

## SOLICITED REVIEW ARTICLE

# Improving the utility of colonoscopy: Recent advances in practice

Crispin J Corte\*<sup>†</sup> and Rupert W Leong\*<sup>†</sup>\*Department of Gastroenterology, Concord Hospital, and <sup>†</sup>Faculty of Medicine, University of New South Wales, Sydney, Australia**Key words**

biology, colon, diagnosis and therapy.

Accepted for publication 24 June 2015.

**Correspondence**

Dr Crispin Corte, Department of Gastroenterology, 1 West, ACE Unit, Concord Hospital, Hospital Rd, Concord, NSW 2139, Australia.

Email: crispcorte@mac.com

**Conflict of interest:** The authors have no relevant disclosures or conflicts of interest.**Introduction**

Colonoscopy is a frequently performed diagnostic and therapeutic test and the primary screening tool in several nationalized bowel cancer screening programs.<sup>1–5</sup> While the cost of colonoscopy varies between countries, it is associated with a large health economic burden.<sup>6,7</sup> However, it remains the gold standard in the identification of colorectal cancer (CRC) and its precursor lesions,<sup>1</sup> as well as in diagnosing and evaluating other colonic mucosal disease such as inflammatory bowel disease (IBD).<sup>8,9</sup> Colonoscopy, screening<sup>10–13</sup>, and colonoscopic polypectomy<sup>14</sup> have been associated with earlier detection of CRC and associated improvements in prevention and CRC survival.<sup>15,16</sup>

The need to optimize the use of colonoscopy is not merely health economic. Poor colonoscopic preparation<sup>17,18</sup> and technique<sup>19–23</sup> are associated with increased miss rates of polyps and CRC precursor lesions and, in turn, interval CRC (CRC appearing between scheduled colonoscopic screening visits). Furthermore, post-colonoscopy complications can occur at increased rates when optimal techniques are not employed.<sup>24–32</sup>

There has been a considerable focus on maximizing the utility of colonoscopy. This has occurred in four key areas. Optimizing patient selection to reduce unnecessary or low yield colonoscopy has offered cost–benefit improvements in population screening. Improving quality assurance, through the development of widely accepted quality metrics for use in individual practice and the research setting, has offered measurable improvements in colonoscopic yield. Significant improvements have been demonstrated in colonoscopic technique, from the administration of preparation to the techniques employed during withdrawal of

**Abstract**

Colonoscopy is a frequently performed diagnostic and therapeutic test and the primary screening tool in several nationalized bowel cancer screening programs. There has been a considerable focus on maximizing the utility of colonoscopy. This has occurred in four key areas: Optimizing patient selection to reduce unnecessary or low yield colonoscopy has offered cost–benefit improvements in population screening. Improving quality assurance, through the development of widely accepted quality metrics for use in individual practice and the research setting, has offered measurable improvements in colonoscopic yield. Significant improvements have been demonstrated in colonoscopic technique, from the administration of preparation to the techniques employed during withdrawal of the colonoscope. Improved techniques to avoid post-procedural complications have also been developed—further maximizing the utility of colonoscopy.

The aim of this review is to summarize the recent evidence-based advances in colonoscopic practice that contribute to the optimal practice of colonoscopy.

the colonoscope. Improved techniques to avoid post-procedural complications have also been developed—further maximizing the utility of colonoscopy.

The aim of this review is to summarize the recent evidence-based advances in colonoscopic practice that contribute to the optimal practice of colonoscopy.

**Patient selection**

Colonoscopy rates vary widely between countries and do not reflect variations in disease incidence.<sup>6</sup> The cost of unnecessary normal colonoscopy is substantial, and the potential cost saving of more efficient triage of colonoscopy is large. There is poor correlation between colonic polyps and lower gastrointestinal (GI) symptoms, and symptom-based screening is not recommended.<sup>33–35</sup> Current screening programs do not stratify for established risk factors other than age; such as family history, sex, smoking status, and body mass index (BMI). Some countries have adopted fecal occult blood test (FOBT) screening to decrease the burden of colonoscopy.<sup>6</sup> This approach is hampered by a low uptake of FOBT<sup>36,37</sup> and false positive and negative results,<sup>38</sup> yet remains effective.<sup>16</sup> Using flexible sigmoidoscopy for screening is being trialled<sup>39</sup> but has the obvious limitation of inability to detect proximal lesions.

Selecting patients to undergo priority screening using scoring systems has been investigated.<sup>40–45</sup> The Asia Pacific Colorectal Screening Score (APCS) developed by Yeoh *et al.*<sup>40</sup> is the most successful (Table 1) and has been validated in both Asian and Western populations. Patients identified as high risk by this score have a 14.9% rate of advanced adenoma at colonoscopy, with

**Table 1** The Asia Pacific Colorectal Screening Score

Risk factor	Criteria	Points
Age (years)	< 50	0
	50–69	2
	≥ 70	3
Gender	Female	0
	Male	1
Family history of CRC in a first degree relative	Absent	0
	Present	2
Smoking	Never	0
	Current or past	1

A score of 0–1 defines average risk, 2–3 moderate risk, and 4–7 high risk. CRC, colorectal cancer.

advanced adenomas in 5.2% of moderate risk and 0.9% of average risk patients. This robust system will undoubtedly become an important feature in resource allocation.

Body mass index is known to be associated with CRC<sup>46–50</sup> and is not included in this scoring system, although has been in others.<sup>41,44</sup> In Western populations, addition of BMI does not augment the discriminative power of the APCS. A low BMI (< 20 kg/m<sup>2</sup>) is associated with a very low incidence of adenoma and may become a useful tool in patient selection.

A significant proportion of colonoscopy in patients with IBD is performed for dysplasia screening. Adherence to guidelines on dysplasia screening is variable and reflects widespread variations in physician knowledge.<sup>51–55</sup> As non-adherence to guidelines typically results in more frequent colonoscopy than is required in this setting, recent data suggesting that increasing physician knowledge<sup>56</sup> has resulted in improvements in adherence to guidelines is reassuring.

## Quality assurance practice

**Development of quality metrics.** Without quality metrics in colonoscopy, there would be no way to measure the success of the practice of colonoscopy or a research intervention. Several quality metrics have evolved to become not only standard endpoints in trials seeking to improve quality of colonoscopy but have become standard practice audit parameters for academic and community endoscopists.<sup>57,58</sup>

The metrics most associated with improving the yield of colonoscopy are caecal intubation rate<sup>59–63</sup> and adenoma detection rate (ADR). Other measures of optimizing colonoscopy technique are covered in the succeeding paragraphs. Caecal intubation rate is the percentage of all colonoscopies attempted where the caecal pole is able to be touched with the tip of the colonoscope.<sup>59–63</sup> Low caecal intubation rates correlate with higher rates of interval cancer.<sup>64</sup> This measure exhibits a learning curve,<sup>65</sup> correlates with endoscopist's annual case volume,<sup>66</sup> and is a measure of competence of the endoscopist, with experienced operators achieving 95% or greater.<sup>67,68</sup>

Adenoma detection rate is the percentage of screening colonoscopies in which at least one adenoma is identified.<sup>69</sup> This measure varies with colonoscopist's experience and training, exhibits a learning curve, and correlates inversely with the incidence of

interval CRC.<sup>64,70–73</sup> ADR is the primary outcome measure in most colonoscopic research.<sup>69</sup>

Alternatives to ADR are the polyp detection rate (PDR)<sup>74–76</sup> or the number of adenomas detected per colonoscopy (APC).<sup>77,78</sup> Neither of these has been as extensively validated and correlated with outcomes as the ADR. Variations in case-mix affect ADR, as well as PDR and APC offering limited advantage of these metrics in this respect.

As ADR can vary with case-mix in non-screening populations, measuring the number of adenomas per patient with adenomas may be a more robust metric in the future. Adoption of this measure would also account for the “one and done” phenomenon wherein an endoscopist undergoing audit may be unconsciously less inclined to identify second and subsequent adenomas as they do not contribute to the ADR.

Adenoma detection rate targets are 25% (30% for males, 20% for females) for screening colonoscopies, although increasing benefit has been shown at all levels of ADR higher than this.<sup>73</sup> ADR does not address sessile serrated lesions (SSA), which do not count toward ADR.<sup>69</sup> Missed sessile serrated and hyperplastic lesions in the proximal colon confer an elevated risk of CRC, and targets relating to these lesions (5% of screening colonoscopies) have been proposed.<sup>79</sup> Ongoing issues with histologic nomenclature make these lesions a less attractive audit tool. Continuous audit of intubation rates, withdrawal times, and ADR is recommended to ensure optimal quality.

**Adoption of audit practices.** Adoption of ADR, withdrawal time, and caecal intubation rate as standard audit parameters is becoming more widespread.<sup>80</sup> Data on the impact of audit are compelling,<sup>80</sup> demonstrating a positive effect on colonoscopy quality merely by applying a self-reported audit that is reproducible and sustainable over years.<sup>81</sup>

Audit adoption has been tied to funding and resource management of bowel cancer screening in several health services—most notably in the National Health Service.<sup>82</sup>

## Technique

The technique of colonoscopy is focused on increasing the amount of visualized mucosa, the amount of time spent examining that mucosa, and adopting mechanical, optical, or imaging strategies to enhance the interpretation of the visualized mucosa. Standardized reporting tools enhance the findings of colonoscopy by making them more reliably communicable, and aid in determination of outcomes of observed lesions.

**Operator.** Multiple studies have demonstrated better colonoscopic technique among gastroenterologists than surgeons, as measured by CRC in follow up.<sup>64,72,83–85</sup> This most likely reflects differences in caecal intubation rate and ADR.

**Preparation.** Among the most obvious requirements for colonoscopy is adequate preparation. Despite the evident necessity of adequate preparation, substantial proportions (at least 15%) of colonoscopies remain inadequately prepared<sup>17</sup>—decreasing the yield of those colonoscopies<sup>17,18,86</sup> and adding cost and inconvenience

to the procedure.<sup>87</sup> Poor preparation can also delay diagnosis of progressive neoplastic lesions.

Split dosing of preparation (administration of the last dose of preparation on the morning of the procedure) has improved overall preparation rates,<sup>88–91</sup> and is now standard care. Adverse preparation related events are less frequent as phosphate-based preparations are now avoided in patients with poor renal function, or at risk for renal dysfunction.<sup>92</sup>

### **Withdrawal technique**

**Withdrawal time.** Withdrawal time is the time taken to withdraw the colonoscope following a completed inspection of the caecum and ileum, to the anus—excluding the time taken to remove the polyps. Longer withdrawal times (independent of other factors) are associated with improved ADR,<sup>20,93,94</sup> although one study has shown no benefit of increasing withdrawal times.<sup>95</sup> Withdrawal time is often standardized in colonoscopy trial protocols as a result of this, and multivariate adjustments for prolonged withdrawal time must be performed as this is an independent factor affecting ADR. Withdrawal time of at least 6 min is advised as an audited quality measure of colonoscopy.<sup>69</sup>

**Position change.** Most colonoscopy is performed in the left lateral position. Improved visualization of the colonic mucosa is obtained by routinely moving the patient during the procedure.<sup>96,97</sup> Not only are small residual pools of effluent moved, but repositioning can bring previously eclipsed portions of bowel into view. Increased visualization results in improved ADR of 34% *versus* 21% ( $P=0.01$ )<sup>98</sup>. Where necessary and possible, position change should be made to improve adenoma detection.

