

# Post-polypectomy colonoscopy surveillance: European Society of Gastrointestinal Endoscopy (ESGE) Guideline



## Authors

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This Guideline is an official statement of the European Society of Gastrointestinal Endoscopy (ESGE). The Grading of Recommendations Assessment, Development, and Evaluation (GRADE) system was adopted to define the strength of recommendations and the quality of evidence.

**Main recommendations:** The following recommendations for post-polypectomy endoscopic surveillance should be applied only after a high quality baseline colonoscopy with complete removal of all detected neoplastic lesions.

**1** In the low risk group (patients with 1–2 tubular adenomas <10 mm with low grade dysplasia), the ESGE recommends participation in existing national screening programmes 10 years after the index colonoscopy. If no screening programme is available, repetition of colonoscopy 10 years after the index colonoscopy is recommended (strong recommendation, moderate quality evidence).

**2** In the high risk group (patients with adenomas with villous histology or high grade dysplasia or ≥10 mm in size, or ≥3 adenomas), the ESGE recommends surveillance colonoscopy 3 years after the index colonoscopy (strong recommendation, moderate quality evidence). Patients with 10 or more adenomas should be referred for genetic counselling (strong recommendation, moderate quality evidence).

**3** In the high risk group, if no high risk adenomas are detected at the first surveillance examination, the ESGE suggests a 5-year interval before a second surveillance colonoscopy (weak recommendation, low quality evidence). If high risk adenomas are detected at first or subsequent surveillance examinations, a 3-year repetition of surveillance colonoscopy is recommended (strong recommendation, low quality evidence).

**4** The ESGE recommends that patients with serrated polyps <10 mm in size with no dysplasia should be classified as low risk (weak recommendation, low quality evidence). The ESGE suggests that patients with large serrated polyps (≥10 mm) or those with dysplasia should be classified as high risk (weak recommendation, low quality evidence).

**5** The ESGE recommends that the endoscopist is responsible for providing a written recommendation for the post-polypectomy surveillance schedule (strong recommendation, low quality evidence).

## Abbreviations

CRC	colorectal cancer
ESGE	European Society of Gastrointestinal Endoscopy
g-FOBT/FIT	guaiac-based faecal occult blood test/ faecal immunochemical test
HGD	high grade dysplasia

## Introduction

Colorectal cancer (CRC) represents a major cause of morbidity and mortality in Western countries [1–3]. CRC screening has been shown to be effective in reducing CRC incidence and/or mortality [4–7], and population-based screening is widely recommended in Europe [8]. The effect of endoscopic screening is conveyed via two mechanisms. First, removal of precancerous adenomatous polyps at the time of the index examination and the detection of CRC at an early stage reduce CRC incidence and/or mortality [4,9–11]. Secondly, stratification based on the endoscopic

findings allows patients at greater risk to benefit from endoscopic surveillance [12–14]. Patients with adenomatous polyps are at greater risk of future development of advanced neoplasia (adenomas  $\geq 10$  mm or with unfavourable histology or cancer) [15–18]. This may be because serious lesions were missed or not radically removed at the initial examination, or because an inherent imbalance of cell proliferation in an individual leads to accelerated carcinogenesis in apparently normal mucosa [16, 19–23].

It is assumed that if patients in whom precancerous polyps have been found are entered into a surveillance programme, then metachronous or recurrent adenomatous lesions and cancer will be detected at an earlier stage. However, no randomized study has directly assessed how much benefit is contributed by the efficacy of post-polypectomy surveillance. The efficacy of endoscopic surveillance has been addressed only in epidemiological series. Such studies have indicated that patients who are not entered into a surveillance programme have a three- to fourfold greater risk of CRC [18, 23].

Screening series have reported an adenoma prevalence of 15%–30% [12, 13, 24, 25]. With the use of high definition colonoscopy equipment, adenomas are found in up to 50% of the population [26, 27]. Thus, an indiscriminate use of post-polypectomy surveillance would represent a substantial burden on endoscopy resources, also resulting in unnecessary costs and longer waiting times for other indications. Currently, close to 20% of endoscopic capacity is occupied by surveillance colonoscopies, approximately the same proportion as primary screening examinations [28–30]. With several European countries initiating population-based screening programmes, the burden of surveillance can be expected to increase in the near future. Although colonoscopy is generally regarded as a safe procedure, a risk of major complications remains [31]. In patients at increased risk of developing cancer, the balance of benefit and risk is generally regarded as favourable. However, the risks, albeit small, may become relevant if the gain associated with surveillance colonoscopies is substantially reduced.

