

## Clinical impact of multidetector computed tomography before double-balloon enteroscopy for obscure gastrointestinal bleeding

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### Abstract

**AIM:** To evaluate the clinical impact of multidetector computed tomography (MDCT) before double-balloon enteroscopy (DBE) for patients with obscure gastrointestinal bleeding (OGIB).

**METHODS:** A retrospective analysis of prospectively collected cases with DBE and MDCT for overt OGIB was conducted from April 2004 to April 2010 at Changhua Christian Hospital. We evaluated the clinical impact of MDCT on the subsequent DBE examinations and the diagnostic yields of both MDCT and DBE respectively.

**RESULTS:** From April 2004 to April 2010, a total of 75 patients underwent DBE for overt OGIB. Thirty one cases received MDCT followed by DBE for OGIB. The overall diagnostic yields of DBE and MDCT was 93.5% and 45.2%. The MDCT had a high diagnostic yield of tumor

vs non-tumor etiology of OGIB (85.7% vs 33.3%,  $P = 0.014$ ). Additionally, the choice of initial route of DBE was correct in those with a positive MDCT vs negative MDCT (100% vs 52.9%,  $P = 0.003$ ).

**CONCLUSION:** This study suggests MDCT as a triage tool may identify patients who will benefit from DBE and aid the endoscopist in choosing the most efficient route.

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**Key words:** Multidetector computed tomography; Capsule endoscopy; Double-balloon endoscopy; Obscure gastrointestinal bleeding

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### INTRODUCTION

Obscure gastrointestinal bleeding (OGIB) is defined by the American Gastroenterological Association (AGA)<sup>[1]</sup> as bleeding of unknown cause after upper or lower endoscopy. OGIB constitutes approximately 5% of patients evaluated for gastrointestinal hemorrhage<sup>[2]</sup>. OGIB can be further classified into two clinical forms: (1) obscure-occult bleeding; and (2) obscure-overt type bleeding (those with hematochezia or melena). The management of overt OGIB is clinically challenging for the gastroenterologist. With the introduction of high yield diagnostic tools such

as capsule endoscopy (CE) and double-balloon endoscopy (DBE) since the year 2000, the AGA recommended the use of CE followed by DBE as the first line diagnostic tool in 2007<sup>[2]</sup>. CE has exhibited some limitations, such as capsule retention, incomplete examination, inability to provide therapy, and high cost<sup>[3]</sup>. Compared with CE, DBE offered a therapeutic ability which is more useful in the setting of overt OGIB and is cost-effective compared with CE<sup>[4]</sup>. Multidetector computed tomography (MDCT) is a non-invasive tool and some recent reports have used it to investigate OGIB<sup>[3,5-7]</sup>. The clinical impact of MDCT prior to DBE for OGIB has not been evaluated. Thus, we performed this study to evaluate, retrospectively, the role of MDCT and DBE in patients with OGIB.

## MATERIALS AND METHODS

### Patients

A retrospective review of the medical records of Changhua Christian hospital from April 2004 to April 2010 identified 75 patients who underwent DBE for OGIB. A retrospective analysis of this prospectively collected database of DBE identified 31 patients who received MDCT within one month prior to DBE for obscure overt gastrointestinal (GI) bleeding and were included for this analysis. OGIB was defined as gastrointestinal bleeding after a non-diagnostic upper and lower endoscopy. All patients provided written consent to undergo MDCT and endoscopy, including endoscopic treatments, such as hemoclip and argon plasma coagulation. All patients were informed that endoscopic examination and treatment are involved in the current standard therapeutic approach used in the evaluation of OGIB. The primary end point of the study was to evaluate the clinical impact of MDCT on the subsequent DBE examinations. The secondary end point of the study was the diagnostic yield for MDCT and DBE for the diagnosis of OGIB.

### MDCT procedure

All the patients received MDCT with non-enhanced and triphasic helical computed tomography (CT) scanning<sup>[8]</sup>. No oral contrast material was given before the examination. First, patients were imaged with a MDCT scanner (LightSpeed Ultra 16, GE Medical Systems, Milwaukee, WI) in a craniocaudal direction beginning at the dome of the liver. A nonionic contrast medium (Optiray 350, Tyco Healthcare, Mansfield, MA) was then administered at a total dose of 100 to 120 mL with an injection rate of 3 mL/s through an antecubital vein. For triphasic acquisitions, scanning was started with a 10 s scan delay for the hepatic arterial phase after the attenuation value of the aorta reached 120 HU. Fifteen seconds after the end point of the hepatic arterial phase, the scans for the portal venous phase were acquired. Delayed-phase images were acquired 80 s after the end of the acquisition of the portal venous phase. Whole scanning was completed in 4 to 8 s with the patients holding their breath.

