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# Dissection-enabled scaffold-assisted resection (DeSCAR): a novel technique for resection of residual or non-lifting gastrointestinal neoplasia of the colon, expanded experience and follow-up





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## **Bibliography**

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#### **ABSTRACT**

**Background and study aims** Colonic lesions may not be amenable to conventional endoscopic mucosal resection (EMR) due to previous manipulation, submucosal invasion, or lesion flatness. In 2018, we described Dissection-enabled Scaffold Assisted Resection (DeSCAR) to be safe for the endoscopic resection of non-lifting or residual colonic

lesions [1] In this study, we expand our original cohort to describe our expanded experience with patients undergoing DeSCAR and assess the efficacy, safety, and feasibility of DeSCAR for endoscopic resection of non-lifting or residual colonic lesions.

Patients and methods We retrospectively reviewed 57 patients from 2015–2019 who underwent DeSCAR for colonic lesions with incomplete lifting and/or previous manipulation. Cases were reviewed for location, prior manipulation, rates of successful resection, adverse events, and endoscopic follow up to assess for residual lesions.

Results Fifty-seven lesions underwent DeSCAR. Of the patients, 51% were female, and average patient age was 69 years. Lesions were located in the cecum (n=16), right colon (n=27), left colon (n=10), and rectum (n=4). Average lesion size was 27.7 mm. Previous manipulation occurred in 54 cases (72% biopsy, 44% resection attempt, 18% intralesional tattoo). The technical success rate for resection of non-lifting lesions was 98%. There were two delayed bleeding episodes (one required endoscopic intervention) and one small perforation (managed by endoscopic hemoclip closure). Endoscopic follow up was available in 31 patients (54%) with no residual adenoma in 28 patients (90% of those surveilled).

**Conclusions** Our expanded experience with DeSCAR demonstrates high safety, feasibility, and effectiveness for the endoscopic management of non-lifting or residual colonic lesions.

# Introduction

Endoscopic resection of superficial colorectal lesions is highly effective for the removal of mucosal lesions with high rates of technical success, relatively low cost, and acceptably low adverse event rates in experienced centers [1–7]. Increased performance in the community setting has led to overall improvement in the en bloc removal of superficial colorectal lesions without the need for surgical intervention. However, for treat-

ment of advanced adenomas (flat/sessile lesions greater than 2 cm in size) and scarred lesions, endoscopic mucosal resection (EMR) is limited given reliance on piecemeal resection techniques in the former and inability to achieve adequate lifting in the latter. This is evidenced by local recurrence rates as high as 10% to 30% [1,2,4–15].

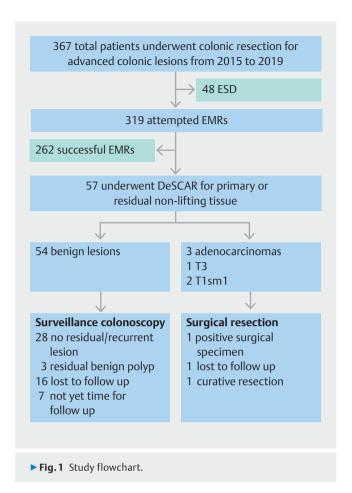
Endoscopic submucosal dissection (ESD) overcomes some of these shortcomings while allowing for en bloc resection of large superficial gastrointestinal neoplasias. The result is improved histopathologic evaluation, assessment of deeper resection margins, and lower tumor recurrence rates with technical success rates of 88% to 96% and curative resection rates approaching 90% reported in high volume centers [8, 16–26]. ESD also has good performance characteristics and technical success rates when applied to advanced adenomas – particularly those with invasion up to the superficial muscularis propria – and scarred, benign lesions not able to be removed by conventional EMR. However, because of the technical complexities of the procedure – which may result in prolonged procedure times and risk of adverse events – ESD implementation in the United States has been limited to expert, high-volume centers and has, thus, not been widely implemented in the community setting, unlike the widespread performance of EMR [4, 22–26].

Hybrid resection methods – including precut EMR-that blend the principles of EMR and ESD have also proven efficacious for en bloc, resection of colorectal neoplasias and as a rescue treatment in cases of difficult ESD [27–29]. Recent comparisons of optimized hybrid versus conventional ESD have shown comparable en bloc resection rates and adverse event profiles with significantly shorter mean procedure times in the hybrid ESD groups, but, like ESD, are limited to expert centers [28, 29].

