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ORIGINAL ARTICLE

Observational Study

Current state of practice for colonic diverticular bleeding in 37 hospitals in Japan: A multicenter questionnaire study

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Abstract

AIM

To clarify the current state of practice for colonic diverticular bleeding (CDB) in Japan.

METHODS

We conducted multicenter questionnaire surveys of the practice for CDB including clinical settings (8 questions), diagnoses (8 questions), treatments (7 questions), and outcomes (4 questions) in 37 hospitals across Japan. The answers were compared between hospitals with high and low number of inpatient beds to investigate which factor influenced the answers.

RESULTS

Endoscopists at all 37 hospitals answered the questions, and the mean number of endoscopists at these hospitals was 12.7. Of all the hospitals, computed tomography was performed before colonoscopy in 67% of the hospitals. The rate of bowel preparation was 46.0%. Early colonoscopy was performed within 24 h in 43.2% of the hospitals. Of the hospitals, 83.8% performed clipping as first-line endoscopic therapy. More than half of the hospitals experienced less than 20% rebleeding events after endoscopic hemostasis. No significant difference was observed in the annual number of patients hospitalized for CDB between high- (≥ 700 beds) and low-volume hospitals. More emergency visits (P = 0.012) and endoscopists (P =0.015), and less frequent participation of nursing staff in early colonoscopy (P = 0.045) were observed in the high-volume hospitals.

CONCLUSION

Some practices unique to Japan were found, such as performing computed tomography before colonoscopy, no bowel preparation, and clipping as first-line therapy. Although, the number of staff differed, the practices for CDB were common irrespective of hospital size.

Key words: Colonic diverticular hemorrhage; Lower gastrointestinal bleeding; Computed tomography; Endoscopy; Bowel preparation

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Core tip: Colonic diverticular bleeding (CDB) is increasing in Asia. There are no practice guidelines for CDB, and it is important to determine which recommendation is acceptable to a majority of hospitals. We conducted multicenter questionnaire surveys of 37 hospitals in Japan regarding management of CDB including clinical settings, diagnosis, treatment, and clinical outcomes, and made comparisons between hospitals with different patient volumes and between hospitals in different regions. Thus, practice styles unique to Japan such as performing computed tomography before colonoscopy, no bowel preparation, and clipping as first-line therapy were identified. However, management of CDB was common among hospitals irrespective of hospital size and region.

Niikura R, Nagata N, Doyama H, Ota R, Ishii N, Mabe K, Nishida T, Hikichi T, Sumiyama K, Nishikawa J, Uraoka T, Kiyotoki S, Fujishiro M, Koike K. Current state of practice for colonic diverticular bleeding in 37 hospitals in Japan: A multicenter questionnaire study. *World J Gastrointest Endosc* 2016; 8(20): 785-794 Available from: URL: http://www.wjgnet.com/1948-5190/full/v8/i20/785.htm DOI: http://dx.doi.org/10.4253/wjge.v8.i20.785

INTRODUCTION

Colonic diverticular bleeding (CDB) is a major cause of lower gastrointestinal bleeding, and is estimated to cause 25% to 40% of all cases of lower gastrointestinal bleeding^[1-3]. In Japan, CDB was found in 427 (1.5%) of 28192 patients who underwent colonoscopy at an emergency hospital^[4]. Its occurrence has increased in Japan as well as in Western countries^[4-7]. CDB results in hemorrhagic shock requiring blood transfusion[8,9], and has a high recurrence rate of 20% within 1 year^[10,11]. As a result, patients are often burdened by the frequent examinations, hospitalization, repeated blood transfusions, and a consequent decrease in their quality of life. Furthermore, these practices for CDB may be different between Western countries and Japan. For example, Western countries perform purged colonoscopy using polyethylene glycol as the first diagnostic procedure, and perform endoscopic hemostasis using clipping^[12]. In contrast, Japanese hospitals have good access to computed tomography (CT)[13] and may select CT as the first diagnostic procedure. In addition, diagnostic tools, endoscopic environment, and treatment strategy may potentially differ among hospitals in Japan. Moreover, the practice for CDB may differ according to hospital patient volume and region, as is seen in the practice for other lower gastrointestinal disease^[14,15]. Some studies have reported significant associations between hospital volume and clinical outcome, and between hospital region and diagnosis methods^[14,15]. Today, there are no practice guidelines for CDB, and it is

important to determine what recommendations would be acceptable to a large number of hospitals.

Therefore, we conducted a multicenter questionnaire survey of the practice for CDB in 37 hospitals across Japan to elucidate the current state of the clinical settings, diagnosis, treatment, and clinical outcomes of patients with CDB, and to compare these findings according to hospital volumes and regions.

MATERIALS AND METHODS

Contents of the questionnaire

First, 1 endoscopist (Doyama H) developed the questionnaire on practice for CDB. Then, 3 endoscopists (Ota R, Niikura R and Nagata N) reviewed and edited the questionnaire regarding the length, clarity, and contents. Finally, 27 survey questions on practice for CDB were developed. The questionnaire consisted of 4 parts (clinical settings, diagnosis, treatment, clinical outcomes) as follows. In part (I), there were 9 questions for clinical settings on: (1) the clinical database for CDB such as gastrointestinal bleeding database, inpatient database, or endoscopy database; (2) institution-specific strategy for CDB; (3) number of CDB admissions; (4) number of emergency ambulance visits; (5) number of endoscopists performing early colonoscopy within 24 h of patient arrival; (6) number of expert endoscopists with hemostatic technical skills; (7) nursing staff who monitored vital signs during bowel preparation; (8) nursing staff assisting early colonoscopy; and (9) use of a water-jet colonoscope. For part (II), there were 8 questions for diagnoses of CBD on (10) the first choice diagnostic examination; (11) early contrast-enhanced CT within 3 h of patient arrival; (12) early colonoscopy; (13) bowel preparation; (14) cap-assisted colonoscopy; (15) how to improve the identification of stigmata of recent hemorrhage (SRH); (16) availability of small bowel examinations in case of negative colonoscopy; and (17) modality for small bowel examinations. For part (III), there were 6 questions for treatment of CDB on (18) first-line endoscopic therapy; (19) selection of non-endoscopic therapy; (20) first-line therapy among non-endoscopic therapies; (21) how to prevent rebleeding; (22) discontinuation of antithrombotic drugs on admission; and (23) strategy for restarting antithrombotic drugs. In part (IV), there were 4 questions for clinical outcomes of CDB on (24) identification rate of SRH; (25) rebleeding rate after endoscopic hemostasis; (26) rebleeding rate after interventional radiology; and (27) rebleeding rate after barium impaction therapy.