The MDCT was reviewed by one radiologist (Dr.

Liu) with 10 years of experience with abdominal imaging. The finding of the presence of small intestinal neoplasm (Figure 1), active contrast extravasation (Figure 2) or hyperdense fluid accumulation was considered to be diagnostic of GI bleeding<sup>[9]</sup>.

### DBE procedure

The double-balloon method was developed as described by Yamamoto *et al*<sup>[10]</sup>. Briefly, the double-balloon endoscope (EN-450P5 or EN-450T5, Fujinon Co., Japan) had two balloons, one attached to the distal end of the scope and the other attached to a transparent overtube (length 140 cm). Through a technique of inflating and deflating the balloons, the 7 m long small intestine can be shrunk to less than 2 m. Therefore, the entire small intestine could be theoretically examined. Fluoroscopy was required to guide the insertion or withdrawal of the endoscope. An oral route or anal route was chosen depending on clinical suspicion of the lesion. The endoscopic examination was stopped when (1) a lesion was found; (2) the endoscope was unable to be inserted or (3) due to patient intolerance. All the patients were admitted and received conscious sedation with intravenous midazolam and meperidine. During the procedure, the patients were monitored with an oximeter, EKG, and blood pressure monitor. Oxygen *via* nasal cannula was provided, as necessary.

### Statistical analysis

All data were analyzed with SPSS 16.0. All quantitative data were expressed as mean  $\pm$  SD. The  $\chi^2$  test was used to compare two categorical data with  $P < 0.05$  being considered statistically significant.

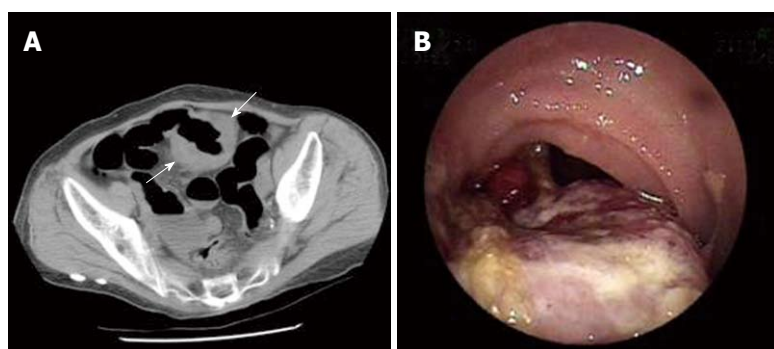
## RESULTS

### Clinical features

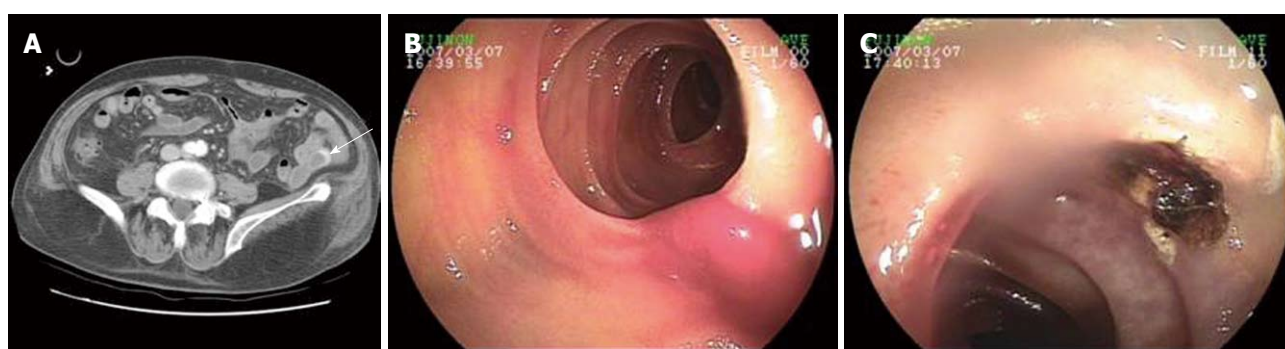
From 2004 to 2010, a total of 75 patients underwent DBE for OGIB and, of these, 31 patients were included in this analysis. These patients had a mean age of 68.6 years. Their mean hemoglobin levels were of 7.95 g/dL. Fourteen patients exhibited shock (defined as systolic blood pressure  $< 90$  mmHg or drop of systolic pressure  $> 40$  mmHg) during the presentation. The median time between the MDCT and DBE procedures was 2 d (range from 0 d to 12 d). The source attributed to OGIB was found in 93.5% of these 31 patients. Two patients had no diagnosis made after both DBE and CT examination and had no further bleeding during the follow-up period (Table 1).