As a result of the above challenges inherent to en bloc and complete removal of these lesions, novel techniques are necessary to address these lesions, ideally ones that could be readily available, easy to adopt, and allow for complete histologic evaluation of the lesion of interest. Dissection-enabled scaffold-assisted resection (DeSCAR) is a new technique that can allow for complete resection of superficial non-lifting lesions that remain after failed lifting and/or snaring attempts because of fibrosis or submucosal invasion [1]. This hybrid, technical variation of ESD combines EMR and ESD electrosurgical principles to allow for en bloc snaring of residual, non-lifting, post-EMR islands for histopathologic review, and our group has previously demonstrated the feasibility, safety and efficacy of this technique. [1] We present our expanded experience and outcomes of the DeSCAR technique.

#### Patients and methods

A retrospective analysis of a prospectively collected database was used to identify patients referred for EMR of colonic lesions. From this database, cases were selected where lifting and/or snaring of the lesion was unsuccessful or incomplete and the DeSCAR technique was undertaken (> Fig. 1). All cases occurred between September 2015 and September 2019. Procedures were performed at the University of Chicago Medicine, a quaternary care, urban academic medical center. Of 367 cases where a complex colonic neoplasm was encountered, a total of 57 patients (16%) underwent DeSCAR. A single experienced endoscopist (I.W.) performed all procedures. Lesions were selected for DeSCAR at the discretion of the endoscopist based on primary non-lifting or incomplete EMR attempts using standard submucosal lifting techniques. Location within the colon was divided into cecum, right side of the colon (including ascending and transverse colon), left side of the colon (including descending and sigmoid colon), and rectum. Exclusion criteria included the use of ESD or EMR alone and primary refer-



ral to surgery based on invasive malignant appearance. Cases were reviewed for patient and polyp characteristics, previous manipulation, technical success rate, and adverse events. Institutional approval for the study was obtained (IRB approval number 11-0721, approved April 2012 with last renewal in June 2019).

## Procedural technique

All procedures were performed on an outpatient basis, and sedation was achieved via monitored anesthesia care. Procedures used high-definition Olympus (Olympus, Tokyo, Japan) model CF190 colonoscopes or Olympus HQ190 gastroscopes. An Olympus 12- or 15-mm translucent cap was placed on the endoscope for better visualization of the mucosa during resection. A standard 15-mm ConMed snare (Utica, New York) was used for endoscopic mucosal resection. Either an electrosurgical DualKnife (Olympus) or the tip of the 15-mm snare was used for submucosal dissection to build the scaffold. Cautery settings were variable and case dependent. Location of the lesion was determined using anatomic landmarks and depth of endoscope insertion as appropriate. Lesion size was estimated visually using comparison with endoscopic snaring devices and/ or direct ex vivo measurement before pathology processing.

In all lesions amenable to an initial EMR attempt, a mixture of methylene blue and hetastarch solution was injected into the submucosa in order to raise the lesion. If lifting was suc-



▶ Fig. 2 Paris Classification IIa + c lesion in the right colon undergoing DeSCAR with histology shown in ▶ Fig. 3.

cessful, complete or piecemeal EMR was attempted. Any suspicious tissue remaining after multiple attempts at snare placement or displaying partial or non-lifting properties were subsequently re-injected with additional methylene blue and hetastarch. A DualKnife or the tip of a 15-mm snare was then used for electrosurgical dissection to create a "scaffold" around the residual island of tissue, circumferentially exposing the lesion to the submucosal level. This scaffold was used to anchor the snaring device in a groove below the level of the scarred mucosa (>Fig. 2). Occasionally, a brief submucosal dissection was performed to further anchor the snare below the scarred or infiltrating portion of the lesion. The residual tissue was then removed by snare en bloc and submitted to pathology in its entirety in a separate jar. Preventative hemostasis and tissue ablation were undertaken on an as-needed basis at the discretion of the endoscopist. Technical success was defined by the endoscopist as complete resection of all endoscopically visible lesions. Adverse events directly related to the procedure were defined as immediate or delayed bleeding requiring hospitalization, transfusion or repeat procedure, and perforation requiring medical and/or procedural interventions. Follow-up endoscopy to detect recurrence was recommended and was performed by the referring physician or at our institution depending on patient preference.