When considering the lack of strong evidence to support post-polypectomy surveillance, and the substantial workload involved, a conservative approach would appear reasonable. It should be remembered that the aim of population-based CRC screening is to reduce the incidence and mortality of CRC, and to do so with a sustainable expenditure of medical and economic resources. For the best balance between the benefits and drawbacks of post-polypectomy surveillance, it should only be offered to patients with a substantial residual risk of CRC. Epidemiological and clinical studies have shown that it is possible to stratify the risk of CRC and to identify a small subgroup of patients with a greater incidence of CRC that persists after baseline polypectomy [21, 32].

The aim of this evidence-based and consensus-based Guideline, commissioned by the European Society of Gastrointestinal Endoscopy (ESGE), is to provide caregivers with a comprehensive review of risk stratification following removal of precancerous neoplastic lesions and with practical recommendations for scheduling endoscopic surveillance. This Guideline does not address surveillance after endoscopic or surgical resection of a malignant polyp, or surveillance in patients affected by hereditary colorectal syndromes.

## Methods



The ESGE commissioned this Guideline. The guideline development process included meetings, telephone conferences, and on-line discussions among members of the guideline committee during February 2012 and February 2013. Subgroups were formed, each in charge of a series of clearly defined key questions (Appendix 1, available online). The committee chairs (C.H., J.M.D.) worked with the subgroup leaders (J.M.D., E.Q., J.R.) to identify pertinent search terms that always included, as a minimum, “post-polypectomy endoscopic surveillance” as well as terms pertinent to specific key questions. Searches were performed in Medline. Articles were first selected by title; their relevance was then confirmed by review of the corresponding manuscripts, and articles with content that was considered irrelevant, including that relating to hereditary colorectal syndromes, were excluded. A repository of selected literature was made available to all members of the guideline development group. Evidence tables were generated for each key question, summarizing the evidence of the available studies. For important outcomes, articles were individually assessed by means of the Grades of Recommendation Assessment, Development, and Evaluation (GRADE) system for grading evidence levels and recommendation strengths (Appendix 2, available online) [33].

Each subgroup developed draft proposals that were presented to the entire group for general discussion during a meeting held in February 2013 (Düsseldorf, Germany). Further details on the methodology of ESGE guidelines have been reported elsewhere [33].

In March 2013, a draft prepared by C.H. was sent to all group members. After agreement on a final version, the manuscript was submitted to *Endoscopy* for publication. The journal subjected the manuscript to peer review, and the manuscript was amended to take into account the reviewers' comments. All authors agreed on the final revised manuscript.

This Guideline was issued in 2013 and will be considered for review in 2018, or sooner if new and relevant evidence becomes available. Any updates to the guideline in the interim will be no-

**Box 1** Main definitions adopted for this Guideline.

Term	Definition
High quality colonoscopy	Complete colonoscopy with a meticulous inspection of adequately cleaned colorectal mucosa. Neoplastic lesions have also been completely removed and retrieved for histological examination.
Index colonoscopy	First high quality colonoscopy on which surveillance strategy is based
Metachronous lesion	Any lesion that is detected at surveillance colonoscopies
Low risk group	1–2 tubular adenomas < 10 mm with low-grade dysplasia; serrated polyps < 10 mm and no dysplasia
High risk group	Adenoma with villous histology or high grade dysplasia or $\geq 10$ mm in size, or $\geq 3$ adenomas; serrated polyps $\geq 10$ mm or with dysplasia
Advanced adenoma	Adenoma with villous histology or high grade dysplasia or $\geq 10$ mm in size
Advanced neoplasia	Adenoma with villous histology or high grade dysplasia or $\geq 10$ mm in size, or colorectal cancer
Serrated polyp	Hyperplastic polyps, sessile serrated polyp, traditional serrated adenomas, and mixed lesions

ted on the ESGE website: <http://www.esge.com/esge-guidelines.html>.

## Recommendations and statements



Evidence statements and recommendations are stated in italics, key evidence statements and recommendations are in bold. See **Box 1** for main definitions for this Guideline.

***The following recommendations for post-polypectomy endoscopic surveillance should only be applied after a high quality baseline colonoscopy with complete removal of all detected neoplastic lesions.***

Colonoscopy that is performed in quality-controlled settings has been associated with a substantial reduction in subsequent CRC incidence and mortality and with a very low risk of metachronous advanced neoplasia [4, 34, 35]. However, large observational studies have recently shown that the protective effect of colonoscopy, whether or not it includes polypectomy, is reduced when it is done in suboptimal conditions [5, 36]. Moreover, studies involving head-to-head colonoscopy and computed tomography (CT) colonography and tandem colonoscopy have demonstrated that colonoscopy misses some polyps [37, 38]. In a systematic review of tandem colonoscopy studies, colonoscopy miss rates for polyps  $\geq 10$  mm, 6–9 mm, and  $< 5$  mm were found to be 2%, 13%, and 26%, respectively [37]. Endoscopist- and centre-related quality factors have been shown to predict a higher risk of interval CRC [39–41]. Endoscopists and endoscopic centres performing low quality examinations, as measured by adenoma/polyp detection rate and cecal intubation rate, have consistently been associated with a higher risk of post-colonoscopy interval CRC [39, 40, 42]. In addition, the incomplete removal of lesions has consistently been shown to increase the subsequent risk of CRC [16, 23, 43].