**Retroflexion.** A large portion of missed polyps, interval cancers, and (harder to identify) flat hyperplastic or SSA are detected in the right colon.<sup>99–101</sup> The anatomy of the right colon (tall haustral folds) potentially decreased maneuverability of the colonoscope and the nature of the lesions in the right colon (more likely to be flat) contribute to this phenomenon. Several studies have examined the effect of routine retroflexion of the colonoscope in the caecum on ADR, and found a positive effect,<sup>102,103</sup> while an older study<sup>104</sup> failed to identify an effect. Where possible and safe (over 95% in one study),<sup>103</sup> retroflexion in the right colon significantly increases adenoma detection.<sup>105</sup>

**Transparent cap.** Transparent caps or hoods can be fitted to the tip of the colonoscope (cap assisted colonoscopy (CAC)) to theoretically improve visualization of colonic mucosa, and aid in navigation past folds. CAC is also used in difficult polypectomy, variceal band ligation, per-oral endoscopic myotomy, and other advanced endoscopic procedures. CAC was found to slightly (48 s) improve caecal intubation time in a meta-analysis.<sup>106</sup> The effect of CAC on caecal intubation has been studied<sup>107–118</sup> and at meta-analysis<sup>106</sup> has not been found to improve caecal intubation rates.

Due to variable reporting, meta-analysis on the effect of CAC on polyp detection is not possible, but seven of the aforementioned 12 trials have identified improvements in polyp detection.<sup>109–113,116,117</sup>

This marginal demonstrated benefit, in association with concerns regarding impairment of complete mucosal inspection due to reduction in field of view with imperfect bowel preparation adhering to the cap, have hampered universal uptake of CAC.

Endocuff (Endocuff AEC120 or AEC140, Arc Medical, Leeds, UK) is a device fitted like a cap to the end of the colonoscope coated with rubber arms.<sup>119,120</sup> It has been demonstrated to improve polyp detection in one randomized trial.<sup>121</sup> As it has similar limitations to the transparent cup, widespread uptake is not yet recommended.

**Chromoendoscopy/image enhancement.** Image enhancement with dye (methylene blue or indigo carmine), results in superior visualization of subtle mucosal lesions. Methylene blue is taken up preferentially by normal colonocytes, and dysplastic mucosa is left variably stained or unstained.<sup>122</sup> Indigo carmine is a non-absorbed dye highlighting mucosal topography.<sup>122</sup> Both of these dyes have been shown to aid in detection of dysplasia in patients with longstanding IBD.<sup>123–126</sup> Targeting biopsies with dye-spray in this setting is clearly superior to random biopsies of colonic mucosa and is now recommended by national societies.<sup>127</sup>

Outside the setting of IBD, the evidence for dye-spray chromoendoscopy is less strong. A marginal increase in small and flat, but not advanced adenomas was found in one study,<sup>78</sup> and this is counterpoised by the prolonged procedure time required for routine chromoendoscopy.<sup>78,128</sup> A systematic review of five studies indicates similar conclusions.<sup>129</sup> The time taken to administer dye-spray could be reduced by a colonic release formulation of methylene blue (Methylene Blue MMX; MMX Cosmo Technologies, Ireland), which has proof of concept in a preliminary trial<sup>130</sup>, but awaits validation.

The use of dye-spray chromoendoscopy in the detection of SSA has more compelling evidence.<sup>131</sup> One study summarizing the impact of chromoendoscopy on right-sided “hyperplastic” polyp detection in the right colon before the clinical relevance of SSA were identified demonstrated an increase in detection rate from 9% to 16% with chromoendoscopy (when looking only at the right colon)<sup>132</sup>. These findings have been echoed in two further studies, demonstrating increase in serrated lesion detection with chromoendoscopy (29.5% *vs* 46.2%,  $P<0.001$ )<sup>128</sup> and an increase in the mean number of non-neoplastic lesions per patient with chromoendoscopy (1.0 *vs* 1.8,  $P<0.001$ )<sup>78</sup>. A clinical trial to evaluate chromoendoscopy to increase yield of serrated polyps in the right colon in the FOBT positive population is now underway (CONSCOP Study; ClinicalTrials.gov identifier: NCT01972451).

Fluorescent probes allowing improved targeting of endoscopic examination to dysplastic areas have been developed and show great clinical promise in Barrett’s oesophagus.<sup>133</sup> While similar results have not yet been obtained in the colon, fluorescent labeling to improve dysplasia detection seems likely to emerge.

Electronic image enhancement uses either limitation of the wavelengths of emitted light (narrow band imaging (NBI); Olympus) or changes the reflected light intensity in processing (flexible spectral imaging color enhancement (FICE); Fujinon, iScan; Pentax). Most research has been performed on NBI, which uses light absorbed by hemoglobin, accentuating vasculature, capillary pattern, and pit-pattern in polyps.

A meta-analysis<sup>134</sup> and systematic review<sup>135</sup> demonstrated no difference in ADR between standard colonoscopy and NBI-assisted

colonoscopy. Technology has since progressed with high-definition colonoscopy becoming commonplace, and a newer generation of NBI (190-NBI) that allows a broader field of view appears to offer some benefit.<sup>136</sup> One recent tandem colonoscopy study of high-definition white light endoscopy (HD-WLE) and 190-NBI demonstrated superior ADR with 190-NBI (48.4% vs 34.4%;  $P=0.01$ ), but no difference in adenoma miss rates between the groups.<sup>136</sup>

Autofluorescence imaging (AFI) detects adenomatous tissue by capturing fluorescence between 500 and 630 nm using an excitation light source of 442 nm using two charge-coupled devices (CCD). AFI increased ADR in a tandem colonoscopy study (26% vs 18%;  $P < 0.05$ ), but the baseline ADR is low in this study (no difference has been found among experienced endoscopists on a sub-analysis).<sup>137</sup>

Electronic image enhancement in the detection of IBD-related dysplasia has been compared with dye-spray chromoendoscopy (as the standard of care). One cross-over trial of 60 patients with chronic colitis comparing dye-spray chromoendoscopy with indigo carmine and NBI (first generation) demonstrated no significant difference in lesion detection, although there was a statistically insignificant higher miss rate in the NBI arm (6/13 vs 2/13;  $P=0.2$ ). Withdrawal times were significantly longer in the indigo carmine arm (16 vs 30 min;  $P < 0.01$ ), and false positives were higher in the NBI arm (196 vs 126;  $P < 0.0001$ ).<sup>138</sup>

A second randomized controlled trial of 108 patients with chronic colitis revealed similar rates of lesion detection in an NBI (first generation) arm and a methylene blue arm (18/112 vs 26/156;  $P=0.39$ ). Withdrawal times were again significantly shorter in the NBI arm (20 vs 27 min;  $P=0.003$ ).<sup>139</sup> A third study<sup>140</sup> compared NBI (first generation) to HD-WLE and demonstrated comparable dysplasia rates only, confirming the results of other previous trials.<sup>141–143</sup>

Two recent studies have examined electronic image enhancement in the detection of IBD-related dysplasia, published only in abstract form,<sup>144,145</sup> both identifying improvements in electronic image enhancement when compared against white light endoscopy. No evidence exists demonstrating superiority of electronic image enhancement over chromoendoscopy for the detection of IBD-related dysplasia. The second generation of NBI may offer some improvements in this area. AFI in IBD-related dysplasia has been examined in one study<sup>143</sup> but there is a technical hurdle in that inflammation and adenoma fluoresce similarly.

**Hyoscine butylbromide.** Hyoscine butylbromide (scopolamine) is an antimuscarinic, anticholinergic, and antispasmodic agent with a quaternary ammonium structure. It has been used for many years and has a good safety profile. Typically, side effects from hyoscine butylbromide are mild and self-limiting.<sup>146,147</sup>

Anaphylaxis is rare,<sup>148,149</sup> and its central muscarinic effects are limited as it does not significantly cross the blood–brain barrier.<sup>146</sup>

Antispasmodic agents (such as hyoscine butylbromide, glucagon, and hyoscyamine) are commonly employed in gastrointestinal endoscopy, in particular, colonoscopy. Use of hyoscine butylbromide in colonoscopy has been examined as it relates to speed and ease of colonoscope insertion,<sup>150–160</sup> patient comfort during and post procedure,<sup>152–154,156,157,159–164</sup> and ileal intubation.<sup>165,166</sup>

Hyoscine butylbromide may increase mucosal view and polyp detection.<sup>167</sup> No fewer than four systematic reviews and meta-analyses<sup>168–171</sup> have been published on the eight trials in this area,<sup>154,172–178</sup> which draw differing conclusions. While meta-analysis does not demonstrate increased polyp detection with hyoscine butylbromide, several of the analyzed studies<sup>154,178</sup> did not have polyp detection or adenoma detection as their primary endpoint, and some are of low quality. An interesting observation from one trial<sup>177</sup> is that antispasmodics may hamper detection of flat polyps, although this has not been noted in other studies. In patients with marked colonic spasm, there is sufficient evidence to recommend the use of hyoscine butylbromide to aid polyp detection.

**New cameras/instruments.** High-definition white light endoscopy, when combined with a high-definition camera, provides superior images of the colon. Five trials have compared HD-WLE with standard WLE with conflicting results. A meta-analysis of these trials including 4422 patients demonstrated an absolute improvement of 3.5% in ADR.<sup>179</sup> HD-WLE has become the standard of care in colonoscopy.

Other technology has focused on measures to overcome the limited field of view in a standard forward viewing colonoscope. Newer generation colonoscopes employ a 170-degree forward-viewing lens, however, even with greater flexibility in the tip of the colonoscope, lesions behind folds can be missed.

The third-eye retroscope (TER; Avantis Medical Systems, Inc., Sunnyvale, CA, USA) is an auxiliary imaging device inserted through the working channel of the colonoscope and fixed into place to inspect at 180 degrees to the tip of the colonoscope. A back to back colonoscopy study of 488 patients demonstrated a 23% increase in ADR with TER.<sup>180</sup> Examinations are typically longer with this device, which needs to be removed prior to using any other devices down the colonoscope (for polypectomy) and inhibits suctioning, a significant problem in the colon. A cost–benefit analysis of this equipment is yet to be performed.

Full spectrum endoscopy (FUSE; Endochoice, Alpharetta, GA, USA) is an endoscopy system offering colonoscopes with two auxiliary cameras perpendicular to the forward-viewing lens and a three-screen viewing platform. The resulting effect is 330 degrees of mucosal view,<sup>181</sup> and a decrease in adenoma miss rate from 41% to 7% ( $P < 0.0001$ ) in one trial of 197 patients.<sup>182</sup> A further trial of FUSE in dysplasia detection reported preliminary results demonstrating dysplasia miss rates of 0% versus 77% ( $P < 0.05$ ), but has only been published in abstract form.<sup>183</sup> These results are yet to be replicated, but they represent a significant advance in adenoma detection, and increasing mucosal view is an ongoing challenge for device developers.