Questionnaire survey

The questionnaire survey was conducted by e-mail that was sent to 1 or 2 endoscopists at each of the 37 hospitals with different numbers of inpatient beds and in different regions in Japan between May 2015 and June 2015. Selection of the hospitals was made by Fujishiro M, who knew that the representative

endoscopists would be interested in this topic from his personal communications. To assess the reproducibility of questionnaire, we conducted a blinded secondary questionnaire survey 2 mo after using the same 16 questionnaire items. Selection of these questionnaire items was made by Niikura R and Nagata N. because these items were found to be related to the practice for CDB. These 37 hospitals were located in East or West Japan and have 100 to 1000 inpatient beds (Appendix).

Statistical analysis

The data from the first questionnaire survey were analyzed, and the intra-observer agreement between the first and second questionnaires was analyzed using kappa statistics. Kappa values were evaluated as follows: > 0.80, excellent agreement; > 0.60 to 0.80, good agreement; > 0.40 to 0.60, moderate; > 0.20 to 0.40, fair; and ≤ 0.20 , poor^[16].

A high-volume hospital was defined as one with over 700 beds, because the median number of beds in our data was 700 beds per hospital. Expert endoscopists were defined as those who were able to perform endoscopic hemostatic treatment by themselves. We evaluated the clinical settings, diagnosis methods, treatment, and outcomes between the groups of hospital separated by hospital volume and region (East Japan, West Japan) using a χ^2 test or Fisher's exact test as appropriate. Continuous variables were compared using the Mann-Whitney U test. We also evaluated the associations of the rates of SRH identification and rebleeding with type of procedure from questionnaire answers using a nonparametric trend test. A P value < 0.05 was considered statistically significant. All statistical analyses were performed using the STATA version 13 software (StataCorp, College Station, TX, United States).

RESULTS

The number of beds per hospital in each region of Japan is shown in Figure 1. There were 18 high-volume hospitals (\geq 700 beds) and 19 low-volume hospitals. Twenty-one of the 37 (56.8%) hospitals were located in East Japan, and 16 hospitals (43.2%) were located in West Japan (Figure 1). All 37 hospitals completed the first questionnaires, and 35 of the hospitals completed the second questionnaires. Intra-observer agreement for each question between the first and second surveys was excellent (mean κ , 0.83, 95% confidence interval 0.78-0.87) (Supplementary Table 1).

Questionnaire items for clinical settings

Questions and answers regarding clinical settings are shown in Table 1. Of all the hospitals, 86.5% answered the questionnaire based on the clinical database of each hospital. Only 13.5% of hospitals had an institution-specific strategy for CDB. The number of CDB patients who received therapy, and the number of emergency ambulance visits, differed among hospitals. The mean



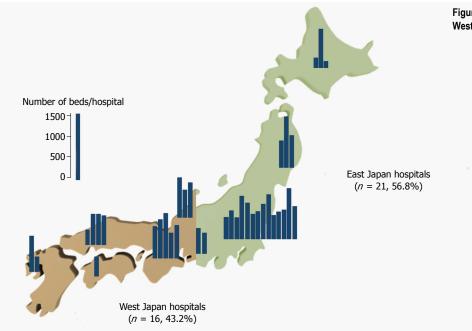


Figure 1 Number of beds per hospital in East and West Japan.

Table 1 Questions and answers regarding clinical settings in 37 hospitals n (%)

No.	Question	Answer (n = 37)	High volume $(n = 18)$	Low volume (n = 19	9) P value	Fact Japan (n = 21)	West Japan (n = 16)	P value
-				•	7) 7 value	Last Japan (// = 21)	West Japan (II = 10)	7 value
1	,	1	ed on a clinical database?					
	Yes	31 (83.8)	16 (51.6)	15 (48.4)		18 (58.1)	13 (41.9)	
	No	6 (16.2)	2 (33.3)	4 (66.7)	0.660	3 (50.0)	3 (50.0)	1.000
2	3	a specific institution	0,					
	Yes	5 (13.5)	0	5 (100)		2 (40.0)	3 (60.0)	
	No	32 (86.5)	18 (56.3)	14 (43.7)	0.046	19 (59.4)	13 (40.6)	0.634
3	, ,	-	ed for CDB annually?					
	1-10	12 (32.5)	7 (58.3)	5 (41.7)		7 (58.3)	5 (41.7)	
	11-20	10 (27.0)	4 (40.0)	6 (60.0)		4 (40.0)	6 (60.0)	
	21-30	5 (13.5)	2 (40.0)	3 (60.0)		1 (20.0)	4 (80.0)	
	≥ 31	10 (27.0)	2 (50.0)	5 (50.0)	0.824	9 (90)	1 (10)	0.035
4	How many en	mergency ambulance	e visits do you receive anr	nually?1				
	< 2000	15 (44.1)	5 (33.3)	10 (66.7)		8 (53.3)	7 (46.7)	
	2000-6000	11 (32.3)	4 (36.4)	7 (63.6)		5 (45.5)	6 (54.5)	
	6000-10000	6 (17.7)	6 (100)	0		3 (50.0)	3 (50.0)	
	≥ 10000	2 (5.9)	0	2 (100)	0.012	2 (100)	0	0.724
5	How many en	ndoscopists perform	early colonoscopy within	24 h after patient arri	val at your	hospital?		
		12.7 ± 9.4	17.0 ± 11.6	8.8 ± 4.4	0.015	10.4 ± 5.7	15.8 ± 12.5	0.296
6	How many as	re expert endoscopis	ts with endoscopic hemos	stasis technical skills a	re there at y	our hospital?		
		10.1 ± 7.5	13.1 ± 9.5	7.3 ± 3.3	0.019	7.9 ± 3.3	13.0 ± 10.2	0.143
7	Do you have	nursing staff who me	onitor the patients' vital s	igns during bowel pre	eparation?			
	Yes	33 (89.2)	17 (51.5)	16 (48.5)		19 (57.6)	14 (42.4)	
	No	4 (10.8)	1 (25.0)	3 (75.0)	0.604	2 (50.0)	2 (50.0)	1.000
8	Do you have	nursing staff for earl	y colonoscopy examinatio	ons within 24 h after p	atient arriva	al at the hospital?		
	Yes	23 (62.2)	8 (34.8)	15 (65.2)		13 (56.5)	10 (43.5)	
	No	14 (37.8)	10 (71.4)	4 (28.6)	0.045	8 (57.1)	6 (42.9)	1.000
9	Do you have	a water-jet colonosco	ppe?					
	Yes	34 (91.9)	17 (50.0)	17 (50.0)		20 (58.9)	14 (41.1)	
	No	3 (8.1)	1 (33.3)	2 (66.7)	1.000	1 (33.3)	2 (66.7)	0.568