For these reasons, widespread implementation of quality assurance programmes is necessary for adequate efficacy of post-polypectomy surveillance. Factors associated with the quality of colonoscopy and of bowel cleansing have been reviewed in specific ESGE Guidelines [44, 45]. There is no evidence that overutilization of endoscopic surveillance can compensate for an initially suboptimal colonoscopy. Moreover, to duplicate an invasive and costly procedure, rather than to address, for example, the quality of bowel cleansing and improved endoscopist training, seems unacceptable from the point of view of cost-effectiveness and patient acceptability. Briefly, high quality colonoscopies should be complete up to the caecum with a meticulous inspection of adequately cleaned colorectal mucosa. Neoplastic lesions should be completely removed (en bloc when feasible) and retrieved for histological examination. This does not contradict early repetition of the colonoscopy if the quality of the initial procedure was suboptimal because of inadequate bowel cleansing or other factors (see below in the specific scenarios, and **Table e1**, available online).

### Appropriate scheduling of surveillance

*The ESGE recommends that the endoscopist is responsible for providing a written recommendation for the post-polypectomy surveillance schedule (strong recommendation, low quality evidence), and that this should be audited (weak recommendation, low quality evidence).*

Surveillance colonoscopies represent a major part of all colonoscopies performed, being nearly 30% in a recent survey [46]. This proportion may increase with the widespread adoption of CRC screening programmes and with improved adenoma detection

related to the use of high resolution colonoscopy and dye-spraying techniques [3, 47, 48]. For these reasons, the required capacity of colonoscopy services is heavily dependent on correct indications and timings for post-polypectomy surveillance [49–51]. Studies have shown that a large proportion of surveillance procedures are inappropriate in both selection of cases and timing of surveillance, representing both over- and underuse of surveillance [46, 51–58]. In a recent survey, 69% of post-polypectomy surveillance procedures were inappropriate regarding either timing or indication [46]. In another study, over 40% of patients with small adenomas had an inappropriately early surveillance examination [52]. Moreover, surveillance is still recommended to patients with clinically irrelevant hyperplastic lesions who do not need any endoscopic surveillance [46, 52–58]. (See also **Table e2**, available online).

Appropriateness of surveillance not only depends on the characteristics and number of completely removed polyps, but also on factors such as the quality of endoscopy, and the patient's age and life-expectancy. For these reasons, the endoscopist should be the professional who advises the patient on the appropriate surveillance interval. Since histology reports become available some time after the polypectomy, we recommend that the endoscopist updates and/or finalizes the endoscopy report after receiving the histology report; the updated colonoscopy report should include a written recommendation on the appropriate surveillance, taking into account all endoscopic, histological, and patient-related factors. Adherence to published surveillance guidelines should be monitored as part of a quality assurance programme [59–61].

### Low risk group

***In the low risk group (patients with 1–2 tubular adenomas  $< 10$  mm with low grade dysplasia), the ESGE recommends participation in existing national screening programmes 10 years after the index colonoscopy. If no screening programme is available, repetition of colonoscopy 10 years after the index colonoscopy is recommended (strong recommendation, moderate quality evidence).***

### Long-term CRC risk in low risk group (see **Table 3a**)

Epidemiological studies have assessed long-term CRC incidence/mortality risk in patients with 1–2 tubular adenomas  $< 10$  mm with low grade dysplasia. In a retrospective study [62], including 1618 patients with adenomas resected by rigid sigmoidoscopy who did not undergo endoscopic surveillance, patients stratified to this low risk group had a similar risk of developing CRC compared with the general population (standardized incidence ratio [SIR] 0.5, 95% confidence interval [CI] 0.1–1.3). The same finding was reported in a registry-based study that included 5779 post-polypectomy patients: the low risk group did not have an increased risk of cancer despite the lack of surveillance (SIR 0.68, 95%CI 0.44–0.99) [18]. Furthermore, case-control studies have also confirmed a low long-term risk of CRC in these patients, with a more profound effect during the 5 years immediately following the index polypectomy [22, 23, 63]. A conservative policy of post-polypectomy endoscopic surveillance was recently tested and found to be adequate in two prospective screening sigmoidoscopy trials [9, 11].