Ultra-magnifying technologies, confocal light endomicroscopy (CLE) and endocytoscopy (EC), have advanced considerably in recent years. CLE is now commercially available (Cellvisio, Mauna Kea Technologies, Paris, France). These technologies allow *in vivo* “histological assessment,”<sup>184</sup> and in colonoscopy may offer most in correct histological classification of polyps prior to resection and retrieval or discard, or in IBD dysplasia screening.

**Validated reporting tools.** After completion of the colonoscopy, the only ongoing information available is photo-documentation and report text. Communication between practitioners and recall of the colonoscopist is enhanced by the use of validated reporting terminology, which has undergone some advances in recent years.<sup>185</sup>

**Paris classification.** Large polyps are often referred to subspecialized colonoscopists for removal. Standard reporting terminology is important for effective preparation for such technically demanding procedures. More importantly, precise phenotyping of polyps using the Paris classification<sup>186</sup> has allowed for accurate correlation with histology, outcomes, and prognosis,<sup>187</sup> to allow for more evidence-based utilization of colonoscopic techniques.

Recent findings suggest that interobserver variability makes the Paris classification unreliable in community gastroenterology practice and that further systems may need to be developed.<sup>188</sup>

**Image enhanced endoscopy.** The histological type of most polyps can be determined with some accuracy by examination of morphology and pit pattern. The classification by Kudo *et al.*<sup>189</sup> has gained the most traction, and interobserver variability is low.<sup>190</sup> Despite this, image enhanced evaluation is reported in only a minority of community gastroenterology centers, and the majority of polyps are sent for definitive histology rather than being resected and discarded. Further, classification systems have increased accuracy by using magnifying colonoscopes (Sano *et al.*,<sup>191</sup> Kanao *et al.*,<sup>192</sup> Wada *et al.*,<sup>193</sup> and Saito *et al.*,<sup>194</sup> however, they are largely used in academic centers with routine access to magnifying colonoscopy. Pit pattern assessment, when accurate, offers the prospect of resecting and discarding benign polyps without the cost of histology and the time taken to retrieve the polyp.<sup>195–197</sup>

Hewett *et al.* propose a simplified classification system for the endoscopic diagnosis of small polyps using NBI<sup>198</sup> (Table 2). This

**Table 2** The NBI International Colorectal Endoscopic Classification

NICE criterion	Type 1	Type 2
Color	Same or lighter than background	Brown relative to background (verify color arises from vessels)
Vessels	None or isolated lacy vessels coursing across the lesion	Brown vessels surrounding white structures*
Surface pattern	Dark or white spots of uniform size, or homogenous absence of pattern	Oval, tubular, or branched white structures* surrounded by brown vessels
Most likely pathology	Hyperplastic	Adenoma

Taken from Hewett *et al.*<sup>198</sup> Can be applied using colonoscopes both with or without optical (zoom) magnification.

\*These structures may represent the pits and the epithelium of the crypt opening.

NBI, narrow band imaging; NICE, NBI International Colorectal Endoscopic Classification.

validated system allowed identification of colonic polyp histology with 89% accuracy, 98% sensitivity, and 95% negative predictive values. Subsequent study has validated an extension of this classification system to identify submucosal invasion in colorectal tumours.<sup>199</sup>

**Ulcerative colitis endoscopic index of severity.** Almost a dozen indices for the endoscopic assessment of ulcerative colitis (UC) have been developed, largely for use in clinical trials.<sup>200</sup> Two studies have reviewed these areas, one study evaluating these indices<sup>200</sup> (without direct comparison), and one comparing a small number of frequently used indices (Simple Clinical Colitis Index (SCCI), Mayo Clinic Index, and Seo Index),<sup>201</sup> only one of which is purely endoscopic (SCCI). The Pediatric Ulcerative Colitis Activity Index has been validated and correlated with symptom severity in children.

The ulcerative colitis endoscopic index of severity (UCEIS) is the first validated index for assessing UC and accounts for 86–88% of the variance between observers in the overall assessment of endoscopic severity.<sup>202,203</sup> This score is compiled at flexible sigmoidoscopy and assessed in the most severely affected colonic segment. The score is determined by assessing mucosal vascular pattern, mucosal bleeding, and erosions or ulceration and assigning the appropriate Likert scale anchor points to each descriptor. These scores are then added (Table 3). The score ranges from 0 (normal) to 8 (worst colitis), and studies have demonstrated accurate prediction across a range of endoscopic severity judged by visual analogue scale.<sup>202,203</sup> The UCEIS is largely unaffected by knowledge of clinical information.<sup>204</sup>

The UCEIS has now been correlated with clinical outcomes in acute severe colitis (ASC), and accurately predicts patients likely to fail first-line therapy (intravenous corticosteroids) and require rescue therapy with cyclosporine or infliximab, or surgery.<sup>205</sup> Of patients with a UCEIS of 7 or 8, 92.9% required rescue therapy, colectomy, or readmission. Fifty percent of those with UCEIS of 5 or greater required rescue therapy *versus* 27% of those with UCEIS less than 5 ( $P < 0.05$ ).<sup>205</sup> The use of a validated tool to reliably report endoscopic appearance in IBD is recommended.

## Reduce complications

Complications are rare events in colonoscopy, however, where they occur, they result in significant morbidity and associated health care costs. The most feared complication of colonoscopy and colonoscopic polypectomy is perforation. Due to the low point incidence of this complication, and indeed the low rate of colonoscopic complications for all non-therapeutic procedures,<sup>206–209</sup> studies of techniques to reduce complications in colonoscopy are rare. Most complications are associated with colonoscopic polypectomy, and the most frequently occurring of these is post-polypectomy hemorrhage (PPH), occurring in between 0.3% and 6.1% of polypectomies.<sup>210–215</sup> PPH is more common in larger polyps.<sup>213,214</sup> Recently, attention has been focused on polypectomy technique and reducing the risk of PPH.

**Hot biopsy/cold snare.** Most polyps encountered at colonoscopy (up to 90%) are diminutive (5 mm or less), or small (6–9 mm).<sup>216–218</sup> Hot biopsy forceps removal of small and

**Table 3** The UCEIS<sup>®</sup> is freely available for use, acknowledging that the copyright of the terminology is registered to Watson Laboratories, NJ, successors in interest to Warner Chilcott Pharmaceuticals

Descriptor (score most severe lesions)	Likert scale anchor points	Definition
Vascular pattern	Normal (0)	Normal vascular pattern with arborisation of capillaries clearly defined or with blurring or patchy loss of capillary margins
	Patchy obliteration (1)	Patchy obliteration of vascular pattern
Bleeding	Obliterated (2)	Complete obliteration of vascular pattern
	None (0)	No visible blood
	Mucosal (1)	Some spots or streaks of coagulated blood on the surface of the mucosa ahead of the scope, which can be washed away
	Luminal mild (2)	Some free liquid blood in the lumen
Erosions & ulcers	Luminal moderate or severe (3)	Frank blood in the lumen ahead of endoscope or visible oozing from mucosa after washing intra-luminal blood, or visible oozing from a hemorrhagic mucosa
	None	Normal mucosa, no visible erosions or ulcers
	Erosions (1)	Tiny ( $\leq 5$ mm) defects in the mucosa, of a white or yellow color with a flat edge
	Superficial ulcer (2)	Larger ( $> 5$ mm) defects in the mucosa, which are discrete fibrin-covered ulcers when compared with erosions, but remain superficial
	Deep ulcer (3)	Deeper excavated defects in the mucosa, with a slightly raised edge

The score is calculated by adding the number corresponding to the appropriate Likert levels for vascular pattern, bleeding, and ulceration to give a total with a range from 0 to 8.<sup>203</sup>

UCEIS, ulcerative colitis endoscopic index of severity.

diminutive polyps relied on the concept that passing electrocautery through the polypectomy site would ablate any residual tissue, and perhaps diathermy any potentially bleeding vessels. Evidence has accumulated recently that this technique is hampered by three issues, and it has fallen out of favor. The quality of the specimens obtained is often poor, having been subjected to electrocautery and being biopsy sized (rather than a complete polypectomy specimen).<sup>219</sup> The risk of thermal injury and the resulting risk of complication is elevated; hot biopsy having been associated with an increased risk of perforation.<sup>218,220</sup> The ablation of remnant polyp is no more successful in hot biopsy than in cold biopsy<sup>221</sup> (remnants of diminutive polyps removed by cold biopsy are common (up to 61% of polypectomies with this technique).<sup>222</sup> and this cold forceps polypectomy is not recommended for any but the smallest of polyps—1–3 mm, able to be removed with one bite of the forceps).<sup>223</sup>

As removal of small and diminutive polyps with forceps has become less popular, cold snare has become the technique of choice. This technique has been described in detail elsewhere.<sup>224</sup> This technique is superior to forceps removal of lesions in that it removes the lesion and a rim of tissue surrounding the lesion to ensure complete removal, and has a shorter procedure time. Histologic eradication of the polyp is superior for polyps over 4 mm.<sup>225</sup>

Cold snare polypectomy has significant advantages over hot snare polypectomy for a significant proportion of lesions. While two studies comparing these techniques demonstrated increased rates of immediate (intraprocedural) bleeding with cold snare polypectomy, this bleeding was typically mild and self-limiting.<sup>226,227</sup> The bleeding related to hot snare is typically delayed and has been shown to be more likely than cold snare in a third comparative study of these two techniques.<sup>228</sup> Bleeding at the time of the procedure has the clear advantage of being amenable to immediate therapy, rather than necessitating re-admission and sedation, and occurring outside of the hospital environment.

It is perhaps for this reason that cold snare polypectomy has been adopted in anticoagulated patients.<sup>228</sup>

**Post-polypectomy hemorrhage prophylaxis.** Post-polypectomy hemorrhage is more common in large polyps.<sup>213,214</sup> Adrenaline injection<sup>229</sup> and mechanical hemostasis<sup>230</sup> are effective in controlling immediate PPH and are widely used. The use of prophylactic measures like adrenaline injection<sup>25,27,28,31,231,30</sup> or mechanical hemostasis<sup>27,31,231</sup> to prevent PPH have been studied, but not widely adopted, or incorporated into guidelines. One meta-analysis examined the use of prophylactic hemostasis to prevent PPH,<sup>32</sup> but included smaller polyps ( $< 10$  mm), which are less likely to bleed.<sup>213,214</sup>

Given the number of small and diminutive polyps encountered, administering prophylactic hemostasis may not be practical, and a second meta-analysis including only polyps greater than 10-mm diameter was performed.<sup>24</sup> Comparing any prophylactic hemostasis to none, the pooled risk ratio (RR) for PPH was 0.35 (0.21–0.57;  $P < 0.0001$ ) number needed to treat (NNT) 11.9, cost to prevent one PPH \$547(USD). Any prophylactic mechanical hemostasis compared with epinephrine injection produced an RR for PPH of 0.28 (0.14–0.57;  $P < 0.0001$ ) NNT 13.9, cost to prevent one PPH \$1547. For polyps over 20 mm in diameter, the pooled RR comparing any mechanical hemostasis with epinephrine alone was 0.33 (0.12–0.90;  $P = 0.03$ ) NNT 16.9, cost to prevent one PPH \$2308.