¹Missing data included. The values in parentheses are percentages, and continuous data are shown as mean ± standard deviation. CDB: Colonic diverticular bleeding.

number of endoscopists and expert endoscopists were 12.7 and 10.1, respectively. Of all the hospitals, 89.2% and 62.2% had nursing staff for monitoring vital signs during bowel preparation and early colonoscopy examination, respectively. Ninety-one percent of hospitals had a water-jet colonoscope.

Comparing hospital with high and low patient volumes, more emergency visits (P=0.012), endoscopists (P=0.015), and expert endoscopists (P=0.019), and less institution-specific management for CDB (P=0.046) and frequent participation of nursing staff in early colonoscopy (P=0.045) were observed



Table 2 Questions and answers regarding diagnosis of colonic diverticular bleeding in 37 hospitals n (%)

No.	Question	Answer $(n = 37)$	High volume $(n = 18)$	Low volume $(n = 19)$	<i>P</i> value	East Japan $(n = 21)$	West Japan $(n = 16)$	P value		
10	What do you use as the first-line diagnostic method for hematochezia and suspected CDB?									
	Non-contrast-enhanced CT	3 (8.1)	0	3 (100)		0	3 (100)			
	Contrast-enhanced CT	22 (59.5)	11 (50.0)	11 (50.0)		12 (54.6)	10 (45.4)			
	Colonoscopy	10 (27.0)	6 (60.0)	4 (40.0)		7 (70.0)	3 (30.0)			
	Contrast-enhanced CT and colonoscopy	2 (5.4)	1 (50.0)	1 (50.0)	0.359	2 (100)	0	0.101		
11	Can you perform urgent contrast-enhanced CT within 3 h after patient arrival at hospital? ²									
	Yes	22 (61.1)	12 (54.6)	10 (45.4)		13 (59.1)	9 (40.9)			
	No	14 (38.9)	6 (42.9)	8 (57.1)	0.494	7 (50.0)	7 (50.0)	0.593		
12	Can you perform early colonoscopy within	24 h after patier	nt arrival at hospit	al?						
	Yes	16 (43.2)	9 (56.3)	7 (43.7)		10 (62.5)	6 (37.5)			
	No	21 (56.8)	9 (42.9)	12 (57.1)	0.419	11 (52.4)	10 (47.6)	0.538		
13	Do you request bowel preparation?									
	Yes	17 (46.0)	6 (35.3)	11 (64.7)		13 (76.5)	4 (23.5)			
	No	3 (8.1)	3 (100)	0		2 (66.7)	1 (33.3)			
	Case by case	17 (45.9)	9 (52.9)	8 (47.1)	0.105	6 (35.3)	11 (64.7)	0.046		
14	Do you use a cap-assisted colonoscopy for early colonoscopy?									
	Yes	24 (64.9)	11 (45.8)	13 (54.2)		15 (62.5)	9 (37.5)			
	No	13 (35.1)	7 (53.9)	6 (46.1)	0.642	6 (46.2)	7 (53.8)	0.338		
15	How do you perform colonoscopy to impro	ve identification	n of SRH?1							
	Cap-assisted colonoscopy	17 (46.0)	10 (58.8)	7 (41.2)	0.254	10 (58.8)	7 (41.2)	0.815		
	Long cap-assisted colonoscopy	13 (35.1)	6 (46.2)	7 (53.8)	0.823	7 (53.9)	6 (46.1)	0.793		
	Inverting diverticulum via suction of	18 (48.7)	11 (61.1)	7 (38.9)	0.140	11 (61.1)	7 (38.9)	0.603		
	colonoscopy									
	Wash out with water	36 (97.3)	18 (50.0)	18 (50.0)	1.000	21 (58.3)	15 (41.7)	0.432		
	Colonoscopy by multiple doctors	3 (8.1)	1 (33.3)	2 (66.7)	1.000	2 (66.7)	1 (33.3)	1.000		
	Colonoscopy under X-ray	3 (8.1)	1 (33.3)	2 (66.7)	1.000	1 (33.3)	2 (66.7)	0.568		
16	Do you examine the small bowel when you	are unable to d	iagnose definite C	DB by colonosco	ppy?	,	` ′			
	Yes	18 (48.7)	11 (61.1)	7 (38.9)	1,	10 (55.6)	8 (44.4)			
	No	7 (18.9)	1 (14.3)	6 (85.7)		4 (57.1)	3 (42.9)			
	Case by case	12 (32.4)	6 (50.0)	6 (50.0)	0.145	7 (58.3)	5 (41.7)	1.000		
17	Which modality do you select for the small bowel examination? ²									
	Capsule endoscopy	29 (85.3)	17 (58.6)	12 (41.4)		18 (62.1)	11 (37.9)			
	Balloon-endoscopy	2 (5.9)	0	2 (100)		1 (50.0)	1 (50.0)			
	Case by case	3 (8.8)	1 (33.3)	2 (66.7)	0.301	1 (33.3)	2 (66.7)	0.776		

¹Duplicated data allowed; ²Missing data included. Parenthesis shows percentage. CT: Computed tomography; CDB: Colonic diverticular bleeding; SRH: Stigmata of recent hemorrhage.

in high-volume hospitals (Table 1). No significant differences were observed in other questionnaire items such as number of patients hospitalized for CDB between the two groups (Table 1). Comparing hospitals in East and West Japan, a higher number of patients hospitalized for CDB was observed in East Japan hospitals (P = 0.035) (Table 1). No significant difference was observed in other questionnaire items between the two groups (Table 1).