### Incidence of metachronous advanced neoplasia in the low risk group

Several cohort studies have compared the incidence of metachronous advanced adenomas between a low risk group and a control

**Table 3** Long-term colorectal cancer risk. **a** Studies reporting standardized incidence ratio (SIR) with 95 % confidence intervals (CI) in low and high risk groups as compared with the general population.

First author	Variable	Low risk	High risk
Atkin [62]	SIR (95 %CI)	0.5 (0.1 – 1.3)	3.6 (2.4 – 5.0)
Cottet [18]	SIR (95 %CI)	0.8 (0.4 – 1.5)	4.3 (2.9 – 6.0)

**Table 3** Long-term colorectal cancer risk. **b** Studies comparing incidence (hazard ratio [HR] or risk ratio [RR]) of metachronous advanced neoplasia between low risk and high risk groups, and patients without neoplasia at baseline examination.

First author	Variable	Low risk	High risk
Yamaji [15]	HR (95 %CI)	Nonadvanced: 2.6 (1.6 – 4.2)	Advanced: 6.6 (3.7 – 12)
Lieberman [35]	RR (95 %CI)	Tubular < 10 mm: 2.6 (0.2 – 5.7)	Tubular > 10 mm: 6.4 (2.7 – 14.9) Villous pattern: 6.1 (2.5 – 14.7) High grade dysplasia: 6.9 (2.6 – 18.1)
Miller [66]	RR (95 %CI)	< 5 mm: 1.1 (0.4 – 3.3) 5 – 9 mm: 1.5 (0.6 – 3.9) Tubular: 0.9 (0.4 – 2.3)	≥ 10 mm: 2.1 (1.3 – 2.7) Villous pattern: 4.2 (1.5 – 11.5)
Chung [64]	HR (95 %CI)	Nonadvanced: 1.1 (0.6 – 2.2)	Advanced: 6.0 (3.7 – 9.7)

CI, confidence interval

group without adenoma at index colonoscopy [15,35,64–66] (Table 3b; see also Table e4, available online). One study found a higher incidence of advanced neoplasia (hazard ratio [HR] 2.6; 95%CI 1.6–4.2) in the low risk group compared with controls [15]. None of the other studies detected a statistically significant difference, either at 5 years [35, 64–66] or at 6–10 years of follow-up [65, 66]. Two randomized controlled trials (RCTs) [35, 67], as well as three cohort studies [64, 66,68], compared the prevalence of advanced neoplasia at different intervals between the index examination and the first surveillance colonoscopy in the low risk group. No statistically significant difference was found when comparing intervals of 2 vs 4 years, 3 vs 5 years, and 3–5 vs 6–10 years [35,64,66,67,69].

### Timing of surveillance/return to screening in low risk group

For individuals without increased risk of CRC (i.e., risk similar to that in the general population) a 10-year interval before undergoing surveillance colonoscopy or returning to a screening programme appears to be justified by the long-term efficacy of lower gastrointestinal endoscopy (i.e., sigmoidoscopy or colonoscopy) as demonstrated in RCTs and case–control studies [7, 11, 70]

### High risk group

**In the high risk group (patients with adenomas with villous histology or high grade dysplasia or ≥ 10 mm in size, or ≥ 3 adenomas), the ESGE recommends surveillance colonoscopy 3 years after the index colonoscopy (strong recommendation, moderate quality evidence). Patients with 10 or more adenomas should be referred for genetic counselling (strong recommendation, moderate quality evidence).**

### Long-term CRC risk in the high risk group (see Table 3a)

Epidemiological studies have indicated that the high risk group is at increased risk of CRC compared with the general population. Patients stratified into the high risk group who were followed for 14 years (without endoscopic surveillance) had a 3.6– to 6.6-fold increase in CRC risk, compared with the general population [62]. Another study found that patients with advanced adenomas who did not undergo endoscopic surveillance had a 4.26 (95%CI 2.89–6.04) times greater risk for CRC [18]. Epidemiological series

also showed a high efficacy of endoscopic surveillance in reducing the CRC risk in the high risk group [22, 23, 63].

