

# Video capsule endoscopy as a tool for evaluation of obscure overt gastrointestinal bleeding in the intensive care unit



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## Bibliography

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## ABSTRACT

**Background and study aims** Video capsule endoscopy (VCE) is a minimally invasive tool that helps visualize the gastrointestinal tract from the esophagus to the right colon

without the need for sedation or preparation. VCE is safe with very few contraindications. However, its role and safety profile in the intensive care unit (ICU) population have not been reported. The aim of this study is to evaluate the safety, efficacy, and feasibility of VCE use in ICU patients.

**Patients and methods** We conducted a single-center retrospective observational study of patients who underwent VCE for evaluation of obscure overt gastrointestinal bleeding in the ICU between 2008 and 2016.

**Results** This study included 48 patients who were admitted to the UMass Memorial Medical Center ICUs for gastrointestinal bleeding. VCE was successfully completed in 43/48 (90%) patients. The entire length of small bowel could be evaluated in 75% and the source of bleeding was identified in 44% of the patients. The most commonly identified source of bleeding included small bowel angioectasias, duodenal erosions/ulcers, and small bowel polyps. No major complications could be attributed to the VCE. Only 1 capsule was retained after 2 wk; however, there was no incidence of bowel obstruction, perforation, or capsule aspiration.

**Conclusions** This observational retrospective study demonstrates that VCE may be a safe, feasible, and effective diagnostic tool in evaluation of gastrointestinal bleeding in the ICU population with few complications. VCE may be a safe diagnostic prelude and be a guide to the correct therapeutic procedure if needed, in the context of patients who are seriously ill.

## Introduction

Patients with severe gastrointestinal bleeding (GIB) who are hemodynamically unstable are usually admitted to an intensive care unit (ICU) [1] for stabilization and possible endoscopic management. By definition, these patients are usually hemodynamically unstable and are at increased risk of endoscopic complications during evaluation [2–6]. Hematemesis localizes bleeding to within reach of a gastroscope. In patients with non-hematemesis GIB, the presence of melena or hematochezia has little localizing value [6], leading to what has become a traditional approach of upper endoscopy followed by colonoscopy

and other procedures if 1 of those 2 are not diagnostic [7]. However, the yield of conventional endoscopic procedures in this context is known to be low. An early diagnostic/endoscopic approach is thought to have higher diagnostic yield in some studies, yet this advantage needs to be balanced with risk of endoscopic complications [8–11].

Video capsule endoscopy (VCE) is a minimally invasive tool that can visualize the gastrointestinal tract from esophagus to the right colon without the need for sedation or preparation. We hypothesize that VCE could serve as an alternative to the traditional approach to GIB in an acutely unstable or high-risk patient. VCE has been investigated as a triaging tool in the

emergency room to determine the population of patients who would benefit from additional diagnostic and therapeutic interventions [12]. However, before the utility of VCE as first-line evaluation of GIB in ICU patients can be established, the safety and efficacy of this technique in ICU patients need to be evaluated further. We therefore investigated the safety and efficacy of VCE use in the ICU in a retrospective case series of patients admitted to the ICU with obscure overt GIB. To our knowledge, there are no prior studies that specifically evaluated the role and safety profile of VCE in the ICU population for evaluation of GIB.