Questionnaire items for diagnoses

Questions and answers regarding diagnosis are shown in Table 2. Of all the hospitals, 59.5% selected contrastenhanced CT as first examination of choice. The rates of urgent CT, early colonoscopy, bowel preparation, capassisted colonoscopy were 61.1%, 43.2%, 46.0%, and 64.9%, respectively. Ninety-one percent of hospitals washed out with water to improve identification of SRH. There was a wide variation among hospitals in small bowel intestinal examination, but 85.3% of hospitals selected capsule endoscopy as the tool of choice when it was unable to diagnose definite CDB.

No significant differences between hospitals with high and low patient volumes were observed in all questionnaire items (Table 2). Comparing hospitals in East and West Japan, East Japan hospitals performed more frequent bowel preparation compared with West Japan hospitals (P=0.046) (Table 2). No significant differences were observed in other questionnaire items between the two groups (Table 2).

Questionnaire items for treatments

Questions and answers regarding treatment are shown in Table 3. In endoscopic treatment, clipping, band ligation, and epinephrine injection were performed as first-line therapy in 83.8%, 13.5%, and 2.7% of hospitals. Seventy-three percent and 67% of hospitals selected non-endoscopic therapy for patients with rebleeding and hemorrhagic shock, and 77.4% of hospitals performed interventional radiology as first-line non-endoscopic therapy. Fifty-nine percent of hospitals discontinued antithrombotic drugs on admission and only 15% of hospitals had a strategy for restarting these drugs.



Table 3 Questions and answers regarding treatments of colonic diverticular bleeding in 37 hospitals n (%)

No.	Question	Answer $(n = 37)$	High volume $(n = 18)$	Low volume (n = 19)	P value	East Japan $(n = 21)$	West Japan (n = 16)	P value	
18	What kind of endoscopic treatment do you perform as first-line therapy?								
	Clipping	31 (83.8)	15 (48.4)	16 (51.6)	1.000	17 (54.8)	14 (45.2)		
	Endoscopic band ligation	5 (13.5)	3 (60.0)	2 (40.0)		3 (60.0)	2 (40.0)		
	Epinephrine injection	1 (2.7)	0	1 (100)		1 (100)	0	1.000	
19	What kinds of patient undergo non-endos	copic therapy	?1						
	Patients with an unidentified bleeding source	18 (48.7)	10 (55.6)	8 (44.4)	0.413	8 (44.4)	10 (55.6)	0.141	
	Patients with rebleeding	27 (73.0)	15 (55.6)	12 (44.4)	0.269	17 (63.0)	10 (37.0)	0.274	
	Patients with hemorrhagic shock	25 (67.6)	15 (60.0)	10 (40.0)	0.079	13 (52.0)	12 (48.0)	0.491	
20	What kind of non-endoscopic therapy do you perform as first-line therapy or when you are unable to identify SRH at endoscopy?								
	IVR	24 (77.4)	2 (50.0)	2 (50.0)	0.253	10 (41.7)	14 (58.3)		
	Surgery	3 (9.7)	14 (58.3)	10 (41.7)		3 (100)	0		
	Barium impaction therapy	4 (12.9)	0	3 (100)		3 (75.0)	1 (25.0)	0.145	
21	What kind of treatment do you perform to	prevent reble	eding? ¹						
	Treatment of diabetes mellitus	0	0	0	NA	0	0	NA	
	Treatment of hypertension	6 (17.1)	2 (33.3)	4 (66.7)	0.658	3 (50.0)	3 (50.0)	1.000	
	Discontinuation NSAIDs	14 (40.0)	7 (50.0)	7 (50.0)	0.890	10 (71.4)	4 (28.6)	0.296	
	Discontinuation antithrombotic drugs	22 (62.9)	11 (50.0)	11 (50.0)	0.826	15 (68.2)	7 (31.8)	0.086	
	Administrating vitamin D	0	0	0	NA	0	0	NA	
	Treatment of constipation	14 (40.0)	9 (64.3)	5 (35.7)	0.129	6 (42.9)	8 (57.1)	0.163	
	Administrating a low fiber diet	5 (14.3)	2 (40.0)	3 (60.0)	1.000	3 (60.0)	2 (40.0)	1.000	
22	Do you discontinue antithrombotic drugs on admission?								
	Yes	22 (59.5)	10 (45.5)	12 (54.5)		12 (54.6)	10 (46.4)		
	No	12 (32.4)	7 (58.3)	5 (41.7)		6 (50.0)	6 (50.0)		
	Case by case	3 (8.1)	1 (33.3)	2 (66.7)	0.693	3 (100)	0	0.398	
23	Do you have a strategy for restarting antithrombotic drugs? ²								
	Yes	4 (15.4)	4 (100)	0		0	4 (100)		
	No	22 (84.6)	6 (27.3)	16 (72.7)	0.014	15 (68.2)	7 (31.8)	0.022	

¹Duplicated data allowed; ²Missing data included. Values in parentheses are percentages. IVR: Interventional radiology; NSAIDs: Non-steroidal anti-inflammatory drugs; SRH: Stigmata of recent bleeding.

Comparing hospitals with high and low patient volume, low-volume hospitals had more strategies for restarting antithrombotic drugs (P=0.014) than low-volume hospitals (Table 3). No significant differences were observed in other questionnaire items between the two groups (Table 3). Comparing hospitals in East and West Japan, East Japan hospitals had less strategies for restarting antithrombotic drugs than West Japan hospitals (P=0.022) (Table 3). No significant differences were observed in other questionnaire items between the two groups (Table 3).

Questionnaire items for clinical outcomes

Questions and answers regarding clinical outcomes are shown in Table 4. The rate of identification of SRH varied widely among hospitals. No significant association between SRH identification rate and type of procedure was observed from questionnaire answers (Table 5). Forty-one percent of hospitals experienced less than 20% rebleeding events after endoscopic hemostasis, interventional radiology, and barium impaction therapy. No significant association was observed between rebleeding rate and endoscopic treatments from questionnaire answers (Table 5). No significant differences between hospitals with high and low patient volumes were observed in all questionnaire items (Table 4). Comparing hospitals in East and West Japan, East

Japan hospitals experienced less rebleeding events after barium impaction therapy than West Japan hospitals (P = 0.005). No significant differences were observed in other questionnaire items between the two groups (Table 4).

