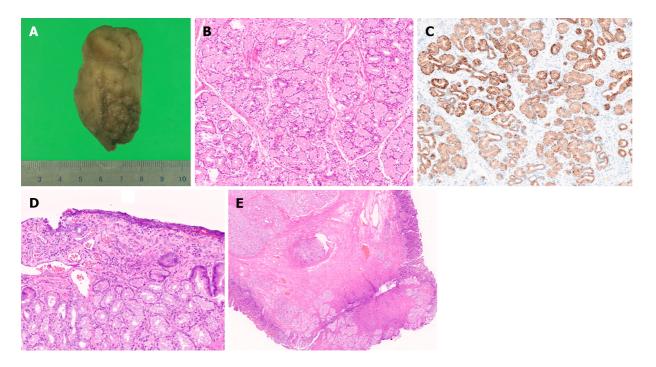


Figure 6 Pathological findings of the lesion. A: The lesion size after formalin fixation was 6 cm × 3.5 cm × 3 cm; B: Histopathological examination revealed nodular proliferation of Brunner's glands without atypia, partitioned by fibrous septa; C: This site exhibited positive staining for MUC6; D: The area observed as a depressed region during EGD showed superficial erosion and regenerating epithelium; E: At the resection margin, thickened muscularis mucosa, fibrosis in the submucosal layer, and normal Brunner's glands were observed.

minimal. From this finding, we concluded that it was possible to safely perform resection with a snare by temporarily increasing the volume of the submucosal layer with hypertonic saline without causing thermal damage to the muscle layer. In practice, during the first snaring attempt, we were unable to achieve satisfactory electrical conduction; however, on the second attempt after repositioning the snare, we successfully achieved effective electrical conduction and were able to cut the lesion. The pathological specimen showed normal Brunner's glands, submucosal fibrosis, and thickening of the muscularis mucosae at the resection margin. The "hardness" of the tissue due to these factors may have prevented adequate tissue strangulation by the snare, resulting in the inability to achieve effective electrical conduction on the first try. Based on this experience, we believe that if prior EUS confirms the absence of obvious muscular layer involvement, even if effective electrical conduction cannot be achieved with initial strangulation by the snare, it may be possible to resect the lesion by changing the snare position, considering the influence of duodenal specific structures such as Brunner's glands.

According to a search in MEDLINE, there were seven reported cases in which EMR or hot snare polypectomy was performed for BGH > 6 cm[15-21] (Table 1). Some of these were described as Brunner's gland hamartomas or adenomas; however, based on the histopathological findings, they were all considered synonymous with BGH. Although some cases only reported the longest axis, the images suggested that the diameter of the base or neck of the lesion was less than 3 cm. EUS was performed in only one other case besides the present case, where the absence of muscle layer invasion was used to aid in the choice of treatment[20]. In cases meeting conditions such as relatively good endoscopic maneuverability, non-broad-based lesions, and lack of significant findings of muscular layer involvement on EUS, EMR could be a viable option even for large lesions.

# CONCLUSION

We encountered a large BGH in an older female patient with anemia that was successfully resected en bloc using EMR. By obtaining information about the internal characteristics of the lesion as well as its base through EUS, we were able to select a minimally invasive treatment option. Because BGH is a benign lesion, the possibility of minimally invasive treatments through imaging diagnosis, such as EUS, should be explored, especially in older adult patients.

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Ref.	Age	Sex	Location	EUS	Size (mm)	Interventional approach
Tai et al[15], 2001	62	M	$1^{st}$	-	105	HSP
Stermer et al[16], 2006	20	F	2 <sup>nd</sup>	-	70	HSP
Itaba et al[17], 2006	75	F	2 <sup>nd</sup> -3 <sup>rd</sup>	-	60×9	HSP
Chen et al[18], 2006	78	F	$1^{st}$	-	60×4×2	HSP following endoclip
Kitagawa et al[19], 2018	64	F	$1^{st}$	-	70×20	EMR following endoclip and endoloop
Pasetti et al[20], 2018	67	M	$1^{st}$	+	60×30×20	HSP
Yi et al[21], 2019	63	F	$1^{st}$	-	70×30×16	HSP following endoloop
Our case	83	F	1 <sup>st</sup>	+	60×35×30	EMR

EUS: Endoscopic ultrasonography; HSP: Hot snare polypectomy; EMR: Endoscopic mucosal resection.

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