### Diagnostic yield of DBE and MDCT

The overall diagnostic yield of DBE was 93.5% and CT was 45.2%. The diagnostic yield of CT depends on the etiology: 100% (2/2) of bleeding angiodysplasia, 85.7% (6/7) of those had bleeding from tumors, 41.7% (5/12) of those from diverticulosis, 14.3% (1/7) of those from ulcers and 0% of those from lymphangiectasia (0/1). One patient had early colon cancer diagnosed by DBE, though missed by CT. The MDCT diagnostic yields were



**Figure 1 Multidetector computed tomography finding of small bowel neoplasm.** A: Abdominal multidetector computed tomography performed after colonoscopy. Abundant bowel gas was observed as a contrast to distend the small intestinal lumen. A focal bowel thickening was discovered (arrows); B: Anal route double-balloon endoscopy disclosed ileal lymphoma accounting for the bowel wall thickening.



**Figure 2 Multidetector computed tomography finding of small bowel bleeding.** A: Multidetector computed tomography disclosed active contrast extravasations in the jejunum (arrow); B: Oral route double-balloon endoscopy disclosed active bleeding angiodysplasia; C: Endoscopic view after control of bleeding.

**Table 1 Clinical features of patients (mean  $\pm$  SD) *n* (%)**

Item	Value
Sex (M/F)	13/18
Age (yr)	68.6 $\pm$ 11.3
Underlying diseases	
Hypertension	23 (74.2)
DM	4 (12.9)
Chronic renal failure	5 (16.1)
Cirrhosis	2 (6.5)
Abdominal operation	1 (3.2)
NSAID/aspirin use	9 (29.0)
Hemoglobin level (g/dL)	7.95 $\pm$ 1.62
Hospital stay (d)	12.8 $\pm$ 10.8
Shock at presentation	14 (45.2)
Median time between DBE and MDCT (d)	2 (0-12)
Final diagnosis	
Ulcers	7
Tumors	7
Angiodysplasia	2
Diverticulosis	10
Lymphangiectasia	1
Undiagnosed	2

DM: Design management; NSAID: Nonsteroidal antiinflammatory drugs; DBE: Double-balloon endoscopy; MDCT: Multidetector computed tomography.

not different according to the hospital stay, shock status, hemoglobin levels and timeframe between presentations to MDCT. The diagnostic yield of MDCT is high com-

**Table 2 Diagnostic yield of double-balloon endoscopy *vs* multidetector computed tomography according to final diagnosis *n* (%)**

Final diagnosis	DBE positive	MDCT positive	Total
Ulcers	7	1	7
Tumors	7	6	7
Angiodysplasia	2	2	2
Diverticulosis	12	5	12
Lymphangiectasia	1	0	1
Undiagnosed	0	0	2
Diagnostic yield	29 (93.5)	14 (45.2)	31

DBE: Double-balloon endoscopy; MDCT: Multidetector computed tomography.

pared to the bleeders from tumor *vs* non-tumor (ulcerative or angiogenic cause) origin (85.7% *vs* 33.3%,  $P = 0.014$ ) (Table 2).

### Clinical impact of MDCT on subsequent DBE

The choice of initial route of DBE examination (for example, oral route or anal route) depended on the clinical suspicion of the lesion by the endoscopist. If previous imaging studies suggested the location of bleeding, the nearest route was chosen; if the patient had a history of hematemesis or tarry stool passage, the oral route was chosen; if the patient had bloody stool passage, the anal route was chosen. Our patients received DBE



**Table 3** Impact of multidetector computed tomography on choice of enteroscopy insertion route

	Route choice right	Route choice wrong
Positive CT	14	0
Negative CT	9	8
Pearson's $\chi^2 = 8.88, P = 0.003$		

CT: Computed tomography.