DISCUSSION

Our questionnaire-based study was the first investigation to evaluate the current practice for CDB such as clinical settings, diagnoses, treatments, and clinical outcomes in 37 hospitals nationwide in Japan. Although the clinical setting such as the number of endoscopists and nursing staff were different between hospitals with high and low patient volumes, the practice for CDB was almost the same throughout Japan, such as performing CT before colonoscopy, various procedures to improve SRH identification rate, and clipping as first-line endoscopic therapy, irrespective of hospital size.

In regard to clinical settings, a high number of emergency visits, endoscopists, and expert endoscopists were observed in high-volume hospitals compared with low-volume hospitals. CDB is a major cause of acute lower gastrointestinal bleeding, and CDB patients experience severe bleeding and require transfusion and intensive care because of their advanced age or comorbidities^[8,17-19]. Therefore, the management of CDB



Table 4 Questions and answers regarding clinical outcomes of colonic diverticular bleeding in 37 hospitals n (%)

No.	Question	Answer $(n = 37)$	High volume $(n = 18)$	Low volume $(n = 19)$	P value	East Japan $(n = 21)$	West Japan $(n = 16)$	P value		
24	How often do you identify SRH in patients who undergo colonoscopy? ¹									
	0%-20%	15 (41.7)	6 (40.0)	9 (60.0)		7 (46.7)	8 (53.3)			
	21%-40%	16 (44.4)	7 (43.8)	9 (56.2)		10 (62.5)	6 (37.5)			
	41%-60%	4 (11.1)	4 (100)	0		3 (75.0)	1 (25.0)			
	61%-80%	1 (2.8)	0	1 (100)		0	1 (100)			
	81%-100%	0	0	0	0.122	0	0	0.658		
25	How often do you ex	perience rebleeding e	vents after endoscopic	hemostasis?1						
	0%-20%	22 (61.1)	10 (45.5)	12 (54.6)		13 (59.1)	9 (40.9)			
	21%-40%	10 (27.8)	4 (40.0)	6 (60.0)		7 (70.0)	3 (30.0)			
	41%-60%	3 (8.3)	2 (66.7)	1 (33.3)		1 (33.3)	2 (66.7)			
	61%-80%	1 (2.8)	1 (100)	0		0	1 (100)			
	81%-100%	0	0	0	0.721	0	0	0.458		
26	How often do you experience rebleeding events after IVR? ¹									
	0%-20%	27 (90.1)	15 (55.6)	12 (44.4)		16 (59.3)	11 (40.7)			
	21%-40%	0	0	0		0	0			
	41%-60%	1 (3.3)	0	1 (100)		0	1 (100)			
	61%-80%	1 (3.3)	1	1 (100)		0	1 (100)			
	81%-100%	1 (3.3)	1 (100)	0	0.448	0	1 (100)	0.090		
27	How often do you ex	perience rebleeding e	vents after barium imp	paction therapy?1						
	0%-20%	10 (71.6)	6 (60.0)	4 (40.0)		9 (90.0)	1 (10.0)			
	21%-40%	1 (7.1)	0	1 (100)		0	1 (100)			
	41%-60%	1 (7.1)	0	1 (100)		0	1 (100)			
	61%-80%	1 (7.1)	1 (100)	0		0	1 (100)			
	81%-100%	1 (7.1)	0	1 (100)	0.559	0	1 (100)	0.005		

¹Missing data included. Values in parentheses are percentages. SRH: Stigmata of recent hemorrhage; IVR: Interventional radiology.

Table 5 Association between procedures and outcomes n(%)

	Answer	SRH identification rate ²								
Procedure ¹ (Question No. 15)	(n = 37)	0%-20% (n = 15)	21%-40% (n = 16)	41%-60% (n = 4)	61%-80% (n = 1)	81%-100% (n = 0)	P for trend			
Cap-assisted colonoscopy	17 (46.0)	4 (25.0)	9 (56.3)	2 (12.5)	1 (6.2)	0	0.081			
Long cap-assisted colonoscopy	13 (35.1)	6 (46.2)	5 (38.5)	2 (15.3)	0	0	0.735			
Inverting diverticulum <i>via</i> suction of colonoscopy	18 (48.7)	5 (29.4)	10 (58.8)	2 (11.8)	0	0	0.588			
Wash out with water	36 (97.3)	14 (40.0)	16 (45.7)	4 (11.4)	1 (2.9)	0	0.323			
Colonoscopy by multiple doctors	3 (8.1)	2 (66.7)	1 (33.3)	0	0	0	0.328			
Colonoscopy under X-ray	3 (8.1)	2 (66.7)	1 (33.3)	0	0	0	0.328			
	Answer			Rebleeding	rate ²					
Endoscopic treatment		0%-20%	21%-40%	41%-60%	61%-80%	81%-100%	P for trend			
(Question No. 18)	(n = 37)	(n = 22)	(n = 10)	(n = 3)	(n = 1)	(n = 0)				
Clipping	31 (83.8)	19 (63.3)	8 (26.7)	3 (10.0)	0	0	0.290			
Endoscopic band ligation	5 (13.5)	2 (40.0)	2 (40.0)	0	1 (20.0)	0	0.142			
Epinephrine injection	1 (2.7)	1 (100)	0	0	0	0	0.489			

¹Duplicated data allowed; ²Missing data included. Values in parentheses are percentages. SRH: Stigmata of recent bleeding.

patients requires an adequate number of medical staff and expert endoscopists, and a careful nursing system during the nighttime and weekend. However, there was no significant difference in the number of CDB patients who received treatment between high- and low-volume hospitals, which indicated that low-volume hospitals also need to treat CDB patients as well as high-volume hospitals regardless of the small number of endoscopists. Therefore, action is needed to handle an increasing number of CDB patients, such as transfer of CDB patients to core hospitals in each region.