## Outcome measurements

The primary outcomes measured were technical success rate, presence or absence of recurrence at follow up, as well as the type and frequency of adverse events. Secondary outcomes included the diagnosis of initial EMR specimens versus diagnosis from residual tissue removed via the DeSCAR technique.

# Results

In addition to our previous cohort of 29 patients who underwent DeSCAR from September 2015 through June 2017 [1], an additional 28 patients underwent DeSCAR for colonic lesions with incomplete lifting and/or snaring from July 2017 through September 2019. Cases were reviewed for location, prior manipulation, rates of successful hybrid resection, adverse events, and endoscopic follow up to assess for residual lesions. Fiftyone percent of the patients were female and 49% were male. The average age was 69 years (range 42−87 years) (▶ Table 1). Lesions were located in the cecum (n=16, 28%), right side of the colon (n=27, 47%), left side of the colon (n=10, 18%), and rectum (n=4, 7%) (▶ Table 1). The mean size of the lesions was 27.7 mm (range 10−100 mm) (▶ Table 2).

Procedural indications included carcinoma (n=2), high grade dysplasia (n=4), tubular adenomas (n=38), and serrated lesions (n=10). Pathology for 3 lesions was not available because they were not sampled before DeSCAR. Previous manipulation of any kind occurred in 95% of cases. A biopsy was pre-

► Table 1 Patient and les	ion characteristics							
Age (years)	68.5 ± 9.6							
Gender (%F/%M)	51/49							
Lesion size (mm)	27.7±16.6							
Size range (mm)	10-100							
Paris classification (n)	IIa – 42							
	lla+c-6							
	Is – 5							
	lla+ls-4							
Lesion location (n)	Cecum/ileocecal valve – 16							
	Right colon – 22							
	Transverse colon – 5							
	Left colon – 7							
	Rectum – 4							
	Other – 3							
Referring diagnosis (n)	Adenocarcinoma – 2							
	Dysplasia – 4							
	Tubular adenoma without advanced dysplasia – 38							
	Serrated adenoma – 10							
	Not available – 3							

viously performed in 72%, resection was attempted in 44%, and intralesion tattoo placement occurred in 18% (**Table 2**). Initial EMR was attempted in 88% of cases (50 patients) before using DeSCAR for residual tissue islands. DeSCAR was used as the initial technique in 12% because of overt visual scarring and/or primary inability to lift the lesion satisfactorily for snare attempts. A DualKnife (n = 32) or the tip of a 15-mm snare (n = 25) were used to create the submucosal scaffold for all cases using the DeSCAR technique.

56 lesions in this cohort were resected endoscopically for a technical success rate of 98% with the lone unresected lesion being a T3 invasive adenocarcinoma (>Fig.3). Prior to resection of the T3 lesion - a lesion previously biopsied by the referring gastroenterologist with histopathology demonstrating tubulovillous adenoma - the 0-lla+c lesion was assessed with high definition endoscopy with narrow band imaging and high magnification endoscopy lesion's pre-procedural histopathology was a tubulovillous adenoma on biopsies; the lesion was non-lifting during initial submucosal injection. On histopathologic review of the completely resected lesions there were two cases of T1sm1 adenocarcinoma (>Fig.2), 40 tubular adenomas, 11 serrated adenomas, and three samples containing only colonic mucosa (> Table 2). No patients experienced upstaging of their diagnosis based on the histology encountered in the residual island of non-lifting tissue. Three lesions demonstrated positive deep margins on histology and were referred for surgery (5%). One of these contained a T3 rectal adenocarcinoma on surgical pathology (as mentioned above); one demonstrated no residual disease in the surgical specimen; and one patient was lost to follow-up.