### Incidence of metachronous advanced neoplasia in the high risk group

In prospective cohort studies, the incidence of metachronous advanced neoplasia was 5–7 times higher in the high risk group compared with individuals without adenomas at the index colonoscopy [15,35,64]. A pooled analysis included individual data on 9167 participants from 8 prospective post-polypectomy trials with a mean follow-up of 47 months. The crude risk of advanced neoplasia during follow-up was 15.5% in the high risk group and 6.9% in the low risk group [32]. In a multivariate analysis, size, multiplicity, and presence of villous component of the baseline lesions appeared to be independent risk factors for metachronous advanced neoplasia, whilst high grade dysplasia was not [32]. These results were largely confirmed by two meta-analyses [71,72]. The risk of metachronous advanced lesions seems to be higher in the high risk groups, but the contribution of each individual unfavourable adenoma feature (size, multiplicity, villous component) was less consistent [32, 71, 72]. Further data on these individual factors in the high risk group are provided in Table e4 and Table e5 (available online).

It has been suggested that individuals with 5 small adenomas, or 3 or more adenomas where at least one was ≥ 10 mm, could benefit from endoscopic surveillance 1 year after the last endoscopy [73]. In a pooled analysis of 4 surveillance studies, including 3226 patients, these individuals had a doubled risk of metachronous advanced lesions compared with those in the high risk group who did not have these characteristics [73]. However, these individuals did not have higher risk of CRC, and there is considerable uncertainty about how this higher risk of advanced neoplasia may translate into CRC risk.

### Timing of surveillance in the high risk group

In the US National Polyp Study, following adenoma resection 1418 patients were randomly allocated to either a 1-year followed by a 3-year surveillance colonoscopy or to a single 3-year surveillance colonoscopy. The incidence of advanced lesions was 3.3% in both groups [74]. In a retrospective observational study,

the cumulative incidence of metachronous advanced neoplasia in the high risk group increased with increasing surveillance interval; after intervals of 1–3, 3–5, 5–10 and 10–20 years, the incidences of metachronous advanced neoplasia were 3.8%, 13.1%, 34.7% and 52%, respectively [68]. In contrast, another observational study found no association between the duration of the surveillance interval (from 0.5 to 10 years) and the risk of metachronous advanced neoplasia in the high risk group (the risk varied between 9.9% and 11.4%) [65]. This finding was confirmed in a case–control study where the risk of CRC was unchanged if the surveillance interval was prolonged from 3 years to 5 years [75]. In line with current recommendations [76], we propose that individuals with 10 or more adenomas be referred for genetic counselling because of the risk of familial adenomatous polyposis (FAP) or other genetic diseases, such as *MYH*-associated polyposis. Tailored surveillance programmes for patients with hereditary colorectal cancer syndromes are outside the scope of this guideline.

***In the high risk group, if no high risk adenomas are detected at the first surveillance examination, the ESGE suggests a 5-year interval before a second surveillance colonoscopy (weak recommendation, low quality evidence). If high risk adenomas are detected at first or subsequent surveillance examinations, a 3-year repetition of surveillance colonoscopy is recommended (strong recommendation, low quality evidence). The ESGE found insufficient evidence to give recommendations in the case where no high risk adenomas are detected during 2 consecutive surveillance colonoscopies. However, intervals longer than 5 years appear reasonable (very low quality evidence).***

Three recent cohort studies have investigated the risk of metachronous advanced lesions at second surveillance colonoscopy, according to the findings at the baseline and first surveillance colonoscopy [65,77,78] (Table e6, available online). The study designs were prone to selection bias because of nonadherence, which might affect generalizability. However, despite heterogeneity in the study populations, results were reassuringly consistent across the studies. In individuals with high risk adenomas at the index colonoscopy and no high risk adenomas at the first surveillance endoscopy, the risk of metachronous advanced neoplasia at the second surveillance colonoscopy was higher than among individuals without high risk adenomas detected at the index colonoscopy [65,77,78]. The absolute risk of metachronous advanced neoplasia at the second surveillance colonoscopy was 5.9%–6.7% among individuals with high risk adenomas at the index colonoscopy, and 3.1%–5.7% among individuals without high risk adenomas [65,77,78]. This supports the recommendation of a second surveillance colonoscopy after 5 years. For individuals with high risk adenomas detected at endoscopic surveillance, the risk of metachronous advanced neoplasia was higher than for individuals without high risk adenomas, regardless of the findings at previous examinations. The absolute risk for metachronous advanced neoplasia at second surveillance endoscopy ranged from 11.5% to 19.3% for individuals with high risk adenomas at first surveillance colonoscopy [65,77,78]. In comparison, the risk varied from 3.1% to 6.7% for individuals without advanced neoplasia at first surveillance [65,77,78]. No study addressed the risk of metachronous advanced neoplasia after two surveillance colonoscopies without high risk adenomas. When considering the progressive decrease in the incidence of such lesions at the first two surveillance colonoscopies, intervals longer than 5 years may appear reasonable.