In regard to diagnostic methods, most Japanese

hospitals performed CT before colonoscopy for CDB diagnosis, and there were no significant differences between the groups separated by hospital volume and region. In contrast, Western countries may perform colonoscopy or scintigraphy, not CT^[20]. This is probably because there were some studies from Japan that showed the usefulness of CT for the diagnosis of CDB, which had a sensitivity of 20.0%-42.9% and specificity of 78.6%-87.5%^[13,21]. Only 46% of hospitals performed bowel preparation, and there was a significant difference between East and West Japan in this respect. This is probably because some physicians are concerned



that bowel preparation potentially increases the risk of aspiration pneumonia, volume overload, and a change in vital signs with blood loss^[22]. However, the presence of colonic diverticula with poor visualization was a risk factor for perforation in screening colonoscopy^[23]. Recent studies have shown that bowel preparation during acute lower gastrointestinal bleeding did not increase adverse events compared with non-gastrointestinal bleeding^[24], and bowel preparation for early colonoscopy was safe as well as for elective colonoscopy^[25]. In addition, bowel preparation contributes to excellent SRH identification rates^[24,26]. Therefore, we may need to expand awareness of the safety of full bowel preparation in CDB diagnosis in Japan. Moreover, the rate of early colonoscopy was 43.2%. Now, we are conducting a randomized control study to resolve these unclarified issues in the diagnostic methods (UMIN 000021129).

In endoscopic treatment, clipping, band ligation, and epinephrine injection were performed as first-line therapy in 83.8%, 13.5%, and 2.7% of cases, which might be different from Western countries^[27]. Some reports have indicated that Western countries usually performed thermal contact therapy^[18,26,28,29]; however, this therapy is not approved in Japan^[30]. Several reports from Western countries showed that clipping was a useful hemostasis treatment^[12,31,32], and clipping may be performed as a common endoscopic treatment for CDB patients. On the other hand, in Japan, endoscopic band ligation was reported as useful for hemostasis in CDB, and therapeutic options for CDB have been expanding in Japan^[33].

There was very limited data on the strategy for antithrombotic drugs in patients with acute gastrointestinal bleeding. The American Society for Gastrointestinal Endoscopy guidelines reported^[34] that endoscopic hemostasis was considered as a procedure with a high risk of bleeding, and recommended that: (1) patients requiring endoscopic hemostasis taking non-steroidal anti-inflammatory drugs or low-dose aspirin continue these medications; (2) those taking thienopyridine should have the medication discontinued; and (3) those taking anticoagulants should consider bridging therapy. In contrast, Japan and European countries have no guidelines on the management of antithrombotic drugs in patients with gastrointestinal bleeding. Only 15% of hospitals have a strategy for antithrombotic drugs, and the timing of discontinuation and restart of antithrombotic drugs were individualized. Physicians considered discontinuation of antithrombotic therapy in patients following a hospitalization for gastrointestinal bleeding^[35,36]. Discontinued use of antithrombotic drugs may decrease the risk of gastrointestinal bleeding, but discontinuation of these drugs was associated with an increased risk of thrombosis and mortality ${}^{[37,38]}$. Although there is no consensus, we believed that patients with antithrombotic drugs need to have these medications continued, or restarted as soon as possible if patients discontinued antithrombotic drugs.

Our study has several strengths. First, our data

were obtained from a large number of hospitals, so the generalizability of the results is high. Second, we evaluated intra-observer agreement, and our data showed a high level of reproducibility. However, our study has limitations. Our study was based on data from a questionnaire, and not based on patient data, so caution should be exercised in the interpretation of our results. In addition, our study has the potential of selection bias.

In conclusion, compared with Western countries, some practice styles unique to Japan such as performing CT before colonoscopy, no bowel preparation, and clipping as first-line endoscopic therapy were found. Although the number of endoscopists and nursing staff were different, the practices for CDB were almost the same, irrespective of the size of the hospital in Japan.

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COMMENTS

Background

Colonic diverticular bleeding (CDB) is increasing in Asia however there are no practice guidelines for CDB. It is important to determine which recommendation is acceptable to a majority of hospitals.

Research frontiers

To clarify the current state of the clinical settings, diagnosis, treatment, and clinical outcomes of patients with CDB.

Innovations and breakthroughs

The authors conducted multicenter questionnaire surveys of 37 hospitals in Japan regarding management of CDB such as the clinical settings, diagnosis, treatment, and clinical outcomes, comparing them between hospitals with different patient volumes and between hospitals in different regions. As a result, some practice styles unique to Japan such as performing computed tomography before colonoscopy, no bowel preparation, and clipping as first-line therapy were found. However, the management of CDB was common among hospitals irrespective of hospital size and region.

Applications

These data were obtained from a large number of hospitals, so the generalizability of the results is high.

Peer-review

This multicenter trial by questionnaire is very useful for assessment of current state of diagnosis and treatment of CDB.