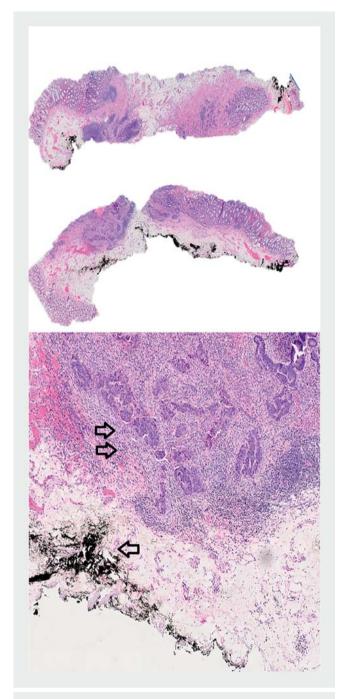
At least one method of preventative hemostasis was attempted for 86% of lesions removed by DeSCAR. This included the use of a bipolar probe (47%), metallic clips (60%), hot biopsy forceps (5%), and coagulation grasping forceps (4%). When metallic clips were placed, the average number of clips deployed was 3.7. 18 patients (32%) required two or more types of preventative hemostasis. APC was not applied in any of the cases for hemostasis or ablation of residual tissue. There were two cases of delayed bleeding (3.5%), one of which was treated endoscopically with bipolar and hot biopsy forceps electrocautery the following day, and the other managed conservatively without a need for endoscopic intervention (> Table 2). There was one case of perforation (1.8%) occurring at the time of DeSCAR, which was closed endoscopically with hemostatic clips (> Table 2). No other significant adverse events reported in any of the patients undergoing procedures using the DeSCAR technique. Follow-up endoscopy was recommended between 6 and 12 months and available for 31 patients (54%). Twentyeight patients (90% of patients with colonoscopic surveillance) had no evidence of residual polyp, and the remaining three patients had residual non-dysplastic adenomatous polyps.

## Discussion

Improved recognition of sessile and flat colorectal lesions has prompted an increase in tertiary care referrals for endoscopic resection [30–32]. However, as a result of submucosal invasion or submucosal fibrosis from previous lesion manipulation (incomplete resection attempts, extensive biopsies, or intralesional tattooing) resections even at expert centers have become more complex, given impeded lifting of those manipulated or more invasive lesions [3, 32, 33].

While generally effective as a first-line technique for the resection of colorectal neoplasia, the efficacy of EMR is best demonstrated in raised, lifting, smaller (≤2cm) lesions in which en bloc resection is achieved [1–7]. With piecemeal approaches for larger, flat lesions, there is a decreased reliability of tissue resection with rates of residual dysplasia and local recurrence as high as 10% to 30% of cases [1,2,4-10]. Thus, lesion selection for EMR is important, as submucosal fibrosis - whether resulting from deep submucosal resection by high-grade benign or malignant lesions or from scarring from previous endoscopic manipulation – can limit complete, adequate resections. Lesion non-lifting during attempted submucosal injection may be a harbinger for submucosal invasion; however, reliance on this "non-lifting sign" alone has limited sensitivity for deep submucosal (sm3) invasion, especially in the setting of previous sampling or injection [2,9,11-15]. High-risk histopathologic features - including poor differentiation, tumor depth exceeding 1000 µm, and lymphovascular invasion – are at higher risk of metastases to locoregional lymph nodes and are better served with surgical resection and nodal sampling [12–15].

	Technical success (#)	16					27					10					3 (Lone failure T3 adeno- carcino- ma)					
	Number	-	0	15			-	1	25			0	0	10			0	0	4			
	Adverse events	Delayed bleeding	Perforation	None			Delayed bleeding	Perforation	None			Delayed bleeding	Perforation	None			Delayed bleeding	Perforation	None			
	Preventa- tive hemo- stasis (#)	4					25					9					4					
	Number	8	3	4	0	-	13	9	9	1-T1sm1	-	7	-	-	0	-	1	0	1	2 (1-T3, 1-T1sm1)	0	
	Resection diagnosis	TA	SSA/SSP	TVA/VA	Adenocarcinoma	Colonic mucosa	TA	SSA/SSP	TVA/VA	Adenocarcinoma	Colonic mucosa	TA	SSA/SSP	TVA/VA	Adenocarcinoma	Colonic mucosa	TA	SSA/SSP	TVA/VA	Adenocarcinoma	Colonic mucosa	
	Number	12	10	-	-		20	6	∞	0		9	2	1	-		3	1	0	-		
	Prior manipu- lation	Biopsy	Resect	Tattoo	N/A		Biopsy	Resect	Tattoo	N/A		Biopsy	Resect	Tattoo	N/A		Biopsy	Resect	Tattoo	N/A		
	Number	12	2	0	2		21	3	1	2		7	2	1	0		2	0	2	0		
	Paris class	lla	ls	lla+c	lla + Is		lla	ls	lla+c	lla + Is		lla	ls	lla+c	lla + Is		lla	ls	lla+c	lla + Is		
ons details.	Size (range, mm)	10-40					10–100					10-40					20-70					
► Table 2 Resected lesions details.	Number	16					27					10					4					
► Table 2	Loca- tion	Cecum					Right					Left					Rectum					



▶ Fig. 3 Polypectomy specimen demonstrating T1sm1 adenocarcinoma (double arrow) after DeSCAR with intralesional tattoo (arrow) seen on histopathology.