### Serrated polyps (see also Table e7, available online)

***The ESGE recommends that patients with serrated polyps < 10 mm in size with no dysplasia should be classified as low risk (weak recommendation, low quality evidence). The ESGE suggests that patients with large serrated polyps (≥ 10 mm) or those with dysplasia should be classified as high risk (weak recommendation, low quality evidence).***

***Patients with 5 or more serrated polyps proximal to the sigmoid, of which 2 or more are sized ≥ 10 mm, or with 20 or more serrated polyps of any size but distributed throughout the colon, meet the World Health Organization criteria for serrated polyposis and should be referred for genetic counselling (strong recommendation, low quality evidence).***

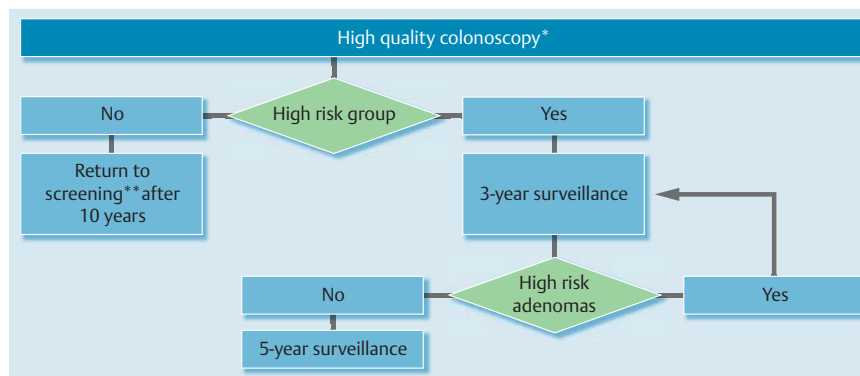
Serrated polyps are classified into different subgroups: (i) hyperplastic polyps, (ii) sessile serrated polyps, (iii) mixed polyps, and (iv) traditional serrated adenomas. No prospective study has yet assessed the long-term risk of CRC in patients with neoplastic and non-neoplastic serrated lesions, leading to uncertainty on the usefulness of endoscopic surveillance.

### Hyperplastic polyps

Observational studies found that in the absence of any neoplasia, hyperplastic polyps are not associated with advanced adenomas [79,80], although a slightly increased risk of adenomas was found [81,82]. The coexistence of hyperplastic polyps with adenomas at index colonoscopy does not increase the risk of adenomas and advanced adenomas at surveillance compared with adenomas alone [81–83]. Indirect evidence of the indolent behavior of hyperplastic polyps is also found in sigmoidoscopy and colonoscopy studies [13,79].

### Sessile serrated polyps (also defined as sessile serrated adenomas/lesions)

One retrospective pathology-based study showed that 15% of patients with sessile serrated polyps at index examination developed advanced neoplasia (CRC/HGD) within approximately 8 years of follow-up, compared with 5.5% of patients with baseline adenomas within 3 years of follow-up [80]. However, the difference in follow-up durations generates some uncertainty about such a comparison [80]. Another study demonstrated that in 50% of patients with sessile serrated polyps at baseline, subsequent sessile serrated polyps were detected within approximately 3 years of follow-up [84]. However, patients with nondysplastic sessile serrated polyps did not present an increased risk of metachronous advanced neoplasia, although size ≥ 10 mm or proximal location were predictors of synchronous advanced neoplasia [85,86]. In particular, large serrated polyps were associated with a higher risk of proximal CRC [86]. Three studies have found an association between type of lesion detected during follow-up and type of lesions found at baseline colonoscopy [84,87,88]. Patients with sessile serrated lesions are more likely to develop further sessile serrated lesions. However, there is no evidence of an increased risk of metachronous CRC [84,87,88]. We recommend that some patients, who fulfil the WHO criteria for serrated polyposis syndrome, should be considered for genetic counselling [89]. This includes: (i) individuals with 5 or more serrated polyps proximal to the sigmoid with 2 or more of those being ≥ 10 mm in diameter, and (ii) individuals with 20 or more serrated polyps of any size distributed throughout the colon (both right- and left-sided).



**Fig. 1** Dichotomization of patients following a high quality colonoscopy in which high risk lesions have or have not been detected. High risk group: patients with an adenoma  $\geq 10$  mm; or with high grade dysplasia; or a villous component or  $\geq 3$  adenomas; serrated polyp  $\geq 10$  mm or with dysplasia. \* Excluding those in whom cancer has already developed. \*\* To a screening programme if available, otherwise to repetition of colonoscopy.

### Mixed polyps and traditional serrated adenomas

Sessile serrated lesions that harbour an adenomatous component are called mixed polyps [90]. These lesions present with a dysplastic component, analogously to the traditional serrated adenomas. No data exist regarding the incidence of metachronous advanced lesions.