*via* oral route only ( $n = 18$ ), anal route only ( $n = 4$ ), and both routes ( $n = 9$ ). Among the 14 cases with a positive CT, the initial route of DBE was correct in all 14 cases. Among those 17 patients with a negative CT, the initial route of DBE was correct in 9 cases. A positive CT led to a correct choice of the route of DBE compared with a negative CT (100% *vs* 52.9%,  $P = 0.003$ ) (Table 3).

## DISCUSSION

The current suggested investigation into OGIB includes CE and DBE by the AGA in 2007<sup>[2]</sup>. Both methods have a variable diagnostic yield from 36% to 80%<sup>[2]</sup>. CE is non-invasive but has some limitations, such as capsule retention and incomplete examination<sup>[3]</sup>. Compared with CE, DBE is more invasive but offers therapeutic ability<sup>[4]</sup>. Unlike routine endoscopy, performing DBE is technically demanding and usually requires a two-physician team<sup>[11]</sup>. An initial approach with CE followed by DBE is ideal for patients with OGIB, but is not cost-effective<sup>[12]</sup>. Both procedures are expensive and not reimbursed by insurance in Taiwan. By contrast, MDCT is a non-invasive tool and had been used to evaluate OGIB<sup>[3,5-7]</sup>. In our previous reports<sup>[9,13,14]</sup>, we determined that MDCT is particularly useful to localize the bleeding for subsequent endoscopy in the setting of active GI bleeding. MDCT is reimbursed by our insurance and is more readily available; therefore, approaching OGIB patients with initial MDCT followed by DBE was suggested as an alternative to the current AGA recommendation<sup>[2]</sup> in our institution since the introduction of DBE<sup>[15]</sup>. There are few studies that compare CE and MDCT in the management of OGIB<sup>[6,16]</sup>. In Zhang's report<sup>[16]</sup>, the combination of MDCT with CE was not superior to CE alone in the diagnosis of OGIB. However, data regarding the role of MDCT before DBE in the setting of OGIB is lacking. Only one study from Chen *et al*<sup>[17]</sup> involving 70 patients suggested the usefulness of MDCT combined with DBE in the assessment of small bowel diseases. Thus, we performed this study to evaluate the role of MDCT prior to DBE in our institution during a 6-year-period.

In our study, we found DBE had higher diagnostic yield compared with MDCT (93.5% *vs* 45.2%) for OGIB. Both the diagnostic yield of DBE<sup>[2]</sup> and MDCT<sup>[3,5-7]</sup> are comparable to previous literature. Most of our patients exhibited severe GI bleeding (45.2% of shock at presentation), accounting for our high diagnostic yield. In this study, we discovered that MDCT is useful in two

ways. Firstly, MDCT has a high diagnostic yield of small intestinal neoplasm compared with other non-tumor origins (ulcerative or angiogenic cause). Among seven patients with a tumor origin accounting for their OGIB, six patients were finally diagnosed to have small bowel tumors and all were diagnosed by MDCT prior to DBE procedure. The failure to diagnose one case of early colon cancer in the study can be explained by the fact that our MDCT protocol did not involve bowel preparation or optimal for colonic examination. The reported diagnostic rate of MDCT for small bowel neoplasm ranges from 67.4%<sup>[16]</sup> to 100%<sup>[5]</sup>. This is particularly useful in the management of patients with OGIB. In two recent large DBE series<sup>[18,19]</sup>, patients with diagnosed neoplasm mostly benefited from the DBE due to the ability for long-term control of bleeding either endoscopically or surgically. By contrast, while a diagnosis of ulcer or vascular lesion was identified, only 40%<sup>[18,19]</sup> of the patients remained free of recurrent bleeding. Thus, the high diagnostic yield of MDCT for small bowel neoplasm allowed the clinician to identify patients that would most benefit from subsequent DBE procedures.