## Summary

Substantial improvements in colonoscopic practice have occurred in recent years. Enhanced patient selection and triage are now available, and providers of colonoscopy should be encouraged to look for more reliable and efficient ways to implement these. During colonoscopy, a number of measures to improve colonoscopic

yield have been developed, and many are incorporated into quality assurance programs—rapidly becoming standard practice in many countries. High-level evidence supports the use of routine audit of completion rate, preparation adequacy adenoma detection rate, and complication rates to improve outcomes and ensure quality. Similarly, improvements such as split preparation, increased withdrawal time, right colon retroflexion, chromoendoscopy for IBD surveillance, and the use of high-definition colonoscopes are supported by high-level evidence. With experience, the use of validated reporting tools and image enhancement provide superior information and can guide management and prognosis. Prevention of complications has largely been focused on polypectomy technique, although high-level evidence now exists demonstrating the efficacy of prophylactic haemostatic measures in large polypectomy.

Spasmolytic agents, AFI, routine chromoendoscopy, the transparent cap, and other devices mentioned are yet to accumulate high-level evidence, and their routine use is controversial. Position change is not yet routine despite substantial evidence in support of this practice, perhaps due to difficulties moving the sedated patient. Screening populations with scores such as the APCS is evidence-based, but so far this tool has not been recommended by government bodies or national gastroenterology societies.

## References

- Winawer S, Fletcher R, Rex D *et al.* Colorectal cancer screening and surveillance: clinical guidelines and rationale—update based on new evidence. *Gastroenterology* 2003; **124**: 544–60.
- Winawer SJ, Fletcher RH, Miller L *et al.* Colorectal cancer screening: clinical guidelines and rationale. *Gastroenterology* 1997; **112**: 594–642.
- Guidelines for the Prevention, Early Detection and Management of Colorectal Cancer*. Sydney: The Cancer Council Australia and Australian Cancer Network, 2005.
- Levin B, Lieberman DA, McFarland B *et al.* Screening and surveillance for the early detection of colorectal cancer and adenomatous polyps, 2008: a joint guideline from the American Cancer Society, the US Multi-Society Task Force on colorectal cancer, and the American College Of Radiology. *Gastroenterology* 2008; **134**: 1570–95.
- Improving Outcomes in Colorectal Cancers*. London: National Institute for Health and Clinical Excellence, 2004.
- Improving Colonoscopy Services in Australia*. Canberra: The National Bowel Cancer Screening Program Quality Working Group, 2009.
- Sonnenberg A, Delcò F, Inadomi JM. Cost-effectiveness of colonoscopy in screening for colorectal cancer. *Ann. Intern. Med.* 2000; **133**: 573–84.
- East JE. Colonoscopic cancer surveillance in inflammatory bowel disease: what's new beyond random biopsy? *Clin Endosc* 2012; **45**: 274–7.
- Dignass A, Eliakim R, Magro F *et al.* Second European evidence-based consensus on the diagnosis and management of ulcerative colitis part 1: definitions and diagnosis. *J. Crohns Colitis* 2012; **6**: 965–90.
- Selby JV, Friedman GD, Quesenberry CP Jr., Weiss NS. A case-control study of screening sigmoidoscopy and mortality from colorectal cancer. *N. Engl. J. Med.* 1992; **326**: 653–7.
- Hardcastle JD, Chamberlain JO, Robinson MH *et al.* Randomised controlled trial of faecal-occult-blood screening for colorectal cancer. *Lancet* 1996; **348**: 1472–7.
- Kronborg O, Fenger C, Olsen J, Jorgensen OD, Sondergaard O. Randomised study of screening for colorectal cancer with faecal-occult-blood test. *Lancet* 1996; **348**: 1467–71.
- Mandel JS, Church TR, Bond JH *et al.* The effect of fecal occult-blood screening on the incidence of colorectal cancer. *N. Engl. J. Med.* 2000; **343**: 1603–7.
- Winawer SJ, Zauber AG, Ho MN *et al.* The national polyp study. *Eur. J. Cancer Prev.* 1993; **2**(Suppl): 283–7.
- Espey DK, Wu XC, Swan J *et al.* Annual report to the nation on the status of cancer, 1975–2004, featuring cancer in American Indians and Alaska natives. *Cancer* 2007; **110**: 2119–52.
- Cole SR, Tucker GR, Osborne JM, Byrne SE, Bampton PA, Fraser RJ, Young GP. Shift to earlier stage at diagnosis as a consequence of the national bowel cancer screening program. *Med. J. Aust.* 2013; **198**: 327–30.
- Froehlich F, Wietlisbach V, Gonvers JJ, Burnand B, Vader JP. Impact of colonic cleansing on quality and diagnostic yield of colonoscopy: the European panel of appropriateness of gastrointestinal endoscopy European multicenter study. *Gastrointest. Endosc.* 2005; **61**: 378–84.
- Harewood GC, Sharma VK, de Garmo P. Impact of colonoscopy preparation quality on detection of suspected colonic neoplasia. *Gastrointest. Endosc.* 2003; **58**: 76–9.
- Rex DK. Colonoscopic withdrawal technique is associated with adenoma miss rates. *Gastrointest. Endosc.* 2000; **51**: 33–6.
- Barclay RL, Vicari JJ, Doughty AS, Johanson JF, Greenlaw RL. Colonoscopic withdrawal times and adenoma detection during screening colonoscopy. *N. Engl. J. Med.* 2006; **355**: 2533–41.
- Simmons DT, Harewood GC, Baron TH, Petersen BT, Wang KK, Boyd-Enders F, Ott BJ. Impact of endoscopist withdrawal speed on polyp yield: implications for optimal colonoscopy withdrawal time. *Aliment. Pharmacol. Ther.* 2006; **24**: 965–71.
- Sanchez W, Harewood GC, Petersen BT. Evaluation of polyp detection in relation to procedure time of screening or surveillance colonoscopy. *Am. J. Gastroenterol.* 2004; **99**: 1941–5.
- Corte CJ, Kim AH, Kaffes AJ. M1417: objective measures of colonoscopic difficulty: correlation with polyp detection and operator perception. *Gastrointest. Endosc.* 2010; **71**: AB215–6.
- Corte CJ, Burger DC, Horgan G, Bailey AA, East JE. Postpolypectomy haemorrhage following removal of large polyps using mechanical haemostasis or epinephrine: a meta-analysis. *United European Gastroenterol. J.* 2014; **2**: 123–30.
- Rohde H, Guenther MW, Budde R, Muhlhofer H. Randomized trial of prophylactic epinephrine-saline injection before snare polypectomy to prevent bleeding. *Endoscopy* 2000; **32**: 1004–5.
- Iishi H, Tatsuta M, Narahara H, Iseki K, Sakai N. Endoscopic resection of large pedunculated colorectal polyps using a detachable snare. *Gastrointest. Endosc.* 1996; **44**: 594–7.
- Di Giorgio P, De Luca L, Calcagno G, Rivellini G, Mandato M, De Luca B. Detachable snare versus epinephrine injection in the prevention of postpolypectomy bleeding: a randomized and controlled study. *Endoscopy* 2004; **36**: 860–3.
- Dobrowolski S, Dobosz M, Babicki A, Dymecki D, Hac S. Prophylactic submucosal saline-adrenaline injection in colonoscopic polypectomy: prospective randomized study. *Surg. Endosc.* 2004; **18**: 990–3.
- Paspatis GA, Paraskeva K, Theodoropoulou A *et al.* A prospective, randomized comparison of adrenaline injection in combination with detachable snare versus adrenaline injection alone in the prevention of postpolypectomy bleeding in large colonic polyps. *Am. J. Gastroenterol.* 2006; **101**.
- Lee SH, Chung IK, Kim SJ *et al.* Comparison of postpolypectomy bleeding between epinephrine and saline submucosal injection for large colon polyps by conventional polypectomy: a prospective