Since most pathologists do not yet correctly classify serrated lesions into the several subtypes, we have preferred not to subclassify such lesions for the purposes of our statement [91].

### Specific scenarios

**In the case of piecemeal resection of adenomas larger than 10 mm, endoscopic follow-up within 6 months is recommended before the patient is entered into a surveillance programme (strong recommendation, moderate quality evidence).**

Incomplete removal of larger neoplastic lesions must be ruled out before an endoscopic surveillance schedule is recommended (Table e1c and d, available online). Recently, inadequate polypectomy has been reported in up to 17% of lesions  $\geq 10$  mm, especially if piecemeal polypectomy had been performed [92]. Incomplete excision of neoplastic lesions has been consistently shown to increase the risk of post-colonoscopy interval CRC [43]. For this reason, an early follow-up of these lesions is recommended within 6 months (Table e1b, available online), even if the resection was apparently complete on the basis of endoscopic and histologic criteria [92, 93]. Normal macroscopic appearance of the polypectomy site and negative scar biopsy specimens at the first follow-up have been shown to be predictive of long-term eradication [93].

*The ESGE found insufficient evidence to provide recommendations on post-polypectomy surveillance based on other potential risk factors, such as age, or family history of CRC (very low quality evidence). However, it seems reasonable to stop endoscopic surveillance at 80 years, or earlier depending on life expectancy (in the case of co-morbidities).*

A pooled analysis showed that age was a strong risk factor for metachronous advanced neoplasia. The risk was almost three times higher among individuals older than 80 years compared with those between 50 and 59 years (OR 2.7; 95%CI 1.3–5.6) [94]. Conversely, there was no significant difference between individuals aged 50 to 59 and those aged 60 to 69 [64]. Older people could be more prone to complications of colonoscopy, and the potential benefit of endoscopic surveillance may be limited by reduced life expectancy, especially when the estimated 10–20-year duration of the traditional adenoma–carcinoma sequence is taken into account. No studies have assessed the optimal age for stopping surveillance. Although statistical simulations indi-

cate that surveillance should be stopped between 75 and 85 years, this needs to be confirmed by future trials [95]. Therefore, individualized recommendations should be based on general health status, comorbidity and the findings at previous colonoscopies [96]. It is likely that individuals with limited life expectancy (i.e., shorter than 10 years) will not benefit from post-polypectomy endoscopic surveillance [95, 96].

A recent meta-analysis reviewed the influence of family history on the incidence of metachronous advanced neoplasia [97]. In all studies, including 21 595 participants, a positive family history was defined as having at least one first-degree relative with CRC (parents, siblings, or children) [64, 94, 98, 99]. None of the studies assessed the influence of family history stratified by age at diagnosis and the number of relatives with CRC. The proportion of participants with a positive family history of CRC ranged between 4.9% and 27.5% [64, 99]. No association was found between first-degree family history of CRC and metachronous advanced neoplasia (OR 1.20, 95%CI 0.96–1.50). Similarly, race/ethnicity did not appear to predict rate of metachronous advanced adenoma at endoscopic surveillance [100].

*The ESGE recommends an early repetition of colonoscopy or a shorter surveillance interval in patients in whom an optimal inspection of colorectal mucosa has been hampered by an inadequate preparation, especially if neoplastic lesions have been detected in the initial examination.*

An inadequate level of bowel preparation has been associated with a reduced detection of neoplastic lesions and, therefore, with a higher risk of missed lesions (Table e1a, available online) [101–103]. It has also been shown to be a strong risk factor for metachronous advanced adenoma at surveillance [104]. Thus, early repetition of colonoscopy seems advisable. For instance, if no high risk lesions have been detected and a sufficient level of mucosal inspection has been achieved (i.e., allowing reasonable exclusion of the presence of lesions  $\geq 5$  mm), rather than 10 years before the subsequent screening colonoscopy, a 5-year interval has been suggested [60]. When repeating colonoscopy or shortening the surveillance interval, all the recommendations for an adequate bowel preparation, including split regimen, must be followed [45].