Secondly, in patients with positive MDCT, the choice of initial route of DBE examination was more likely to be correct compared to a negative MDCT (100% *vs* 52.9%,  $P = 0.003$ ). As DBE usually requires examination *via* either the oral or anal route, the choice of correct insertion route of DBE is critical for allowing rapid approaches to the bleeding<sup>[20]</sup>. No recommended standard for selection of the insertion route of DBE currently exists. Although clinical presentation such as hematemesis and stool color are useful for the endoscopist to choose the route of insertion, the initial route of DBE is not reliable, especially in the setting of massive bleeding<sup>[20]</sup>. Few studies<sup>[20,21]</sup> have utilized the finding of CE to guide the insertion route of subsequent DBE. In our study, we demonstrated that a positive MDCT is useful for guiding the insertion route of subsequent DBE.

Several methods of MDCT are described in the literature, including MDCT with oral saline as a contrast<sup>[16,22]</sup> or multi-phase CT-enterography<sup>[3,5,7]</sup>. The reported diagnostic yield for OGIB ranges from 30%<sup>[16]</sup> to 83%<sup>[22]</sup> with different MDCT protocols. No comparative study comparing the diagnostic yield of different CT protocols exists. In our institution, the triphasic MDCT protocol without oral contrast<sup>[8]</sup> utilized in our institution demonstrates a similar diagnostic yield as compared with previous reports. Although the use of fluid contrast can distend the intestinal lumen effectively and increase the diagnostic ability, especially for mucosal lesions<sup>[3,16,22]</sup>, this may delay the subsequent DBE examination. In our institution, an emergent MDCT can be arranged as quickly as 2 h after a non-diagnostic endoscopy. If previous endoscopic studies suggested bleeding in the small bowel, the endoscopist would withdraw the endoscope without air suction, which leaves air in the bowel as a contrast (Figure 1). This approach is simple and similar to the recently described virtual enteroscopy<sup>[23]</sup>. *Via* this approach, once MDCT is positive, a DBE can be arranged early without the need to wait for

gastric emptying due to the use of an oral fluid contrast.

This study has some limitations. The first is in the retrospective nature of the study. Although CE or DBE are recommended for the diagnosis of OGIB<sup>[24]</sup>, both of these procedures are expensive and neither is covered by health insurance providers in our region. In addition, MDCT is more readily available compared with CE and DBE and less invasive than conventional angiography. Thus, MDCT is used as the diagnostic and triage tool of choice for OGIB at our institution<sup>[9]</sup>. Therefore, our findings of the high diagnostic yields of both DBE and MDCT may be an overestimation because some patients with small intestinal tumors or those who have self-limited bleeding may not undergo subsequent DBE and were not included in this analysis. Secondly, the use of MDCT still has some disadvantages<sup>[4]</sup>. The procedure results in substantial radiation exposure and carries the risk of contrast nephrotoxicity that may limit its use in elderly patients, who constitute the main population of OGIB patients.

Our study demonstrates that MDCT provides a reasonable diagnostic yield for OGIB in comparison with previous reports<sup>[3,7,16,22]</sup>. The use of MDCT as a triage tool prior to DBE may aid the endoscopist in identifying patients who will benefit from the examination, and may aid the endoscopist in choosing the most efficient route of DBE examination. However, our study is limited to a retrospective nature and only a small number of patients were enrolled in this study. Further prospective studies with more patients are required to confirm our observation.

## COMMENTS

### Background

Capsule endoscopy followed by double-balloon endoscopy is the current standard approach for obscure gastrointestinal bleeding. Multidetector computed tomography (MDCT) has been used recently to investigate obscure gastrointestinal bleeding, but its role in current standard approach to obscure gastrointestinal bleeding (OGIB) has not been evaluated.

### Research frontiers

This study analyzed the clinical impact of MDCT in a subset of patients with OGIB in a single medical center.

### Innovations and breakthroughs

This study is novel in that the researchers analyzed the clinical utility of MDCT before double-balloon enteroscopy (DBE) for the indication of OGIB.

### Applications

MDCT provides a reasonable diagnostic yield in this study. The use of MDCT as a triage tool prior to DBE may aid the endoscopist in identifying patients who will benefit from the examination and may aid the endoscopist in choosing the most efficient route of DBE examination.

### Peer review

The authors demonstrated that MDCT may provide a reasonable diagnostic yield in a select group of patients with OGIB by a retrospective study.

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