ESD has demonstrated particular effectiveness for the resection of lesions that are otherwise not amenable to en bloc snare removal by EMR with improved histologic diagnosis, lower recurrence rates and reasonably comparable safety outcomes for the management of tumors with submucosal involvement. This approach has continued to diffuse into major American and European institutions with expert center complete resection rates as high as 88% to 96% and curative resection near 90%, outperforming the same performance outcomes of EMR [2, 8,16–24].

While other prospective and European studies have been less optimistic, when applied to well-selected lesions, en bloc and curative resections remain high and compare favorably to EMR and surgical interventions [16,17,22,23,25]. Although the adoption of advanced endoscopic dissection techniques represents an important advancement in the field of therapeutic endoscopy, in Western centers these procedures are limited to specialized, high-volume centers due to the technical complexities, prolonged procedure times, and risk of adverse events [2, 4, 21–24, 26].

Hybrid endoscopic resection techniques that combine the ease and safety profile of EMR with the complete resection potential of ESD are also effective strategies for en bloc resection of colorectal lesions and offer a reasonable rescue treatment for cases of difficult ESD [27]. A more recent prospective, randomized controlled study comparing optimized hybrid ESD to standard ESD demonstrated similar en bloc resection rates among both groups (94.1% vs 100%; P=.493) with shorter procedure duration in the hybrid ESD group while maintaining a comparable adverse event rate profile [28, 29].

Endoscopic full-thickness resection (EFTR) is a newer technique wherein an over-the-scope clip is deployed over an adequately grasped/suctioned lesion (typically through a transparent cap), and the lesion is subsequently removed with a snare [34-39]. The technical success of EFTR for benign, recurrent adenomas with non-lifting sign, advanced histopathological findings, or submucosal involvement approached 75% to 100% and histologically complete resection (R0) rates of 80% to 93% has led to curative resection of advanced benign and malignant lesions, obviating the need for surgery, particularly for lesions limited to the superficial submucosa without evidence of lymphovascular invasion or poor grade of differentiation [35-39]. Applied to previously manipulated lesions, R0 resection rates approach 89% [35–39]. However, while surgery can be avoided in many cases in which EFTR is used, R0 resection rates are lower for lesions greater than 2 cm in size (33 %-87%) and those that contain adenocarcinoma (67%), and a total complication rate of 5%-15% was observed in studies published to date [34-39]. A hybrid EMR-EFTR technique - wherein the lifting portions of the lesion are removed by EMR and the nonlifting portions removed by EFTR – was developed to address the size limitation of EFTR and was applied to colonic lesions with a median size of 35.5 mm with 100% R0 resection rate, no residual or recurrent adenoma at 3-month follow up, and no reported complications [40]. However, this was a limited cohort of 10 patients, and this approach warrants further study.

Argon plasma coagulation (APC) is a less invasive endoscopic method for managing residual neoplasia after unsuccessful snare removal during EMR and is attractive because of its safety profile, ease of use, and ubiquitous availability at most endoscopy centers [41]. However, data are mixed on the efficacy as a primary therapy for colorectal neoplasia, as multiple studies have suggested that APC is an independent risk factor for recurrent neoplasia, occurring in as high as59% of cases [6, 10, 42].

Cold or hot avulsion techniques – that is, grasping of nonlifting tissue using forceps with or without the application high-frequency cutting current and mechanical traction – have been described as methods to remove difficult to resect neoplastic tissue and has demonstrated more favorable rates of local recurrence when compared to APC in some studies [42–46]. This technique is particularly useful for residual lesions unable to be removed with conventional EMR whether due to difficult locations or inability to capture the lesion with the snare. Incomplete histologic evaluation of the non-lifting central scardue to tissue fragmentation and sample degradation due to cautery effects, the latter seen in the case of hot avulsion – as well as a non-negligible tumor recurrence rate remaining over 10% are the main limitations of these techniques [46].