## Patients and methods

We conducted a single-center retrospective observational study (case-series) of patients who underwent VCE for evaluation of obscure overt GIB in our medical, cardiac, and surgical ICUs between 2008 and 2016 at UMass Memorial Medical Center, a large tertiary referral center in central Massachusetts. Inclusion criteria included age more than 18 y, a diagnosis of GIB during that admission, and VCE during an ICU stay. Patients were identified through billing codes, and data collected included patients' demographics, comorbidities, VCE results, additional diagnostic/therapeutic procedures, and laboratory data. Standard descriptive statistics were used for demographic data, VCE findings, and complications using Microsoft Excel 2007 (Microsoft, Redmond, WA, USA). Additionally, we used Student's t-test for comparing continuous variables, and chi-squared test was used for binary variables. The patients were investigated using either the M<sub>2</sub>A or PillCam SB2 capsules (GivenImaging, Jocqneam, Israel). The capsules were read on a workstation using Rapid software versions 3 to 8. All VCE images were read by expert endoscopists with many years of experience in interpreting VCEs. All patients had been screened for contraindications of VCE including but not limited to history of bowel obstruction. In a few patients with high risk of gastric retention (e.g., prior history) or evidence of retention on the real-time viewer, a dose of intravenous metoclopramide or erythromycin was administered before or during the capsule study. VCE was deemed successful if the entire length of the small bowel could be evaluated or if significant length of the small bowel was evaluated without significant device malfunction, missing frames, or gastric retention. The retrospective chart review was approved by the UMass Memorial Institutional Review Board.

## Results

This study included 48 patients who were admitted to the ICU for GIB and who had negative esophagogastroduodenoscopy (EGD) and in many cases colonoscopy (69%) and were categorized as having obscure overt GIB on further investigation. Average age of patients was  $70 \pm 16.5$  y with 58% male (► **Table 1**). The majority of patients (92%) were Caucasian. More than half (56%) were on mono or dual antiplatelet therapy, and 31% were anticoagulated. About one-fifth of patients (19%) had international normalized ratio (INR) >3, and 29% required 2 L or more

► **Table 1** Demographics.

	n (%)
Male	28 (58)
Female	20 (42)
Age (y)	70.4 ± 16.5
BMI	28.3 ± 7.5
Caucasian	44 (92)
Current smoker	13 (27)
Former smoker	26 (54)
Alcohol use	15 (31)
NSAIDs use	5 (10)
Aspirin alone	22 (46)
Dual antiplatelet therapy	5 (10)
Anticoagulated	15 (31)
<b>Presenting symptom</b>	
Hematemesis	6 (13)
Hematochezia	15 (31)
Melena	23 (48)
Anemia	4 (8)
<b>Comorbidities</b>	
Coronary artery disease	28 (58)
Recent MI	4 (8)
COPD	11 (23)
Pneumonia	8 (17)
Severe aortic stenosis	6 (13)
CHF	13 (27)
ESRD	6 (13)
Cirrhosis	7 (15)
History of IBD	0 (0)
Prior abdominal surgeries	23 (48)
History of small bowel obstruction	0 (0)
INR >3	9 (19)
Hypoxia (>2 L NC)	14 (29)
Vasopressors required	3 (6)
Need for transfusion	45 (94)
Lowest HGB (average)	6.39 ± 1.14
Lowest HGB (range)	3.8–8.5
Units of blood transfused	5.9 ± 4.8

BMI: body mass index; COPD: chronic obstructive pulmonary disease; CHF: congestive heart failure; ESRD: end-stage renal disease; IBD: inflammatory bowel disease; MI: myocardial infarction; NSAID: nonsteroidal anti-inflammatory drug; HGB: hemoglobin; NC: nasal canula

of supplemental oxygen. Three patients required vasopressor support for hypotension. Twenty-three patients (48%) initially presented with melena, 15 (31%) with hematochezia, and 6 (13%) with hematemesis. The most common underlying comorbidities included coronary artery disease, congestive heart failure, and chronic obstructive pulmonary disease. Nearly half of patients (48%) had history of intra-abdominal surgery, but none had history of prior small bowel obstruction, inflammatory bowel disease, or fistulas. The mean lowest hemoglobin was  $6.39 \pm 1.14$  mg/dL (range: 3.8–8.5 mg/dL) on or within 2 d of admission. Almost all patients 45 (94%) required blood transfusions with a mean of  $5.8 \pm 4.8$  units transfused. Transfusion requirements were higher in patients whose bleeding source could be identified as compared to those with unidentified source ( $7.9 \pm 5.7$  vs.  $4.3 \pm 4.8$  units transfused,  $P < 0.01$ ). The mean lengths of hospital and ICU stays were  $10.4 \pm 9.6$  d and  $5.5 \pm 3.6$  d, respectively.