- randomized, multicenter study. *World J. Gastroenterol.* 2007; **13**: 2973–7.
- 31 Kouklakis G, Mpoumpouris A, Gatopoulou A, Efraimidou E, Manolas K, Lirantzopoulos N. Endoscopic resection of large pedunculated colonic polyps and risk of postpolypectomy bleeding with adrenaline injection versus endoloop and hemoclip: a prospective, randomized study. *Surg. Endosc.* 2009; **23**: 2732–7.
  - 32 Li LY, Liu QS, Li L, Cao YJ, Yuan Q, Liang SW, Qu CM. A meta-analysis and systematic review of prophylactic endoscopic treatments for postpolypectomy bleeding. *Int. J. Colorectal Dis.* 2011; **26**: 709–19.
  - 33 Adelstein BA, Irwig L, Macaskill P, Turner RM, Chan SF, Katelaris PH. Who needs colonoscopy to identify colorectal cancer? Bowel symptoms do not add substantially to age and other medical history. *Aliment. Pharmacol. Ther.* 2010; **32**: 270–81.
  - 34 Adelstein BA, Macaskill P, Chan SF, Katelaris PH, Irwig L. Most bowel cancer symptoms do not indicate colorectal cancer and polyps: a systematic review. *BMC Gastroenterol.* 2011; **11**: 65.
  - 35 Adelstein BA, Macaskill P, Turner RM, Katelaris PH, Irwig L. The value of age and medical history for predicting colorectal cancer and adenomas in people referred for colonoscopy. *BMC Gastroenterol.* 2011; **11**: 97.
  - 36 Rees CJ, Bevan R. The National Health Service bowel cancer screening program: the early years. *Expert Rev. Gastroenterol. Hepatol.* 2013; **7**: 421–37.
  - 37 Christou A, Katzenellenbogen JM, Thompson SC. Australia's national bowel cancer screening program: does it work for indigenous Australians? *BMC Public Health* 2010; **10**: 373.
  - 38 Hewitson P, Glasziou Paul P, Irwig L, Towler B, Watson E: Screening for colorectal cancer using the faecal occult blood test, hemoccult. In: *Cochrane Database of Syst. Rev.* Oxford: John Wiley & Sons, Ltd, 2007.
  - 39 Bevan R, Rubin G, Sofianopoulou E, Patnick J, Rees CJ. Implementing a national flexible sigmoidoscopy screening program: results of the english early pilot. *Endoscopy* 2015; **47**: 225–31.
  - 40 Yeoh KG, Ho KY, Chiu HM *et al.* The Asia-pacific colorectal screening score: a validated tool that stratifies risk for colorectal advanced neoplasia in asymptomatic Asian subjects. *Gut* 2011; **60**: 1236–41.
  - 41 Wang JY, Li ZT, Zhu YM, Wang WC, Ma Y, Liu YL. Utility of the Asia-pacific colorectal screening scoring system and the presence of metabolic syndrome components in screening for sporadic colorectal cancer. *World J. Gastroenterol.* 2014; **20**: 11394–9.
  - 42 Levitzky BE, Brown CC, Heeren TC, Schroy PC, 3rd. Performance of a risk index for advanced proximal colorectal neoplasia among a racially/ethnically diverse patient population (risk index for advanced proximal neoplasia). *Am. J. Gastroenterol.* 2011; **106**: 1099–106.
  - 43 Schroy PC 3rd, Coe AM, Mylvaganam SR *et al.* The your disease risk index for colorectal cancer is an inaccurate risk stratification tool for advanced colorectal neoplasia at screening colonoscopy. *Cancer prevention research* 2012; **5**: 1044–52.
  - 44 Kim DH, Cha JM, Shin HP, Joo KR, Lee JI, Park DI. Development and validation of a risk stratification-based screening model for predicting colorectal advanced neoplasia in Korea. *J. Clin. Gastroenterol.* 2014; **49**: 41–9.
  - 45 Wong MC, Lam TY, Tsoi KK *et al.* A validated tool to predict colorectal neoplasia and inform screening choice for asymptomatic subjects. *Gut* 2014; **63**: 1130–6.
  - 46 Liu CS, Hsu HS, Li CI *et al.* Central obesity and atherogenic dyslipidemia in metabolic syndrome are associated with increased risk for colorectal adenoma in a Chinese population. *BMC Gastroenterol.* 2010; **10**: 51.
  - 47 Shen Z, Wang S, Ye Y, Yin M, Yang X, Jiang K, Liu Y. Clinical study on the correlation between metabolic syndrome and colorectal carcinoma. *ANZ J. Surg.* 2010; **80**: 331–6.
  - 48 Minor RK, Allard JS, Younts CM, Ward TM, de Cabo R. Dietary interventions to extend life span and health span based on calorie restriction. *J. Gerontol. A Biol. Sci. Med. Sci.* 2010; **65**: 695–703.
  - 49 Meynet O, Ricci JE. Caloric restriction and cancer: molecular mechanisms and clinical implications. *Trends Mol. Med.* 2014; **20**: 419–27.
  - 50 Kitahara CM, Berndt SI, de Gonzalez AB, Coleman HG, Schoen RE, Hayes RB, Huang WY. Prospective investigation of body mass index, colorectal adenoma, and colorectal cancer in the prostate, lung, colorectal, and ovarian cancer screening trial. *J. Clin. Oncol.* 2013; **31**: 2450–9.
  - 51 Bernstein CN, Weinstein WM, Levine DS, Shanahan F. Physicians' perceptions of dysplasia and approaches to surveillance colonoscopy in ulcerative colitis. *Am. J. Gastroenterol.* 1995; **90**: 2106–14.
  - 52 Spiegel BM, Ho W, Esrailian E *et al.* Controversies in ulcerative colitis: a survey comparing decision making of experts versus community gastroenterologists. *Clin. Gastroenterol. Hepatol.* 2009; **7**: 168–74, 174.e161.
  - 53 Eaden JA, Ward BA, Mayberry JF. How gastroenterologists screen for colonic cancer in ulcerative colitis: an analysis of performance. *Gastrointest. Endosc.* 2000; **51**: 123–8.
  - 54 van Rijn AF, Fockens P, Siersema PD, Oldenburg B. Adherence to surveillance guidelines for dysplasia and colorectal carcinoma in ulcerative and Crohn's colitis patients in the Netherlands. *World J. Gastroenterol.* 2009; **15**: 226–30.
  - 55 Geary RB, Wakeman CJ, Barclay ML, Chapman BA, Collett JA, Burt MJ, Frizelle FA. Surveillance for dysplasia in patients with inflammatory bowel disease: a national survey of colonoscopic practice in New Zealand. *Dis. Colon Rectum* 2004; **47**: 314–22.
  - 56 Leong R, Perry J, Campbell B, Koo J, Turner I, Corte C, Fok I. Knowledge and predictors of dysplasia surveillance performance in inflammatory bowel diseases in Australia. *Gastrointest. Endosc.* 2015; **82**: 708–14.
  - 57 Rex DK, Petrini JL, Baron TH *et al.* Quality indicators for colonoscopy. *Am. J. Gastroenterol.* 2006; **101**: 873–85.
  - 58 Faigel DO, Pike IM, Baron TH *et al.* Quality indicators for gastrointestinal endoscopic procedures: an introduction. *Am. J. Gastroenterol.* 2006; **101**: 866–72.
  - 59 Rex DK. Quality in colonoscopy: cecal intubation first, then what? *Am. J. Gastroenterol.* 2006; **101**: 732–4.
  - 60 Chilton A, Rutter M. *Quality Assurance Guidelines for Colonoscopy.* Sheffield: NHS Cancer Screening Programmes; 2011.
  - 61 Rembacken B, Hassan C, Riemann JF *et al.* Quality in screening colonoscopy: position statement of the European Society of Gastrointestinal Endoscopy (ESGE). *Endoscopy* 2012; **44**: 957–68.
  - 62 Marshall JB, Barthel JS. The frequency of total colonoscopy and terminal ileal intubation in the 1990s. *Gastrointest. Endosc.* 1993; **39**: 518–20.
  - 63 Valori R, Rey JF, Atkin WS *et al.* European guidelines for quality assurance in colorectal cancer screening and diagnosis. First edition—quality assurance in endoscopy in colorectal cancer screening and diagnosis. *Endoscopy* 2012; **44**(Suppl 3): Se88–105.
  - 64 Baxter NN, Sutradhar R, Forbes SS, Paszat LF, Saskin R, Rabeneck L. Analysis of administrative data finds endoscopist quality measures associated with postcolonoscopy colorectal cancer. *Gastroenterology* 2011; **140**: 65–72.
  - 65 Lee SH, Chung IK, Kim SJ *et al.* An adequate level of training for technical competence in screening and diagnostic colonoscopy: a

- prospective multicenter evaluation of the learning curve. *Gastrointest. Endosc.* 2008; **67**: 683–9.
- 66 Shah HA, Paszat LF, Saskin R, Stukel TA, Rabeneck L. Factors associated with incomplete colonoscopy: a population-based study. *Gastroenterology* 2007; **132**: 2297–303.
- 67 Rathgaber SW, Wick TM. Colonoscopy completion and complication rates in a community gastroenterology practice. *Gastrointest. Endosc.* 2006; **64**: 556–62.
- 68 Niv Y, Hazazi R, Levi Z, Fraser G. Screening colonoscopy for colorectal cancer in asymptomatic people: a meta-analysis. *Dig. Dis. Sci.* 2008; **53**: 3049–54.
- 69 Rex DK, Schoenfeld PS, Cohen J *et al.* Quality indicators for colonoscopy. *Gastrointest. Endosc.* 2015; **81**: 31–53.
- 70 Pohl H, Robertson DJ. Colorectal cancers detected after colonoscopy frequently result from missed lesions. *Clin. Gastroenterol. Hepatol.* 2010; **8**: 858–64.
- 71 Kaminski MF, Regula J, Kraszewska E *et al.* Quality indicators for colonoscopy and the risk of interval cancer. *New Engl J Med* 2010; **362**: 1795–803.
- 72 Baxter NN, Warren JL, Barrett MJ, Stukel TA, Doria-Rose VP. Association between colonoscopy and colorectal cancer mortality in a us cohort according to site of cancer and colonoscopist specialty. *J. Clin. Oncol.* 2012; **30**: 2664–9.
- 73 Corley DA, Jensen CD, Marks AR *et al.* Adenoma detection rate and risk of colorectal cancer and death. *New Engl J Med* 2014; **370**: 1298–306.
- 74 Williams JE, Le TD, Faigel DO. Polypectomy rate as a quality measure for colonoscopy. *Gastrointest. Endosc.* 2011; **73**: 498–506.
- 75 Williams JE, Holub JL, Faigel DO. Polypectomy rate is a valid quality measure for colonoscopy: results from a national endoscopy database. *Gastrointest. Endosc.* 2012; **75**: 576–82.
- 76 Francis DL, Rodriguez-Correa DT, Buchner A, Harewood GC, Wallace M. Application of a conversion factor to estimate the adenoma detection rate from the polyp detection rate. *Gastrointest. Endosc.* 2011; **73**: 493–7.
- 77 Rex DK, Helbig CC. High yields of small and flat adenomas with high-definition colonoscopes using either white light or narrow band imaging. *Gastroenterology* 2007; **133**: 42–7.
- 78 Kahi CJ, Anderson JC, Waxman I, Kessler WR, Imperiale TF, Li X, Rex DK. High-definition chromocolonoscopy vs. high-definition white light colonoscopy for average-risk colorectal cancer screening. *Am. J. Gastroenterol.* 2010; **105**: 1301–7.
- 79 Kahi CJ, Li X, Eckert GJ, Rex DK. High colonoscopic prevalence of proximal colon serrated polyps in average-risk men and women. *Gastrointest. Endosc.* 2012; **75**: 515–20.
- 80 Kahi CJ, Ballard D, Shah AS, Mears R, Johnson CS. Impact of a quarterly report card on colonoscopy quality measures. *Gastrointest. Endosc.* 2013; **77**: 925–31.
- 81 Fraser AG, Gamble GD, Rose TR, Dunn JP. Colonoscopy audit over 10 years—what can be learnt? *N. Z. Med. J.* 2013; **126**: 25–35.
- 82 *Nhs Public Health Functions Agreement 2015–16 Service Specification No.26 Bowel Cancer Screening Programme.* UK: Department of Health, 2015.
- 83 Singh H, Nugent Z, Demers AA, Kliever EV, Mahmud SM, Bernstein CN. The reduction in colorectal cancer mortality after colonoscopy varies by site of the cancer. *Gastroenterology* 2010; **139**: 1128–37.
- 84 Rex DK, Rahmani EY, Haseman JH, Lemmel GT, Kaster S, Buckley JS. Relative sensitivity of colonoscopy and barium enema for detection of colorectal cancer in clinical practice. *Gastroenterology* 1997; **112**: 17–23.
- 85 Rabeneck L, Paszat LF, Saskin R. Endoscopist specialty is associated with incident colorectal cancer after a negative colonoscopy. *Clin. Gastroenterol. Hepatol.* 2010; **8**: 275–9.
- 86 Leibold B, Kastrinos F, Glick M, Rosenbaum AJ, Wang T, Neugut AI. The impact of suboptimal bowel preparation on adenoma miss rates and the factors associated with early repeat colonoscopy. *Gastrointest. Endosc.* 2011; **73**: 1207–14.
- 87 Rex DK, Imperiale TF, Latinovich DR, Bratcher LL. Impact of bowel preparation on efficiency and cost of colonoscopy. *Am. J. Gastroenterol.* 2002; **97**: 1696–700.
- 88 Di Palma JA, Rodriguez R, McGowan J, Cleveland M. A randomized clinical study evaluating the safety and efficacy of a new, reduced-volume, oral sulfate colon-cleansing preparation for colonoscopy. *Am. J. Gastroenterol.* 2009; **104**: 2275–84.
- 89 Kilgore TW, Abdinoor AA, Szary NM *et al.* Bowel preparation with split-dose polyethylene glycol before colonoscopy: a meta-analysis of randomized controlled trials. *Gastrointest. Endosc.* 2011; **73**: 1240–5.
- 90 Rex DK, Katz PO, Bertiger G, Vanner S, Hookey LC, Alderfer V, Joseph RE. Split-dose administration of a dual-action, low-volume bowel cleanser for colonoscopy: the see clear i study. *Gastrointest. Endosc.* 2013; **78**: 132–41.
- 91 Varughese S, Kumar AR, George A, Castro FJ. Morning-only one-gallon polyethylene glycol improves bowel cleansing for afternoon colonoscopies: a randomized endoscopist-blinded prospective study. *Am. J. Gastroenterol.* 2010; **105**: 2368–74.
- 92 Belsey J, Epstein O, Heresbach D. Systematic review: adverse event reports for oral sodium phosphate and polyethylene glycol. *Aliment. Pharmacol. Ther.* 2009; **29**: 15–28.
- 93 Rex DK. Colonoscopic withdrawal technique is associated with adenoma miss rates. *Gastrointest. Endosc.* 2000; **51**: 33–6.
- 94 Lee RH, Tang RS, Muthusamy VR *et al.* Quality of colonoscopy withdrawal technique and variability in adenoma detection rates (with videos). *Gastrointest. Endosc.* 2011; **74**: 128–34.
- 95 Sawhney MS, Cury MS, Neeman N *et al.* Effect of institution-wide policy of colonoscopy withdrawal time  $\geq 7$  minutes on polyp detection. *Gastroenterology* 2008; **135**: 1892–8.
- 96 East JE, Suzuki N, Arebi N, Bassett P, Saunders BP. Position changes improve visibility during colonoscope withdrawal: a randomized, blinded, crossover trial. *Gastrointest. Endosc.* 2007; **65**: 263–9.
- 97 East JE, Stavrandis M, Thomas-Gibson S, Guenther T, Tekkis PP, Saunders BP. A comparative study of standard vs. high definition colonoscopy for adenoma and hyperplastic polyp detection with optimized withdrawal technique. *Aliment. Pharmacol. Ther.* 2008; **28**: 768–76.
- 98 East JE, Bassett P, Arebi N, Thomas-Gibson S, Guenther T, Saunders BP. Dynamic patient position changes during colonoscope withdrawal increase adenoma detection: a randomized, crossover trial. *Gastrointest. Endosc.* 2011; **73**: 456–63.
- 99 Gill P, Rafferty H, Munday D *et al.* Proximal colon cancer and serrated adenomas—hunting the missing 10%. *Clin. Med. (London, England)* 2013; **13**: 557–61.
- 100 Kahi CJ, Hewett DG, Norton DL, Eckert GJ, Rex DK. Prevalence and variable detection of proximal colon serrated polyps during screening colonoscopy. *Clin. Gastroenterol. Hepatol.* 2011; **9**: 42–6.
- 101 Brenner H, Hoffmeister M, Arndt V, Stegmaier C, Altenhofen L, Haug U. Protection from right- and left-sided colorectal neoplasms after colonoscopy: population-based study. *J. Natl. Cancer Inst.* 2010; **102**: 89–95.
- 102 Hewett DG, Rex DK. Miss rate of right-sided colon examination during colonoscopy defined by retroflexion: an observational study. *Gastrointest. Endosc.* 2011; **74**: 246–52.
- 103 Chandran S, Parker F, Vaughan R *et al.* Right-sided adenoma detection with retroflexion versus forward-view colonoscopy. *Gastrointest. Endosc.* 2014; **81**: 608–13.
- 104 Harrison M, Singh N, Rex DK. Impact of proximal colon retroflexion on adenoma miss rates. *Am. J. Gastroenterol.* 2004; **99**: 519–22.