REFERENCES

- Steinmuller DR, Graneto D, Swift C, Novick AC, Streem SB, Cunningham RJ, Hodge E, Bretan P. Use of intravenous immunoglobulin prophylaxis for primary cytomegalovirus infection post living-related-donor renal transplantation. *Transplant Proc* 1989; 21: 2069-2071 [PMID: 2540554]
- Nagata N, Niikura R, Aoki T, Shimbo T, Kishida Y, Sekine K, Tanaka S, Okubo H, Watanabe K, Sakurai T, Yokoi C, Akiyama J, Yanase M, Mizokami M, Uemura N. Lower GI bleeding risk of nonsteroidal anti-inflammatory drugs and antiplatelet drug use alone and the effect of combined therapy. *Gastrointest Endosc* 2014; 80: 1124-1131 [PMID: 25088922 DOI: 10.1016/j.gie.2014.06.039]
- 3 Tsuruoka N, Iwakiri R, Hara M, Shirahama N, Sakata Y, Miyahara K, Eguchi Y, Shimoda R, Ogata S, Tsunada S, Sakata H, Fujimoto K. NSAIDs are a significant risk factor for colonic diverticular hemorrhage in elder patients: evaluation by a case-control study. *J Gastroenterol Hepatol* 2011; 26: 1047-1052 [PMID: 21198829 DOI: 10.1111/j.1440-1746.2010.06610.x]
- 4 Nagata N, Niikura R, Aoki T, Shimbo T, Itoh T, Goda Y, Suda R, Yano H, Akiyama J, Yanase M, Mizokami M, Uemura N. Increase in colonic diverticulosis and diverticular hemorrhage in an aging society: lessons from a 9-year colonoscopic study of 28,192 patients in Japan. *Int J Colorectal Dis* 2014; 29: 379-385 [PMID: 24317937 DOI: 10.1007/s00384-013-1808-4]
- 5 Song JH, Kim YS, Lee JH, Ok KS, Ryu SH, Lee JH, Moon JS. Clinical characteristics of colonic diverticulosis in Korea: a prospective study. *Korean J Intern Med* 2010; 25: 140-146 [PMID: 20526386 DOI: 10.3904/kjim.2010.25.2.140]
- 6 Sharara AI, El-Halabi MM, Mansour NM, Malli A, Ghaith OA, Hashash JG, Maasri K, Soweid A, Barada K, Mourad FH, El Zahabi L. Alcohol consumption is a risk factor for colonic diverticulosis. J Clin Gastroenterol 2013; 47: 420-425 [PMID: 23164685 DOI: 10.1097/MCG.0b013e31826be847]
- Peery AF, Barrett PR, Park D, Rogers AJ, Galanko JA, Martin CF, Sandler RS. A high-fiber diet does not protect against asymptomatic diverticulosis. *Gastroenterology* 2012; 142: 266-72.e1 [PMID: 22062360 DOI: 10.1053/j.gastro.2011.10.035]
- 8 McGuire HH. Bleeding colonic diverticula. A reappraisal of natural history and management. *Ann Surg* 1994; 220: 653-656 [PMID: 7979613 DOI: 10.1097/00000658-199411000-00008]
- 9 Niikura R, Yasunaga H, Yamaji Y, Horiguchi H, Fushimi K, Yamada A, Hirata Y, Koike K. Factors affecting in-hospital mortality in patients with lower gastrointestinal tract bleeding: a retrospective study using a national database in Japan. *J Gastroenterol* 2015; 50: 533-540 [PMID: 25181990 DOI: 10.1007/s00535-014-0994-3]
- Niikura R, Nagata N, Yamada A, Akiyama J, Shimbo T, Uemura N. Recurrence of colonic diverticular bleeding and associated risk factors. *Colorectal Dis* 2012; 14: 302-305 [PMID: 21692963 DOI: 10.1111/j.1463-1318.2011.02611.x]
- Okamoto T, Watabe H, Yamada A, Hirata Y, Yoshida H, Koike K. The association between arteriosclerosis related diseases and diverticular bleeding. *Int J Colorectal Dis* 2012; 27: 1161-1166 [PMID: 22584295 DOI: 10.1007/s00384-012-1491-x]
- 12 Kaltenbach T, Watson R, Shah J, Friedland S, Sato T, Shergill A, McQuaid K, Soetikno R. Colonoscopy with clipping is useful in the diagnosis and treatment of diverticular bleeding. *Clin Gastroenterol Hepatol* 2012; 10: 131-137 [PMID: 22056302 DOI: 10.1016/j.cgh.2011.10.029]
- Nagata N, Niikura R, Aoki T, Moriyasu S, Sakurai T, Shimbo T, Shinozaki M, Sekine K, Okubo H, Watanabe K, Yokoi C, Yanase M, Akiyama J, Uemura N. Role of urgent contrast-enhanced multidetector computed tomography for acute lower gastrointestinal bleeding in patients undergoing early colonoscopy. *J Gastroenterol* 2015; 50: 1162-1172 [PMID: 25812518 DOI: 10.1007/s00535-015-1069-9]
- 14 Finlayson EV, Goodney PP, Birkmeyer JD. Hospital volume and operative mortality in cancer surgery: a national study. *Arch Surg* 2003; 138: 721-725; discussion 726 [PMID: 12860752 DOI: 10.1001/archsurg.138.7.721]
- 15 Schreuders EH, Ruco A, Rabeneck L, Schoen RE, Sung JJ, Young