*The ESGE recommends against the use of interval faecal occult blood tests (FOBTs) for post-polypectomy surveillance (strong recommendation, low quality evidence). In the case of an unplanned positive FOBT, the decision to repeat colonoscopy should be based on clinical judgment (weak recommendation, low quality evidence).*

The risk of metachronous CRC in patients following polypectomy is stratified by the findings at the index colonoscopy. For this reason, the attempt to re-stratify risk of CRC by applying a guaiac-faecal occult blood test/faecal immunochemical test (g-FOBT/FIT) would appear to be mere duplication. Although interval CRC may be detected by g-FOBT/FIT, the expected low prevalence of disease would result in a high false-positive rate and a substantial burden on personal, endoscopic, and economic resources. In two nonrandomized studies including high risk individuals, a total of 1856 participants underwent at least one interval FIT during a colonoscopy-based CRC screening programme [105,106]. Colonoscopy was performed in 454 FIT-positive individuals; it led to detection of 18 CRCs, giving a positive predictive value of 4% which is dramatically lower than in a primary FIT screening setting [107]. Unplanned FOBT, although recommended against, may turn out to be positive. The decision whether or not to repeat a colonoscopy should depend on careful clinical evaluation, including the quality of the latest colonoscopy, and the time interval between the latest colonoscopy and FOBT.

*The ESGE suggests that individuals with symptoms in the surveillance interval should be managed as clinically indicated (weak recommendation, low quality evidence).*

Patients under appropriate surveillance are at low risk of CRC but interval CRC may develop, whether polypectomy has been done or not [5,16]. Thus, repetition of colonoscopy should be considered if there is clinical suspicion of interval CRC.

## Discussion

Following a high quality colonoscopy with no detection of CRC, patients may be simply dichotomized according to the presence or absence of high risk adenomatous and serrated colorectal lesions (● Fig. 1). Endoscopic surveillance is recommended for individuals in the high risk group (● Box 1). Surveillance is not indicated for individuals in the low risk group, as with individuals with normal colonoscopy findings, for whom return to screening after 10 years is recommended. This simple approach eliminates confusion about the timing of surveillance colonoscopy, and optimizes the utilization of endoscopic resources. Nevertheless, it offers intensive surveillance, i.e., 3 colonoscopies over 10 years, to individuals who are the most likely to benefit from this.

The main difference between the ESGE and the recent US Multi-Society Task Force (MSTF) post-polypectomy Guidelines is the American recommendation for 5–10-year endoscopic surveillance in the low risk group [60]. The main reason for the 5-year US-MSTF recommendation is the possibility of inadequate preparation or poor quality endoscopic examination. We excluded low quality colonoscopy from the scope of our main recommendations. However, in the specific scenarios, we also allowed the possibility of shortening the interval to the next screening colonoscopy in the case of inadequate bowel preparation. There is insufficient evidence regarding the appropriate surveillance interval after a suboptimal colonoscopy, and we want to emphasize the need to repeat colonoscopy as soon as is practicable in the case of a suboptimal examination. In contrast to the ESGE recommendations, the European quality assurance guidelines propose that the first surveillance in patients with 5 polyps or more or with adenoma  $\geq 20$ mm should be after 1 year rather than 3 years. However, the evidence to underpin this advice does not appear firm [61]. In particular, these patients do not show a high-

er risk of incident CRC, whilst it is unlikely that the moderate increase in the risk of advanced adenoma may represent a significant cause of morbidity/mortality in the subsequent 2 years of follow-up [32,61]. Moreover, our Guideline already recommends that patients with an adenoma  $\geq 10$ mm that was removed piecemeal should have a 6-month surveillance, according to our guideline.

Overall, discrepancies among the main recommendations of different societies seem to be related to the quality of the supporting studies. It should be remembered that most of these studies were carried out before the advent of high resolution colonoscopy and before quality assurance had been incorporated into clinical practice [32].

There is little consistency in the surveillance recommendations for patients with serrated polyps [60,61], because of the lack of firm data on the risk of subsequent polyps and CRC in these patients. However, because of the consistently higher risk of synchronous advanced neoplasia in patients with large serrated polyps, we preferred to recommend a prudent approach until more definitive evidence becomes available.

The ESGE Guideline provides an evidence-based risk-stratification strategy for post-polypectomy surveillance, limiting surveillance to patients with a greater CRC risk. This approach husbands resources whilst maximizing benefits. Such an approach seems of critical importance when the progressive implementation of CRC screening programs throughout Europe is considered. Further studies in this field, especially dealing with serrated lesions, are needed (● Table e8, available online).

ESGE guidelines represent a consensus of best practice based on the available evidence at the time of preparation. They may not apply in all situations and should be interpreted in the light of specific clinical situations and resource availability. Further controlled clinical studies may be needed to clarify aspects of these statements, and revision may be necessary as new data appear. Clinical consideration may justify a course of action at variance to these recommendations. ESGE guidelines are intended to be an educational device to provide information that may assist endoscopists in providing care to patients. They are not rules and should not be construed as establishing a legal standard of care or as encouraging, advocating, requiring, or discouraging any particular treatment.

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## Appendix e1 and e2, Table e1–e8,

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