- 105 Dik VK, Moons LM, Siersema PD. Endoscopic innovations to increase the adenoma detection rate during colonoscopy. *World J. Gastroenterol.* 2014; **20**: 2200–11.
- 106 Morgan J, Thomas K, Lee-Robichaud H, Nelson RL, Braungart S. Transparent cap colonoscopy versus standard colonoscopy to improve caecal intubation. *Cochrane Database Syst. Rev.* 2012: Cd008211.
- 107 Tee HP, Corte C, Al-Ghamdi H *et al.* Prospective randomized controlled trial evaluating cap-assisted colonoscopy vs standard colonoscopy. *World J. Gastroenterol.* 2010; **16**: 3905–10.
- 108 Shida T, Katsuura Y, Teramoto O, Kaiho M, Takano S, Yoshidome H, Miyazaki M. Transparent hood attached to the colonoscope: does it really work for all types of colonoscopes? *Surg. Endosc.* 2008; **22**: 2654–8.
- 109 Rastogi A, Bansal A, Rao DS *et al.* Higher adenoma detection rates with cap-assisted colonoscopy: a randomised controlled trial. *Gut* 2012; **61**: 402–8.
- 110 Park S-Y, Kim H-S, Yoon K-W *et al.* Usefulness of cap-assisted colonoscopy during colonoscopic emr: a randomized, controlled trial. *Gastrointest. Endosc.* 2011; **74**: 869–75.
- 111 Matsushita M, Hajiro K, Okazaki K, Takakuwa H, Tominaga M. Efficacy of total colonoscopy with a transparent cap in comparison with colonoscopy without the cap. *Endoscopy* 1998; **30**: 444–7.
- 112 Lee YT, Lai LH, Hui AJ *et al.* Efficacy of cap-assisted colonoscopy in comparison with regular colonoscopy: a randomized controlled trial. *Am. J. Gastroenterol.* 2009; **104**: 41–6.
- 113 Kondo S, Yamaji Y, Watabe H *et al.* A randomized controlled trial evaluating the usefulness of a transparent hood attached to the tip of the colonoscope. *Am. J. Gastroenterol.* 2007; **102**: 75–81.
- 114 Choi DH, Shin HK, Lee YC, Lim CH, Jeong SK, Lee S-H, Yang HK. Efficacy of transparent cap-attached colonoscopy: does it improve the quality of colonoscopy? *J Korean Soc Coloproctol* 2010; **26**: 116–22.
- 115 de Wijkerslooth TR, Stoop EM, Bossuyt PM *et al.* Adenoma detection with cap-assisted colonoscopy versus regular colonoscopy: a randomised controlled trial. *Gut* 2012; **61**: 1426–34.
- 116 Horiuchi A, Nakayama Y. Improved colorectal adenoma detection with a transparent retractable extension device. *Am. J. Gastroenterol.* 2008; **103**: 341–5.
- 117 Hewett DG, Rex DK. Cap-fitted colonoscopy: a randomized, tandem colonoscopy study of adenoma miss rates. *Gastrointest. Endosc.* 2010; **72**: 775–81.
- 118 Harada Y, Hirasawa D, Fujita N *et al.* Impact of a transparent hood on the performance of total colonoscopy: a randomized controlled trial. *Gastrointest. Endosc.* 2009; **69**(3 Pt 2): 637–44.
- 119 Lenze F, Beyna T, Lenz P, Heinzow HS, Hengst K, Ullerich H. Endocuff-assisted colonoscopy: a new accessory to improve adenoma detection rate? Technical aspects and first clinical experiences. *Endoscopy* 2014; **46**: 610–4.
- 120 Biecker E, Floer M, Heinecke A, Strobel P, Bohme R, Schepke M, Meister T. Novel endocuff-assisted colonoscopy significantly increases the polyp detection rate: a randomized controlled trial. *J. Clin. Gastroenterol.* 2015; **49**: 413–8.
- 121 Floer M, Biecker E, Fitzlaff R *et al.* Higher adenoma detection rates with endocuff-assisted colonoscopy—a randomized controlled multicenter trial. *PLoS One* 2014; **9**: e114267.
- 122 Wong Kee Song LM, Adler DG, Chand B *et al.* Chromoendoscopy. *Gastrointest. Endosc.* 2007; **66**: 639–49.
- 123 Kiesslich R, Fritsch J, Holtmann M *et al.* Methylene blue-aided chromoendoscopy for the detection of intraepithelial neoplasia and colon cancer in ulcerative colitis. *Gastroenterology* 2003; **124**: 880–8.
- 124 Rutter MD, Saunders BP, Schofield G, Forbes A, Price AB, Talbot IC. Pancolonic indigo carmine dye spraying for the detection of dysplasia in ulcerative colitis. *Gut* 2004; **53**: 256–60.
- 125 Hurlstone DP, Sanders DS, Lobo AJ, McAlindon ME, Cross SS. Indigo carmine-assisted high-magnification chromoscopic colonoscopy for the detection and characterisation of intraepithelial neoplasia in ulcerative colitis: a prospective evaluation. *Endoscopy* 2005; **37**: 1186–92.
- 126 Marion JF, Wayne JD, Present DH *et al.* Chromoendoscopy-targeted biopsies are superior to standard colonoscopic surveillance for detecting dysplasia in inflammatory bowel disease patients: a prospective endoscopic trial. *Am. J. Gastroenterol.* 2008; **103**: 2342–9.
- 127 Cairns SR, Scholefield JH, Steele RJ *et al.* Guidelines for colorectal cancer screening and surveillance in moderate and high risk groups (update from 2002). *Gut* 2010; **59**: 666–89.
- 128 Pohl J, Schneider A, Vogell H, Mayer G, Kaiser G, Ell C. Pancolonic chromoendoscopy with indigo carmine versus standard colonoscopy for detection of neoplastic lesions: a randomised two-centre trial. *Gut* 2011; **60**: 485–90.
- 129 Brown SR, Baraza W. Chromoscopy versus conventional endoscopy for the detection of polyps in the colon and rectum. *Cochrane Database Syst. Rev.* 2010: Cd006439.
- 130 Repici A, Di Stefano AF, Radicioni MM, Jas V, Moro L, Danese S. Methylene blue mmx tablets for chromoendoscopy. Safety tolerability and bioavailability in healthy volunteers. *Contemp. Clin. Trials* 2012; **33**: 260–7.
- 131 East J, Vieth M, Rex D. Serrated lesions in colorectal cancer screening: detection, resection, pathology and surveillance. *Gut* 2015; **64**: 991–1000.
- 132 East JE, Saunders BP, Jass JR. Sporadic and syndromic hyperplastic polyps and serrated adenomas of the colon: classification, molecular genetics, natural history, and clinical management. *Gastroenterol. Clin. North Am.* 2008; **37**: 25–46, v.
- 133 Bird-Lieberman EL, Neves AA, Lao-Sirieix P *et al.* Molecular imaging using fluorescent lectins permits rapid endoscopic identification of dysplasia in Barrett's esophagus. *Nat. Med.* 2012; **18**: 315–21.
- 134 Dinesen L, Chua TJ, Kaffes AJ. Meta-analysis of narrow-band imaging versus conventional colonoscopy for adenoma detection. *Gastrointest. Endosc.* 2012; **75**: 604–11.
- 135 van den Broek FJ, Reitsma JB, Curvers WL, Fockens P, Dekker E. Systematic review of narrow-band imaging for the detection and differentiation of neoplastic and nonneoplastic lesions in the colon (with videos). *Gastrointest. Endosc.* 2009; **69**: 124–35.
- 136 Leung WK, Lo OS, Liu KS *et al.* Detection of colorectal adenoma by narrow band imaging (hq190) vs. high-definition white light colonoscopy: a randomized controlled trial. *Am. J. Gastroenterol.* 2014; **109**: 855–63.
- 137 Moriichi K, Fujiya M, Sato R *et al.* Back-to-back comparison of autofluorescence imaging (afi) versus high resolution white light colonoscopy for adenoma detection. *BMC Gastroenterol.* 2012; **1275**.
- 138 Pellise M, Lopez-Ceron M, Rodriguez de Miguel C *et al.* Narrow-band imaging as an alternative to chromoendoscopy for the detection of dysplasia in long-standing inflammatory bowel disease: a prospective, randomized, crossover study. *Gastrointest. Endosc.* 2011; **74**: 840–8.
- 139 Bisschops R, Bessissow T, Baert F *et al.* Chromo-endoscopy versus narrow band imaging in ulcerative colitis: a prospective randomized controlled trial. *Gastrointest. Endosc.* 2012; **75**: 148.
- 140 Ignjatovic A, East JE, Subramanian V *et al.* Narrow band imaging for detection of dysplasia in colitis: a randomized controlled trial. *Am. J. Gastroenterol.* 2012; **107**: 885–90.
- 141 Dekker E, van den Broek FJ, Reitsma JB *et al.* Narrow-band imaging compared with conventional colonoscopy for the detection of dysplasia in patients with longstanding ulcerative colitis. *Endoscopy* 2007; **39**: 216–21.
- 142 van den Broek FJ, Fockens P, van Eeden S, Stokkers PC, Ponsioen CY, Reitsma JB, Dekker E. Narrow-band imaging versus high-