- GP, Kuipers EJ. Colorectal cancer screening: a global overview of existing programmes. *Gut* 2015; **64**: 1637-1649 [PMID: 26041752 DOI: 10.1136/gutjnl-2014-309086]
- Sim J, Wright CC. The kappa statistic in reliability studies: use, interpretation, and sample size requirements. *Phys Ther* 2005; 85: 257-268 [PMID: 15733050]
- Nagata N, Niikura R, Aoki T, Shimbo T, Sekine K, Okubo H, Watanabe K, Sakurai T, Yokoi C, Akiyama J, Yanase M, Mizokami M, Uemura N. Impact of discontinuing non-steroidal antiinflammatory drugs on long-term recurrence in colonic diverticular bleeding. World J Gastroenterol 2015; 21: 1292-1298 [PMID: 25632204 DOI: 10.3748/wjg.v21.i4.1292]
- Foutch PG. Diverticular bleeding: are nonsteroidal anti-inflammatory drugs risk factors for hemorrhage and can colonoscopy predict outcome for patients? Am J Gastroenterol 1995; 90: 1779-1784 [PMID: 7572894]
- 19 Strate LL, Orav EJ, Syngal S. Early predictors of severity in acute lower intestinal tract bleeding. *Arch Intern Med* 2003; 163: 838-843 [PMID: 12695275 DOI: 10.1001/archinte.163.7.838]
- 20 Pasha SF, Shergill A, Acosta RD, Chandrasekhara V, Chathadi KV, Early D, Evans JA, Fisher D, Fonkalsrud L, Hwang JH, Khashab MA, Lightdale JR, Muthusamy VR, Saltzman JR, Cash BD. The role of endoscopy in the patient with lower GI bleeding. *Gastrointest Endosc* 2014; 79: 875-885 [PMID: 24703084 DOI: 10.1016/j.gie.2013.10.039]
- 21 Obana T, Fujita N, Sugita R, Hirasawa D, Sugawara T, Harada Y, Oohira T, Maeda Y, Koike Y, Suzuki K, Yamagata T, Kusaka J, Masu K. Prospective evaluation of contrast-enhanced computed tomography for the detection of colonic diverticular bleeding. *Dig Dis Sci* 2013; 58: 1985-1990 [PMID: 23504354 DOI: 10.1007/s10620-013-2629-6]
- 22 Lhewa DY, Strate LL. Pros and cons of colonoscopy in management of acute lower gastrointestinal bleeding. *World J Gastroenterol* 2012; 18: 1185-1190 [PMID: 22468081 DOI: 10.3748/wjg.v18.i11.1185]
- 23 Loffeld RJ, Engel A, Dekkers PE. Incidence and causes of colonoscopic perforations: a single-center case series. *Endoscopy* 2011; 43: 240-242 [PMID: 21165826 DOI: 10.1055/s-0030-1255939]
- Niikura R, Nagata N, Aoki T, Shimbo T, Tanaka S, Sekine K, Kishida Y, Watanabe K, Sakurai T, Yokoi C, Yanase M, Akiyama J, Mizokami M, Uemura N. Predictors for identification of stigmata of recent hemorrhage on colonic diverticula in lower gastrointestinal bleeding. *J Clin Gastroenterol* 2015; 49: e24-e30 [PMID: 24859714 DOI: 10.1097/MCG.000000000000140]
- Nagata N, Niikura R, Sakurai T, Shimbo T, Aoki T, Moriyasu S, Sekine K, Okubo H, Imbe K, Watanabe K, Yokoi C, Yanase M, Akiyama J, Uemura N. Safety and Effectiveness of Early Colonoscopy in Management of Acute Lower Gastrointestinal Bleeding on the Basis of Propensity Score Matching Analysis. Clin Gastroenterol Hepatol 2016; 14: 558-564 [PMID: 26492844 DOI: 10.1016/j.cgh.2015.10.011]
- Jensen DM, Machicado GA, Jutabha R, Kovacs TO. Urgent colonoscopy for the diagnosis and treatment of severe diverticular hemorrhage. N Engl J Med 2000; 342: 78-82 [PMID: 10631275 DOI: 10.1056/NEJM200001133420202]
- 27 Strate LL, Naumann CR. The role of colonoscopy and radiological procedures in the management of acute lower intestinal bleeding. *Clin Gastroenterol Hepatol* 2010; 8: 333-343; quiz e44 [PMID: 20036757 DOI: 10.1016/j.cgh.2009.12.017]
- 28 Bloomfeld RS, Rockey DC, Shetzline MA. Endoscopic therapy of acute diverticular hemorrhage. Am J Gastroenterol 2001; 96: 2367-2372 [PMID: 11513176 DOI: 10.1111/j.1572-0241.2001.04048.x]
- Green BT, Rockey DC, Portwood G, Tarnasky PR, Guarisco S, Branch MS, Leung J, Jowell P. Urgent colonoscopy for evaluation and management of acute lower gastrointestinal hemorrhage: a randomized controlled trial. *Am J Gastroenterol* 2005; 100: 2395-2402 [PMID: 16279891 DOI: 10.1111/j.1572-0241.2005.00306.x]
- Yamada A, Niikura R, Yoshida S, Hirata Y, Koike K. Endoscopic management of colonic diverticular bleeding. *Dig Endosc* 2015; 27: 720-725 [PMID: 26258405 DOI: 10.1111/den.12534]
- 1 Yen EF, Ladabaum U, Muthusamy VR, Cello JP, McQuaid KR,



- Shah JN. Colonoscopic treatment of acute diverticular hemorrhage using endoclips. *Dig Dis Sci* 2008; **53**: 2480-2485 [PMID: 18157637 DOI: 10.1007/s10620-007-0151-4]
- 32 Simpson PW, Nguyen MH, Lim JK, Soetikno RM. Use of endoclips in the treatment of massive colonic diverticular bleeding. *Gastrointest Endosc* 2004; 59: 433-437 [PMID: 14997150 DOI: 10.1016/S0016-5107(03)02711-1]
- 33 Ishii N, Setoyama T, Deshpande GA, Omata F, Matsuda M, Suzuki S, Uemura M, Iizuka Y, Fukuda K, Suzuki K, Fujita Y. Endoscopic band ligation for colonic diverticular hemorrhage. *Gastrointest Endosc* 2012; 75: 382-387 [PMID: 21944311 DOI: 10.1016/j.gie.2011.07.030]
- 34 Anderson MA, Ben-Menachem T, Gan SI, Appalaneni V, Banerjee S, Cash BD, Fisher L, Harrison ME, Fanelli RD, Fukami N, Ikenberry SO, Jain R, Khan K, Krinsky ML, Lichtenstein DR, Maple JT, Shen B, Strohmeyer L, Baron T, Dominitz JA. Management of antithrombotic agents for endoscopic procedures. *Gastrointest Endosc* 2009; 70: 1060-1070 [PMID: 19889407 DOI: 10.1016/j.gie.2009.09.040]
- White RH, McKittrick T, Takakuwa J, Callahan C, McDonell M, Fihn S. Management and prognosis of life-threatening bleeding during warfarin therapy. National Consortium of Anticoagulation Clinics. *Arch Intern Med* 1996; 156: 1197-1201 [PMID: 8639014 DOI: 10.1001/archinte.1996.00440100095011]
- 36 Guerrouij M, Uppal CS, Alklabi A, Douketis JD. The clinical impact of bleeding during oral anticoagulant therapy: assessment of morbidity, mortality and post-bleed anticoagulant management. *J Thromb Thrombolysis* 2011; 31: 419-423 [PMID: 21181236 DOI: 10.1007/s11239-010-0536-7]
- Witt DM, Delate T, Garcia DA, Clark NP, Hylek EM, Ageno W, Dentali F, Crowther MA. Risk of thromboembolism, recurrent hemorrhage, and death after warfarin therapy interruption for gastrointestinal tract bleeding. *Arch Intern Med* 2012; 172: 1484-1491 [PMID: 22987143 DOI: 10.1001/archinternmed.2012.4261]
- 38 Rodríguez LA, Cea-Soriano L, Martín-Merino E, Johansson S. Discontinuation of low dose aspirin and risk of myocardial infarction: case-control study in UK primary care. *BMJ* 2011; 343: d4094 [PMID: 21771831 DOI: 10.1136/bmj.d4094]

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