Most patients 44/48 (92%) had EGD within 24 h of admission. Two patients had prior negative EGDs within the past 30 d, and 1 patient had EGD at day 8 of admission due to presentation with hematochezia on day 1 and repeat bleeding on day 7. Average time to colonoscopy was  $1.9 \pm 0.9$  d in the two-thirds of patients who underwent this procedure (► **Table 2**). One patient with prior history of small bowel bleeding underwent VCE as first-line evaluation of GIB. Sixteen patients (33%) underwent VCE as second-line evaluation. The remainder of patients underwent VCE as third- or fourth-line evaluation for obscure overt GIB. VCE was successfully completed in 43/48 (90%) patients. The majority of capsules were swallowed by the patient (75%) while the other 25% were placed endoscopically via EGD, most commonly during negative EGD as the next step in the patient's evaluation (► **Table 2**). The entire length of small bowel could be evaluated in 36/48 (75%). The source of bleeding was identified in 21/48 (44%) patients. However, the source was identified more commonly when VCE was performed within 48 h of suspected bleeding (63% within 48 h vs. 31% with VCE after 48 h,  $P = 0.03$ ). The most commonly identified source of bleeding included small bowel angioectasias, duodenal erosions/ulcers, and small bowel polyps. The most commonly identified incidental lesions in the small bowel included angioectasias (nonbleeding and no stigmata of recent bleeding, portal hypertensive gastropathy/small bowel varices, and erosions. Additional diagnostic interventions (tagged Red Blood Cell scan, angiography, and deep enteroscopy) was performed in 17% of patients while 23% required additional therapeutic interventions.

No major complications could be attributed to the VCE. Only 1 capsule was retained on abdominal imaging after 2 wk; however, there was no incidence of bowel obstruction, perforation, or capsule aspiration (► **Table 3**). Two additional capsules did not pass through the stomach to enter the small bowel within the 8 h of recorded images but were excreted on follow-up imaging within 2 wk. Two earlier studies also included missing frames or poor image quality within the small bowel, which limited the interpretation of the small bowel images.

► **Table 2** Outcomes.

	n (%)
Successful completion of VCE	43 (90)
Entire small bowel visualized	36 (75)
Anatomic source of bleeding identified	21 (44)
▪ VCE within 48 h	12/19 (63)
▪ VCE after 48 h	9/29 (31)
▪ p-value	0.03
Actively bleeding lesions	9 (19)
Common sources of bleeding identified	
Angioectasias	10 (21)
Small intestinal erosions/ulcers	2 (4)
Small bowel polyps	2 (4)
Portal hypertensive gastropathy	2 (4)
Other	5 (11)
Most common incidental findings	
Angioectasias (not clear source of bleed)	7 (15)
Portal hypertensive gastropathy / small intestinal varices	4 (8)
Erosions	2 (4)
Mode of capsule delivery	
Swallowed capsule	36 (75)
Endoscopically placed	12 (25)
Timing of initial diagnostic procedures	
EGD within 24 h	44 (92)
Prior EGD within 30 d	2 (4)
Colonoscopies during the same admission	33 (69)
Days to colonoscopy (d)	$1.9 \pm 0.9$
Capsule as first-line	1 (2)
Days to VCE (d)	$3.47 \pm 2.3$
Need for additional procedures	19 (40)
Diagnostic	8 (17)
Therapeutic	11 (23)

## Discussion

This retrospective case series demonstrates that VCE may be a safe, feasible, and effective diagnostic tool in the evaluation of obscure overt GIB in the critically ill ICU population. The overall success rate of the procedure was 90%, with the source of bleeding identified in nearly half of the patients. Almost all patients in this study had obscure overt GIB and were previously evaluated via other modalities. There were no major complications attributed to the VCE.