- definition endoscopy for the diagnosis of neoplasia in ulcerative colitis. *Endoscopy* 2011; **43**: 108–15.
- 143 van den Broek FJ, Fockens P, van Eeden S, Reitsma JB, Hardwick JC, Stokkers PC, Dekker E. Endoscopic tri-modal imaging for surveillance in ulcerative colitis: randomised comparison of high-resolution endoscopy and autofluorescence imaging for neoplasia detection; and evaluation of narrow-band imaging for classification of lesions. *Gut* 2008; **57**: 1083–9.
- 144 Iacucci M, Fort Gasia M, Urbanski S, *et al.* Detection and characterization of colonic dysplastic lesions in ibd surveillance colonoscopy—a randomised comparison of high definition alone with high definition dye spraying and electronic virtual chromoendoscopy using iscan. *J. Crohns Colitis* 2015; **9**: s149–150.
- 145 Cassinotti A, Ardizzone S, Buffoli F *et al.* Virtual chromoendoscopy with fice is superior to standard colonoscopic surveillance for flat visible dysplastic lesions and raised lesions (polyps and pseudopolyps) evaluation in long-standing ulcerative colitis: a prospective, randomized, trial. *J. Crohns Colitis* 2015; **9**: S1205.
- 146 Tytgat GN. Hyoscine butylbromide—a review on its parenteral use in acute abdominal spasm and as an aid in abdominal diagnostic and therapeutic procedures. *Curr. Med. Res. Opin.* 2008; **24**: 3159–73.
- 147 Grainger SL, Smith SE. Dose-response relationships of intravenous hyoscine butylbromide and atropine sulphate on heart rate in healthy volunteers. *Br. J. Clin. Pharmacol.* 1983; **16**: 623–6.
- 148 Gonzalez-Mendiola R, Sanchez Fernandez C, Prieto Montano P, Cuevas M, Cena Delgado M, Sanchez CM. Acute urticaria induced by hyoscine butylbromide. *Allergy* 2004; **59**: 787–8.
- 149 Treweeke P, Barrett NK. Allergic reaction to buscopan. *Br. J. Radiol.* 1987; **60**: 417–8.
- 150 Altintas E, Ucbilek E, Sezgin O, Sayici Y. Alverine citrate plus simethicone reduces cecal intubation time in colonoscopy—a randomized study. *Turk. J. Gastroenterol.* 2008; **19**: 174–9.
- 151 Chaptini LA, Janec EM, Seltzer G, Peikin S, Elfant AB. Sublingual hyoscyamine spray as premedication for colonoscopy: a randomized double-blinded placebo-controlled trial. *Am. J. Surg.* 2008; **196**: 51–5.
- 152 Dumot JA, Verzola E, Nicol S, Easley KA, Vargo JJ, van Stolk RU. Sublingual hyoscyamine for patient comfort during screening sigmoidoscopy: a randomized, double-blind, placebo-controlled clinical trial. *Gastrointest. Endosc.* 1998; **48**: 283–6.
- 153 Marshall JB, Patel M, Mahajan RJ, Early DS, King PD, Banerjee B. Benefit of intravenous antispasmodic (hyoscyamine sulfate) as premedication for colonoscopy. *Gastrointest. Endosc.* 1999; **49**: 720–6.
- 154 Mui LM, Ng EK, Chan KC *et al.* Randomized, double-blinded, placebo-controlled trial of intravenously administered hyoscine n-butyl bromide in patients undergoing colonoscopy with patient-controlled sedation. *Gastrointest. Endosc.* 2004; **59**: 22–7.
- 155 Norfleet RG. Premedication for colonoscopy: randomized, double-blind study of glucagon versus placebo. *Gastrointest. Endosc.* 1978; **24**: 164–5.
- 156 Norfleet RG, Saviage K. Atropine premedication for colonoscopy: a randomized double-blind study. *Gastrointest. Endosc.* 1983; **29**: 157.
- 157 Waxman I, Mathews J, Gallagher J *et al.* Limited benefit of atropine as premedication for colonoscopy. *Gastrointest. Endosc.* 1991; **37**: 329–31.
- 158 Yoong KY, Perkin D, Portal J, Strickland I, Heymann T. Intravenous hyoscine as a premedication for colonoscopy: a randomized double-blind controlled trial. *Endoscopy* 2004; **36**: 720–2.
- 159 Yoshikawa I, Yamasaki M, Taguchi M *et al.* Comparison of glucagon and scopolamine butylbromide as premedication for colonoscopy in unsedated patients. *Dis. Colon Rectum* 2006; **49**: 1393–8.
- 160 Cutler CS, Rex DK, Hawes RH, Lehman GA. Does routine intravenous glucagon administration facilitate colonoscopy? A randomized trial. *Gastrointest. Endosc.* 1995; **42**: 346–50.
- 161 Bond JH, Chally CH, Blackwood WD. A controlled trial of premedication with dicyclomine hydrochloride (bentyl) in colonoscopy. *Gastrointest. Endosc.* 1974; **21**: 61.
- 162 Elphick DA, Donnelly MT, Smith KS, Riley SA. Factors associated with abdominal discomfort during colonoscopy: a prospective analysis. *Eur. J. Gastroenterol. Hepatol.* 2009; **21**: 1076–82.
- 163 Lee YC, Wang HP, Chiu HM *et al.* Factors determining post-colonoscopy abdominal pain: prospective study of screening colonoscopy in 1000 subjects. *J. Gastroenterol. Hepatol.* 2006; **21**: 1575–80.
- 164 Takahashi Y, Tanaka H, Kinjo M, Sakumoto K. Prospective evaluation of factors predicting difficulty and pain during sedation-free colonoscopy. *Dis. Colon Rectum* 2005; **48**: 1295–300.
- 165 Ansari A, Soon SY, Saunders BP, Sanderson JD. A prospective study of the technical feasibility of ileoscopy at colonoscopy. *Scand. J. Gastroenterol.* 2003; **38**: 1184–6.
- 166 Misra SP, Dwivedi M. Role of intravenously administered hyoscine butyl bromide in retrograde terminal ileoscopy: a randomized, double-blinded, placebo-controlled trial. *World J. Gastroenterol.* 2007; **13**: 1820–3.
- 167 Froehlich F. Colonoscopy: antispasmodics not only for premedication, but also during endoscope withdrawal? *Gastrointest. Endosc.* 2000; **51**: 379.
- 168 Rondonotti E, Zolk O, Amato A, Paggi S, Baccarin A, Spinzi G, Radaelli F. The impact of hyoscine-n-butylbromide on adenoma detection during colonoscopy: meta-analysis of randomized, controlled studies. *Gastrointest. Endosc.* 2014; **80**: 1103–1112.e1102.
- 169 Madhoun MF, Ali T, Tierney WM, Maple JT. Effect of hyoscine n-butylbromide on adenoma detection rate: meta-analysis of randomized clinical trials. *Dig. Endosc.* 2015; **27**: 354–60.
- 170 Cui PJ, Yao J, Han HZ, Zhao YJ, Yang J. Does hyoscine butylbromide really improve polyp detection during colonoscopy? a meta-analysis of randomized controlled trials. *World J. Gastroenterol.* 2014; **20**: 7034–9.
- 171 Ashraf I, Ashraf S, Siddique S, Nguyen DL, Choudhary A, Bechtold ML. Hyoscine for polyp detection during colonoscopy: a meta-analysis and systematic review. *World J Gastrointest Endosc* 2014; **6**: 549–54.
- 172 Lee JM, Cheon JH, Park JJ, Moon CM, Kim ES, Kim TI, Kim WH. Effects of hyosine n-butyl bromide on the detection of polyps during colonoscopy. *Hepatogastroenterology* 2010; **57**: 90–4.
- 173 Byun TJ, Han DS, Ahn SB *et al.* Role of intravenous hyoscine n-butyl bromide at the time of colonoscopic withdrawal for polyp detection rates: a randomized, double-blinded, placebo-controlled trial. *Gastrointest. Endosc.* 2009; **69**: AB229.
- 174 Corte C, Dahlenburg L, Selby W, Griffin S, Byrne C, Chua T, Kaffes A. Hyoscine butylbromide administered at the cecum increases polyp detection: a randomized double-blind placebo-controlled trial. *Endoscopy* 2012; **44**: 917–22.
- 175 Brouwer EJ, Arbouw ME, van der Zwet WC, van Herwaarden MA, Ledebor M, Jansman FG, Ter Borg F. Hyoscine n-butylbromide does not improve polyp detection during colonoscopy: a double-blind, randomized, placebo-controlled, clinical trial. *Gastrointest. Endosc.* 2012; **75**: 835–40.
- 176 Han DS, Song SC, Park JY *et al.* A clinical usefulness of premedication with hyoscine-n-butyl bromide (buscopan(r)) in colonoscopy: a randomized, double blinded, prospective study. *Korean J Gastrointest Endosc* 1997; **17**: 346–50.
- 177 Rondonotti E, Radaelli F, Paggi S *et al.* Hyoscine n-butylbromide for adenoma detection during colonoscopy: a randomized, double-blind, placebo-controlled study. *Dig. Liver Dis.* 2013; **45**: 663–8.
- 178 Saunders BP, Williams CB. Premedication with intravenous antispasmodic speeds colonoscope insertion. *Gastrointest. Endosc.* 1996; **43**: 209–11.