While the above techniques used to endoscopically remove scarred, non-lifting colorectal lesions are feasible strategies for resection of these historically difficult-to-resect colorectal lesions, the improved recognition of sessile and flat colorectal lesions; limitations of EMR for large, sessile colorectal lesions; complexities associated with standard ESD and EMR-ESD techniques; lack of widespread adoption of ESD or EFTR; and lack of consistent efficacy of ablative or avulsion therapies have unsurprisingly led to an increase in tertiary care referrals for endoscopic resection of colorectal lesions [3, 31, 32]. However, as a result of submucosal invasion or submucosal fibrosis from previous lesion manipulation (incomplete resection attempts, extensive biopsies, or intralesional tattooing) resections even at expert centers have become more complex, given impeded lifting of those manipulated or more invasive lesions [3, 33, 41-43].

DeSCAR is a hybrid endoscopic resection technique blending the ease and speed of EMR with a minimal submucosal dissection, allowing for en bloc, intact pathologic specimens. This results in improved diagnostic accuracy for the identification of advanced disease while assessing the adequacy of resection. This improvement in removal of non-lifting lesions also has the potential to increase rates of curative resection over piecemeal EMR. Similar to other techniques utilizing pre-cut mucosal and submucosal tracts - techniques utilizing biopsy forceps, grasp or snare devices through the second channel of a dual channel endoscope, or tissue anchoring/traction-assisted methods the achieved rates of complete resection were slightly above conventional EMR with comparable safety and limited additional procedural time [47-52]. However, these techniques have not resulted in equivalent complete resection rates as ESD [50-52]. Also, none of these studies were applied in patients with residual invasive and/or fibrosed areas of tissue after primary piecemeal EMR attempts.

In our study, the technical success rate for resection of residual or primary non-lifting tissue using DeSCAR was 98%, demonstrating its feasibility in expert hands and continued technical success in our additional patient cohort. It is important to note that our observed complete resection rates are noticeably higher than those reported in EMR among non-lifting, scarred lesions, which have been historically difficult lesions to resect endoscopically and were often referred for surgery in the past. This observation is likely a result of the margination of the lesion with a scaffold that extends to the level of the submucosa, allowing the snare to capture the entire lesion at a deeper level

than what is capable with conventional EMR. As a result, the recurrent adenoma rate compares superiorly to piecemeal EMR and EMR for non-lifting, scarred or deeply penetrating lesions and is comparable to published data for both ESD and EFTR. Our data are limited by our small overall patient cohort of 57 patients and our limited 12-month follow up of 31 patients (54% of our total cohort). Anecdotally, our procedure times were shorter than conventional ESD for similarly sized lesions, but we did not directly record our DeSCAR procedure lengths. Further experience and follow up are needed in order to comment further on the efficacy of DeSCAR, but our results with an expanded cohort continues to support the feasibility and efficacy of this technique for resection of non-lifting lesions.

There were only three complications observed (5%), including two delayed bleeding events – one of which was promptly treated with standard endoscopic hemostatic therapies; the other was managed conservatively without the need for endoscopic therapy – and one case of perforation at the time of DeSCAR that was treated promptly with endoscopic hemoclip closure. This demonstrates a promising safety profile with delayed bleeding and perforation rates comparable with current EMR techniques. Three patients who continued in surveillance at our institution (10% of patients undergoing surveillance) had residual, non-dysplastic adenomatous tissue, which is comparable to previously published ESD data.

# Conclusion

For removal of large sessile gastrointestinal lesions, piecemeal EMR continues to be a commonly practiced technique but is limited in the setting of previously manipulated or deeply invasive neoplasias. Without universal availability or expertise, ESD is also limited in the Western hemisphere and may not be technically feasible in many cases. While application of DeSCAR in this study was limited to a small number of cases that were evaluated in a retrospective fashion, our expanded experience with DeSCAR continues to demonstrate a strong safety profile and feasibility for en bloc removal of non-lifting colorectal neoplasias and offers the possibility for more complete eradication of neoplastic tissue with improved histopathologic evaluation. Further high-quality research is needed to determine if this method can offer decreased recurrence rates and higher diagnostic accuracy as compared to other ablative modalities as well. Improved awareness within the GI community regarding avoidance of partial mucosal resection and appropriate tattoo location before specialist referral remains necessary.

#### Competing interests

Dr. Waxman is affiliated with Boston Scientific, Cook Medical, Medtronic, and Olympus. Dr. Siddiqui is affiliated with Boston Scientific, ConMed, Medtronic, and Olympus. Dr. Chapman is affiliated with Apollo Endosurgery, Boston Scientific, and Olympus

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