► **Table 3** Safety data.

Major complications	0 (0)
Minor complications	1 (1)
Capsule retention at 2 wk	1 (1)
Aspiration	0 (0)
Bowel obstruction	0 (0)
Length of hospital stay (d)	10.4 ± 9.5
Length of ICU stay (d)	5.5 ± 3.6
Need for intubation for GIB or tracheostomy	6 (12.5)
Recurrence of bleeding at 6 mo	12 (25)

While the current study does not address the utility of VCE as a first-line diagnostic tool for evaluation of GIB, as a minimally invasive tool, VCE may be a safe diagnostic alternative to standard endoscopic procedures in patients at risk for significant complications from the more invasive procedures, such as those with elevated INR, hypoxemia, or recent myocardial infarction [3]. This study demonstrates that VCE is safe in the most critically ill patients with obscure overt GIB who do not otherwise have other major contraindications to VCE. However, the role of VCE as the first-line diagnostic modality in this setting needs to be evaluated further. Inpatient data shows the earlier VCE is used the better the diagnostic yield [8]. This observation is supported by similar findings in the current study as patients who underwent VCE within 48 h of admission had twice the likelihood of bleeding source identification. Larger prospective studies, particularly with early deployment, are necessary to further evaluate the safety and efficacy of this tool in these settings.

Newer video capsule models allow live visualization of the upper gastrointestinal tract in real time to determine the presence of active bleeding in the esophagus or stomach and confirm passage of capsule into the small intestine. This may help stratify high-risk patients regarding urgency of more definite therapeutic interventions while offering additional diagnostic information regarding possible sources of bleeding in the small intestine or presence of blood in the right colon. This can, in turn, potentially minimize the number of invasive procedures, decrease the length of hospitalization, and minimize complications in the critically ill patients.

In our cohort, we observed a few cases of gastric retention of the capsule and delayed transit. Other published and unpublished studies confirm the occurrence of delayed transit in bed-ridden patients, which are common in the ICU setting. Several medications including opioids, commonly used in the ICU, may also alter gut motility and lead to delayed gastric emptying. The only other study of VCE in ICU patients, to our knowledge, investigated transit time of the VCE in a small cohort of ICU patients [13]. While this study was not able to show a statistically significant difference in transit time between ICU patients compared to ambulatory patients (most likely due to the very small sample size), there was a large degree of variability in

transit times in the critically ill population. In our practice, we have most recently developed an algorithm for passage into the small intestine for swallowed capsules at about an hour after capsule ingestion. If the capsule is not in the small intestine at that point, we have used prokinetic agents such as metoclopramide or erythromycin with good success. This increases the yield of VCE in such select patients.

Another rare technical difficulty was missing frames or poor image quality in 2 (4%) of our earlier patients. This is most likely due to electrical interference from nearby devices, distance from the receiver, or the patient's excessive movement. Newer devices with higher-quality signals and images have mostly resolved these issues in our most recent cohorts of ICU and non-ICU patients. However, wireless interference from ventricular assist devices on image quality of VCE continues to remain a limitation reported in the literature [14].

The main limitations of this study include the small sample size and the limitations of the retrospective design. The sample size is limited by the number of VCEs already performed at our institution, but to our knowledge, this is the largest cohort of VCEs studied in the ICU setting. A prospective design will increase the power of the study and make the study more generalizable. The retrospective design also adds its limitations as there is no control arm and neither the patient nor the clinicians are blinded. In addition, this study by its nature is self-selected for patients with obscure GIB. Therefore, further larger prospective studies are necessary before generalizing the result to the general ICU population with GIB. Other limitations may include variation in patient acuity and severity of disease. Our center is a large tertiary/quaternary with high-acuity patients, but further studies may help generalize the result to smaller community sites.

In summary, VCE is safe for use in ICU patients and may be useful in patients with GIB in detecting the site of active bleeding and providing guidance for the most appropriate therapeutic procedure, if needed.

### Competing interests

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