- 179 Subramanian V, Mannath J, Hawkey CJ, Raguath K. High definition colonoscopy vs. standard video endoscopy for the detection of colonic polyps: a meta-analysis. *Endoscopy* 2011; **43**: 499–505.
- 180 Leufkens AM, DeMarco DC, Rastogi A *et al.* Effect of a retrograde-viewing device on adenoma detection rate during colonoscopy: the terrace study. *Gastrointest. Endosc.* 2011; **73**: 480–9.
- 181 Gralnek IM, Segol O, Suissa A *et al.* A prospective cohort study evaluating a novel colonoscopy platform featuring full-spectrum endoscopy. *Endoscopy* 2013; **45**: 697–702.
- 182 Gralnek IM, Siersema PD, Halpern Z *et al.* Standard forward-viewing colonoscopy versus full-spectrum endoscopy: an international, multicentre, randomised, tandem colonoscopy trial. *Lancet Oncol.* 2014; **15**: 353–60.
- 183 Ooi M, Panetta J, Zhu M, Corte C, Leong R. Forward-viewing colonoscopy versus full-spectrum endoscopy (FUSE) for dysplasia detection with and without chromoendoscopy in inflammatory bowel diseases: a prospective, randomized-order, crossover tandem surveillance colonoscopy study. *J. Gastroenterol. Hepatol.* 2014; **29** (Suppl. 2): 115–6.
- 184 Nakai Y, Isayama H, Shinoura S, Iwashita T, Samarasena JB, Chang KJ, Koike K. Confocal laser endomicroscopy in gastrointestinal and pancreatobiliary diseases. *Dig. Endosc.* 2014; **26**(Suppl): 186–94.
- 185 Aabakken L, Barkun AN, Cotton PB *et al.* Standardized endoscopic reporting. *J. Gastroenterol. Hepatol.* 2014; **29**: 234–40.
- 186 Update on the Paris classification of superficial neoplastic lesions in the digestive tract. *Endoscopy* 2005; **37**: 570–8.
- 187 Moss A, Bourke MJ, Williams SJ *et al.* Endoscopic mucosal resection outcomes and prediction of submucosal cancer from advanced colonic mucosal neoplasia. *Gastroenterology* 2011; **140**: 1909–18.
- 188 van Doorn SC, Hazewinkel Y, East JE *et al.* Polyp morphology: an interobserver evaluation for the Paris classification among international experts. *Am. J. Gastroenterol.* 2015; **110**: 180–7.
- 189 Kudo S, Rubio CA, Teixeira CR, Kashida H, Kogure E. Pit pattern in colorectal neoplasia: endoscopic magnifying view. *Endoscopy* 2001; **33**: 367–73.
- 190 Huang Q, Fukami N, Kashida H *et al.* Interobserver and intra-observer consistency in the endoscopic assessment of colonic pit patterns. *Gastrointest. Endosc.* 2004; **60**: 520–6.
- 191 Sano Y, Horimatsu T, Fu KI, Katagiri A, Muto M, Ishikawa H. Magnifying observation of microvascular architecture of colorectal lesions using a narrow-band imaging system. *Dig. Endosc.* 2006; **18**: S44–51.
- 192 Kanao H, Tanaka S, Oka S, Hirata M, Yoshida S, Chayama K. Narrow-band imaging magnification predicts the histology and invasion depth of colorectal tumors. *Gastrointest. Endosc.* 2009; **69**(3 Pt 2): 631–6.
- 193 Wada Y, Kudo SE, Kashida H *et al.* Diagnosis of colorectal lesions with the magnifying narrow-band imaging system. *Gastrointest. Endosc.* 2009; **70**: 522–31.
- 194 Saito S, Tajiri H, Ohya T, Nikami T, Aihara H, Ikegami M. Imaging by magnifying endoscopy with NBI implicates the remnant capillary network as an indication for endoscopic resection in early colon cancer. *Int. J. Surg. Oncol.* 2011; **2011**: 242608–10.
- 195 Hassan C, Pickhardt PJ, Rex DK. A resect and discard strategy would improve cost-effectiveness of colorectal cancer screening. *Clin. Gastroenterol. Hepatol.* 2010; **8**: 865–9, 869.e861–863.
- 196 Hattori S, Iwatate M, Sano W *et al.* Narrow-band imaging observation of colorectal lesions using nice classification to avoid discarding significant lesions. *World J Gastrointest. Endosc.* 2014; **6**: 600–5.
- 197 Takeuchi Y, Hanafusa M, Kanzaki H, Ohta T, Hanaoka N. Proposal of a new 'resect and discard' strategy using magnifying narrow band imaging: pilot study of diagnostic accuracy. *Dig. Endosc.* 2014; **26** (Suppl) 290–7.
- 198 Hewett DG, Kaltenbach T, Sano Y *et al.* Validation of a simple classification system for endoscopic diagnosis of small colorectal polyps using narrow-band imaging. *Gastroenterology* 2012; **143**: 599–607.e591.
- 199 Hayashi N, Tanaka S, Hewett DG *et al.* Endoscopic prediction of deep submucosal invasive carcinoma: validation of the narrow-band imaging international colorectal endoscopic (nice) classification. *Gastrointest. Endosc.* 2013; **78**: 625–32.
- 200 D'Haens G, Sandborn WJ, Feagan BG *et al.* A review of activity indices and efficacy end points for clinical trials of medical therapy in adults with ulcerative colitis. *Gastroenterology* 2007; **132**: 763–86.
- 201 Walsh AJ, Ghosh A, Brain AO *et al.* Comparing disease activity indices in ulcerative colitis. *J. Crohns Colitis* 2014; **8**: 318–25.
- 202 Travis SP, Schnell D, Krzeski P *et al.* Developing an instrument to assess the endoscopic severity of ulcerative colitis: the ulcerative colitis endoscopic index of severity (UCEIS). *Gut* 2012; **61**: 535–42.
- 203 Travis SP, Schnell D, Krzeski P *et al.* Reliability and initial validation of the ulcerative colitis endoscopic index of severity. *Gastroenterology* 2013; **145**: 987–95.
- 204 Travis S, Schnell P, Feagan B *et al.* The impact of clinical information on the assessment of endoscopic activity: characteristics of the ulcerative colitis endoscopic index of severity [UCEIS]. *J Crohns Colitis* 2015; **9**: 607–16.
- 205 Corte C, Fernandopulle A, Catuneanu A *et al.* Association between the ulcerative colitis endoscopic index of severity (UCEIS) and outcomes in acute severe ulcerative colitis. *Journal of Crohn's and Colitis* 2015; **9**: 376–81.
- 206 Ko CW, Dominitz JA. Complications of colonoscopy: magnitude and management. *Gastrointest. Endosc. Clin. N. Am.* 2010; **20**: 659–71.
- 207 Ko CW, Riffle S, Michaels L *et al.* Serious complications within 30 days of screening and surveillance colonoscopy are uncommon. *Clin. Gastroenterol. Hepatol.* 2010; **8**: 166–73.
- 208 Panteris V, Haringsma J, Kuipers EJ. Colonoscopy perforation rate, mechanisms and outcome: from diagnostic to therapeutic colonoscopy. *Endoscopy* 2009; **41**: 941–51.
- 209 Rabeneck L, Paszat LF, Hilsden RJ *et al.* Bleeding and perforation after outpatient colonoscopy and their risk factors in usual clinical practice. *Gastroenterology* 2008; **135**: 1899–906, 1906.e1891.
- 210 Wayne JD, Lewis BS, Yessayan S. Colonoscopy: a prospective report of complications. *J. Clin. Gastroenterol.* 1992; **15**: 347–51.
- 211 Gibbs DH, Opelka FG, Beck DE, Hicks TC, Timmcke AE, Gathright JB Jr. Postpolypectomy colonic hemorrhage. *Dis. Colon Rectum* 1996; **39**: 806–10.
- 212 Wayne JD, Kahn O, Auerbach ME. Complications of colonoscopy and flexible sigmoidoscopy. *Gastrointest. Endosc. Clin. N. Am.* 1996; **6**: 343–77.
- 213 Rosen L, Bub DS, Reed JF, 3rd, Nastase SA. Hemorrhage following colonoscopic polypectomy. *Dis. Colon Rectum* 1993; **36**: 1126–31.
- 214 Sorbi D, Norton I, Conio M, Balm R, Zinsmeister A, Gostout CJ. Postpolypectomy lower GI bleeding: descriptive analysis. *Gastrointest. Endosc.* 2000; **51**: 690–6.
- 215 Levin TR, Zhao W, Conell C, Seeff LC, Manninen DL, Shapiro JA, Schulman J. Complications of colonoscopy in an integrated health care delivery system. *Ann. Intern. Med.* 2006; **145**: 880–6.
- 216 von Karsa L, Patnick J, Segnan N. European guidelines for quality assurance in colorectal cancer screening and diagnosis. First edition—executive summary. *Endoscopy* 2012; **44**(Suppl 3): Se1–8.
- 217 Regula J, Rupinski M, Kraszewska E *et al.* Colonoscopy in colorectal-cancer screening for detection of advanced neoplasia. *N. Engl. J. Med.* 2006; **355**: 1863–72.
- 218 Anderloni A, Jovani M, Hassan C, Repici A. Advances, problems, and complications of polypectomy. *Clin Exp Gastroenterol* 2014; **7**: 285–96.

- 219 Monkemuller KE, Fry LC, Jones BH, Wells C, Mikolaenko I, Eloubeidi M. Histological quality of polyps resected using the cold versus hot biopsy technique. *Endoscopy* 2004; **36**: 432–6.
- 220 Metz AJ, Moss A, McLeod D, Tran K, Godfrey C, Chandra A, Bourke MJ. A blinded comparison of the safety and efficacy of hot biopsy forceps electrocauterization and conventional snare polypectomy for diminutive colonic polypectomy in a porcine model. *Gastrointest. Endosc.* 2013; **77**: 484–90.
- 221 Peluso F, Goldner F. Follow-up of hot biopsy forceps treatment of diminutive colonic polyps. *Gastrointest. Endosc.* 1991; **37**: 604–6.
- 222 Efthymiou M, Taylor AC, Desmond PV, Allen PB, Chen RY. Biopsy forceps is inadequate for the resection of diminutive polyps. *Endoscopy* 2011; **43**: 312–6.
- 223 Jung YS, Park JH, Kim HJ *et al.* Complete biopsy resection of diminutive polyps. *Endoscopy* 2013; **45**: 1024–9.
- 224 Hewett DG. Colonoscopic polypectomy: current techniques and controversies. *Gastroenterol. Clin. North Am.* 2013; **42**: 443–58.
- 225 Lee CK, Shim JJ, Jang JY. Cold snare polypectomy vs. cold forceps polypectomy using double-biopsy technique for removal of diminutive colorectal polyps: a prospective randomized study. *Am. J. Gastroenterol.* 2013; **108**: 1593–600.
- 226 Paspatis GA, Tribonias G, Konstantinidis K *et al.* A prospective randomized comparison of cold vs hot snare polypectomy in the occurrence of postpolypectomy bleeding in small colonic polyps. *Colorectal Dis.* 2011; **13**: e345–8.
- 227 Ichise Y, Horiuchi A, Nakayama Y, Tanaka N. Prospective randomized comparison of cold snare polypectomy and conventional polypectomy for small colorectal polyps. *Digestion* 2011; **84**: 78–81.
- 228 Horiuchi A, Nakayama Y, Kajiyama M, Tanaka N, Sano K, Graham DY. Removal of small colorectal polyps in anticoagulated patients: a prospective randomized comparison of cold snare and conventional polypectomy. *Gastrointest. Endosc.* 2014; **79**: 417–23.
- 229 Conio M, Repici A, Demarquay JF, Blanche S, Dumas R, Filiberti R. Emr of large sessile colorectal polyps. *Gastrointest. Endosc.* 2004; **60**: 234–41.
- 230 Parra-Blanco A, Kaminaga N, Kojima T *et al.* Hemoclipping for postpolypectomy and postbiopsy colonic bleeding. *Gastrointest. Endosc.* 2000; **51**: 37–41.
- 231 Paspatis GA, Paraskeva K, Theodoropoulou A *et al.* A prospective, randomized comparison of adrenaline injection in combination with detachable snare versus adrenaline injection alone in the prevention of postpolypectomy bleeding in large colonic polyps: CME. *Am. J. Gastroenterol.* 2006; **101**: 2805–9.